

THE OFFICIAL
PATIENT'S SOURCEBOOK

on

ADULT
PRIMARY LIVER
CANCER



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AND PHILIP M. PARKER, PH.D., EDITORS

ICON Health Publications
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Dedication

To the healthcare professionals dedicating their time and efforts to the study of adult primary liver cancer.

Acknowledgements

The collective knowledge generated from academic and applied research summarized in various references has been critical in the creation of this sourcebook which is best viewed as a comprehensive compilation and collection of information prepared by various official agencies which directly or indirectly are dedicated to adult primary liver cancer. All of the *Official Patient's Sourcebooks* draw from various agencies and institutions associated with the United States Department of Health and Human Services, and in particular, the Office of the Secretary of Health and Human Services (OS), the Administration for Children and Families (ACF), the Administration on Aging (AOA), the Agency for Healthcare Research and Quality (AHRQ), the Agency for Toxic Substances and Disease Registry (ATSDR), the Centers for Disease Control and Prevention (CDC), the Food and Drug Administration (FDA), the Healthcare Financing Administration (HCFA), the Health Resources and Services Administration (HRSA), the Indian Health Service (IHS), the institutions of the National Institutes of Health (NIH), the Program Support Center (PSC), and the Substance Abuse and Mental Health Services Administration (SAMHSA). In addition to these sources, information gathered from the National Library of Medicine, the United States Patent Office, the European Union, and their related organizations has been invaluable in the creation of this sourcebook. Some of the work represented was financially supported by the Research and Development Committee at INSEAD. This support is gratefully acknowledged. Finally, special thanks are owed to Tiffany LaRochelle for her excellent editorial support.

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Table of Contents

INTRODUCTION.....	1
<i>Overview</i>	1
<i>Organization</i>	3
<i>Scope</i>	3
<i>Moving Forward</i>	4
PART I: THE ESSENTIALS	7
CHAPTER 1. THE ESSENTIALS ON ADULT PRIMARY LIVER CANCER:	
GUIDELINES	9
<i>Overview</i>	9
<i>What Is Adult Primary Liver Cancer?</i>	11
<i>Stages of Adult Primary Liver Cancer</i>	12
<i>How Is Adult Primary Liver Cancer Treated?</i>	13
<i>Treatment by Stage</i>	15
<i>To Learn More</i>	17
<i>About PDQ</i>	18
<i>More Guideline Sources</i>	19
<i>Vocabulary Builder</i>	24
CHAPTER 2. SEEKING GUIDANCE	29
<i>Overview</i>	29
<i>Associations and Adult Primary Liver Cancer</i>	29
<i>Cancer Support Groups</i>	31
<i>The Cancer Information Service</i>	33
<i>Finding Cancer Resources in Your Community</i>	36
<i>Finding Doctors Who Specialize in Cancer Care</i>	39
<i>Selecting Your Doctor</i>	42
<i>Working with Your Doctor</i>	43
<i>Finding a Cancer Treatment Facility</i>	44
<i>Additional Cancer Support Information</i>	46
<i>Vocabulary Builder</i>	46
CHAPTER 3. CLINICAL TRIALS AND ADULT PRIMARY LIVER CANCER	
.....	49
<i>Overview</i>	49
<i>Recent Trials on Adult Primary Liver Cancer</i>	52
<i>Benefits and Risks</i>	76
<i>Clinical Trials and Insurance Coverage</i>	79
<i>Clinical Trials and Medicare Coverage</i>	82
<i>Increasing the Likelihood of Insurance Coverage for Trials</i>	83
<i>If Your Insurance Claim Is Denied after the Trial Has Begun</i>	85
<i>Government Initiatives to Expand Insurance Coverage for Trials</i>	88
<i>Keeping Current on Clinical Trials</i>	89

<i>General References</i>	90
<i>Vocabulary Builder</i>	91

PART II: ADDITIONAL RESOURCES AND ADVANCED MATERIAL..... 97

CHAPTER 4. STUDIES ON ADULT PRIMARY LIVER CANCER	99
<i>Overview</i>	99
<i>The Combined Health Information Database</i>	99
<i>Federally-Funded Research on Adult Primary Liver Cancer</i>	101
<i>E-Journals: PubMed Central</i>	103
<i>The National Library of Medicine: PubMed</i>	104
<i>Vocabulary Builder</i>	123
CHAPTER 5. BOOKS ON ADULT PRIMARY LIVER CANCER	129
<i>Overview</i>	129
<i>The National Library of Medicine Book Index</i>	129
<i>Chapters on Adult Primary Liver Cancer</i>	133
<i>General Home References</i>	133
<i>Vocabulary Builder</i>	135
CHAPTER 6. MULTIMEDIA ON ADULT PRIMARY LIVER CANCER	137
<i>Overview</i>	137
<i>Bibliography: Multimedia on Adult Primary Liver Cancer</i>	137
<i>Vocabulary Builder</i>	139
CHAPTER 7. PHYSICIAN GUIDELINES AND DATABASES	141
<i>Overview</i>	141
<i>NIH Guidelines</i>	141
<i>What Is Adult Primary Liver Cancer?</i>	142
<i>Cellular Classification</i>	144
<i>Stage Information</i>	145
<i>TNM Definitions</i>	145
<i>AJCC Stage Groupings</i>	146
<i>Treatment Option Overview</i>	148
<i>NIH Databases</i>	153
<i>Other Commercial Databases</i>	161
<i>The Genome Project and Adult Primary Liver Cancer</i>	162
<i>Specialized References</i>	166
<i>Vocabulary Builder</i>	167

PART III. APPENDICES 169

APPENDIX A. RESEARCHING YOUR MEDICATIONS	171
<i>Overview</i>	171
<i>Your Medications: The Basics</i>	172
<i>Learning More about Your Medications</i>	174
<i>Commercial Databases</i>	175

<i>Drug Development and Approval</i>	176
<i>Understanding the Approval Process for New Cancer Drugs</i>	177
<i>The Role of the Federal Drug Administration (FDA)</i>	178
<i>Getting Drugs to Patients Who Need Them</i>	182
<i>Contraindications and Interactions (Hidden Dangers)</i>	184
<i>A Final Warning</i>	185
<i>General References</i>	185
<i>Vocabulary Builder</i>	186
APPENDIX B. RESEARCHING ALTERNATIVE MEDICINE	187
<i>Overview</i>	187
<i>What Is CAM?</i>	188
<i>What Are the Domains of Alternative Medicine?</i>	189
<i>Finding CAM References on Adult Primary Liver Cancer</i>	194
<i>Additional Web Resources</i>	197
<i>General References</i>	204
<i>Vocabulary Builder</i>	205
APPENDIX C. RESEARCHING NUTRITION	207
<i>Overview</i>	207
<i>Food and Nutrition: General Principles</i>	208
<i>Finding Studies on Adult Primary Liver Cancer</i>	212
<i>Federal Resources on Nutrition</i>	213
<i>Additional Web Resources</i>	214
<i>Vocabulary Builder</i>	215
APPENDIX D. FINDING MEDICAL LIBRARIES	217
<i>Overview</i>	217
<i>Preparation</i>	217
<i>Finding a Local Medical Library</i>	218
<i>Medical Libraries Open to the Public</i>	218
APPENDIX E. YOUR RIGHTS AND INSURANCE	225
<i>Overview</i>	225
<i>Your Rights as a Patient</i>	225
<i>Patient Responsibilities</i>	229
<i>Choosing an Insurance Plan</i>	230
<i>Medicare and Medicaid</i>	233
<i>Financial Assistance for Cancer Care</i>	236
<i>NORD's Medication Assistance Programs</i>	238
<i>Additional Resources</i>	239
<i>Vocabulary Builder</i>	239
ONLINE GLOSSARIES	241
<i>Online Dictionary Directories</i>	247
ADULT PRIMARY LIVER CANCER GLOSSARY ...	249
<i>General Dictionaries and Glossaries</i>	265

INDEX..... 267

INTRODUCTION

Overview

Dr. C. Everett Koop, former U.S. Surgeon General, once said, “The best prescription is knowledge.”¹ The Agency for Healthcare Research and Quality (AHRQ) of the National Institutes of Health (NIH) echoes this view and recommends that every patient incorporate education into the treatment process. According to the AHRQ:

Finding out more about your condition is a good place to start. By contacting groups that support your condition, visiting your local library, and searching on the Internet, you can find good information to help guide your treatment decisions. Some information may be hard to find – especially if you don’t know where to look.²

As the AHRQ mentions, finding the right information is not an obvious task. Though many physicians and public officials had thought that the emergence of the Internet would do much to assist patients in obtaining reliable information, in March 2001 the National Institutes of Health issued the following warning:

The number of Web sites offering health-related resources grows every day. Many sites provide valuable information, while others may have information that is unreliable or misleading.³

¹ Quotation from <http://www.drkoop.com>.

² The Agency for Healthcare Research and Quality (AHRQ):
<http://www.ahrq.gov/consumer/diaginfo.htm>.

³ Adapted from the NIH, National Cancer Institute (NCI):
<http://cancertrials.nci.nih.gov/beyond/evaluating.html>.

Since the late 1990s, physicians have seen a general increase in patient Internet usage rates. Patients frequently enter their doctor's offices with printed Web pages of home remedies in the guise of latest medical research. This scenario is so common that doctors often spend more time dispelling misleading information than guiding patients through sound therapies. *The Official Patient's Sourcebook on Adult Primary Liver Cancer* has been created for patients who have decided to make education and research an integral part of the treatment process. The pages that follow will tell you where and how to look for information covering virtually all topics related to adult primary liver cancer, from the essentials to the most advanced areas of research.

The title of this book includes the word "official." This reflects the fact that the sourcebook draws from public, academic, government, and peer-reviewed research. Selected readings from various agencies are reproduced to give you some of the latest official information available to date on adult primary liver cancer.

Given patients' increasing sophistication in using the Internet, abundant references to reliable Internet-based resources are provided throughout this sourcebook. Where possible, guidance is provided on how to obtain free-of-charge, primary research results as well as more detailed information via the Internet. E-book and electronic versions of this sourcebook are fully interactive with each of the Internet sites mentioned (clicking on a hyperlink automatically opens your browser to the site indicated). Hard copy users of this sourcebook can type cited Web addresses directly into their browsers to obtain access to the corresponding sites. Since we are working with ICON Health Publications, hard copy *Sourcebooks* are frequently updated and printed on demand to ensure that the information provided is current.

In addition to extensive references accessible via the Internet, every chapter presents a "Vocabulary Builder." Many health guides offer glossaries of technical or uncommon terms in an appendix. In editing this sourcebook, we have decided to place a smaller glossary within each chapter that covers terms used in that chapter. Given the technical nature of some chapters, you may need to revisit many sections. Building one's vocabulary of medical terms in such a gradual manner has been shown to improve the learning process.

We must emphasize that no sourcebook on adult primary liver cancer should affirm that a specific diagnostic procedure or treatment discussed in a research study, patent, or doctoral dissertation is "correct" or your best option. This sourcebook is no exception. Each patient is unique. Deciding on

appropriate options is always up to the patient in consultation with their physician and healthcare providers.

Organization

This sourcebook is organized into three parts. Part I explores basic techniques to researching adult primary liver cancer (e.g. finding guidelines on diagnosis, treatments, and prognosis), followed by a number of topics, including information on how to get in touch with organizations, associations, or other patient networks dedicated to adult primary liver cancer. It also gives you sources of information that can help you find a doctor in your local area specializing in treating adult primary liver cancer. Collectively, the material presented in Part I is a complete primer on basic research topics for patients with adult primary liver cancer.

Part II moves on to advanced research dedicated to adult primary liver cancer. Part II is intended for those willing to invest many hours of hard work and study. It is here that we direct you to the latest scientific and applied research on adult primary liver cancer. When possible, contact names, links via the Internet, and summaries are provided. It is in Part II where the vocabulary process becomes important as authors publishing advanced research frequently use highly specialized language. In general, every attempt is made to recommend “free-to-use” options.

Part III provides appendices of useful background reading for all patients with adult primary liver cancer or related disorders. The appendices are dedicated to more pragmatic issues faced by many patients with adult primary liver cancer. Accessing materials via medical libraries may be the only option for some readers, so a guide is provided for finding local medical libraries which are open to the public. Part III, therefore, focuses on advice that goes beyond the biological and scientific issues facing patients with adult primary liver cancer.

Scope

While this sourcebook covers adult primary liver cancer, your doctor, research publications, and specialists may refer to your condition using a variety of terms. Therefore, you should understand that adult primary liver cancer is often considered a synonym or a condition closely related to the following:

- Cancer Liver
- Cholangiocarcinoma
- Fibrolamellar Carcinoma
- Hepatocellular Carcinoma
- Hepatoma
- Liver Cancer
- Primary Liver Cell Carcinoma
- Tumor Liver

In addition to synonyms and related conditions, physicians may refer to adult primary liver cancer using certain coding systems. The International Classification of Diseases, 9th Revision, Clinical Modification (ICD-9-CM) is the most commonly used system of classification for the world's illnesses. Your physician may use this coding system as an administrative or tracking tool. The following classification is commonly used for adult primary liver cancer:⁴

- 155.0 hepatocellular carcinoma
- 155.0 malignant neoplasm of liver, primary

For the purposes of this sourcebook, we have attempted to be as inclusive as possible, looking for official information for all of the synonyms relevant to adult primary liver cancer. You may find it useful to refer to synonyms when accessing databases or interacting with healthcare professionals and medical librarians.

Moving Forward

Since the 1980s, the world has seen a proliferation of healthcare guides covering most illnesses. Some are written by patients or their family members. These generally take a layperson's approach to understanding and coping with an illness or disorder. They can be uplifting, encouraging, and highly supportive. Other guides are authored by physicians or other

⁴ This list is based on the official version of the World Health Organization's 9th Revision, International Classification of Diseases (ICD-9). According to the National Technical Information Service, "ICD-9CM extensions, interpretations, modifications, addenda, or errata other than those approved by the U.S. Public Health Service and the Health Care Financing Administration are not to be considered official and should not be utilized. Continuous maintenance of the ICD-9-CM is the responsibility of the federal government."

healthcare providers who have a more clinical outlook. Each of these two styles of guide has its purpose and can be quite useful.

As editors, we have chosen a third route. We have chosen to expose you to as many sources of official and peer-reviewed information as practical, for the purpose of educating you about basic and advanced knowledge as recognized by medical science today. You can think of this sourcebook as your personal Internet age reference librarian.

Why “Internet age”? All too often, patients diagnosed with adult primary liver cancer will log on to the Internet, type words into a search engine, and receive several Web site listings which are mostly irrelevant or redundant. These patients are left to wonder where the relevant information is, and how to obtain it. Since only the smallest fraction of information dealing with adult primary liver cancer is even indexed in search engines, a non-systematic approach often leads to frustration and disappointment. With this sourcebook, we hope to direct you to the information you need that you would not likely find using popular Web directories. Beyond Web listings, in many cases we will reproduce brief summaries or abstracts of available reference materials. These abstracts often contain distilled information on topics of discussion.

While we focus on the more scientific aspects of adult primary liver cancer, there is, of course, the emotional side to consider. Later in the sourcebook, we provide a chapter dedicated to helping you find peer groups and associations that can provide additional support beyond research produced by medical science. We hope that the choices we have made give you the most options available in moving forward. In this way, we wish you the best in your efforts to incorporate this educational approach into your treatment plan.

The Editors

PART I: THE ESSENTIALS

ABOUT PART I

Part I has been edited to give you access to what we feel are “the essentials” on adult primary liver cancer. The essentials of a disease typically include the definition or description of the disease, a discussion of who it affects, the signs or symptoms associated with the disease, tests or diagnostic procedures that might be specific to the disease, and treatments for the disease. Your doctor or healthcare provider may have already explained the essentials of adult primary liver cancer to you or even given you a pamphlet or brochure describing adult primary liver cancer. Now you are searching for more in-depth information. As editors, we have decided, nevertheless, to include a discussion on where to find essential information that can complement what your doctor has already told you. In this section we recommend a process, not a particular Web site or reference book. The process ensures that, as you search the Web, you gain background information in such a way as to maximize your understanding.

CHAPTER 1. THE ESSENTIALS ON ADULT PRIMARY LIVER CANCER: GUIDELINES

Overview

Official agencies, as well as federally funded institutions supported by national grants, frequently publish a variety of guidelines on adult primary liver cancer. These are typically called “Fact Sheets” or “Guidelines.” They can take the form of a brochure, information kit, pamphlet, or flyer. Often they are only a few pages in length. The great advantage of guidelines over other sources is that they are often written with the patient in mind. Since new guidelines on adult primary liver cancer can appear at any moment and be published by a number of sources, the best approach to finding guidelines is to systematically scan the Internet-based services that post them.

The National Institutes of Health (NIH)⁵

The National Institutes of Health (NIH) is the first place to search for relatively current patient guidelines and fact sheets on adult primary liver cancer. Originally founded in 1887, the NIH is one of the world’s foremost medical research centers and the federal focal point for medical research in the United States. At any given time, the NIH supports some 35,000 research grants at universities, medical schools, and other research and training institutions, both nationally and internationally. The rosters of those who have conducted research or who have received NIH support over the years include the world’s most illustrious scientists and physicians. Among them are 97 scientists who have won the Nobel Prize for achievement in medicine.

⁵ Adapted from the NIH: <http://www.nih.gov/about/NIHoverview.html>.

There is no guarantee that any one Institute will have a guideline on a specific disease, though the National Institutes of Health collectively publish over 600 guidelines for both common and rare diseases. The best way to access NIH guidelines is via the Internet. Although the NIH is organized into many different Institutes and Offices, the following is a list of key Web sites where you are most likely to find NIH clinical guidelines and publications dealing with adult primary liver cancer and associated conditions:

- Office of the Director (OD); guidelines consolidated across agencies available at <http://www.nih.gov/health/consumer/conkey.htm>
- National Library of Medicine (NLM); extensive encyclopedia (A.D.A.M., Inc.) with guidelines available at <http://www.nlm.nih.gov/medlineplus/healthtopics.html>
- National Cancer Institute (NCI); guidelines available at http://cancernet.nci.nih.gov/pdq/pdq_treatment.shtml

Among the above, the National Cancer Institute (NCI) is particularly noteworthy. The NCI coordinates the National Cancer Program, which conducts and supports research, training, health information dissemination, and other programs with respect to the cause, diagnosis, prevention, and treatment of cancer, rehabilitation from cancer, and the continuing care of cancer patients and the families of cancer patients.⁶ Specifically, the Institute:

- Supports and coordinates research projects conducted by universities, hospitals, research foundations, and businesses throughout this country and abroad through research grants and cooperative agreements.
- Conducts research in its own laboratories and clinics.
- Supports education and training in fundamental sciences and clinical disciplines for participation in basic and clinical research programs and treatment programs relating to cancer through career awards, training grants, and fellowships.
- Supports research projects in cancer control.
- Supports a national network of cancer centers.
- Collaborates with voluntary organizations and other national and foreign institutions engaged in cancer research and training activities.
- Encourages and coordinates cancer research by industrial concerns where such concerns evidence a particular capability for programmatic research.
- Collects and disseminates information on cancer.

⁶ This paragraph has been adapted from the NCI: <http://www.nci.nih.gov/>. “Adapted” signifies that a passage has been reproduced exactly or slightly edited for this book.

- Supports construction of laboratories, clinics, and related facilities necessary for cancer research through the award of construction grants.

The NCI, established under the National Cancer Act of 1937, is the Federal Government's principal agency for cancer research and training. The National Cancer Act of 1971 broadened the scope and responsibilities of the NCI and created the National Cancer Program. Over the years, legislative amendments have maintained the NCI authorities and responsibilities and added new information dissemination mandates as well as a requirement to assess the incorporation of state-of-the-art cancer treatments into clinical practice. Information dissemination is made possible through the NCI Online at www.cancer.gov. Cancer.gov offers to the public and physicians up-to-date information on the latest cancer research, current and upcoming clinical trials, statistics, research programs, and research funding.

The following patient guideline was recently published by the NCI on adult primary liver cancer.

What Is Adult Primary Liver Cancer?⁷

Adult primary liver cancer is a disease in which cancer (malignant) cells start to grow in the tissues of the liver. The liver is one of the largest organs in the body, filling the upper right side of the abdomen and protected by the rib cage. The liver has many functions. It has an important role in making food into energy and also filters and stores blood.

People who have hepatitis B or C (viral infections of the liver) or a disease of the liver called cirrhosis are more likely than other people to get adult primary liver cancer. Primary liver cancer is different from cancer that has spread from another place in the body to the liver.

A doctor should be seen if the following symptoms appear:

- A hard lump just below the rib cage on the right side where the liver has swollen,
- Discomfort in the upper abdomen on the right side,
- Pain around the right shoulder blade, or
- Yellowing of the skin (jaundice).

⁷ The following guidelines appeared on the NCI website on Aug. 26, 2002. The text was last modified in April 2002. The text has been adapted for this sourcebook.

If there are symptoms, a doctor may order special x-rays, such as a computed tomographic scan or a liver scan. If a lump is seen on an x-ray, a doctor may use a needle inserted into the abdomen to remove a small amount of tissue from the liver. This procedure is called a needle biopsy, and a doctor usually will use an x-ray for guidance. The doctor will have the tissue looked at under a microscope to see if there are any cancer cells. Before the test, a patient will be given a local anesthetic (a drug that causes loss of feeling for a short period of time) in the area so that no pain is felt.

A doctor may also want to look at the liver with an instrument called a laparoscope, which is a small tube-shaped instrument with a light on the end. For this test, a small cut is made in the abdomen so that the laparoscope can be inserted. The doctor may also take a small piece of tissue (biopsy specimen) during the laparoscopy and look at it under the microscope to see if there are any cancer cells. An anesthetic will be given so no pain is felt.

A doctor may also order an examination called an angiography. During this examination, a tube (catheter) is inserted into the main blood vessel that takes blood to the liver. Dye is then injected through the tube so that the blood vessels in the liver can be seen on an x-ray. Angiography can help a doctor tell whether the cancer is primary liver cancer or cancer that has spread from another part of the body. This test is usually done in the hospital.

Certain blood tests (such as alpha-fetoprotein, or AFP) may also help a doctor diagnose primary liver cancer.

Prognosis

The chance of recovery (prognosis) and choice of treatment depend on the stage of the cancer (whether it is just in the liver or has spread to other places) and the patient's general state of health.

Stages of Adult Primary Liver Cancer

Once adult primary liver cancer is found, more tests will be done to find out if the cancer cells have spread to other parts of the body (staging). The following stages are used for adult primary liver cancer:

Localized Resectable

Cancer is found in one place in the liver and can be totally removed in an operation.

Localized Unresectable

Cancer is found only in one part of the liver, but the cancer cannot be totally removed.

Advanced

Cancer has spread through much of the liver or to other parts of the body.

Recurrent

Recurrent disease means that the cancer has come back (recurred) after it has been treated. It may come back in the liver or in another part of the body.

How Is Adult Primary Liver Cancer Treated?

There are treatments for all patients with adult primary liver cancer. Three kinds of treatment are used:

- Surgery (taking out the cancer in an operation)
- Radiation therapy (using high-dose x-rays to kill cancer cells)
- Chemotherapy (using drugs to kill cancer cells)

Surgery

Surgery may be used to take out the cancer or to replace the liver.

- Resection of the liver takes out the part of the liver where the cancer is found.
- A liver transplant is the removal of the entire liver and replacement with a healthy liver donated from someone else. Very few patients with liver cancer are eligible for this procedure.

- Cryosurgery is a type of surgery that kills cancer by freezing it.

Radiation Therapy

Radiation therapy is the use of x-rays or other high-energy rays to kill cancer cells and shrink tumors. Radiation may come from a machine outside the body (external-beam radiation therapy) or from putting materials that contain radiation through thin plastic tubes (internal radiation therapy) in the area where the cancer cells are found. Drugs may be given with the radiation therapy to make the cancer cells more sensitive to radiation (radiosensitization).

Radiation may also be given by attaching radioactive substances to antibodies (radiolabeled antibodies) that search out certain cells in the liver. Antibodies are made by the body to fight germs and other harmful things; each antibody fights specific cells.

Chemotherapy

Chemotherapy is the use of drugs to kill cancer cells. Chemotherapy for liver cancer is usually put into the body by inserting a needle into a vein or artery. This type of chemotherapy is called a systemic treatment because the drug enters the bloodstream, travels through the body, and can kill cancer cells outside the liver. In another type of chemotherapy called regional chemotherapy, a small pump containing drugs is placed in the body. The pump puts drugs directly into the blood vessels that go to the tumor.

Chemoembolization of the hepatic artery involves blocking the hepatic artery (the major artery that supplies blood to the liver) and then injecting chemotherapy drugs between the blockage and the liver, using the liver's arteries to deliver the chemotherapy throughout the liver.

If a doctor removes all the cancer that can be seen at the time of the operation, the patient may be given chemotherapy after surgery to kill any remaining cells. Chemotherapy that is given after surgery to remove the cancer is called adjuvant chemotherapy.

Treatments under Clinical Trial

Hyperthermia (warming the body to kill cancer cells) and biological therapy (using the body's immune system to fight cancer) are being tested in clinical trials.

Hyperthermia therapy is the use of a special machine to heat the body for a certain period of time to kill cancer cells. Because cancer cells are often more sensitive to heat than normal cells, the cancer cells die and the tumor shrinks.

Biological therapy uses the body's immune system to fight cancer. Materials made by the body or made in a laboratory are used to boost, direct, or restore the body's natural defenses against disease. Biological therapy is sometimes called biological response modifier therapy or immunotherapy.

Treatment by Stage

Treatments for adult primary liver cancer depend on the stage of the disease, the condition of the liver, and the patient's age and general health. Standard treatment may be considered, based on its effectiveness in patients in past studies, or participation into a clinical trial. Many patients are not cured with standard therapy, and some standard treatments may have more side effects than are desired. For these reasons, clinical trials are designed to find better ways to treat cancer patients and are based on the most up-to-date information. Clinical trials are ongoing in most parts of the country for most stages of adult liver cancer. For more information, call the Cancer Information Service at 1-800-4-CANCER (1-800-422-6237); TTY at 1-800-332-8615.

Localized Resectable Adult Primary Liver Cancer

Treatment is usually surgery (resection). Liver transplantation may be done in certain patients. Clinical trials are testing adjuvant systemic or regional chemotherapy following surgery.

Localized Unresectable Adult Primary Liver Cancer

Treatment may be one of the following:

- Blocking the hepatic artery and then injecting chemotherapy drugs into the artery and liver (chemoembolization), surgery to freeze and kill the tumor (cryosurgery), injection of ethanol into the tumor, or use of highly focused radio waves designed to destroy the tumor.
- Liver transplantation.
- Regional chemotherapy, including injecting the chemotherapy directly into the tumor.
- Systemic chemotherapy.
- Surgery with or without chemotherapy possibly followed by radiation therapy.
- Injection of alcohol directly into the tumor.
- Radiation therapy plus special drugs that make the tumor more susceptible to the radiation.
- Highly focused radio waves designed to destroy the tumor

Advanced Adult Primary Liver Cancer

There is no standard treatment for patients with advanced adult primary liver cancer. Patients may wish to consider taking part in a clinical trial. Some clinical trials may also help reduce or relieve symptoms.

Recurrent Adult Primary Liver Cancer

Treatment of recurrent adult primary liver cancer depends on what treatment a patient has already received, the part of the body where the cancer has come back, whether the liver has cirrhosis, and other factors. Patients may wish to consider taking part in a clinical trial.

To Learn More

Call

For more information, U.S. residents may call the National Cancer Institute's (NCI's) Cancer Information Service toll-free at 1-800-4-CANCER (1-800-422-6237), Monday through Friday from 9:00 a.m. to 4:30 p.m. Deaf and hard-of-hearing callers with TTY equipment may call 1-800-332-8615. The call is free and a trained Cancer Information Specialist is available to answer your questions.

Web Sites and Organizations

The NCI's Cancer.gov Web site (<http://cancer.gov>) provides online access to information on cancer, clinical trials, and other Web sites and organizations that offer support and resources for cancer patients and their families. There are also many other places where people can get materials and information about cancer treatment and services. Local hospitals may have information on local and regional agencies that offer information about finances, getting to and from treatment, receiving care at home, and dealing with problems associated with cancer treatment.

Publications

The NCI has booklets and other materials for patients, health professionals, and the public. These publications discuss types of cancer, methods of cancer treatment, coping with cancer, and clinical trials. Some publications provide information on tests for cancer, cancer causes and prevention, cancer statistics, and NCI research activities. NCI materials on these and other topics may be ordered online or printed directly from the NCI Publications Locator (<http://cancer.gov/publications>). These materials can also be ordered by telephone from the Cancer Information Service toll-free at 1-800-4-CANCER (1-800-422-6237), TTY at 1-800-332-8615.

LiveHelp

The NCI's LiveHelp service, a program available on several of the Institute's Web sites, provides Internet users with the ability to chat online with an Information Specialist. The service is available from 9:00 a.m. to 7:30 p.m. Eastern time, Monday through Friday. Information Specialists can help

Internet users find information on NCI Web sites and answer questions about cancer.

Write

For more information from the NCI, please write to this address:

National Cancer Institute
Office of Communications
31 Center Drive, MSC 2580
Bethesda, MD 20892-2580

About PDQ

PDQ Is a Comprehensive Cancer Database Available on Cancer.gov

PDQ is the National Cancer Institute's (NCI's) comprehensive cancer information database. Most of the information contained in PDQ is available online at Cancer.gov (<http://cancer.gov>), the NCI's Web site. PDQ is provided as a service of the NCI. The NCI is part of the National Institutes of Health, the federal government's focal point for biomedical research.

PDQ Contains Cancer Information Summaries

The PDQ database contains summaries of the latest published information on cancer prevention, detection, genetics, treatment, supportive care, and complementary and alternative medicine. Most summaries are available in two versions. The health professional versions provide detailed information written in technical language. The patient versions are written in easy-to-understand, non-technical language. Both versions provide current and accurate cancer information.

The PDQ cancer information summaries are developed by cancer experts and reviewed regularly. Editorial Boards made up of experts in oncology and related specialties are responsible for writing and maintaining the cancer information summaries. The summaries are reviewed regularly and changes are made as new information becomes available. The date on each summary ("Date Last Modified") indicates the time of the most recent change.

PDQ Contains Information on Clinical Trials

Before starting treatment, patients may want to think about taking part in a clinical trial. A clinical trial is a study to answer a scientific question, such as whether one treatment is better than another. Trials are based on past studies and what has been learned in the laboratory. Each trial answers certain scientific questions in order to find new and better ways to help cancer patients. During treatment clinical trials, information is collected about new treatments, the risks involved, and how well they do or do not work. If a clinical trial shows that a new treatment is better than one currently being used, the new treatment may become “standard.”

Listings of clinical trials are included in PDQ and are available online at Cancer.gov (http://cancer.gov/clinical_trials). Descriptions of the trials are available in health professional and patient versions. Many cancer doctors who take part in clinical trials are also listed in PDQ. For more information, call the Cancer Information Service at 1-800-4-CANCER (1-800-422-6237); TTY at 1-800-332-8615.

More Guideline Sources

The guideline above on adult primary liver cancer is only one example of the kind of material that you can find online and free of charge. The remainder of this chapter will direct you to other sources which either publish or can help you find additional guidelines on topics related to adult primary liver cancer. Many of the guidelines listed below address topics that may be of particular relevance to your specific situation or of special interest to only some patients with adult primary liver cancer. Due to space limitations these sources are listed in a concise manner. Do not hesitate to consult the following sources by either using the Internet hyperlink provided, or, in cases where the contact information is provided, contacting the publisher or author directly.

Topic Pages: MEDLINEplus

For patients wishing to go beyond guidelines published by specific Institutes of the NIH, the National Library of Medicine has created a vast and patient-oriented healthcare information portal called MEDLINEplus. Within this Internet-based system are “health topic pages.” You can think of a health topic page as a guide to patient guides. To access this system, log on to <http://www.nlm.nih.gov/medlineplus/healthtopics.html>. From there you

can either search using the alphabetical index or browse by broad topic areas. Recently, MEDLINEplus listed the following as being relevant to adult primary liver cancer:

- Guides On Adult Primary Liver Cancer

Liver Cancer

<http://www.nlm.nih.gov/medlineplus/livercancer.html>

- Other Guides

Jaundice-associated conditions

<http://www.nlm.nih.gov/medlineplus/ency/article/000210.htm>

Within the health topic page dedicated to adult primary liver cancer, the following was recently recommended to patients:

- Diagnosis/Symptoms

Liver and Hepatobiliary Imaging

Source: Society of Nuclear Medicine

<http://www.snm.org/nuclear/liver.html>

Liver Biopsy

Source: National Digestive Diseases Information Clearinghouse

<http://www.niddk.nih.gov/health/digest/pubs/liverbiopsy/liverbiopsy.htm>

Ultrasound-Abdomen

Source: American College of Radiology, Radiological Society of North America

<http://www.radiologyinfo.org/content/ultrasound-abdomen.htm>

- Treatment

Adult Primary Liver Cancer (PDQ): Treatment

Source: National Cancer Institute

http://www.cancer.gov/cancer_information/doc_pdq.aspx?version=patient&viewid=d1e6c99a-3331-40c7-8006-07887a129d29

Questions and Answers About Cryosurgery in Cancer Treatment

Source: National Cancer Institute

http://cis.nci.nih.gov/fact/7_34.htm

Radiofrequency Ablation

Source: National Institutes of Health, Clinical Center

<http://www.cc.nih.gov/drd/rfa/frame-patient.html>

Radiofrequency Catheter Ablation (RFA)

Source: Society of Interventional Radiology

<http://www.sirweb.org/patPub/radiofrequencyAblation.shtml>**Treatments: Chemoembolization, Tumor Ablation, and More**

Source: Society of Interventional Radiology

<http://www.sirweb.org/patPub/cancerTreatments.shtml>

- Specific Conditions/Aspects

Oral Contraceptives and Cancer Risk

Source: National Cancer Institute

http://cis.nci.nih.gov/fact/3_13.htm

- Children

Childhood Liver Cancer (PDQ): Treatment

Source: National Cancer Institute

http://www.cancer.gov/cancer_information/doc_pdq.aspx?version=patient&viewid=3593e766-131b-4499-b167-5691b0fd8538

- From the National Institutes of Health

What You Need to Know About Liver Cancer

Source: National Cancer Institute

http://www.cancer.gov/cancer_information/doc_wyntk.aspx?viewid=2ab11798-ed80-4ee0-9a0e-1708724db2b9

- Latest News

New Technique Cuts Blood Loss in Liver Surgery

Source: 11/04/2002, Reuters Health

http://www.nlm.nih.gov/medlineplus/news/fullstory_10188.html

- Organizations

American Cancer Society<http://www.cancer.org/>**National Cancer Institute**<http://www.cancer.gov/>

- Prevention/Screening

- **ALT (Alanine Aminotransferase) Test**

- Source: American Association for Clinical Chemistry

- <http://www.labtestsonline.org/understanding/analytes/alt/test.html>

- **Hepatocellular Cancer (PDQ): Screening**

- Source: National Cancer Institute

- http://www.cancer.gov/cancer_information/doc_pdq.aspx?version=patient&viewid=3541b0cd-fa15-43a0-be0e-601ec30ec97b

- **Liver Panel**

- Source: American Association for Clinical Chemistry

- http://www.labtestsonline.org/understanding/analytes/liver_panel/glance.html

- Research

- **Chemoembolization May Help Some Patients with Inoperable Liver Cancer**

- Source: National Cancer Institute

- <http://www.cancer.gov/clinicaltrials/results/chemoembolization0602>

- **Radiofrequency Energy Sizzles Tumors Without Surgery**

- Source: National Institutes of Health, Clinical Center

- <http://www.nih.gov/news/pr/nov2001/wgmcc-26.htm>

- **Ritalin and Liver Cancer**

- Source: Mayo Foundation for Medical Education and Research

- <http://www.mayoclinic.com/invoke.cfm?id=AN00132>

If you do not find topics of interest when browsing health topic pages, then you can choose to use the advanced search utility of MEDLINEplus at <http://www.nlm.nih.gov/medlineplus/advancedsearch.html>. This utility is similar to the NIH Search Utility, with the exception that it only includes material linked within the MEDLINEplus system (mostly patient-oriented information). It also has the disadvantage of generating unstructured results. We recommend, therefore, that you use this method only if you have a very targeted search.

The National Guideline Clearinghouse™

The National Guideline Clearinghouse™ offers hundreds of evidence-based clinical practice guidelines published in the United States and other countries. You can search their site located at <http://www.guideline.gov> by using the keyword “adult primary liver cancer” or synonyms.

Healthfinder™

Healthfinder™ is an additional source sponsored by the U.S. Department of Health and Human Services which offers links to hundreds of other sites that contain healthcare information. This Web site is located at <http://www.healthfinder.gov>. Again, keyword searches can be used to find guidelines. The following was recently found in this database:

- **Adult Primary Liver Cancer (PDQ®): Treatment**

Summary: General overview and treatment options for adult primary liver cancer are discussed in this brochure. Treatment choice depends on the stage of the cancer and the patient's general state of health.

Source: National Cancer Institute, National Institutes of Health

<http://www.healthfinder.gov/scripts/recordpass.asp?RecordType=0&RecordID=6135>

- **Adult Primary Liver Cancer (PDQ®): Treatment Information for Professionals**

Summary: Treatment information for liver cancer from NCI's PDQ® database, intended for use by doctors and other health care professionals.

Source: National Cancer Institute, National Institutes of Health

<http://www.healthfinder.gov/scripts/recordpass.asp?RecordType=0&RecordID=4355>

The NIH Search Utility

After browsing the references listed at the beginning of this chapter, you may want to explore the NIH Search Utility. This allows you to search for documents on over 100 selected Web sites that comprise the NIH-WEB-SPACE. Each of these servers is “crawled” and indexed on an ongoing basis. Your search will produce a list of various documents, all of which will relate in some way to adult primary liver cancer. The drawbacks of this approach

are that the information is not organized by theme and that the references are often a mix of information for professionals and patients. Nevertheless, a large number of the listed Web sites provide useful background information. We can only recommend this route, therefore, for relatively rare or specific disorders, or when using highly targeted searches. To use the NIH search utility, visit the following Web page: <http://search.nih.gov/index.html>.

Additional Web Sources

A number of Web sites that often link to government sites are available to the public. These can also point you in the direction of essential information. The following is a representative sample:

- AOL: <http://search.aol.com/cat.adp?id=168&layer=&from=subcats>
- drkoop.com[®]: <http://www.drkoop.com/conditions/ency/index.html>
- Family Village: <http://www.familyvillage.wisc.edu/specific.htm>
- Google:
http://directory.google.com/Top/Health/Conditions_and_Diseases/
- Med Help International: <http://www.medhelp.org/HealthTopics/A.html>
- Open Directory Project:
http://dmoz.org/Health/Conditions_and_Diseases/
- Yahoo.com: http://dir.yahoo.com/Health/Diseases_and_Conditions/
- WebMD[®]Health: http://my.webmd.com/health_topics

Vocabulary Builder

The material in this chapter may have contained a number of unfamiliar words. The following Vocabulary Builder introduces you to terms used in this chapter that have not been covered in the previous chapter:

Abdomen: The part of the body that contains the pancreas, stomach, intestines, liver, gallbladder, and other organs. [NIH]

Abdominal: Having to do with the abdomen, which is the part of the body between the chest and the hips that contains the pancreas, stomach, intestines, liver, gallbladder, and other organs. [NIH]

Adjuvant: A substance which aids another, such as an auxiliary remedy; in immunology, nonspecific stimulator (e.g., BCG vaccine) of the immune response. [EU]

AFP: Alpha-fetoprotein. A protein normally produced by a developing fetus. AFP levels are usually undetectable in the blood of healthy nonpregnant adults. An elevated level of AFP suggests the presence of either a primary liver cancer or germ cell tumor. [NIH]

Alanine: A non-essential amino acid that occurs in high levels in its free state in plasma. It is produced from pyruvate by transamination. It is involved in sugar and acid metabolism, increases immunity, and provides energy for muscle tissue, brain, and the central nervous system. [NIH]

American Cancer Society: A voluntary organization concerned with the prevention and treatment of cancer through education and research. [NIH]

Angiography: A procedure to x-ray blood vessels. The blood vessels can be seen because of an injection of a dye that shows up in the x-ray pictures. [NIH]

Antibody: A type of protein made by certain white blood cells in response to a foreign substance (antigen). Each antibody can bind to only a specific antigen. The purpose of this binding is to help destroy the antigen. Antibodies can work in several ways, depending on the nature of the antigen. Some antibodies destroy antigens directly. Others make it easier for white blood cells to destroy the antigen. [NIH]

Arteries: The vessels carrying blood away from the heart. [NIH]

Biopsy: The removal of cells or tissues for examination under a microscope. When only a sample of tissue is removed, the procedure is called an incisional biopsy or core biopsy. When an entire tumor or lesion is removed, the procedure is called an excisional biopsy. When a sample of tissue or fluid is removed with a needle, the procedure is called a needle biopsy or fine-needle aspiration. [NIH]

Carcinoma: Cancer that begins in the skin or in tissues that line or cover internal organs. [NIH]

Cardiology: The study of the heart, its physiology, and its functions. [NIH]

Catheter: A flexible tube used to deliver fluids into or withdraw fluids from the body. [NIH]

Cell: The individual unit that makes up all of the tissues of the body. All living things are made up of one or more cells. [NIH]

Chemoembolization: A procedure in which the blood supply to the tumor is blocked surgically or mechanically, and anticancer drugs are administered directly into the tumor. This permits a higher concentration of drug to be in contact with the tumor for a longer period of time. [NIH]

Chemotherapy: Treatment with anticancer drugs. [NIH]

Cirrhosis: A type of chronic, progressive liver disease. [NIH]

Colorectal: Having to do with the colon or the rectum. [NIH]

Contraceptive: An agent that diminishes the likelihood of or prevents conception. [EU]

Cryosurgery: Treatment performed with an instrument that freezes and destroys abnormal tissues. This procedure is a form of cryotherapy. [NIH]

Endocrinologist: A doctor that specializes in diagnosing and treating hormone disorders. [NIH]

Endocrinology: A subspecialty of internal medicine concerned with the metabolism, physiology, and disorders of the endocrine system. [NIH]

Ethanol: A clear, colorless liquid rapidly absorbed from the gastrointestinal tract and distributed throughout the body. It has bactericidal activity and is used often as a topical disinfectant. It is widely used as a solvent and preservative in pharmaceutical preparations as well as serving as the primary ingredient in alcoholic beverages. [NIH]

Genitourinary: Pertaining to the genital and urinary organs; urogenital; urinosexual. [EU]

Hepatic: Refers to the liver. [NIH]

Hepatitis: Inflammation of the liver. [NIH]

Hepatobiliary: Pertaining to the liver and the bile or the biliary ducts. [EU]

Hyperthermia: A type of treatment in which body tissue is exposed to high temperatures to damage and kill cancer cells or to make cancer cells more sensitive to the effects of radiation and certain anticancer drugs. [NIH]

Immunodeficiency: The decreased ability of the body to fight infection and disease. [NIH]

Immunotherapy: Treatment to stimulate or restore the ability of the immune system to fight infection and disease. Also used to lessen side effects that may be caused by some cancer treatments. Also called biological therapy or biological response modifier (BRM) therapy. [NIH]

Infarction: 1. the formation of an infarct. 2. an infarct. [EU]

Inoperable: Not suitable to be operated upon. [EU]

Jaundice: A condition in which the skin and the whites of the eyes become yellow, urine darkens, and stool becomes clay colored. Jaundice occurs when the liver is not working properly or when a bile duct is blocked. [NIH]

Laparoscopy: The insertion of a thin, lighted tube (called a laparoscope) through the abdominal wall to inspect the inside of the abdomen and remove tissue samples. [NIH]

Liver: A large, glandular organ located in the upper abdomen. The liver cleanses the blood and aids in digestion by secreting bile. [NIH]

Malignant: Cancerous; a growth with a tendency to invade and destroy

nearby tissue and spread to other parts of the body. [NIH]

Oncology: The study of cancer. [NIH]

Ophthalmology: A surgical specialty concerned with the structure and function of the eye and the medical and surgical treatment of its defects and diseases. [NIH]

Radioactive: Giving off radiation. [NIH]

Radiofrequency ablation: The use of electrical current to destroy tissue. [NIH]

Radiolabeled: Any compound that has been joined with a radioactive substance. [NIH]

Radiology: The use of radiation (such as x-rays) or other imaging technologies (such as ultrasound and magnetic resonance imaging) to diagnose or treat disease. [NIH]

Radiosensitization: The use of a drug that makes tumor cells more sensitive to radiation therapy. [NIH]

Resection: Removal of tissue or part or all of an organ by surgery. [NIH]

Screening: Checking for disease when there are no symptoms. [NIH]

Staging: Performing exams and tests to learn the extent of the cancer within the body, especially whether the disease has spread from the original site to other parts of the body. [NIH]

Systemic: Affecting the entire body. [NIH]

Thyroid: A gland located near the windpipe (trachea) that produces thyroid hormone, which helps regulate growth and metabolism. [NIH]

Transplantation: The replacement of an organ with one from another person. [NIH]

Unresectable: Unable to be surgically removed. [NIH]

Venereal: Pertaining or related to or transmitted by sexual contact. [EU]

Viral: Pertaining to, caused by, or of the nature of virus. [EU]

Virus: Submicroscopic organism that causes infectious disease. In cancer therapy, some viruses may be made into vaccines that help the body build an immune response to, and kill, tumor cells. [NIH]

CHAPTER 2. SEEKING GUIDANCE

Overview

Some patients are comforted by the knowledge that a number of organizations dedicate their resources to helping people with adult primary liver cancer. These associations can become invaluable sources of information and advice. Many associations offer aftercare support, financial assistance, and other important services. Furthermore, healthcare research has shown that support groups often help people to better cope with their conditions.⁸ In addition to support groups, your physician can be a valuable source of guidance and support. Therefore, finding a physician that can work with your unique situation is a very important aspect of your care.

In this chapter, we direct you to resources that can help you find patient organizations and medical specialists. We begin by describing how to find associations and peer groups that can help you better understand and cope with adult primary liver cancer. The chapter ends with a discussion on how to find a doctor that is right for you.

Associations and Adult Primary Liver Cancer

As mentioned by the Agency for Healthcare Research and Quality, sometimes the emotional side of an illness can be as taxing as the physical side.⁹ You may have fears or feel overwhelmed by your situation. Everyone has different ways of dealing with disease or physical injury. Your attitude, your expectations, and how well you cope with your condition can all

⁸ Churches, synagogues, and other houses of worship might also have groups that can offer you the social support you need.

⁹ This section has been adapted from <http://www.ahcpr.gov/consumer/diaginf5.htm>.

influence your well-being. This is true for both minor conditions and serious illnesses. For example, a study on female breast cancer survivors revealed that women who participated in support groups lived longer and experienced better quality of life when compared with women who did not participate. In the support group, women learned coping skills and had the opportunity to share their feelings with other women in the same situation. There are a number of directories that list additional medical associations that you may find useful. While not all of these directories will provide different information, by consulting all of them, you will have nearly exhausted all sources for patient associations.

The National Cancer Institute (NCI)

The National Cancer Institute (NCI) has compiled a list of national organizations that offer services to people with cancer and their families. To view the list, see the NCI fact sheet online at the following Web address: http://cis.nci.nih.gov/fact/8_1.htm. The name of each organization is accompanied by its contact information and a brief explanation of its services.

The National Health Information Center (NHIC)

The National Health Information Center (NHIC) offers a free referral service to help people find organizations that provide information about adult primary liver cancer. For more information, see the NHIC's Web site at <http://www.health.gov/NHIC/> or contact an information specialist by calling 1-800-336-4797.

DIRLINE

A comprehensive source of information on associations is the DIRLINE database maintained by the National Library of Medicine. The database comprises some 10,000 records of organizations, research centers, and government institutes and associations which primarily focus on health and biomedicine. DIRLINE is available via the Internet at the following Web site: <http://dirline.nlm.nih.gov/>. Simply type in "adult primary liver cancer" (or a synonym) or the name of a topic, and the site will list information contained in the database on all relevant organizations.

The Combined Health Information Database

Another comprehensive source of information on healthcare associations is the Combined Health Information Database. Using the “Detailed Search” option, you will need to limit your search to “Organizations” and “adult primary liver cancer”. Type the following hyperlink into your Web browser: **<http://chid.nih.gov/detail/detail.html>**. To find associations, use the drop boxes at the bottom of the search page where “You may refine your search by.” For publication date, select “All Years.” Then, select your preferred language and the format option “Organization Resource Sheet.” By making these selections and typing in “adult primary liver cancer” (or synonyms) into the “For these words:” box, you will only receive results on organizations dealing with adult primary liver cancer. You should check back periodically with this database since it is updated every 3 months.

The National Organization for Rare Disorders, Inc.

The National Organization for Rare Disorders, Inc. has prepared a Web site that provides, at no charge, lists of associations organized by specific diseases. You can access this database at the following Web site: **<http://www.rarediseases.org/cgi-bin/nord/searchpage>**. Select the option called “Organizational Database (ODB)” and type “adult primary liver cancer” (or a synonym) in the search box.

Cancer Support Groups¹⁰

People diagnosed with cancer and their families face many challenges that may leave them feeling overwhelmed, afraid, and alone. It can be difficult to cope with these challenges or to talk to even the most supportive family members and friends. Often, support groups can help people affected by cancer feel less alone and can improve their ability to deal with the uncertainties and challenges that cancer brings. Support groups give people who are affected by similar diseases an opportunity to meet and discuss ways to cope with the illness.

¹⁰ This section has been adapted from the NCI: **http://cis.nci.nih.gov/fact/8_8.htm**.

How Can Support Groups Help?

People who have been diagnosed with cancer sometimes find they need assistance coping with the emotional as well as the practical aspects of their disease. In fact, attention to the emotional burden of cancer is sometimes part of a patient's treatment plan. Cancer support groups are designed to provide a confidential atmosphere where cancer patients or cancer survivors can discuss the challenges that accompany the illness with others who may have experienced the same challenges. For example, people gather to discuss the emotional needs created by cancer, to exchange information about their disease—including practical problems such as managing side effects or returning to work after treatment—and to share their feelings. Support groups have helped thousands of people cope with these and similar situations.

Can Family Members and Friends Participate in Support Groups?

Family and friends are affected when cancer touches someone they love, and they may need help in dealing with stresses such as family disruptions, financial worries, and changing roles within relationships. To help meet these needs, some support groups are designed just for family members of people diagnosed with cancer; other groups encourage families and friends to participate along with the cancer patient or cancer survivor.

How Can People Find Support Groups?

Many organizations offer support groups for people diagnosed with cancer and their family members or friends. The NCI fact sheet *National Organizations That Offer Services to People with Cancer and Their Families* lists many cancer-concerned organizations that can provide information about support groups. This fact sheet is available at http://cis.nci.nih.gov/fact/8_1.htm on the Internet, or can be ordered from the Cancer Information Service at 1-800-4-CANCER (1-800-422-6237). Some of these organizations provide information on their Web sites about contacting support groups.

Doctors, nurses, or hospital social workers who work with cancer patients may also have information about support groups, such as their location, size, type, and how often they meet. Most hospitals have social services departments that provide information about cancer support programs.

Additionally, many newspapers carry a special health supplement containing information about where to find support groups.

What Types of Support Groups Are Available?

Several kinds of support groups are available to meet the individual needs of people at all stages of cancer treatment, from diagnosis through follow-up care. Some groups are general cancer support groups, while more specialized groups may be for teens or young adults, for family members, or for people affected by a particular disease. Support groups may be led by a professional, such as a psychiatrist, psychologist, or social worker, or by cancer patients or survivors. In addition, support groups can vary in approach, size, and how often they meet. Many groups are free, but some require a fee (people can contact their health insurance company to find out whether their plan will cover the cost). It is important for people to find an atmosphere that is comfortable and meets their individual needs.

Online Support Groups

In addition to support groups, commercial Internet service providers offer forums and chat rooms for people with different illnesses and conditions. WebMD[®], for example, offers such a service at their Web site: **<http://boards.webmd.com/roundtable>**. These online self-help communities can help you connect with a network of people whose concerns are similar to yours. Online support groups are places where people can talk informally. If you read about a novel approach, consult with your doctor or other healthcare providers, as the treatments or discoveries you hear about may not be scientifically proven to be safe and effective.

The Cancer Information Service¹¹

The Cancer Information Service (CIS) is a program of the National Cancer Institute (NCI), the Nation's lead agency for cancer research. As a resource for information and education about cancer, the CIS is a leader in helping people become active participants in their own health care by providing the latest information on cancer in understandable language. Through its network of regional offices, the CIS serves the United States, Puerto Rico, the U.S. Virgin Islands, and the Pacific Islands.

¹¹ This section has been adapted from the NCI: **http://cis.nci.nih.gov/fact/2_5.htm**.

For 25 years, the Cancer Information Service has provided the latest and most accurate cancer information to patients and families, the public, and health professionals by:

- Interacting with people one-on-one through its Information Service,
- Working with organizations through its Partnership Program,
- Participating in research efforts to find the best ways to help people adopt healthier behaviors,
- Providing access to NCI information over the Internet.

How Does the CIS Assist the Public?

Through the CIS toll-free telephone service (1-800-4-CANCER), callers speak with knowledgeable, caring staff who are experienced at explaining medical information in easy-to-understand terms. CIS information specialists answer calls in English and Spanish. They also provide cancer information to deaf and hard of hearing callers through the toll-free TTY number (1-800-332-8615). CIS staff have access to comprehensive, accurate information from the NCI on a range of cancer topics, including the most recent advances in cancer treatment. They take as much time as each caller needs, provide thorough and personalized attention, and keep all calls confidential.

The CIS also provides live, online assistance to users of NCI Web sites through LiveHelp, an instant messaging service that is available from 9:00 a.m. to 7:30 p.m. Eastern time, Monday through Friday. Through LiveHelp, information specialists provide answers to questions about cancer and help in navigating Cancer.gov, the NCI's Web site.

Through the telephone numbers or LiveHelp service, CIS users receive:

- Answers to their questions about cancer, including ways to prevent cancer, symptoms and risks, diagnosis, current treatments, and research studies;
- Written materials from the NCI;
- Referrals to clinical trials and cancer-related services, such as treatment centers, mammography facilities, or other cancer organizations;
- Assistance in quitting smoking from information specialists trained in smoking cessation counseling.

What Kind of Assistance Does the CIS Partnership Program Offer?

Through its Partnership Program, the CIS collaborates with established national, state, and regional organizations to reach minority and medically underserved audiences with cancer information. Partnership Program staff provide assistance to organizations developing programs that focus on breast and cervical cancer, clinical trials, tobacco control, and cancer awareness for special populations. To reach those in need, the CIS:

- Helps bring cancer information to people who do not traditionally seek health information or who may have difficulties doing so because of educational, financial, cultural, or language barriers;
- Provides expertise to organizations to help strengthen their ability to inform people they serve about cancer; and
- Links organizations with similar goals and helps them plan and evaluate programs, develop coalitions, conduct training on cancer-related topics, and use NCI resources.

How Do CIS Research Efforts Assist the Public?

The CIS plays an important role in research by studying the most effective ways to communicate with people about healthy lifestyles; health risks; and options for preventing, diagnosing, and treating cancer. The ability to conduct health communications research is a unique aspect of the CIS. Results from these research studies can be applied to improving the way the CIS communicates about cancer and can help other programs communicate more effectively.

How Do People Reach the Cancer Information Service?

- To speak with a CIS information specialist call 1-800-4-CANCER (1-800-422-6237), 9:00 a.m. to 4:30 p.m. local time, Monday through Friday. Deaf or hard of hearing callers with TTY equipment may call 1-800-332-8615.
- To obtain online assistance visit the NCI's Cancer Information Web site at http://cancer.gov/cancer_information and click on the LiveHelp link between 9:00 a.m. and 7:30 p.m. Eastern time, Monday through Friday.
- For information 24 hours a day, 7 days a week call 1-800-4-CANCER and select option 4 to hear recorded information at any time.
- Visit NCI's Web site at <http://cancer.gov> on the Internet.
- Visit the CIS Web site at <http://cancer.gov/cis> on the Internet.

Finding Cancer Resources in Your Community¹²

If you have cancer or are undergoing cancer treatment, there are places in your community to turn to for help. There are many local organizations throughout the country that offer a variety of practical and support services to people with cancer. However, people often don't know about these services or are unable to find them. National cancer organizations can assist you in finding these resources, and there are a number of things you can do for yourself.

Whether you are looking for a support group, counseling, advice, financial assistance, transportation to and from treatment, or information about cancer, most neighborhood organizations, local health care providers, or area hospitals are a good place to start. Often, the hardest part of looking for help is knowing the right questions to ask.

What Kind of Help Can I Get?

Until now, you probably never thought about the many issues and difficulties that arise with a diagnosis of cancer. There are support services to help you deal with almost any type of problem that might occur. The first step in finding the help you need is knowing what types of services are available. The following pages describe some of these services and how to find them.

- **Information on Cancer.** Most national cancer organizations provide a range of information services, including materials on different types of cancer, treatments, and treatment-related issues.
- **Counseling.** While some people are reluctant to seek counseling, studies show that having someone to talk to reduces stress and helps people both mentally and physically. Counseling can also provide emotional support to cancer patients and help them better understand their illness. Different types of counseling include individual, group, family, self-help (sometimes called peer counseling), bereavement, patient-to-patient, and sexuality.
- **Medical Treatment Decisions.** Often, people with cancer need to make complicated medical decisions. Many organizations provide hospital and physician referrals for second opinions and information on clinical trials (research studies with people), which may expand treatment options.

¹² Adapted from the NCI: http://cis.nci.nih.gov/fact/8_9.htm.

- **Prevention and Early Detection.** While cancer prevention may never be 100 percent effective, many things (such as quitting smoking and eating healthy foods) can greatly reduce a person's risk for developing cancer. Prevention services usually focus on smoking cessation and nutrition. Early detection services, which are designed to detect cancer when a person has no symptoms of disease, can include referrals for screening mammograms, Pap tests, or prostate exams.
- **Home Health Care.** Home health care assists patients who no longer need to stay in a hospital or nursing home, but still require professional medical help. Skilled nursing care, physical therapy, social work services, and nutrition counseling are all available at home.
- **Hospice Care.** Hospice is care focused on the special needs of terminally ill cancer patients. Sometimes called *palliative care*, it centers around providing comfort, controlling physical symptoms, and giving emotional support to patients who can no longer benefit from curative treatment. Hospice programs provide services in various settings, including the patient's home, hospice centers, hospitals, or skilled nursing facilities. Your doctor or social worker can provide a referral for these services.
- **Rehabilitation.** Rehabilitation services help people adjust to the effects of cancer and its treatment. Physical rehabilitation focuses on recovery from the physical effects of surgery or the side effects associated with chemotherapy. Occupational or vocational therapy helps people readjust to everyday routines, get back to work, or find employment.
- **Advocacy.** Advocacy is a general term that refers to promoting or protecting the rights and interests of a certain group, such as cancer patients. Advocacy groups may offer services to assist with legal, ethical, medical, employment, legislative, or insurance issues, among others. For instance, if you feel your insurance company has not handled your claim fairly, you may want to advocate for a review of its decision.
- **Financial.** Having cancer can be a tremendous financial burden to cancer patients and their families. There are programs sponsored by the government and nonprofit organizations to help cancer patients with problems related to medical billing, insurance coverage, and reimbursement issues. There are also sources for financial assistance, and ways to get help collecting entitlements from Medicaid, Medicare, and the Social Security Administration.
- **Housing/Lodging.** Some organizations provide lodging for the family of a patient undergoing treatment, especially if it is a child who is ill and the parents are required to accompany the child to treatment.

- **Children's Services.** A number of organizations provide services for children with cancer, including summer camps, make-a-wish programs, and help for parents seeking child care.

How to Find These Services

Often, the services that people with cancer are looking for are right in their own neighborhood or city. The following is a list of places where you can begin your search for help.

- The hospital, clinic, or medical center where you see your doctor, received your diagnosis, or where you undergo treatment should be able to give you information. Your doctor or nurse may be able to tell you about your specific medical condition, pain management, rehabilitation services, home nursing, or hospice care.
- Most hospitals also have a social work, home care, or discharge planning department. This department may be able to help you find a support group, a nonprofit agency that helps people who have cancer, or the government agencies that oversee Social Security, Medicare, and Medicaid. While you are undergoing treatment, be sure to ask the hospital about transportation, practical assistance, or even temporary child care. Talk to a hospital financial counselor in the business office about developing a monthly payment plan if you need help with hospital expenses.
- The public library is an excellent source of information, as are patient libraries at many cancer centers. A librarian can help you find books and articles through a literature search.
- A local church, synagogue, YMCA or YWCA, or fraternal order may provide financial assistance, or may have volunteers who can help with transportation and home care. Catholic Charities, the United Way, or the American Red Cross may also operate local offices. Some of these organizations may provide home care, and the United Way's information and referral service can refer you to an agency that provides financial help. To find the United Way serving your community, visit their online directory at <http://www.unitedway.org> on the Internet or look in the White Pages of your local telephone book.
- Local or county government agencies may offer low-cost transportation (sometimes called para-transit) to individuals unable to use public transportation. Most states also have an Area Agency on Aging that offers low-cost services to people over 60. Your hospital or community social worker can direct you to government agencies for entitlements,

including Social Security, state disability, Medicaid, income maintenance, and food stamps. (Keep in mind that most applications to entitlement programs take some time to process.) The Federal government also runs the Hill-Burton program (1-800-638-0742), which funds certain medical facilities and hospitals to provide cancer patients with free or low-cost care if they are in financial need.

Getting the Most From a Service: What To Ask

No matter what type of help you are looking for, the only way to find resources to fit your needs is to ask the right questions. When you are calling an organization for information, it is important to think about what questions you are going to ask before you call. Many people find it helpful to write out their questions in advance, and to take notes during the call. Another good tip is to ask the name of the person with whom you are speaking in case you have follow-up questions. Below are some of the questions you may want to consider if you are calling or visiting a new agency and want to learn about how they can help:

- How do I apply [for this service]?
- Are there eligibility requirements? What are they?
- Is there an application process? How long will it take? What information will I need to complete the application process? Will I need anything else to get the service?
- Do you have any other suggestions or ideas about where I can find help?

The most important thing to remember is that you will rarely receive help unless you ask for it. In fact, asking can be the hardest part of getting help. Don't be afraid or ashamed to ask for assistance. Cancer is a very difficult disease, but there are people and services that can ease your burdens and help you focus on your treatment and recovery.

Finding Doctors Who Specialize in Cancer Care¹³

One of the most important aspects of your treatment will be the relationship between you and your doctor or specialist. All patients with adult primary liver cancer must go through the process of selecting a physician. A common way to find a doctor who specializes in cancer care is to ask for a referral

¹³ Adapted from the NCI: http://cis.nci.nih.gov/fact/7_47.htm.

from your primary care physician. Sometimes, you may know a specialist yourself, or through the experience of a family member, coworker, or friend.

The following resources may also be able to provide you with names of doctors who specialize in treating specific diseases or conditions. However, these resources may not have information about the quality of care that the doctors provide.

- Your local hospital or its patient referral service may be able to provide you with a list of specialists who practice at that hospital.
- Your nearest National Cancer Institute (NCI)-designated cancer center can provide information about doctors who practice at that center. The NCI fact sheet *The National Cancer Institute Cancer Centers Program* describes and gives contact information, including Web sites, for NCI-designated cancer treatment centers around the country. Many of the cancer centers' Web sites have searchable directories of physicians who practice at each facility. The NCI's fact sheet is available at http://cis.nci.nih.gov/fact/1_2.htm on the Internet, or by calling the Cancer Information Service (CIS) at 1-800-4-CANCER (1-800-422-6237).
- The American Board of Medical Specialties (ABMS) publishes a list of board-certified physicians. The *Official ABMS Directory of Board Certified Medical Specialists* lists doctors' names along with their specialty and their educational background. This resource is available in most public libraries. The ABMS also has a Web site that can be used to verify whether a specific physician is board-certified. This free service is located at <http://www.abms.org/newsearch.asp> on the Internet. Verification of a physician's board certification can also be obtained by calling the ABMS at 1-866-275-2267 (1-866-ASK-ABMS).
- The American Medical Association (AMA) provides an online service called AMA Physician Select that offers basic professional information on virtually every licensed physician in the United States and its possessions. The database can be searched by doctor's name or by medical specialty. The AMA Physician Select service is located at <http://www.ama-assn.org/aps/amahg.htm> on the Internet.
- The American Society of Clinical Oncologists (ASCO) provides an online list of doctors who are members of ASCO. The member database has the names and affiliations of over 15,000 oncologists worldwide. It can be searched by doctor's name, institution's name, location, and/or type of board certification. This service is located at http://www.asco.org/people/db/html/m_db.htm on the Internet.

- The American College of Surgeons (ACOS) Fellowship Database is an online list of surgeons who are Fellows of the ACOS. The list can be searched by doctor's name, geographic location, or medical specialty. This service is located at <http://web.facs.org/acsdire/default.htm> on the Internet. The ACOS can be contacted at 633 North Saint Clair Street, Chicago, IL 60611-3211; or by telephone at 312-202-5000.
- Local medical societies may maintain lists of doctors in each specialty.
- Public and medical libraries may have print directories of doctors' names, listed geographically by specialty.
- Your local Yellow Pages may have doctors listed by specialty under "Physicians."

The Agency for Healthcare Research and Quality (AHRQ) offers *Your Guide to Choosing Quality Health Care*, which has information for consumers on choosing a health plan, a doctor, a hospital, or a long-term care provider. The Guide includes suggestions and checklists that you can use to determine which doctor or hospital is best for you. This resource is available at <http://www.ahrq.gov/consumer/qntool.htm> on the Internet. You can also order the Guide by calling the AHRQ Publications Clearinghouse at 1-800-358-9295.

If you are a member of a health insurance plan, your choice may be limited to doctors who participate in your plan. Your insurance company can provide you with a list of participating primary care doctors and specialists. It is important to ask your insurance company if the doctor you choose is accepting new patients through your health plan. You also have the option of seeing a doctor outside your health plan and paying the costs yourself. If you have a choice of health insurance plans, you may first wish to consider which doctor or doctors you would like to use, then choose a plan that includes your chosen physician(s).

The National Comprehensive Cancer Network (NCCN) Physician Directory lists specialists who practice in the NCCN's 19 member institutions across the U.S. To access the directory, go to <http://www.nccn.org/> and click on "Physician Directory". To use this service, you will be required to scroll to the bottom of the page and select "I agree." Enter your search criteria and select "Find" at the bottom of the page. To obtain more information on a physician or institution, contact the institution's Physician Referral Department or the NCCN Patient Information and Referral Service at 1-888-909-NCCN or patientinformation@nccn.org.

If the previous sources did not meet your needs, you may want to log on to the Web site of the National Organization for Rare Disorders (NORD) at <http://www.rarediseases.org/>. NORD maintains a database of doctors with expertise in various rare diseases. The Metabolic Information Network (MIN), 800-945-2188, also maintains a database of physicians with expertise in various metabolic diseases.

Selecting Your Doctor¹⁴

There are many factors to consider when choosing a doctor. To make the most informed decision, you may wish to speak with several doctors before choosing one. When you meet with each doctor, you might want to consider the following:

- Does the doctor have the education and training to meet my needs?
- Does the doctor use the hospital that I have chosen?
- Does the doctor listen to me and treat me with respect?
- Does the doctor explain things clearly and encourage me to ask questions?
- What are the doctor's office hours?
- Who covers for the doctor when he or she is unavailable? Will that person have access to my medical records?
- How long does it take to get an appointment with the doctor?

If you are choosing a surgeon, you may wish to ask additional questions about the surgeon's background and experience with specific procedures. These questions may include:

- Is the surgeon board-certified?¹⁵
- Has the surgeon been evaluated by a national professional association of surgeons, such as the American College of Surgeons (ACOS)?
- At which treatment facility or facilities does the surgeon practice?
- How often does the surgeon perform the type of surgery I need?

¹⁴ This section has been adapted from the AHRQ:
<http://www.ahrq.gov/consumer/qntascii/qntdr.htm>

¹⁵ While board certification is a good measure of a doctor's knowledge, it is possible to receive quality care from doctors who are not board certified.

- How many of these procedures has the surgeon performed? What was the success rate?

It is important for you to feel comfortable with the specialist that you choose, because you will be working closely with that person to make decisions about your cancer treatment. Trust your own observations and feelings when deciding on a doctor for your medical care.

Other health professionals and support services may also be important during cancer treatment. The National Cancer Institute fact sheet *Your Health Care Team: Your Doctor Is Only the Beginning* has information about these providers and services, and how to locate them. This fact sheet is located at http://cis.nci.nih.gov/fact/8_10.htm on the Internet, or can be obtained by calling the CIS at 1-800-4-CANCER (1-800-422-6237).

Working with Your Doctor¹⁶

Research has shown that patients who have good relationships with their doctors tend to be more satisfied with their care and have better results. Here are some tips to help you and your doctor become partners:

- You know important things about your symptoms and your health history. Tell your doctor what you think he or she needs to know.
- It is important to tell your doctor personal information, even if it makes you feel embarrassed or uncomfortable.
- Bring a “health history” list with you (and keep it up to date).
- Always bring any medications you are currently taking with you to the appointment, or you can bring a list of your medications including dosage and frequency information. Talk about any allergies or reactions you have had to your medications.
- Tell your doctor about any natural or alternative medicines you are taking.
- Bring other medical information, such as x-ray films, test results, and medical records.
- Ask questions. If you don’t, your doctor will assume that you understood everything that was said.

¹⁶ This section has been adapted from the AHRQ:
www.ahrq.gov/consumer/qntascii/qntdr.htm.

- Write down your questions before your visit. List the most important ones first to make sure that they are addressed.
- Consider bringing a friend with you to the appointment to help you ask questions. This person can also help you understand and/or remember the answers.
- Ask your doctor to draw pictures if you think that this would help you understand.
- Take notes. Some doctors do not mind if you bring a tape recorder to help you remember things, but always ask first.
- Let your doctor know if you need more time. If there is not time that day, perhaps you can speak to a nurse or physician assistant on staff or schedule a telephone appointment.
- Take information home. Ask for written instructions. Your doctor may also have brochures and audio and videotapes that can help you.
- After leaving the doctor's office, take responsibility for your care. If you have questions, call. If your symptoms get worse or if you have problems with your medication, call. If you had tests and do not hear from your doctor, call for your test results. If your doctor recommended that you have certain tests, schedule an appointment to get them done. If your doctor said you should see an additional specialist, make an appointment.

By following these steps, you will enhance the relationship you will have with your physician.

Finding a Cancer Treatment Facility¹⁷

Choosing a treatment facility is another important consideration for getting the best medical care possible. Although you may not be able to choose which hospital treats you in an emergency, you can choose a facility for scheduled and ongoing care. If you have already found a doctor for your cancer treatment, you may need to choose a facility based on where your doctor practices. Your doctor may be able to recommend a facility that provides quality care to meet your needs. You may wish to ask the following questions when considering a treatment facility:

- Has the facility had experience and success in treating my condition?

¹⁷ Adapted from the NCI: http://cis.nci.nih.gov/fact/7_47.htm. At this Web site, information on how to find treatment facilities is also available for patients living outside the U.S.

- Has the facility been rated by state, consumer, or other groups for its quality of care?
- How does the facility check and work to improve its quality of care?
- Has the facility been approved by a nationally recognized accrediting body, such as the American College of Surgeons (ACOS) and/or the Joint Commission on Accredited Healthcare Organizations (JCAHO)?
- Does the facility explain patients' rights and responsibilities? Are copies of this information available to patients?
- Does the treatment facility offer support services, such as social workers and resources to help me find financial assistance if I need it?
- Is the facility conveniently located?

If you are a member of a health insurance plan, your choice of treatment facilities may be limited to those that participate in your plan. Your insurance company can provide you with a list of approved facilities. Although the costs of cancer treatment can be very high, you have the option of paying out-of-pocket if you want to use a treatment facility that is not covered by your insurance plan. If you are considering paying for treatment yourself, you may wish to discuss the potential costs with your doctor beforehand. You may also want to speak with the person who does the billing for the treatment facility. In some instances, nurses and social workers can provide you with more information about coverage, eligibility, and insurance issues.

The following resources may help you find a hospital or treatment facility for your care:

- The NCI fact sheet *The National Cancer Institute Cancer Centers Program* describes and gives contact information for NCI-designated cancer treatment centers around the country.
- The ACOS accredits cancer programs at hospitals and other treatment facilities. More than 1,400 programs in the United States have been designated by the ACOS as Approved Cancer Programs. The ACOS Web site offers a searchable database of these programs at <http://web.facs.org/cpm/default.htm> on the Internet. The ACOS can be contacted at 633 North Saint Clair Street, Chicago, IL 60611-3211; or by telephone at 312-202-5000.
- The JCAHO is an independent, not-for-profit organization that evaluates and accredits health care organizations and programs in the United States. It also offers information for the general public about choosing a

treatment facility. The JCAHO Web site is located at <http://www.jcaho.org> on the Internet. The JCAHO is located at One Renaissance Boulevard, Oakbrook Terrace, IL 60181-4294. The telephone number is 630-792-5800.

- The JCAHO offers an online Quality Check service that patients can use to determine whether a specific facility has been accredited by the JCAHO and view the organization's performance reports. This service is located at <http://www.jcaho.org/qualitycheck/directry/directry.asp> on the Internet.
- The AHRQ publication *Your Guide To Choosing Quality Health Care* has suggestions and checklists for choosing the treatment facility that is right for you.

Additional Cancer Support Information

In addition to the references above, the NCI has set up guidance Web sites that offers information on issues relating to cancer. These include:

- Facing Forward - A Guide for Cancer Survivors:
http://www.cancer.gov/cancer_information/doc_img.aspx?viewid=cc93a843-6fc0-409e-8798-5c65afc172fe
- Taking Time: Support for People With Cancer and the People Who Care About Them:
http://www.cancer.gov/cancer_information/doc_img.aspx?viewid=21a46445-a5c8-4fee-95a3-d9d0d665077a
- When Cancer Recurs: Meeting the Challenge:
http://www.cancer.gov/cancer_information/doc_img.aspx?viewid=9e13d0d2-b7de-4bd6-87da-5750300a0dab
- Your Health Care Team: Your Doctor Is Only the Beginning:
http://cis.nci.nih.gov/fact/8_10.htm

Vocabulary Builder

The following vocabulary builder provides definitions of words used in this chapter that have not been defined in previous chapters:

Bereavement: Refers to the whole process of grieving and mourning and is associated with a deep sense of loss and sadness. [NIH]

Cervical: Relating to the neck, or to the neck of any organ or structure.

Cervical lymph nodes are located in the neck; cervical cancer refers to cancer of the uterine cervix, which is the lower, narrow end (the "neck") of the uterus. [NIH]

Charities: Social welfare organizations with programs designed to assist individuals in times of need. [NIH]

Curative: Tending to overcome disease and promote recovery. [EU]

Mammogram: An x-ray of the breast. [NIH]

Mammography: The use of x-rays to create a picture of the breast. [NIH]

Oncologist: A doctor who specializes in treating cancer. Some oncologists specialize in a particular type of cancer treatment. For example, a radiation oncologist specializes in treating cancer with radiation. [NIH]

Palliative: 1. affording relief, but not cure. 2. an alleviating medicine. [EU]

Pap test: The collection of cells from the cervix for examination under a microscope. It is used to detect changes that may be cancer or may lead to cancer, and can show noncancerous conditions, such as infection or inflammation. Also called a Pap smear. [NIH]

Prostate: A gland in males that surrounds the neck of the bladder and the urethra. It secretes a substance that liquifies coagulated semen. It is situated in the pelvic cavity behind the lower part of the pubic symphysis, above the deep layer of the triangular ligament, and rests upon the rectum. [NIH]

CHAPTER 3. CLINICAL TRIALS AND ADULT PRIMARY LIVER CANCER

Overview

Very few medical conditions have a single treatment. The basic treatment guidelines that your physician has discussed with you, or those that you have found using the techniques discussed in Chapter 1, may provide you with all that you will require. For some patients, current treatments can be enhanced with new or innovative techniques currently under investigation. In this chapter, we will describe how clinical trials work and show you how to keep informed of trials concerning adult primary liver cancer.

What Is a Clinical Trial?¹⁸

Clinical trials involve the participation of people in medical research. Most medical research begins with studies in test tubes and on animals. Treatments that show promise in these early studies may then be tried with people. The only sure way to find out whether a new treatment is safe, effective, and better than other treatments for adult primary liver cancer is to try it on patients in a clinical trial.

¹⁸ The discussion in this chapter has been adapted from the NIH and the NEI: www.nei.nih.gov/netrials/ctivr.htm.

What Kinds of Clinical Trials Are There?

Clinical trials are carried out in three phases:

- **Phase I.** Researchers first conduct Phase I trials with small numbers of patients and healthy volunteers. If the new treatment is a medication, researchers also try to determine how much of it can be given safely.
- **Phase II.** Researchers conduct Phase II trials in small numbers of patients to find out the effect of a new treatment on adult primary liver cancer.
- **Phase III.** Finally, researchers conduct Phase III trials to find out how new treatments for adult primary liver cancer compare with standard treatments already being used. Phase III trials also help to determine if new treatments have any side effects. These trials--which may involve hundreds, perhaps thousands, of people--can also compare new treatments with no treatment.

How Is a Clinical Trial Conducted?

Various organizations support clinical trials at medical centers, hospitals, universities, and doctors' offices across the United States. The "principal investigator" is the researcher in charge of the study at each facility participating in the clinical trial. Most clinical trial researchers are medical doctors, academic researchers, and specialists. The "clinic coordinator" knows all about how the study works and makes all the arrangements for your visits.

All doctors and researchers who take part in the study on adult primary liver cancer carefully follow a detailed treatment plan called a protocol. This plan fully explains how the doctors will treat you in the study. The "protocol" ensures that all patients are treated in the same way, no matter where they receive care.

Clinical trials are controlled. This means that researchers compare the effects of the new treatment with those of the standard treatment. In some cases, when no standard treatment exists, the new treatment is compared with no treatment. Patients who receive the new treatment are in the treatment group. Patients who receive a standard treatment or no treatment are in the "control" group. In some clinical trials, patients in the treatment group get a new medication while those in the control group get a placebo. A placebo is a harmless substance, a "dummy" pill, that has no effect on adult primary liver cancer. In other clinical trials, where a new surgery or device (not a medicine) is being tested, patients in the control group may receive a "sham

treatment.” This treatment, like a placebo, has no effect on adult primary liver cancer and does not harm patients.

Researchers assign patients “randomly” to the treatment or control group. This is like flipping a coin to decide which patients are in each group. If you choose to participate in a clinical trial, you will not know which group you will be appointed to. The chance of any patient getting the new treatment is about 50 percent. You cannot request to receive the new treatment instead of the placebo or sham treatment. Often, you will not know until the study is over whether you have been in the treatment group or the control group. This is called a “masked” study. In some trials, neither doctors nor patients know who is getting which treatment. This is called a “double masked” study. These types of trials help to ensure that the perceptions of the patients or doctors will not affect the study results.

Natural History Studies

Unlike clinical trials in which patient volunteers may receive new treatments, natural history studies provide important information to researchers on how adult primary liver cancer develops over time. A natural history study follows patient volunteers to see how factors such as age, sex, race, or family history might make some people more or less at risk for adult primary liver cancer. A natural history study may also tell researchers if diet, lifestyle, or occupation affects how a disease or disorder develops and progresses. Results from these studies provide information that helps answer questions such as: How fast will a disease or disorder usually progress? How bad will the condition become? Will treatment be needed?

What Is Expected of Patients in a Clinical Trial?

Not everyone can take part in a clinical trial for a specific disease or disorder. Each study enrolls patients with certain features or eligibility criteria. These criteria may include the type and stage of disease or disorder, as well as, the age and previous treatment history of the patient. You or your doctor can contact the sponsoring organization to find out more about specific clinical trials and their eligibility criteria. If you are interested in joining a clinical trial, your doctor must contact one of the trial’s investigators and provide details about your diagnosis and medical history.

If you participate in a clinical trial, you may be required to have a number of medical tests. You may also need to take medications and/or undergo

surgery. Depending upon the treatment and the examination procedure, you may be required to receive inpatient hospital care. Or, you may have to return to the medical facility for follow-up examinations. These exams help find out how well the treatment is working. Follow-up studies can take months or years. However, the success of the clinical trial often depends on learning what happens to patients over a long period of time. Only patients who continue to return for follow-up examinations can provide this important long-term information.

Recent Trials on Adult Primary Liver Cancer

The National Institutes of Health and other organizations sponsor trials on various diseases and disorders. Because funding for research goes to the medical areas that show promising research opportunities, it is not possible for the NIH or others to sponsor clinical trials for every disease and disorder at all times. The following lists recent trials dedicated to adult primary liver cancer.¹⁹ If the trial listed by the NIH is still recruiting, you may be eligible. If it is no longer recruiting or has been completed, then you can contact the sponsors to learn more about the study and, if published, the results. Further information on the trial is available at the Web site indicated. Please note that some trials may no longer be recruiting patients or are otherwise closed. Before contacting sponsors of a clinical trial, consult with your physician who can help you determine if you might benefit from participation.

- **An Evaluation of Chronic Thalidomide Administration in Patients Undergoing Chemoembolization for Unresectable Hepatocellular Cancer**

Condition(s): Liver Cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Center for Research Resources (NCRR)

Purpose - Excerpt: This is a clinical trial to test the safety and efficacy of the drug thalidomide in combination with a procedure called chemoembolization in patients with inoperable liver cancer. Chemoembolization is the process by which chemotherapy is instilled directly into the blood vessels feeding the tumor, so that the blood vessels feeding the tumor may be blocked. Chemoembolization consists of two separate procedures. It will be done by infusing chemotherapy with the drug doxorubicin through the hepatic artery into the liver and then by infusing collagen to cut off the blood supply to the tumor. A catheter will be inserted at various times to allow for these infusions. The objectives

¹⁹ These are listed at www.ClinicalTrials.gov.

are to investigate the feasibility and potential activity of chronic administration of thalidomide in patients with unresectable hepatocellular cancer who receive chemoembolization to predominant tumor masses. The toxicity of thalidomide in these patients will be evaluated. Overall safety will also be assessed. Serum levels of angiogenic cytokines such as VEGF, bFGF, and TNF- α , that are believed to have a role in hepatocellular carcinoma, will be collected.

Phase(s): Phase II

Study Type: Interventional

Contact(s): Madeline Peyton 1-212-263-6485; New York; Kaplan Comprehensive Cancer Center, New York, New York, 10016, United States; Recruiting; Madeline Peyton 212-263-6485

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00006198;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Antineoplaston Therapy in Treating Patients With Primary Liver Cancer**

Condition(s): recurrent childhood liver cancer; recurrent adult primary liver cancer; stage IV childhood liver cancer; advanced adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): Burzynski Research Institute

Purpose - Excerpt: Rationale: Antineoplastons are naturally occurring substances found in urine. Antineoplastons may inhibit the growth of cancer cells. Purpose: Phase II trial to study the effectiveness of antineoplaston therapy in treating patients who have primary liver cancer.

Phase(s): Phase II

Study Type: Treatment

Contact(s): Texas; Burzynski Research Institute, Houston, Texas, 77055, United States; Recruiting; Stanislaw R. Burzynski 713-335-5697. Study chairs or principal investigators: Stanislaw R. Burzynski, Study Chair; Burzynski Research Institute

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00003530;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **BMS-247550 in Treating Patients With Liver or Gallbladder Cancer**

Condition(s): recurrent adult primary liver cancer; localized gallbladder cancer; advanced adult primary liver cancer; cholangiocarcinoma of the gallbladder; localized resectable adult primary liver cancer; adult primary hepatocellular carcinoma; localized extrahepatic bile duct cancer; recurrent gallbladder cancer; unresectable gallbladder cancer; localized unresectable adult primary liver cancer; cholangiocarcinoma of the extrahepatic bile duct; adult primary cholangiocellular carcinoma; recurrent extrahepatic bile duct cancer; unresectable extrahepatic bile duct cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); University of Chicago Cancer Research Center

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Purpose: Phase II trial to study the effectiveness of BMS-247550 in treating patients who have liver or gallbladder cancer.

Phase(s): Phase II

Study Type: Treatment

Contact(s): see Web site below

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00023946;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Capecitabine Combined with Cisplatin in Treating Patients With Locally Advanced or Metastatic Solid Tumors**

Condition(s): lung cancer; pancreatic cancer; gastric cancer; lip and oral cavity cancer; esophageal cancer; head and neck cancer; adult primary liver cancer; oropharyngeal cancer; breast cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Kaplan Cancer Center

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Combining more than one chemotherapy drug may kill more tumor cells. Purpose: Phase I trial to study the effectiveness of capecitabine combined with cisplatin in treating patients who have locally advanced or metastatic solid tumors.

Phase(s): Phase I

Study Type: Treatment

Contact(s): New York; NYU School of Medicine's Kaplan Comprehensive Cancer Center, New York, New York, 10016, United States; Recruiting; Franco M. Muggia 212-263-6485. Study chairs or principal investigators: Franco M. Muggia, Study Chair; Kaplan Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00010023;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Chemotherapy in Treating Children With Liver Cancer**

Condition(s): stage III childhood liver cancer; stage II childhood liver cancer; stage I childhood liver cancer; recurrent childhood liver cancer; stage IV childhood liver cancer; childhood hepatoblastoma; childhood hepatocellular carcinoma

Study Status: This study is currently recruiting patients.

Sponsor(s): Societe Internationale d'Oncologie Pediatrique

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. It is not yet known which chemotherapy regimen is more effective in treating children with liver cancer. Purpose: Randomized phase III trial to study the effectiveness of cisplatin with or without doxorubicin and the effectiveness of combining cisplatin, carboplatin, and doxorubicin in treating children who have liver cancer.

Phase(s): Phase III

Study Type: Treatment

Contact(s): Italy; University of Padua, Padua, 35128, Italy; Recruiting; Giorgio Perilongo 0039-49-8213517; United Kingdom, England; Saint Bartholomew's Hospital, London, England, EC1A 7BE, United Kingdom; Recruiting; Liz Shafford 44-207-377-7796. Study chairs or principal investigators: Giorgio Perilongo, Study Chair; Societe Internationale d'Oncologie Pediatrique

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00003912;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Chemotherapy in Treating Patients With Recurrent or Unresectable Liver Cancer**

Condition(s): recurrent adult primary liver cancer; advanced adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): Zarix

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. It is not yet known which chemotherapy regimen is more effective for liver cancer. Purpose: Randomized phase III trial to compare the effectiveness of two different chemotherapy regimens in treating patients who have recurrent or unresectable liver cancer.

Phase(s): Phase III

Study Type: Treatment

Contact(s): see Web site below

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00012324;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Combination Chemotherapy and Tamoxifen in Treating Patients With Solid Tumors**

Condition(s): childhood soft tissue sarcoma; thyroid cancer; childhood liver cancer; adult soft tissue sarcoma; brain tumor; head and neck cancer; adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): Ottawa Regional Cancer Centre

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Estrogen can stimulate the growth of tumor cells. Hormone therapy using tamoxifen may fight cancer by blocking the uptake of estrogen. Combining tamoxifen and chemotherapy may kill more tumor cells. Purpose: Phase II trial to study the effectiveness of combination chemotherapy consisting of cisplatin and doxorubicin plus tamoxifen in treating patients who have solid tumors.

Phase(s): Phase II

Study Type: Treatment

Contact(s): Canada, Ontario; Ottawa Regional Cancer Centre, Ottawa, Ontario, K1H 1C4, Canada; Recruiting; Stan Z. Gertler 613-737-7700 ext. 56764. Study chairs or principal investigators: Stan Z. Gertler, Study Chair; Ottawa Regional Cancer Centre

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00002608;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Erlotinib in Treating Patients With Liver Cancer That Cannot be Surgically Removed**

Condition(s): advanced adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); M.D. Anderson Cancer Center

Purpose - Excerpt: Rationale: Erlotinib may stop the growth of tumor cells by blocking the enzymes necessary for tumor cell growth. Purpose: Phase II trial to study the effectiveness of erlotinib in treating patients who have liver cancer that cannot be surgically removed.

Phase(s): Phase II

Study Type: Treatment

Contact(s): Texas; University of Texas - MD Anderson Cancer Center, Houston, Texas, 77030-4009, United States; Recruiting; James L. Abbruzzese 713-792-2828. Study chairs or principal investigators: James L. Abbruzzese, Study Chair; M.D. Anderson Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00047333;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Erlotinib in Treating Patients With Unresectable Liver Cancer and Liver Dysfunction**

Condition(s): recurrent adult primary liver cancer; advanced adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); M.D. Anderson Cancer Center

Purpose - Excerpt: Rationale: Biological therapies such as erlotinib may interfere with the growth of tumor cells and slow the growth of the tumor. Purpose: Phase I trial to study the effectiveness of erlotinib in treating patients who have unresectable liver cancer and liver dysfunction.

Phase(s): Phase I

Study Type: Treatment

Contact(s): Texas; University of Texas - MD Anderson Cancer Center, Houston, Texas, 77030-4009, United States; Recruiting; James L. Abbruzzese 713-792-2828. Study chairs or principal investigators: James L. Abbruzzese, Study Chair; M.D. Anderson Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00047346;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Hepatic Arterial Infusion Plus Internal Radiation Therapy in Treating Patients With Liver Cancer That Cannot Be Removed By Surgery**

Condition(s): recurrent adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): University of Pittsburgh Cancer Institute

Purpose - Excerpt: Rationale: Hepatic arterial infusion uses a catheter to deliver anticancer substances directly into the liver. Internal radiation uses radioactive material placed directly into the tumor to kill tumor cells and cause less damage to normal tissue. Purpose: Phase II trial to study the effectiveness of combining hepatic arterial infusion with internal radiation therapy in treating patients who have liver cancer that cannot be removed by surgery.

Phase(s): Phase II

Study Type: Treatment

Contact(s): Pennsylvania; University of Pittsburgh Cancer Institute, Pittsburgh, Pennsylvania, 15213-3489, United States; Recruiting; Brian I. Carr 412-624-6672 (office). Study chairs or principal investigators: Brian I. Carr, Study Chair; University of Pittsburgh Cancer Institute

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00039078;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **High-Dose Combination Chemotherapy Plus Peripheral Stem Cell Transplantation in Treating Patients With Advanced Cancer**

Condition(s): melanoma; ovarian epithelial cancer; pancreatic cancer; colon cancer; gastric cancer; adult soft tissue sarcoma; rectal cancer; breast cancer; esophageal cancer; colorectal cancer; kidney tumor; adult primary liver cancer; bone cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Beckman Research Institute

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Peripheral stem cell transplantation may allow the doctor to give higher doses of chemotherapy drugs and kill more tumor cells. Purpose: Phase I trial to study the effectiveness of combination chemotherapy plus peripheral stem cell transplantation in treating patients who have advanced cancer.

Phase(s): Phase I

Study Type: Treatment

Contact(s): California; Cancer Center and Beckman Research Institute, City of Hope, Duarte, California, 91010-3000, United States; Recruiting; George Somlo 626-359-8111. Study chairs or principal investigators: George Somlo, Study Chair; Beckman Research Institute

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00002854;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Liver Perfusion for Patients with Inoperable Liver Cancer**

Condition(s): Liver Cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI)

Purpose - Excerpt: This study will determine the maximum safe dose and side effects of the drug melphalan given through the DelCath system to treat liver cancer. In this system, the chemotherapy is delivered to the liver through special catheters (flexible plastic tubes) inserted through small puncture holes in the groin area into the blood vessels going into and out of the liver. This allows the drug to be administered directly to the liver in amounts that would not be safe if it circulated throughout the body. Patients 14 years of age or older with cancer that has arisen in or spread mainly to the liver and cannot be removed surgically may be eligible for this study. Candidates will be screened with a physical examination, blood tests, electrocardiogram (EKG), chest X-ray, and imaging studies, such as X-ray studies, CT scan of the chest, abdomen and pelvis and MRI scan of the liver, to evaluate the tumor. Other tests may be done if medically indicated. Participants will be admitted to the hospital for treatment under sedation. The groin area is numbed and catheters are placed into the large artery and vein that supply blood to and from the liver, creating a separate circulation for the liver, similar to the heart-lung bypass in open-heart surgery. Once the catheters are in

place, melphalan is perfused into the liver for 30 minutes. The circulation to the liver remains separate and the blood is filtered through the DelCath system for another 30 minutes after the drug is given to remove as much of it as possible from the blood coming out of the liver. Following the procedure, patients stay in the intensive care unit for 24 hours for observation. Patients who enter the study early receive lower doses of melphalan. The dose is increased as the study progresses to determine the highest safe and effective dose. Patients will be informed of the progress of the study at the point they enter. Lower doses may be less effective against the tumor, and higher doses may have a greater risk of side effects. Patients stay in the hospital for 3 to 5 days during the perfusion part of the study and return to the outpatient clinic after 3 weeks to evaluate recovery. If recovery is satisfactory, a second treatment will be scheduled. About 4 weeks after the second treatment, the tumor will be evaluated to assess the response to therapy. Patients whose tumor has shrunk or stayed the same will be offered another two treatments. When the treatments are finished, patients will return for evaluation every 3 months for 2 years and then every 4 months for a third year. During these visits they will have a physical examination, imaging scans, X-rays and blood tests to evaluate disease status. If the liver tumor grows or spreads beyond the liver during this time, the patient will be taken off the study and other alternatives for care will be discussed.

Phase(s): Phase I

Study Type: Interventional

Contact(s): Maryland; National Cancer Institute (NCI), 9000 Rockville Pike Bethesda, Maryland, 20892, United States; Recruiting; Cynthia Helsabeck, R.N. 3014954012 cynthia_helsabeck@nih.gov

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00021606;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Liver Perfusion of Melphalan in Inoperable Liver Cancer**

Condition(s): Liver Neoplasm; Neoplasm Metastasis

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI)

Purpose - Excerpt: This study is testing whether administration of a drug called melphalan directly into the liver (isolated hepatic perfusion) can shrink tumors in patients with inoperable cancer whose tumor is confined to the liver. Blood tests and special X-ray studies will be done before treatment and over a period of several months after treatment to

evaluate the treatment's effectiveness. Similar perfusion studies with melphalan have been done in patients with cancer (who received another drug together with melphalan) and perfusion of the extremity has been performed in patients with a serious type of skin cancer called melanoma. In both studies tumors shrank, some up to less than half the original size. In some patients, tumors went away completely. Isolated liver perfusion is a major operation, done under general anesthesia. In this procedure, catheters (plastic tubes) are placed into the large blood vessels that deliver blood to and remove blood from the liver, creating a circulation to the liver separate from the rest of the body. During the operation, the liver receives oxygen from an external pump similar to the heart-lung machine used in open-heart surgery. For one hour, melphalan is administered through the pump in large doses that would not be safe if it reached all parts of the body in the normal blood circulation. The entire operation lasts between six and eight hours.

Phase(s): Phase II

Study Type: Interventional

Contact(s): Maryland; National Cancer Institute (NCI), 9000 Rockville Pike Bethesda, Maryland, 20892, United States; Recruiting; Clinical Studies Support Center/NCI 1-888-624-1937 ncicssc@mail.nih.gov

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00001820;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Long Term Interferon for Patients Who Did Not Clear Hepatitis C Virus with Standard Treatment**

Condition(s): Chronic Hepatitis C; Cirrhosis, Liver; Fibrosis, Liver; Hepatic Cirrhosis

Study Status: This study is currently recruiting patients.

Sponsor(s): National Institute of Diabetes and Digestive and Kidney Diseases (NIDDK); National Institute of Allergy and Infectious Diseases (NIAID); National Center on Minority Health and Health Disparities (NCMHD); National Cancer Institute (NCI); Hoffmann-La Roche Ltd

Purpose - Excerpt: The HALT-C Trial is a National Institute of Diabetes and Digestive and Kidney Diseases sponsored, randomized clinical trial of long-term use of Peginterferon alfa-2a (pegylated interferon) in patients who failed to respond to prior interferon treatment. All patients who enter the trial will be treated for 6 months with Peginterferon alfa-2a and Ribavirin. Patients who respond to this 6 month treatment will continue to be treated for an additional 6 months. Patients who do not

respond to this treatment will be eligible for the long-term maintenance phase of this study where patients will be randomly selected to be treated with Peginterferon alfa-2a or to discontinue treatment for 3.5 years. Patients in both arms of this study will be followed closely with quarterly study visits. The combination of peginterferon plus ribavirin has recently been approved by the FDA for treatment of chronic HCV. Patients who remain HCV-RNA positive after being treated for at least 6 months with peginterferon and ribavirin outside of this study may be eligible to directly enter the randomized portion of the HALT-C Trial. The HALT-C study is designed to determine if continuing interferon long-term over several years will suppress Hepatitis C virus, prevent progression to cirrhosis, prevent liver cancer and reduce the need for liver transplantation.

Phase(s): Phase III

Study Type: Interventional

Contact(s): see Web site below

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00006164;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Magnetic-Targeted Doxorubicin in Treating Patients with Cancer Metastatic to the Liver**

Condition(s): Metastases, Neoplasm; Colorectal Neoplasms; Esophageal Neoplasms; Stomach Neoplasms; Pancreatic Neoplasms; Breast Neoplasms; Melanoma; Sarcoma; Gastrointestinal Neoplasms; Lung Neoplasms; Liver Neoplasms; Cholangiocarcinoma

Study Status: This study is currently recruiting patients.

Sponsor(s): FeRx

Purpose - Excerpt: MTC-DOX is Doxorubicin or DOX, a chemotherapy drug, that is adsorbed, or made to "stick", to magnetic beads (MTCs). MTCs are tiny, microscopic particles of iron and carbon. When DOX is added to MTCs, DOX attaches to the carbon part of the MTCs. MTC-DOX is directed to and deposited in the area of a tumor, where it is thought that it then "leaks" through the blood vessel walls. Once in the surrounding tissues, it is thought that Doxorubicin becomes "free from" the magnetic beads and will then be able to act against the tumor cells. The iron component of the particle has magnetic properties, making it possible to direct MTC-DOX to specific tumor sites in the liver by placing a magnet on the body surface. It is hoped that MTC-DOX used with the

magnet may target the chemotherapy drug directly to liver tumors and provide a treatment to patients with cancers that have spread to the liver.

Phase(s): Phase I; Phase II

Study Type: Interventional

Contact(s): California; UCSF Cancer Center, San Francisco, California, 94143, United States; Recruiting; Jean Anne Donnell, RN 415-353-7067; Alan P Venook, MD, Principal Investigator; Scripps Stevens Cancer Division, San Diego, California, 92037, United States; Recruiting; Michelle Gould, RN 858-882-8334; Cheryl Kosits, RN 858-882-8320; Richard G. Just, MD, Principal Investigator. Study chairs or principal investigators: Joy Koda, PhD, Study Chair

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00041808;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Megestrol in Treating Patients With Liver Cancer That Cannot Be Removed By Surgery**

Condition(s): advanced adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): NMRC Asia-Pacific Hepatocellular Carcinoma Trials Group

Purpose - Excerpt: Rationale: Estrogen can stimulate the growth of cancer cells. Hormone therapy using megestrol may fight liver cancer by blocking the uptake of estrogen. It is not yet known if megestrol is an effective treatment for liver cancer. Purpose: Randomized phase III trial to determine the effectiveness of megestrol in treating patients who have liver cancer that cannot be removed by surgery.

Phase(s): Phase III

Study Type: Treatment

Contact(s): see Web site below

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00041275;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Ondansetron With or Without Dexamethasone to Prevent Vomiting in Patients Receiving Radiation Therapy to the Upper Abdomen**

Condition(s): pancreatic cancer; gastric cancer; colorectal cancer; adult primary liver cancer; testicular cancer; ovarian epithelial cancer; colon cancer; cervical cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): NCIC-Clinical Trials Group

Purpose - Excerpt: Rationale: Antiemetic drugs may help to reduce or prevent vomiting in patients treated with radiation therapy. It is not yet known if ondansetron is more effective with or without dexamethasone in preventing vomiting caused by radiation therapy. Purpose: Randomized phase III trial to compare the effectiveness of ondansetron with or without dexamethasone in preventing vomiting in patients with cancer who are receiving radiation therapy to the upper abdomen.

Phase(s): Phase III

Study Type: Supportive Care

Contact(s): see Web site below

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00016380;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Radiation Therapy and Fluorouracil Before Surgery in Treating Patients With Primary or Recurrent Bile Duct Cancer**

Condition(s): recurrent adult primary liver cancer; localized resectable adult primary liver cancer; localized extrahepatic bile duct cancer; recurrent extrahepatic bile duct cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): EORTC Chronotherapy Study Group

Purpose - Excerpt: Rationale: Radiation therapy uses high-energy x-rays to damage tumor cells. Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Combining radiation therapy with chemotherapy before surgery may shrink the tumor so that it can be removed during surgery. Purpose: Phase II trial to study the effectiveness of combining radiation therapy with fluorouracil before surgery in treating patients who have primary or recurrent bile duct cancer.

Phase(s): Phase II

Study Type: Treatment

Contact(s): Virginia; Cancer Center at the University of Virginia, Charlottesville, Virginia, 22908, United States; Recruiting; Tyvin Andrew Rich 804-924-9412. Study chairs or principal investigators: Tyvin Andrew Rich, Study Chair; EORTC Chronotherapy Study Group

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00030511;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Radioactive Iodine in Treating Patients Who Have Undergone Surgery for Liver Cancer**

Condition(s): localized resectable adult primary liver cancer; adult primary hepatocellular carcinoma

Study Status: This study is currently recruiting patients.

Sponsor(s): NMRC Asia-Pacific Hepatocellular Carcinoma Trials Group

Purpose - Excerpt: Rationale: Radioactive iodine may be effective in reducing the rate of recurrence of liver cancer after surgery to remove the tumor. It is not yet known if radioactive iodine is more effective than no further treatment after surgery. Purpose: Randomized phase III trial to determine the effectiveness of radioactive iodine in treating patients who have undergone surgery for liver cancer.

Phase(s): Phase III

Study Type: Treatment

Contact(s): Singapore; Changi General Hospital, Singapore, 529889, Singapore; Recruiting; Tay Khoon Hean; National Cancer Centre - Singapore, Singapore, 169610, Singapore; Recruiting; Pierce Chow +65 321-4051; National University Hospital, Singapore, 119074, Singapore; Recruiting; Leow Chon Kar 7724350. Study chairs or principal investigators: London Lucien Ooi Peng Jin, Study Chair; NMRC Asia-Pacific Hepatocellular Carcinoma Trials Group

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00027768;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Rebeccamycin Analogue in Treating Patients With Advanced Liver and/or Biliary Cancer**

Condition(s): advanced adult primary liver cancer; cholangiocarcinoma of the gallbladder; adult primary hepatocellular carcinoma; unresectable gallbladder cancer; localized unresectable adult primary liver cancer;

cholangiocarcinoma of the extrahepatic bile duct; adult primary cholangiocellular carcinoma; unresectable extrahepatic bile duct cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Ireland Cancer Center

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Purpose: Phase II trial to study the effectiveness of rebeccamycin analogue in treating patients who have advanced liver and/or biliary cancer.

Phase(s): Phase II

Study Type: Treatment

Contact(s): see Web site below

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00005997;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Safety and Efficacy of Doxorubicin Adsorbed to Magnetic Beads vs. IV Doxorubicin in Treating Liver Cancer**

Condition(s): Carcinoma, Hepatocellular

Study Status: This study is currently recruiting patients.

Sponsor(s): FeRx

Purpose - Excerpt: MTC-DOX is Doxorubicin or DOX, a chemotherapy drug, that is adsorbed, or made to "stick", to magnetic beads (MTCs). MTCs are tiny, microscopic particles of iron and carbon. When DOX is added to MTCs, DOX attaches to the carbon part of the MTCs. MTC-DOX is directed to and deposited in the area of a tumor, where it is thought that it then "leaks" through the blood vessel walls. Once in the surrounding tissues, it is thought that Doxorubicin becomes "free from" the magnetic beads and will then be able to act against the tumor cells. The iron component of the particle has magnetic properties, making it possible to direct MTC-DOX to specific tumor sites in the liver by placing a magnet on the body surface. It is hoped that MTC-DOX used with the magnet may target the chemotherapy directly to liver tumors and provide a treatment to patients with liver cancer. To be sure of the effect of MTC-DOX on liver cancer, it will be compared to the effect of Doxorubicin given through the vein. The study treatments will be administered every three weeks, (which is considered a study treatment cycle), until you complete six treatment cycles, the tumor grows, disappears, or you experience a side effect, which may cause you to leave the study. Follow-up visits will occur on Days 3, 10, and 21 following

treatment in the first cycle and Days 7 and 21 for the remaining cycles, and also 60 days after you receive your last treatment cycle. Therefore, the purpose of this Phase 2/3 study is to evaluate safety, tolerance, and efficacy (survival time) of an MTC-DOX dosing strategy where the DOX dose is determined by tumor size

Phase(s): Phase II; Phase III

Study Type: Interventional

Contact(s): see Web site below

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00034333;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Sho-Saiko-To Following Removal of Liver Cancer By Embolization in Treating Patients With Liver Cancer That Cannot Be Surgically Removed**

Condition(s): recurrent adult primary liver cancer; advanced adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Memorial Sloan-Kettering Cancer Center

Purpose - Excerpt: Rationale: The Chinese herbal medicine Sho-saiko-to contains ingredients that may slow the growth of tumor cells and stimulate a person's immune system to help kill tumor cells. This may be an effective treatment following hepatic artery embolization. Purpose: Phase II trial to study the effectiveness of Sho-saiko-to following hepatic artery embolization in treating patients who have liver cancer.

Phase(s): Phase II

Study Type: Treatment

Contact(s): New York; Memorial Sloan-Kettering Cancer Center, New York, New York, 10021, United States; Recruiting; Yuman Fong 212-639-2016. Study chairs or principal investigators: Yuman Fong, Study Chair; Memorial Sloan-Kettering Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00040898;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Stem Cell Transplantation for Metastatic Solid Tumors**

Condition(s): Cholangiocarcinoma; Colon/Rectal Ca; Bladder Ca; Breast Ca; Basal Cell Ca; Adrenal Ca; Esophageal/Gastric Ca; Hepatocellular Ca; Ovarian Ca; Prostate Ca; Small Cell Lung Ca; Non Small Cell Lung Ca; Adenocarcinoma, Unk origin; Pancreatic Ca; Bony/Soft Tissue Sarcoma

Study Status: This study is currently recruiting patients.

Sponsor(s): National Heart, Lung, and Blood Institute (NHLBI)

Purpose - Excerpt: The goal of this research study is to identify other types of cancer (malignant neoplasms) that may be treatable with stem cell transplantation (allogenic peripheral blood stem cell transplantation. Patients with a variety of different types of cancerous tumors that have spread (metastasized) and whose conditions have not improved with stand therapy, will be eligible to participate. Those patients selected to participate in the study will undergo a procedure known as a "mini-transplant". The mini-transplant is a transplantation of stem-cells collected from a sibling (brother or sister) of the patient. Unlike traditional bone marrow transplants, the mini-transplant does not require intense chemotherapy or radiation therapy. Because of this, patients experience fewer and less severe side effects. This study is open to patients diagnosed with a variety of metastatic solid tumors including esophageal, gastric (stomach), colon, rectal, liver tumors (hepatoma), cancer of the biliary system (cholangiocarcinoma), cancer of the pancreas, lung, breast, prostate, bone (sarcoma), adrenal basal cell, bladder, and adenocarcinomas of unk primary origin.

Phase(s): Phase II

Study Type: Interventional

Contact(s): Maryland; National Heart, Lung and Blood Institute (NHLBI), 9000 Rockville Pike Bethesda, Maryland, 20892, United States; Recruiting; Patient Recruitment and Public Liaison Office 1-800-411-1222 prpl@mail.cc.nih.gov; TTY 1-866-411-1010

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00001880;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Study of T900607-sodium in chemotherapy naive patients with hepatocellular carcinoma.**

Condition(s): Hepatocellular Carcinoma

Study Status: This study is currently recruiting patients.

Sponsor(s): Tularik

Purpose - Excerpt: The purpose of the study is to determine whether T900607-sodium is effective and safe in treating hepatocellular carcinoma, a type of liver cancer.

Phase(s): Phase II

Study Type: Interventional

Contact(s): see Web site below

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00043433;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Thalidomide and Chemoembolization With Doxorubicin in Treating Patients With Liver Cancer That Cannot be Removed by Surgery**

Condition(s): advanced adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Kaplan Cancer Center

Purpose - Excerpt: Rationale: Thalidomide may stop the growth of liver cancer by stopping blood flow to the tumor. Chemoembolization kills tumor cells by delivering drugs directly into the tumor and then blocking the blood flow to the tumor. Combining thalidomide with chemoembolization may kill more tumor cells. Purpose: Phase II trial to study the effectiveness of combining thalidomide and chemoembolization in treating patients who have liver cancer that cannot be removed by surgery.

Phase(s): Phase II

Study Type: Treatment

Contact(s): New York; Mount Sinai School of Medicine, New York, New York, 10029, United States; Recruiting; Max W. Sung 212-241-7902; NYU School of Medicine's Kaplan Comprehensive Cancer Center, New York, New York, 10016, United States; Recruiting; Franco M. Muggia 212-263-6485. Study chairs or principal investigators: Alec S. Goldenberg, Study Chair; Kaplan Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00006016;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Thalidomide Plus Interferon alfa in Treating Patients With Progressive Liver Cancer That Cannot be Surgically Removed**

Condition(s): recurrent adult primary liver cancer; advanced adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Kaplan Cancer Center

Purpose - Excerpt: Rationale: Thalidomide may stop the growth of liver cancer by stopping blood flow to the tumor. Interferon alfa may interfere with the growth of the cancer cells. Combining thalidomide and interferon alfa may kill more tumor cells. Purpose: Phase II trial to study the effectiveness of thalidomide plus interferon alfa in treating patients who have progressive liver cancer that cannot be surgically removed.

Phase(s): Phase II

Study Type: Treatment

Contact(s): New York; Mount Sinai School of Medicine, New York, New York, 10029, United States; Recruiting; Jonathan Schwartz 212-241-3984; NYU School of Medicine's Kaplan Comprehensive Cancer Center, New York, New York, 10016, United States; Recruiting; Franco M. Muggia 212-263-6485. Study chairs or principal investigators: Matthew D. Volm, Study Chair; Kaplan Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00006006;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Treatment of hepatocellular carcinoma with Tetrathiomolybdate**

Condition(s): Carcinoma, Hepatocellular

Study Status: This study is currently recruiting patients.

Sponsor(s): National Center for Research Resources (NCRR)

Purpose - Excerpt: Hepatocellular carcinoma (HCC) is a deadly tumor for which the incidence is increasing in the United States, primarily due to prevalence of hepatitis C infection. An important aspect of the development of HCC is that it occurs in patients who have underlying cirrhosis of the liver, thereby limiting the therapeutic options. There is potential curative treatment for these patients, such as resection of the tumor lesion and liver transplantation, but these treatments are feasible in a small percent of patients only. Furthermore, the majority of the patients with HCC are also not candidates for palliative treatments such as percutaneous ablation of the tumor, chemotherapy or radiation.

Additionally, it has been shown that these palliative treatment modalities do not alter survival, and are associated with significant risks. Therefore, there are no treatment options for most patients with HCC. A new theory has emerged in the fight against cancer through inhibition of angiogenesis (development of new blood vessels). The hypothesis being that if there is no blood supply "feeding" the tumor cells cannot divide or survive. One such approach, pioneered in this institution by Drs. George Brewer and Sofia Merajver, is the anticopper approach using the medication tetrathiomolybdate (TM). By creating a mild copper deficiency state, several pathways required for angiogenesis are inhibited. They performed a Phase I trial in which patients with metastatic cancer were treated with TM resulting in decrease tumor vascularity. TM had excellent safety profile in this patient population. HCC is well known to be a hypervascular tumor. An antiangiogenesis approach might provide a novel treatment for this HCC. This is a pilot study of 10 patients with HCC who are not candidates for curative surgical therapy with resection or liver transplantation, nor for ablative techniques. Patients seen in the General Liver clinic and Liver Transplant clinic who have an overall good performance status, with an expected survival of more than 6 months will be enrolled. After an initial evaluation, they will be given 120 mg/day of TM in divided doses for one year. The size and vascularity of the tumor will be evaluated by magnetic resonance imaging (MRI). The primary outcome of this study is to prevent tumor progression.

Phase(s): Phase II

Study Type: Interventional

Contact(s): Jorge A. Marrero, M.D. 1-734-936-4780; Michigan; 3912 Taubman Center, Ann Arbor, Michigan, 48109-0362, United States; Recruiting; Jorge A. Marrero, M.D. 734-936-4780

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00006332;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **UCN-01 and Irinotecan in Treating Patients With Advanced Solid Tumors**

Condition(s): lung cancer; pancreatic cancer; gastric cancer; esophageal cancer; colorectal cancer; adult primary liver cancer; ovarian epithelial cancer; colon cancer; rectal cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Sidney Kimmel Cancer Center

Purpose - Excerpt: Rationale: UCN-01 may stop the growth of tumor cells by blocking the enzymes necessary for tumor cell growth. Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Combining UCN-01 with chemotherapy may kill more tumor cells. Purpose: Phase I trial to study the effectiveness of combining UCN-01 with irinotecan in treating patients who have advanced solid tumors.

Phase(s): Phase I

Study Type: Treatment

Contact(s): Maryland; Sidney Kimmel Comprehensive Cancer Center at Johns Hopkins, Baltimore, Maryland, 21231-2410, United States; Recruiting; Ross C. Donehower 410-955-8838. Study chairs or principal investigators: Ross C. Donehower, Study Chair; Sidney Kimmel Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00047242;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **UCN-01 and Irinotecan in Treating Patients With Metastatic or Unresectable Solid Tumors**

Condition(s): lung cancer; pancreatic cancer; gastric cancer; lip and oral cavity cancer; endometrial cancer; esophageal cancer; head and neck cancer; colorectal cancer; adult primary liver cancer; ovarian sarcoma; ovarian epithelial cancer; colon cancer; prostate cancer; cervical cancer; oropharyngeal cancer; rectal cancer; breast cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Washington University School of Medicine

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. UCN-01 may help irinotecan kill more cancer cells by making tumor cells more sensitive to the drug. Purpose: Phase I trial to study the effectiveness of combining UCN-01 with irinotecan in treating patients who have metastatic or unresectable solid tumors.

Phase(s): Phase I

Study Type: Treatment

Contact(s): Missouri; Washington University School of Medicine, Saint Louis, Missouri, 63110, United States; Recruiting; Paula M. Fracasso 314-454-8817. Study chairs or principal investigators: Paula M. Fracasso, Study Chair; Washington University School of Medicine

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00031681;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Vaccine Therapy in Treating Patients With Liver Cancer**

Condition(s): recurrent adult primary liver cancer; advanced adult primary liver cancer; localized resectable adult primary liver cancer; adult primary hepatocellular carcinoma; localized unresectable adult primary liver cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Jonsson Comprehensive Cancer Center

Purpose - Excerpt: Rationale: Vaccines made from a person's white blood cells mixed with tumor proteins may make the body build an immune response to kill tumor cells. Purpose: Phase I/II trial to study the effectiveness of vaccine therapy in treating patients who have liver cancer.

Phase(s): Phase I; Phase II

Study Type: Treatment

Contact(s): California; Jonsson Comprehensive Cancer Center, UCLA, Los Angeles, California, 90095-1781, United States; Recruiting; John A. Glaspy 310-794-1274. Study chairs or principal investigators: James S. Economou, Study Chair; Jonsson Comprehensive Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00022334;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Vaccine Therapy With or Without Sargramostim in Treating Patients With Advanced or Metastatic Cancer**

Condition(s): lung cancer; pancreatic cancer; gastric cancer; salivary gland cancer; head and neck cancer; colorectal cancer; adult primary liver cancer; testicular cancer; ovarian epithelial cancer; colon cancer; thyroid cancer; rectal cancer; breast cancer

Study Status: This study is currently recruiting patients.

Sponsor(s): National Cancer Institute (NCI); Fox Chase Cancer Center

Purpose - Excerpt: Rationale: Vaccines may make the body build an immune response to kill tumor cells. Colony-stimulating factors such as sargramostim may increase the number of immune cells found in bone marrow or peripheral blood. Combining vaccine therapy with

sargramostim may make tumor cells more sensitive to the vaccine and may kill more tumor cells. Purpose: Phase I trial to study the effectiveness of vaccine therapy with or without sargramostim in treating patients who have advanced or metastatic cancer.

Phase(s): Phase I

Study Type: Treatment

Contact(s): Pennsylvania; Fox Chase Cancer Center, Philadelphia, Pennsylvania, 19111, United States; Recruiting; Margaret von Mehren 215-728-3545. Study chairs or principal investigators: Margaret von Mehren, Study Chair; Fox Chase Cancer Center

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00028496;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Herbal Treatment of Hepatitis C in Methadone Maintained Patients**

Condition(s): Hepatitis C

Study Status: This study is no longer recruiting patients.

Sponsor(s): National Center for Complementary and Alternative Medicine (NCCAM)

Purpose - Excerpt: Hepatitis C (HCV) is a chronic viral illness leading to progressive liver damage that has emerged as a major public health issue in the United States. While HCV affects all population groups, individuals with a history of intravenous drug use form the largest known risk group. Between 90 and 100 percent of long term intravenous drug use will eventually test positive for HCV, and there is substantial risk that even short term experimentation will result in infection. Studies suggest that HCV will be the major cause of cirrhosis and liver cancer in the next century. Currently, approved therapy includes recombinant interferons, which lead to sustained remission in a minority of patients. However, patients abusing other substances, including alcohol, are not eligible for interferon therapy. The need for investigation into other potential therapies is clear. Current practice patterns in the Far East include the use of traditional herbal remedies for symptomatic chronic viral hepatitis. This study is intended to examine the effect of commonly used herbal remedies for the treatment of symptomatic HCV.

Phase(s): Phase II

Study Type: Interventional

Contact(s): Minnesota; Hennepin County Medical Center, Minneapolis, Minnesota, United States. Study chairs or principal investigators: Jeff

Albrecht, MD, Principal Investigator; Hennepin County Medical Center - Minneapolis

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00010816;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Gemcitabine and Capecitabine in Treating Patients With Unresectable Locally Advanced or Metastatic Cancer of the Gallbladder or Bile Duct**

Condition(s): adenocarcinoma with squamous metaplasia of the gallbladder; adenocarcinoma of the extrahepatic bile duct; squamous cell carcinoma of the gallbladder; cholangiocarcinoma of the gallbladder; recurrent gallbladder cancer; unresectable gallbladder cancer; cholangiocarcinoma of the extrahepatic bile duct; adult primary cholangiocellular carcinoma; adenocarcinoma of the gallbladder; recurrent extrahepatic bile duct cancer; unresectable extrahepatic bile duct cancer

Study Status: This study is not yet open for patient recruitment.

Sponsor(s): National Cancer Institute (NCI); Southwest Oncology Group

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Combining more than one drug may kill more tumor cells. Purpose: Phase II trial to study the effectiveness of combining gemcitabine with capecitabine in treating patients who have locally advanced or metastatic cancer of the gallbladder or bile duct.

Phase(s): Phase II

Study Type: Treatment

Contact(s): Charles A. Coltman, Jr. 210-616-5580. Study chairs or principal investigators: Charles A. Coltman, Jr., Study Chair; Southwest Oncology Group

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00033540;jsessionid=9BD291013B26CDD4524E13722A4347FA>

- **Oblimersen and Doxorubicin in Treating Patients With Advanced Liver Cancer or Other Solid Tumor**

Condition(s): recurrent adult primary liver cancer; advanced adult primary liver cancer; adult primary hepatocellular carcinoma; unspecified adult solid tumor, protocol specific; localized unresectable adult primary liver cancer

Study Status: This study is not yet open for patient recruitment.

Sponsor(s): National Cancer Institute (NCI); Princess Margaret Hospital

Purpose - Excerpt: Rationale: Drugs used in chemotherapy use different ways to stop tumor cells from dividing so they stop growing or die. Oblimersen may increase the effectiveness of doxorubicin by making tumor cells more sensitive to the drugs. Purpose: Phase I/II trial to study the effectiveness of combining oblimersen with doxorubicin in treating patients who have locally advanced, recurrent, or metastatic liver cancer or other solid tumor.

Phase(s): Phase I; Phase II

Study Type: Treatment

Contact(s): Jennifer Knox 416-946-2399. Study chairs or principal investigators: Jennifer Knox, Study Chair; Princess Margaret Hospital

Web Site:

<http://clinicaltrials.gov/ct/gui/show/NCT00047229;jsessionid=9BD291013B26CDD4524E13722A4347FA>

Benefits and Risks²⁰

What Are the Benefits of Participating in a Clinical Trial?

If you are interested in a clinical trial, it is important to realize that your participation can bring many benefits to you and society at large:

- A new treatment could be more effective than the current treatment for adult primary liver cancer. Although only half of the participants in a clinical trial receive the experimental treatment, if the new treatment is proved to be more effective and safer than the current treatment, then those patients who did not receive the new treatment during the clinical trial may be among the first to benefit from it when the study is over.
- If the treatment is effective, then it may improve health or prevent diseases or disorders.
- Clinical trial patients receive the highest quality of medical care. Experts watch them closely during the study and may continue to follow them after the study is over.

²⁰ This section has been adapted from ClinicalTrials.gov, a service of the National Institutes of Health:
http://www.clinicaltrials.gov/ct/gui/c/a1r/info/whatis?JServSessionIdzone_ct=9jmun6f291.

- People who take part in trials contribute to scientific discoveries that may help other people with adult primary liver cancer. In cases where certain diseases or disorders run in families, your participation may lead to better care or prevention for your family members.

The Informed Consent

Once you agree to take part in a clinical trial, you will be asked to sign an “informed consent.” This document explains a clinical trial’s risks and benefits, the researcher’s expectations of you, and your rights as a patient.

What Are the Risks?

Clinical trials may involve risks as well as benefits. Whether or not a new treatment will work cannot be known ahead of time. There is always a chance that a new treatment may not work better than a standard treatment. There is also the possibility that it may be harmful. The treatment you receive may cause side effects that are serious enough to require medical attention.

How Is Patient Safety Protected?

Clinical trials can raise fears of the unknown. Understanding the safeguards that protect patients can ease some of these fears. Before a clinical trial begins, researchers must get approval from their hospital’s Institutional Review Board (IRB), an advisory group that makes sure a clinical trial is designed to protect patient safety. During a clinical trial, doctors will closely watch you to see if the treatment is working and if you are experiencing any side effects. All the results are carefully recorded and reviewed. In many cases, experts from the Data and Safety Monitoring Committee carefully monitor each clinical trial and can recommend that a study be stopped at any time. You will only be asked to take part in a clinical trial as a volunteer giving informed consent.

What Are a Patient's Rights in a Clinical Trial?

If you are eligible for a clinical trial, you will be given information to help you decide whether or not you want to participate. As a patient, you have the right to:

- Information on all known risks and benefits of the treatments in the study.
- Know how the researchers plan to carry out the study, for how long, and where.
- Know what is expected of you.
- Know any costs involved for you or your insurance provider.
- Know before any of your medical or personal information is shared with other researchers involved in the clinical trial.
- Talk openly with doctors and ask any questions.

After you join a clinical trial, you have the right to:

- Leave the study at any time. Participation is strictly voluntary. However, you should not enroll if you do not plan to complete the study.
- Receive any new information about the new treatment.
- Continue to ask questions and get answers.
- Maintain your privacy. Your name will not appear in any reports based on the study.
- Know whether you participated in the treatment group or the control group (once the study has been completed).

What Should You Ask before Deciding to Join a Clinical Trial?

Questions you should ask when thinking about joining a clinical trial include the following:

- What is the purpose of the clinical trial?
- What are the standard treatments for adult primary liver cancer? Why do researchers think the new treatment may be better? What is likely to happen to me with or without the new treatment?
- What tests and treatments will I need? Will I need surgery? Medication? Hospitalization?

- How long will the treatment last? How often will I have to come back for follow-up exams?
- What are the treatment's possible benefits to my condition? What are the short- and long-term risks? What are the possible side effects?
- Will the treatment be uncomfortable? Will it make me feel sick? If so, for how long?
- How will my health be monitored?
- Where will I need to go for the clinical trial? How will I get there?
- How much will it cost to be in the study? What costs are covered by the study? How much will my health insurance cover?
- Will I be able to see my own doctor? Who will be in charge of my care?
- Will taking part in the study affect my daily life? Do I have time to participate?
- How do I feel about taking part in a clinical trial? Are there family members or friends who may benefit from my contributions to new medical knowledge?

Clinical Trials and Insurance Coverage²¹

As you consider enrolling in a clinical trial, you will face the critical issue of how to cover the costs of care. Even if you have health insurance, your coverage may not include some or all of the patient care costs associated with a clinical trial. This is because some health plans define clinical trials as “experimental” or “investigational” procedures.

Because lack of coverage for these costs can keep people from enrolling in trials, the National Cancer Institute is working with major health plans and managed care groups to find solutions. In the meantime, there are strategies that may help you deal with cost and coverage barriers. This section answers frequently asked questions about insurance coverage for clinical trial participation and directs you to additional information resources.

The material here is mainly concerned with treatment clinical trials, since other types of trials (prevention, screening, etc.) are newer and generally not covered by health insurance at all. However, this guide may become more

²¹ Adapted from the NCI:

http://www.cancer.gov/clinical_trials/doc_header.aspx?viewid=1d92be79-8748-4bda-8005-2a56d332463b.

relevant for prevention and other types of trials as these trials grow more common.

If you do not have any health insurance, you may find this section helpful for understanding some of the costs that trials involve.

What Costs Do Trials Involve? Who Is Usually Responsible for Paying Them?

There are two types of costs associated with a trial: patient care costs and research costs.

Patient care costs fall into two categories:

- Usual care costs, such as doctor visits, hospital stays, clinical laboratory tests, x-rays, etc., which occur whether you are participating in a trial or receiving standard treatment. These costs have usually been covered by a third-party health plan, such as Medicare or private insurance.
- Extra care costs associated with clinical trial participation, such as the additional tests that may or may not be fully covered by the clinical trial sponsor and/or research institution.

The sponsor and the participant's health plan need to resolve coverage of these costs for particular trials.

Research costs are those associated with conducting the trial, such as data collection and management, research physician and nurse time, analysis of results, and tests purely performed for research purposes. Such costs are usually covered by the sponsoring organization, such as NCI or a pharmaceutical company.

Criteria Used by Health Plans to Make Reimbursement Decisions about Trials

Health insurance companies and managed care companies decide which health care services they will pay for by developing coverage policy regarding the specific services. In general, the most important factor determining whether something is covered is a health plan's judgment as to whether the service is established or investigational. Health plans usually designate a service as established if there is a certain amount of scientific data to show that it is safe and effective. If the health plan does not think that

such data exist in sufficient quantity, the plan may label the service as investigational.

Health care services delivered within the setting of a clinical trial are very often categorized as investigational and not covered. This is because the health plan thinks that the major reason to perform the clinical trial is that there is not enough data to establish the safety and effectiveness of the service being studied. Thus, for some health plans, any mention of the fact that the patient is involved in a clinical trial results in a denial of payment.

Your health plan may define specific criteria that a trial must meet before extending coverage, such as the following:

Sponsorship

Some plans may only cover costs of trials sponsored by organizations whose review and oversight of the trial is careful and scientifically rigorous, according to standards set by the health plan.

Trial Phase and Type

Some plans may cover patient care costs only for the clinical trials they judge to be “medically necessary” on a case-by-case basis. Trial phase may also affect coverage; for example, while a plan may be willing to cover costs associated with Phase III trials, which include treatments that have already been successful with a certain number of people, the plan may require some documentation of effectiveness before covering a Phase I or II trial.

While health plans are interested in efforts to improve prevention and screening, they currently seem less likely to have a review process in place for these trials. Therefore, it may be more difficult to get coverage for the care costs associated with them.

Some plans, especially smaller ones, will not cover any costs associated with a clinical trial. Policies vary widely, but in most cases your best bet is to have your doctor initiate discussions with the health plan.

Cost “Neutrality”

Some health plans may limit coverage to trials they consider cost-neutral (i.e., not significantly more expensive than the treatments considered standard).

Lack of Standard Therapy

Some plans limit coverage of trials to situations in which no standard therapy is available.

Facility and Personnel Qualifications

A health plan may require that the facility and medical staff meet specific qualifications to conduct a trial involving unique services, especially intensive therapy such as a bone marrow transplant (high-dose chemotherapy with bone marrow/ stem cell rescue).

Clinical Trials and Medicare Coverage

For up-to-date information about Medicare coverage of clinical trials, go to the Web site for the Centers for Medicaid & Medicare (<http://www.hcfa.gov/coverage/8d.htm>; formerly the Health Care Financing Administration). As of January 2001, the following information was accurate²²:

What Will Medicare Pay?

- Anything normally covered is still covered when it is part of a clinical trial. This includes test, procedures, and doctor visits that are ordinarily covered.
- Anything normally covered even if it is a service or item associated with the experimental treatment. For example, Medicare will pay for the

²² On June 7, 2000, Present Clinton announced that Medicare would revise its payment policy to reimburse the routine patient care costs of clinical trials. The announcement is available for public viewing at the following Web address:
http://www.cancer.gov/clinical_trials/doc.aspx?viewid=320DD013-BA7A-4177-A000-2011089F34A0.

intravenous administration of a new chemotherapy drug being tested in a trial, including any therapy to prevent side effects from the new drug.

- Anything normally covered even if it resulted from your being in the clinical trial. For example, a test or hospitalization resulting from a side effect of the new treatment that Medicare would ordinarily cover.

What Costs Are Not Covered?

- Investigational items or services being tested in a trial. Sponsors of clinical trials often provide the new drug free, but make sure you ask your doctor before you begin.
- Items or services used solely for the data collection needs of the trial.
- Anything being provided free by the sponsor of the trial.

What Kinds of Clinical Trials Are Covered?

NCI's Cancer Information Service has provided a fact sheet for Medicare beneficiaries at the following Web site: http://cis.nci.nih.gov/fact/8_14.htm. In general, cancer treatment and diagnosis trials are covered if:

- They are funded by the National Cancer Institute (NCI), NCI-Designated Cancer Centers, NCI-Sponsored Clinical Trials Cooperative Groups and all other Federal agencies that fund cancer research. Other trials may be eligible for coverage and doctors can ask Medicare to pay the patients' costs. Ask your doctor about this before you begin.
- They are designed to treat or diagnose your cancer.
- The purpose or subject of the trial is within a Medicare benefit category. For example, cancer diagnosis and treatment are Medicare benefits, so these trials are covered. Cancer prevention trials are not currently covered.

Increasing the Likelihood of Insurance Coverage for Trials²³

There are several steps you can follow to deal with coverage issues up front when deciding to enter a clinical trial. Along the way, enlist the help of

²³ This section has been adapted from the NCI:
http://www.cancer.gov/clinical_trials/doc_header.aspx?viewid=1d92be79-8748-4bda-8005-2a56d332463b&docid=0df4397a-eccb-465f-bd33-a89e7a708c46.

family members and your doctor or other health professionals. You may find the following checklist useful:

Understand the Costs Associated with the Trial

Ask your doctor or the trial's contact person about the costs that must be covered by you or your health plan. Are these costs significantly higher than those associated with standard care? Also, inquire about the experience of other patients in the trial. Have their plans paid for their care? Have there been any persistent problems with coverage? How often have the trial's administrators been successful in getting plans to cover patient care costs?

Understand Your Health Plan

Be sure you know what's in your policy; request and carefully review the actual contract language. If there's a specific exclusion for "experimental treatment," look closely at the policy to see how the plan defines such treatment and under what conditions it might be covered. If it is not clearly defined, call the plan's customer service line, consult their Web site, and/or write to them. Ask for specific information about clinical trials coverage.

Work Closely with Your Doctor

Talk with your doctor about the paperwork he or she submits to your health plan. If there have been problems with coverage in the past, you might ask your doctor or the hospital to send an information package to the plan that includes studies supporting the procedure's safety, benefits, and medical appropriateness. This package might include:

- Publications from peer-reviewed literature about the proposed therapy that demonstrate patient benefits;
- A letter that uses the insurance contract's own language to explain why the treatment, screening method, or preventive measure should be covered;
- Letters from researchers that explain the clinical trial;
- Support letters from patient advocacy groups.

Be sure to keep your own copy of any materials that the doctor sends to your health plan for future reference.

Work Closely with Your Company's Benefits Manager

This person may be helpful in enlisting the support of your employer to request coverage by the health plan.

Give Your Health Plan a Deadline

Ask the hospital or cancer center to set a target date for the therapy. This will help to ensure that coverage decisions are made promptly.

Know Your Rights²⁴

A number of state governments are addressing the question of whether insurance companies ought to cover the costs associated with patients' participation in clinical trials. Lack of such coverage is a significant barrier to many patients who might otherwise benefit from enrolling in a trial. Lack of coverage also makes it harder for researchers to successfully conduct trials that could improve prevention and treatment options. Information on State initiatives and legislation concerning cancer-related clinical trials is available at <http://www.cancer.gov/ClinicalTrials/insurancelaws>. By conducting your own research and learning about your rights, you may increase the likelihood that your insurance company will cover the costs of a trial.

If Your Insurance Claim Is Denied after the Trial Has Begun

If a claim is denied, read your policy to find out what steps you can follow to make an appeal. In "What Cancer Survivors Need to Know about Health Insurance", the National Coalition for Cancer Survivorship suggests that you and your doctor demonstrate to the health plan that:

- The therapy is not just a research study, but also a valid procedure that benefits patients;
- Your situation is similar to that of other patients who are participating in clinical trials as part of a covered benefit;
- Possible complications have been anticipated and can be handled effectively.

²⁴ Adapted from Cancer.gov: <http://www.cancer.gov/ClinicalTrials/insurancelaws>.

You also may wish to contact your state insurance counseling hotline or insurance department for more help, or write your state insurance commissioner describing the problem.

Where Else Can I Turn for Assistance?

It's never easy to deal with financial issues when you or a loved one faces cancer. Unfortunately, costs can present a significant barrier to clinical trials participation. The range of insurance issues and health plan contracts makes it impossible to deal with all of them here. You may wish to consult this partial list of publications, organizations, and Web sites for more information:

Publications

What Cancer Survivors Need to Know about Health Insurance

National Coalition of Cancer Survivorship
1010 Wayne Avenue, 5th floor
Silver Spring, MD 20910
(301) 650-8868
<http://www.cansearch.org/>

Cancer Treatments Your Insurance Should Cover

The Association of Community Cancer Centers
11600 Nebel Street, Suite 201
Rockville, MD 20852
(301) 984-9496
<http://www.accc-cancer.org/main2001.shtml>

The Managed Care Answer Guide

Patient Advocate Foundation
739 Thimble Shoals Boulevard, Suite 704
Newport News, VA 23606
(757) 873-6668
E-mail: ndepaf@pinn.net

1998 Guide to Health Insurance for People with Medicare, The Medicare Handbook

Medicare Helpline: 1-800-444-4606

Health Care Financing Administration: <http://www.hcfa.gov/>

New Medicare site: <http://www.medicare.gov/>

Assistance Programs

Candlelighters Childhood Cancer Foundation

Ombudsman Program

910 Woodmont Avenue, #4607

Bethesda, MD 20814

(301) 657-8401; 1-800-366-2223 (toll-free)

E-mail: info@candlelighters.org

<http://www.candlelighters.org>

The Ombudsman Program helps families of children with cancer and survivors of childhood cancer resolve a range of problems, including insurance coverage difficulties. Local groups appoint a Parent Advocate who works with the treatment center on behalf of families.

Medical Care Management Corporation

5272 River Road, Suite 650

Bethesda, MD 20816-1405

(301) 652-1818

email: mcman@mcman.com

<http://www.mcman.com/>

Working for a range of clients, including health plans, employers, and patients, MCMC conducts independent, objective reviews of high-technology medical care cases to assist in decision-making. While it does charge for its services, MCMC also offers a volunteer program for those who cannot afford to pay.

More Information Resources

OncoLink

A service of the University of Pennsylvania Cancer Center.

<http://www.oncolink.com/>

In addition to general cancer information, this web site features a section on financial information for patients. Among the topics: viatical settlements, life insurance, a glossary of financial and medical terms, and news about billing and insurance.

American Association of Health Plans

1129 20th Street, NW, Suite 600

Washington, DC 20036-3421

(202) 778-3200

<http://www.aahp.org/>

The Web site section “For Consumers” includes a fact sheet on clinical research that describes various health plans’ efforts to support research initiatives and collaborate with academic health centers and universities.

Health Insurance Association of America

555 13th Street, NW

Washington, DC 20004

(202) 824-1600

- Home page: **<http://www.hiaa.org/>**
- Consumer Information: **<http://www.hiaa.org/consumer/>**
- Insurance Counseling Hotlines by State:
http://www.hiaa.org/consumer/insurance_counsel.cfm
- State Insurance Departments:
http://www.hiaa.org/consumer/state_insurance.cfm

Government Initiatives to Expand Insurance Coverage for Trials²⁵

The good news is that there has been a recent effort in the U.S. to assure clinical trials coverage, with NCI involved in several new initiatives as described below:

NCI-Department of Defense Agreement

An innovative 1996 agreement between NCI and the Department of Defense (DoD) has given thousands of DoD cancer patients more options for care and greater access to state-of-the-art treatments. Patients who are beneficiaries of TRICARE/CHAMPUS, the DoD’s health program, are covered for NCI-sponsored Phase II and Phase III clinical treatment trials. NCI and DoD are refining a system that allows physicians and patients to determine quickly what current trials meet their needs and where they are taking place.

²⁵ Adapted from the NCI:

http://www.cancer.gov/clinical_trials/doc_header.aspx?viewid=1d92be79-8748-4bda-8005-2a56d332463b&docid=d8092601-daf9-4794-8536-3be2712eb6b9.

NCI-Department of Veterans Affairs Agreement

A 1997 agreement with the Department of Veterans Affairs provides coverage for eligible veterans of the armed services to participate in NCI-sponsored prevention, diagnosis, and treatment studies nationwide. For additional information, see the VA/DoD Beneficiaries Digest Page at <http://www.va.gov/cancer.htm>.

Midwest Health Plans Agreement

Some NCI Cooperative Groups have reached agreements with several insurers in Wisconsin and Minnesota to provide more than 200,000 people with coverage. This coverage is allocated for patient care costs if they participate in a cooperative group-sponsored trial.

Pediatric Cancer Care Network

This network, a cooperative agreement among the Children's Cancer Group, the Pediatric Oncology Group, and the Blue Cross Blue Shield System Association (BCBS) nationwide, will ensure that children of BCBS subscribers receive care at designated centers of cancer care excellence and may promote the enrollment of children in Cooperative Group clinical trials.

Keeping Current on Clinical Trials

Various government agencies maintain databases on trials. The U.S. National Institutes of Health, through the National Library of Medicine, has developed ClinicalTrials.gov to provide patients, family members, and physicians with current information about clinical research across the broadest number of diseases and conditions.

The site was launched in February 2000 and currently contains approximately 5,700 clinical studies in over 59,000 locations worldwide, with most studies being conducted in the United States. ClinicalTrials.gov receives about 2 million hits per month and hosts approximately 5,400 visitors daily. To access this database, simply go to their Web site (www.clinicaltrials.gov) and search by "adult primary liver cancer" (or synonyms).

While ClinicalTrials.gov is the most comprehensive listing of NIH-supported clinical trials available, not all trials are in the database. The database is updated regularly, so clinical trials are continually being added. The following is a list of specialty databases affiliated with the National Institutes of Health that offer additional information on trials:

- For clinical studies at the Warren Grant Magnuson Clinical Center located in Bethesda, Maryland, visit their Web site:
<http://clinicalstudies.info.nih.gov/>
- For clinical studies conducted at the Bayview Campus in Baltimore, Maryland, visit their Web site:
<http://www.jhbmc.jhu.edu/studies/index.html>
- For cancer trials, visit the National Cancer Institute:
<http://cancertrials.nci.nih.gov/>

General References

The following references describe clinical trials and experimental medical research. They have been selected to ensure that they are likely to be available from your local or online bookseller or university medical library. These references are usually written for healthcare professionals, so you may consider consulting with a librarian or bookseller who might recommend a particular reference. The following includes some of the most readily available references (sorted alphabetically by title; hyperlinks provide rankings, information and reviews at Amazon.com):

- **A Guide to Patient Recruitment : Today's Best Practices & Proven Strategies** by Diana L. Anderson; Paperback - 350 pages (2001), CenterWatch, Inc.; ISBN: 1930624115;
<http://www.amazon.com/exec/obidos/ASIN/1930624115/icongroupinterna>
- **A Step-By-Step Guide to Clinical Trials** by Marilyn Mulay, R.N., M.S., OCN; Spiral-bound - 143 pages Spiral edition (2001), Jones & Bartlett Pub; ISBN: 0763715697;
<http://www.amazon.com/exec/obidos/ASIN/0763715697/icongroupinterna>
- **The CenterWatch Directory of Drugs in Clinical Trials** by CenterWatch; Paperback - 656 pages (2000), CenterWatch, Inc.; ISBN: 0967302935;
<http://www.amazon.com/exec/obidos/ASIN/0967302935/icongroupinterna>
- **The Complete Guide to Informed Consent in Clinical Trials** by Terry Hartnett (Editor); Paperback - 164 pages (2000), PharmSource Information Services, Inc.; ISBN: 0970153309;
<http://www.amazon.com/exec/obidos/ASIN/0970153309/icongroupinterna>

- **Dictionary for Clinical Trials** by Simon Day; Paperback - 228 pages (1999), John Wiley & Sons; ISBN: 0471985961;
<http://www.amazon.com/exec/obidos/ASIN/0471985961/icongroupinterna>
- **Extending Medicare Reimbursement in Clinical Trials** by Institute of Medicine Staff (Editor), et al; Paperback 1st edition (2000), National Academy Press; ISBN: 0309068886;
<http://www.amazon.com/exec/obidos/ASIN/0309068886/icongroupinterna>
- **Handbook of Clinical Trials** by Marcus Flather (Editor); Paperback (2001), Remedica Pub Ltd; ISBN: 1901346293;
<http://www.amazon.com/exec/obidos/ASIN/1901346293/icongroupinterna>

Vocabulary Builder

The following vocabulary builder gives definitions of words used in this chapter that have not been defined in previous chapters:

Adenocarcinoma: Cancer that begins in cells that line certain internal organs and that have glandular (secretory) properties. [NIH]

Anesthesia: Loss of feeling or awareness. Local anesthetics cause loss of feeling in a part of the body. General anesthetics put the person to sleep. [NIH]

Angiogenesis: Blood vessel formation. Tumor angiogenesis is the growth of blood vessels from surrounding tissue to a solid tumor. This is caused by the release of chemicals by the tumor. [NIH]

Antiangiogenesis: Prevention of the growth of new blood vessels. [NIH]

Antiemetic: An agent that prevents or alleviates nausea and vomiting. Also antinauseant. [EU]

Antineoplastons: Substances isolated from normal human blood and urine being tested as a type of treatment for some tumors and AIDS. [NIH]

Arterial: Pertaining to an artery or to the arteries. [EU]

Bile: A fluid made by the liver and stored in the gallbladder. Bile is excreted into the small intestine where it helps digest fat. [NIH]

Bypass: A surgical procedure in which the doctor creates a new pathway for the flow of body fluids. [NIH]

Capecitabine: An anticancer drug that belongs to the family of drugs called antimetabolites. [NIH]

Carboplatin: An anticancer drug that belongs to the family of drugs called platinum compounds. [NIH]

Chronic: A disease or condition that persists or progresses over a long

period of time. [NIH]

Chronotherapy: The adaptation of the administration of drugs to circadian rhythms. The concept is based on the response of biological functions to time-related events, such as the low point in epinephrine levels between 10 p.m. and 4 a.m. or the elevated histamine levels between midnight and 4 a.m. The treatment is aimed at supporting normal rhythms or modifying therapy based on known variations in body rhythms. While chronotherapy is commonly used in cancer chemotherapy, it is not restricted to cancer therapy or to chemotherapy. [NIH]

Cisplatin: An anticancer drug that belongs to the family of drugs called platinum compounds. [NIH]

Collagen: A fibrous protein found in cartilage and other connective tissue. [NIH]

Colon: The long, coiled, tubelike organ that removes water from digested food. The remaining material, solid waste called stool, moves through the colon to the rectum and leaves the body through the anus. [NIH]

Cytokines: A class of substances that are produced by cells of the immune system and can affect the immune response. Cytokines can also be produced in the laboratory by recombinant DNA technology and given to people to affect immune responses. [NIH]

Dexamethasone: A synthetic steroid (similar to steroid hormones produced naturally in the adrenal gland). Dexamethasone is used to treat leukemia and lymphoma and may be used to treat some of the problems caused by other cancers and their treatment. [NIH]

Doxorubicin: An anticancer drug that belongs to the family of drugs called antitumor antibiotics. It is an anthracycline. [NIH]

Endometrial: Having to do with the endometrium (the layer of tissue that lines the uterus). [NIH]

Enzyme: A protein that speeds up chemical reactions in the body. [NIH]

Epithelial: Refers to the cells that line the internal and external surfaces of the body. [NIH]

Esophageal: Having to do with the esophagus, the muscular tube through which food passes from the throat to the stomach. [NIH]

Extremity: A limb; an arm or leg (membrum); sometimes applied specifically to a hand or foot. [EU]

Fibrosis: The growth of fibrous tissue. [NIH]

Fluorouracil: An anticancer drug that belongs to the family of drugs called antimetabolites. [NIH]

Gallbladder: The pear-shaped organ that sits below the liver. Bile is

concentrated and stored in the gallbladder. [NIH]

Gastric: Having to do with the stomach. [NIH]

Gastrointestinal: Refers to the stomach and intestines. [NIH]

Gemcitabine: An anticancer drug that belongs to the family of drugs called antimetabolites. [NIH]

Groin: The area where the thigh meets the abdomen. [NIH]

Hepatoblastoma: A type of liver tumor that occurs in infants and children. [NIH]

Hepatoma: A liver tumor. [NIH]

Hypervascular: Having a large number of blood vessels. [NIH]

Infusion: A method of putting fluids, including drugs, into the bloodstream. Also called intravenous infusion. [NIH]

Interferon: A biological response modifier (a substance that can improve the body's natural response to disease). Interferons interfere with the division of cancer cells and can slow tumor growth. There are several types of interferons, including interferon-alpha, -beta, and -gamma. These substances are normally produced by the body. They are also made in the laboratory for use in treating cancer and other diseases. [NIH]

Intravenous: IV. Into a vein. [NIH]

Iodine: A nonmetallic element of the halogen group that is represented by the atomic symbol I, atomic number 53, and atomic weight of 126.90. It is a nutritionally essential element, especially important in thyroid hormone synthesis. In solution, it has anti-infective properties and is used topically. [NIH]

Irinotecan: An anticancer drug that belongs to a family of anticancer drugs called topoisomerase inhibitors. It is a camptothecin analogue. Also called CPT 11. [NIH]

Lesion: An area of abnormal tissue change. [NIH]

Megestrol: A drug that belongs to the group of hormones called progestins, used as hormone therapy to block estrogen and to suppress the effects of estrogen and androgens. It is also used to stimulate the appetite in people with cancer. [NIH]

Melanoma: A form of skin cancer that arises in melanocytes, the cells that produce pigment. Melanoma usually begins in a mole. [NIH]

Melphalan: An anticancer drug that belongs to the family of drugs called alkylating agents. [NIH]

Metaplasia: A change of cells to a form that does not normally occur in the tissue in which it is found. [NIH]

Metastasis: The spread of cancer from one part of the body to another.

Tumors formed from cells that have spread are called "secondary tumors" and contain cells that are like those in the original (primary) tumor. The plural is metastases. [NIH]

Metastatic: Having to do with metastasis, which is the spread of cancer from one part of the body to another. [NIH]

MRI: Magnetic resonance imaging (mag-NET-ik REZ-o- nans IM-a-jing). A procedure in which a magnet linked to a computer is used to create detailed pictures of areas inside the body. [NIH]

Neoplasm: A new growth of benign or malignant tissue. [NIH]

Ondansetron: A drug that prevents or reduces nausea and vomiting. [NIH]

Pancreas: A glandular organ located in the abdomen. It makes pancreatic juices, which contain enzymes that aid in digestion, and it produces several hormones, including insulin. The pancreas is surrounded by the stomach, intestines, and other organs. [NIH]

Pancreatic: Having to do with the pancreas. [NIH]

Particle: A tiny mass of material. [EU]

Pelvis: The lower part of the abdomen, located between the hip bones. [NIH]

Percutaneous: Performed through the skin, as injection of radiopaque material in radiological examination, or the removal of tissue for biopsy accomplished by a needle. [EU]

Perfusion: Bathing an organ or tissue with a fluid. In regional perfusion, a specific area of the body (usually an arm or a leg) receives high doses of anticancer drugs through a blood vessel. Such a procedure is performed to treat cancer that has not spread. [NIH]

Prevalence: The total number of cases of a given disease in a specified population at a designated time. It is differentiated from incidence, which refers to the number of new cases in the population at a given time. [NIH]

Progression: Increase in the size of a tumor or spread of cancer in the body. [NIH]

Progressive: Advancing; going forward; going from bad to worse; increasing in scope or severity. [EU]

Proteins: Polymers of amino acids linked by peptide bonds. The specific sequence of amino acids determines the shape and function of the protein. [NIH]

Randomized: Describes an experiment or clinical trial in which animal or human subjects are assigned by chance to separate groups that compare different treatments. [NIH]

Rebexamycin: An anticancer drug that belongs to the family of drugs called antineoplastic antibiotics. [NIH]

Recombinant: 1. a cell or an individual with a new combination of genes not found together in either parent; usually applied to linked genes. [EU]

Recurrence: The return of cancer, at the same site as the original (primary) tumor or in another location, after the tumor had disappeared. [NIH]

Regimen: A treatment plan that specifies the dosage, the schedule, and the duration of treatment. [NIH]

Remission: A decrease in or disappearance of signs and symptoms of cancer. In partial remission, some, but not all, signs and symptoms of cancer have disappeared. In complete remission, all signs and symptoms of cancer have disappeared, although there still may be cancer in the body. [NIH]

Ribavirin: A drug used to treat respiratory syncytial virus (RSV) infection in the lungs. [NIH]

Sarcoma: A cancer of the bone, cartilage, fat, muscle, blood vessels or other connective or supportive tissue. [NIH]

Sargramostim: A colony-stimulating factor that stimulates the production of blood cells, especially platelets, during chemotherapy. It is a cytokine that belongs to the family of drugs called hematopoietic (blood-forming) agents. Also called GM-CSF. [NIH]

Serum: The clear liquid part of the blood that remains after blood cells and clotting proteins have been removed. [NIH]

Squamous: Scaly, or platelike. [EU]

Stomach: An organ that is part of the digestive system. It helps in the digestion of food by mixing it with digestive juices and churning it into a thin liquid. [NIH]

Symptomatic: Having to do with symptoms, which are signs of a condition or disease. [NIH]

Tamoxifen: An anticancer drug that belongs to the family of drugs called antiestrogens. Tamoxifen blocks the effects of the hormone estrogen in the body. It is used to prevent or delay the return of breast cancer or to control its spread. [NIH]

Testicular: Pertaining to a testis. [EU]

Thalidomide: A drug that belongs to the family of drugs called angiogenesis inhibitors. It prevents the growth of new blood vessels into a solid tumor. [NIH]

Tolerance: 1. the ability to endure unusually large doses of a drug or toxin. 2. acquired drug tolerance; a decreasing response to repeated constant doses of a drug or the need for increasing doses to maintain a constant response. [EU]

Toxicity: The quality of being poisonous, especially the degree of virulence

of a toxic microbe or of a poison. [EU]

Urine: Fluid containing water and waste products. Urine is made by the kidneys, stored in the bladder, and leaves the body through the urethra. [NIH]

Vaccine: A substance or group of substances meant to cause the immune system to respond to a tumor or to microorganisms, such as bacteria or viruses. [NIH]

PART II: ADDITIONAL RESOURCES AND ADVANCED MATERIAL

ABOUT PART II

In Part II, we introduce you to additional resources and advanced research on adult primary liver cancer. All too often, patients who conduct their own research are overwhelmed by the difficulty in finding and organizing information. The purpose of the following chapters is to provide you an organized and structured format to help you find additional information resources on adult primary liver cancer. In Part II, as in Part I, our objective is not to interpret the latest advances on adult primary liver cancer or render an opinion. Rather, our goal is to give you access to original research and to increase your awareness of sources you may not have already considered. In this way, you will come across the advanced materials often referred to in pamphlets, books, or other general works. Once again, some of this material is technical in nature, so consultation with a professional familiar with adult primary liver cancer is suggested.

CHAPTER 4. STUDIES ON ADULT PRIMARY LIVER CANCER

Overview

Every year, academic studies are published on adult primary liver cancer or related conditions. Broadly speaking, there are two types of studies. The first are peer reviewed. Generally, the content of these studies has been reviewed by scientists or physicians. Peer-reviewed studies are typically published in scientific journals and are usually available at medical libraries. The second type of studies is non-peer reviewed. These works include summary articles that do not use or report scientific results. These often appear in the popular press, newsletters, or similar periodicals.

In this chapter, we will show you how to locate peer-reviewed references and studies on adult primary liver cancer. We will begin by discussing research that has been summarized and is free to view by the public via the Internet. We then show you how to generate a bibliography on adult primary liver cancer and teach you how to keep current on new studies as they are published or undertaken by the scientific community.

The Combined Health Information Database

The Combined Health Information Database summarizes studies across numerous federal agencies. To limit your investigation to research studies and adult primary liver cancer, you will need to use the advanced search options. First, go to <http://chid.nih.gov/index.html>. From there, select the "Detailed Search" option (or go directly to that page with the following hyperlink: <http://chid.nih.gov/detail/detail.html>). The trick in extracting studies is found in the drop boxes at the bottom of the search page where "You may refine your search by." Select the dates and language you prefer,

and the format option “Journal Article.” At the top of the search form, select the number of records you would like to see (we recommend 100) and check the box to display “whole records.” We recommend that you type in “adult primary liver cancer” (or synonyms) into the “For these words:” box. Consider using the option “anywhere in record” to make your search as broad as possible. If you want to limit the search to only a particular field, such as the title of the journal, then select this option in the “Search in these fields” drop box. The following is a sample of what you can expect from this type of search:

- **Impact Factors on Development of Cirrhosis and Subsequent Hepatocellular Carcinoma**

Source: Compendium of Continuing Education in Dentistry. 22(3): 19-33. July 2001.

Contact: Available from Dental Learning Systems. 241 Forsgate Drive, Jamesburg, NJ 08831. (800) 926-7636.

Summary: Hepatocellular carcinoma (HCC, liver cancer) on the rise in many countries, is of multifactorial etiology (caused by many factors). This article reviews the development of cirrhosis and subsequent HCC. The causes of liver cancer differ between populations at high and low risk. Africans and Chinese have the highest incidence of HCC, but other affected groups include African Americans, Japanese, and Native Americans. Chronic infection by hepatitis B and hepatitis C viruses are major risk factors worldwide, although mechanisms through which the infections cause liver cancer are yet to be explained. Other documented risk factors have been proposed and include dietary exposure, cigarette smoking, alcohol consumption, diabetes, oral infection, and oral contraceptive use. In addition, many naturally occurring and synthetic chemicals to which humans are exposed via accidental contamination of food or water have been shown to induce liver cancer in experimental animals. Consequently, assessment of possible human liver cancer risk associated with such exposures is complex. Early diagnosis and transplantation are the best treatments presently, although transplantation is not widely available due to donor shortage. The author stresses that every effort should be directed toward the prevention of HCC, through the treatment and prevention of hepatitis and oral infections, prevention of chronic hepatitis progressing to cirrhosis (liver scarring), and prevention of the cirrhotic liver from developing HCC through chemopreventive modalities. The article is designed as a continuing education curriculum for dentists and other oral health care professionals. 3 figures. 4 tables. 85 references.

Federally-Funded Research on Adult Primary Liver Cancer

The U.S. Government supports a variety of research studies relating to adult primary liver cancer and associated conditions. These studies are tracked by the Office of Extramural Research at the National Institutes of Health.²⁶ CRISP (Computerized Retrieval of Information on Scientific Projects) is a searchable database of federally-funded biomedical research projects conducted at universities, hospitals, and other institutions. Visit the CRISP Web site at http://commons.cit.nih.gov/crisp3/CRISP.Generate_Ticket. You can perform targeted searches by various criteria including geography, date, as well as topics related to adult primary liver cancer and related conditions.

For most of the studies, the agencies reporting into CRISP provide summaries or abstracts. As opposed to clinical trial research using patients, many federally-funded studies use animals or simulated models to explore adult primary liver cancer and related conditions. In some cases, therefore, it may be difficult to understand how some basic or fundamental research could eventually translate into medical practice. The following sample is typical of the type of information found when searching the CRISP database for adult primary liver cancer:

- **Project Title: Clinical Applications of Modulation of DNA Repair Pathways**

Principal Investigator & Institution: Cornetta, Kenneth G.; Professor; Indiana Univ-Purdue Univ at Indianapolis 355 N Lansing Indianapolis, in 46202

Timing: Fiscal Year 2000

Summary: The clinical studies outlined in this specific aim target the manipulation of 06-methylguanine DNA methyltransferase (MGMT) in humans and are the result of extensive pre-clinical work demonstrating the efficacy of the approaches in animal models. One project proposes to increase the expression of MGMT in hematopoietic cells in an effort to diminish the cumulative myelosuppression commonly encountered with chloroethylnitrosoureas (CENUs). This project utilizes a recombinant retroviral vector extensively tested in murine studies and produced by the National Gene Vector Laboratory at Indiana University for human clinical trials. The clinical study builds on a current pilot study at Indiana

²⁶ Healthcare projects are funded by the National Institutes of Health (NIH), Substance Abuse and Mental Health Services (SAMHSA), Health Resources and Services Administration (HRSA), Food and Drug Administration (FDA), Centers for Disease Control and Prevention (CDCP), Agency for Healthcare Research and Quality (AHRQ), and Office of Assistant Secretary of Health (OASH).

University, designed by Dr. Regina Jakacki and supported in part by NCI funding, which utilizes peripheral blood stem-progenitor cell infusions to decrease hematopoietic toxicities and allow schedule compression of an extensively used brain treatment protocol called "PCV" (procarbazine, CCNU, vincristine). The second project is designed to diminish expression of MGMT in tumor cells. Based on pre-clinical work by Dr. Leonard Erickson, MGMT can be effectively depleted from tumor cell lines by the sequential treatment with agents that produce the natural substrate for MGMT, 06-methylguanine, or act as a substrate for MGMT directly. One such agent, 06-benzylguanine (6-BG), is currently in phase I trials at other institutions. The specific aims of the study are: 1) To conduct a pilot study of dose-intensified procarbazine, CCNU, vincristine (PCV) for poor prognosis pediatric and adult brain tumor utilizing fibronectin-assisted, retroviral-mediated modification of CD34+ peripheral blood cells with 06-methylguanine DNA methyltransferase (MGMT). 2) To conduct a phase I trial to determine the toxicity of the combination of 06-benzylguanine and BCNU, and to examine the inhibition of MGMT activity in tumor biopsies in patients treated with this combination chemotherapy for relapsed B cell malignancies.

Website: http://commons.cit.nih.gov/crisp3/CRISP.Generate_Ticket

- **Project Title: Inhibition of NA-H Exchanger Selectively Kills Gliomas**

Principal Investigator & Institution: Gorin, Fredric A.; Neurology; University of California Davis 1 Shields Ave Davis, Ca 95616

Timing: Fiscal Year 2001; Project Start 1-JUN-2001; Project End 0-APR-2004

Summary: High grade astrocytomas (, malignant gliomas) are the most commonly occurring type of lethal adult brain-tumor with an individual's average life expectancy being less than 2 years from the time of diagnosis. Neither radiation therapy nor chemotherapy has significantly improved quality or length of survival. Since Warburg's initial observation, it has been recognized that most transformed tumor cells have high rates of glycolytic metabolism and consequent H⁺ production. Given the optimal alkaline pH dependence of key glycolytic enzymes, such as phosphofructokinase and hexokinase, it is essential that tumor cells employ an effective means of removing free cytosolic H⁺ to maintain metabolism. We have determined that intracellular pH in rat and human gliomas are significantly above that of normal astrocytes (0.2-0.6 pH units) despite the tumor's high rates of metabolic H⁺ production. This intracellular alkalosis appears to result from persistent activation of the type 1 isoform of the Na⁺-H⁺ exchanger (NHE 1). Our preliminary investigations have determined that this altered regulation of NHE 1 is

most probably posttranslational and does not result from alterations of the NHE1 gene or proteins expressed in these highly malignant astrocytomas. Unexpectedly, we found that inhibition of NHE I in rat and human glioma cell lines with the diuretic drug, amiloride, or with its derivatives, HOE 694 and EIPA, cause a 70-100 percent cell death within 48-72 hours. By contrast, primary astrocyte cultures were unaffected by NHE 1 inhibition. Cell culture and in vivo analyses indicate that glioma death following NHE 1 inhibition appears to be predominantly non-apoptotic and independent of preceding caspase activation. Rat C6 gliomas were implanted into rat brains and allowed to establish for 4 days. Amiloride infusion into the cerebrospinal fluid for 8 days produced a 73 percent reduction in tumor volume. Amiloride is an oral diuretic that is approved for human use. Preliminarily, this novel intrathecal administration of amiloride in rats does not appear to cause behavioral or neuropathological alterations. Amiloride produced a dose-dependent decrease in intracellular pH in malignant gliomas, but not astrocytes. We have pilot data indicating that this ApHi initiates the selective tumor death. We propose to (1) identify the intracellular mechanisms mediating glioma death; (2) use brain implanted tumor models to more thoroughly delineate selective glioma death by NHE I inhibitors; and (3) study the pharmacological and H⁺ regulatory properties of surviving gliomas. NHE I inhibitors may represent a new class of pharmacological agents that are useful for treatment of these highly aggressive and lethal brain tumors.

Website: http://commons.cit.nih.gov/crisp3/CRISP.Generate_Ticket

E-Journals: PubMed Central²⁷

PubMed Central (PMC) is a digital archive of life sciences journal literature developed and managed by the National Center for Biotechnology Information (NCBI) at the U.S. National Library of Medicine (NLM).²⁸ Access to this growing archive of e-journals is free and unrestricted.²⁹ To search, go to <http://www.pubmedcentral.nih.gov/index.html#search>, and type "adult

²⁷ Adapted from the National Library of Medicine:

<http://www.pubmedcentral.nih.gov/about/intro.html>.

²⁸ With PubMed Central, NCBI is taking the lead in preservation and maintenance of open access to electronic literature, just as NLM has done for decades with printed biomedical literature. PubMed Central aims to become a world-class library of the digital age.

²⁹ The value of PubMed Central, in addition to its role as an archive, lies the availability of data from diverse sources stored in a common format in a single repository. Many journals already have online publishing operations, and there is a growing tendency to publish material online only, to the exclusion of print.

primary liver cancer” (or synonyms) into the search box. This search gives you access to full-text articles. The following is a sample of items found for adult primary liver cancer in the PubMed Central database:

- **Chlorophyllin intervention reduces aflatoxin --DNA adducts in individuals at high risk for liver cancer** by Patricia A. Egner, Jin-Bing Wang, Yuan-Rong Zhu, Bao-Chu Zhang, Yan Wu, Qi-Nan Zhang, Geng-Sun Qian, Shuang-Yuan Kuang, Stephen J. Gange, Lisa P. Jacobson, Kathy J. Helzlsouer, George S. Bailey, John D. Groopman, and Thomas W. Kensler; 2001 December 4
<http://www.pubmedcentral.nih.gov/articlerender.fcgi?artid=64728>

The National Library of Medicine: PubMed

One of the quickest and most comprehensive ways to find academic studies in both English and other languages is to use PubMed, maintained by the National Library of Medicine. The advantage of PubMed over previously mentioned sources is that it covers a greater number of domestic and foreign references. It is also free to the public.³⁰ If the publisher has a Web site that offers full text of its journals, PubMed will provide links to that site, as well as to sites offering other related data. User registration, a subscription fee, or some other type of fee may be required to access the full text of articles in some journals.

To generate your own bibliography of studies dealing with adult primary liver cancer, simply go to the PubMed Web site at www.ncbi.nlm.nih.gov/pubmed. Type “adult primary liver cancer” (or synonyms) into the search box, and click “Go.” The following is the type of output you can expect from PubMed for “adult primary liver cancer” (hyperlinks lead to article summaries):

- **188rhenium-TDD-lipiodol in treatment of inoperable primary hepatocellular carcinoma—a case report.**
 Author(s): Sundram FX, Yu SW, Jeong JM, Somanesan S, Premaraj J, Saw MM, Tan BS.

³⁰ PubMed was developed by the National Center for Biotechnology Information (NCBI) at the National Library of Medicine (NLM) at the National Institutes of Health (NIH). The PubMed database was developed in conjunction with publishers of biomedical literature as a search tool for accessing literature citations and linking to full-text journal articles at Web sites of participating publishers. Publishers that participate in PubMed supply NLM with their citations electronically prior to or at the time of publication.

Source: Ann Acad Med Singapore. 2001 September; 30(5): 542-5.
http://www.ncbi.nlm.nih.gov/80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11603144&dopt=Abstract

- **A case of hepatocellular carcinoma rupturing after angiography.**
 Author(s): Kinoshita H, Sato S, Hashimoto M, Hashino K, Kawabata M, Furukawa S, Nishimura K, Kodama T, Nagashima J, Okuda K, Imayama H, Aoyagi S.
 Source: Kurume Med J. 2001; 48(3): 241-5.
http://www.ncbi.nlm.nih.gov/80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11680942&dopt=Abstract
- **A case-control study of hepatocellular carcinoma in Hyogo Prefecture.**
 Author(s): Inoue Y, Shiraki T, Doi H, Yamanaka N, Sato S.
 Source: The Kobe Journal of Medical Sciences. 2000 August; 46(4): 181-8.
http://www.ncbi.nlm.nih.gov/80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11354929&dopt=Abstract
- **A new staging system for mass-forming intrahepatic cholangiocarcinoma: analysis of preoperative and postoperative variables.**
 Author(s): Okabayashi T, Yamamoto J, Kosuge T, Shimada K, Yamasaki S, Takayama T, Makuuchi M.
 Source: Cancer. 2001 November 1; 92(9): 2374-83.
http://www.ncbi.nlm.nih.gov/80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11745293&dopt=Abstract
- **A successful resection and long-term survival of a patient with intrahepatic recurrences of combined hepatocellular-cholangiocarcinoma: report of a case.**
 Author(s): Eguchi H, Nagano H, Sakon M, Miyamoto A, Kondo M, Arai I, Morimoto O, Dono K, Umeshita K, Nakamori S, Wakasa K, Monden M.
 Source: Surgery Today. 2002; 32(8): 742-6.
http://www.ncbi.nlm.nih.gov/80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=12181730&dopt=Abstract
- **Adult-type hepatocellular carcinoma in the center of a fibrolamellar hepatocellular carcinoma.**
 Author(s): Seitz G, Zimmermann A, Friess H, Buchler MW.

Source: Hum Pathol. 2002 July; 33(7): 765-9.
http://www.ncbi.nlm.nih.gov/80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=12196930&dopt=Abstract

- **alpha-Fetoprotein mRNA in the circulation as a predictor of postsurgical recurrence of hepatocellular carcinoma: a prospective study.**

Author(s): Ijichi M, Takayama T, Matsumura M, Shiratori Y, Omata M, Makuuchi M.

Source: Hepatology. 2002 April; 35(4): 853-60.

http://www.ncbi.nlm.nih.gov/80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11915031&dopt=Abstract

- **Analysis of tumor characteristics and survival in liver transplant recipients with incidentally diagnosed hepatocellular carcinoma.**

Author(s): Cho CS, Knechtle SJ, Heisey DM, Hermina M, Armbrust M, D'Alessandro AM, Musat AI, Kalayoglu M.

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- **Androgen receptor in primary hepatocellular carcinoma and its clinical significance.**

Author(s): Zhang X, He L, Lu Y, Liu M, Huang X.

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- **Arterial embolisation or chemoembolisation versus symptomatic treatment in patients with unresectable hepatocellular carcinoma: a randomised controlled trial.**

Author(s): Llovet JM, Real MI, Montana X, Planas R, Coll S, Aponte J, Ayuso C, Sala M, Muchart J, Sola R, Rodes J, Bruix J.

Source: Lancet. 2002 May 18; 359(9319): 1734-9.

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- **Arterial embolization of unresectable hepatocellular carcinoma with use of cyanoacrylate and lipiodol.**

Author(s): Loewe C, Cejna M, Schoder M, Thurnher MM, Lammer J, Thurnher SA.

Source: Journal of Vascular and Interventional Radiology : Jvir. 2002 January; 13(1): 61-9.

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- **Biallelic inactivation of the APC gene is associated with hepatocellular carcinoma in familial adenomatous polyposis coli.**

Author(s): Su LK, Abdalla EK, Law CH, Kohlmann W, Rashid A, Vauthey JN.

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- **Bilateral adrenal metastases from hepatocellular carcinoma after liver transplantation.**

Author(s): Castroagudin JF, Gonzalez-Quintela A, Martinez J, Tome S, Forteza J, Varo E.

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- **Biliary dysplasia as a marker of cholangiocarcinoma in primary sclerosing cholangitis.**

Author(s): Fleming KA, Boberg KM, Glaumann H, Bergquist A, Smith D, Clausen OP.

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- **Budd-Chiari syndrome secondary to intracardiac extension of hepatocellular carcinoma. Two cases treated by radical resection.**

Author(s): Saisse J, Hardwigsen J, Castellani P, Caus T, Le Treut YP.

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- **Calciphylaxis associated with cholangiocarcinoma treated with low-molecular-weight heparin and vitamin K.**

Author(s): Riegert-Johnson DL, Kaur JS, Pfeifer EA.

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Author(s): Jarnagin WR, Weber S, Tickoo SK, Koea JB, Obiekwe S, Fong Y, DeMatteo RP, Blumgart LH, Klimstra D.

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 Author(s): Pang E, Wong N, Lai PB, To KF, Lau WY, Johnson PJ.
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Author(s): Huang GW, Yang LY.
Source: World J Gastroenterol. 2002 August; 8(4): 650-3.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=12174372&dopt=Abstract

- **Metastatic hepatocellular carcinoma presenting as epidural hematoma: case report.**
Author(s): McIver JL, Scheithauer BW, Rydberg CH, Atkinson JL.
Source: Neurosurgery. 2001 August; 49(2): 447-9.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11504122&dopt=Abstract
- **Mitochondrial D-loop mutations as clonal markers in multicentric hepatocellular carcinoma and plasma.**
Author(s): Nomoto S, Yamashita K, Koshikawa K, Nakao A, Sidransky D.
Source: Clinical Cancer Research : an Official Journal of the American Association for Cancer Research. 2002 February; 8(2): 481-7.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11839667&dopt=Abstract
- **Molecular cloning and characterization of a novel gene which is highly expressed in hepatocellular carcinoma.**
Author(s): Zeng JZ, Wang HY, Chen ZJ, Ullrich A, Wu MC.
Source: Oncogene. 2002 July 25; 21(32): 4932-43.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=12118372&dopt=Abstract
- **Multi locular presentation of hepatocellular carcinoma.**
Author(s): Khokhar N.
Source: J Pak Med Assoc. 2001 November; 51(11): 407-8. No Abstract Available.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11840609&dopt=Abstract
- **Multimodality treatment resulting in long-term survival in hepatocellular carcinoma.**
Author(s): Gasztonyi B, Par A, Battyany I, Hegedus G, Molnar TF, Horvath L, Mozsik G.
Source: Journal of Physiology, Paris. 2001 January-December; 95(1-6): 413-6.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11595468&dopt=Abstract
- **Multiple genetic alterations involved in the tumorigenesis of human cholangiocarcinoma: a molecular genetic and clinicopathological study.**
Author(s): Cong WM, Bakker A, Swalsky PA, Raja S, Woods J, Thomas S, Demetris AJ, Finkelstein SD.

Source: Journal of Cancer Research and Clinical Oncology. 2001; 127(3): 187-92.

http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11260864&dopt=Abstract

- **Pathology and pathogenesis of hepatocellular carcinoma.**
Author(s): Rocken C, Carl-McGrath S.
Source: Digestive Diseases (Basel, Switzerland). 2001; 19(4): 269-78. Review.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11935086&dopt=Abstract
- **Percutaneous microwave coagulation therapy for primary or recurrent hepatocellular carcinoma: long-term results.**
Author(s): Itamoto T, Katayama K, Fukuda S, Fukuda T, Yano M, Nakahara H, Okamoto Y, Sugino K, Marubayashi S, Asahara T.
Source: Hepatogastroenterology. 2001 September-October; 48(41): 1401-5.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11677974&dopt=Abstract
- **Percutaneous US-guided radiofrequency ablation of hepatocellular carcinomas: results in 15 patients.**
Author(s): Poggi G, Gatti C, Cupella F, Fiori M, Avanza F, Baldi M.
Source: Anticancer Res. 2001 January-February; 21(1B): 739-42.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11299836&dopt=Abstract
- **Phase II study of 4-ipomeanol, a naturally occurring alkylating furan, in patients with advanced hepatocellular carcinoma.**
Author(s): Lakhanpal S, Donehower RC, Rowinsky EK.
Source: Investigational New Drugs. 2001; 19(1): 69-76.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11291834&dopt=Abstract
- **Primary paraganglioma strictly confined to the liver and mimicking hepatocellular carcinoma: an immunohistochemical and in situ hybridization study.**
Author(s): Corti B, D'Errico A, Pierangeli F, Fiorentino M, Altimari A, Grigioni WF.

Source: The American Journal of Surgical Pathology. 2002 July; 26(7): 945-9.

http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=12131164&dopt=Abstract

- **Rapid progression of hepatocellular carcinoma after transcatheter arterial chemoembolization and percutaneous radiofrequency ablation in the primary tumour region.**
 Author(s): Seki T, Tamai T, Ikeda K, Imamura M, Nishimura A, Yamashiki N, Nakagawa T, Inoue K.
 Source: European Journal of Gastroenterology & Hepatology. 2001 March; 13(3): 291-4.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11293452&dopt=Abstract
- **Recombinant interferon alfa 2b therapy in a patient with metastatic hepatocellular carcinoma.**
 Author(s): Yuen MF, Hon C, Hui CK, Siu CW, Lai CL.
 Source: Journal of Clinical Gastroenterology. 2002 September; 35(3): 272-5.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=12192207&dopt=Abstract
- **Repeat hepatectomy is the most useful treatment for recurrent hepatocellular carcinoma.**
 Author(s): Sugimachi K, Maehara S, Tanaka S, Shimada M, Sugimachi K.
 Source: Journal of Hepato-Biliary-Pancreatic Surgery. 2001; 8(5): 410-6.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11702249&dopt=Abstract
- **Repeat liver resection for hepatocellular carcinoma.**
 Author(s): Nakajima Y, Ko S, Kanamura T, Nagao M, Kanehiro H, Hisanaga M, Aomatsu Y, Ikeda N, Nakano H.
 Source: Journal of the American College of Surgeons. 2001 March; 192(3): 339-44.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11245376&dopt=Abstract
- **Risk factors for distant recurrence of hepatocellular carcinoma in the liver after complete coagulation by microwave or radiofrequency ablation.**

Author(s): Izumi N, Asahina Y, Noguchi O, Uchihara M, Kanazawa N, Itakura J, Himeno Y, Miyake S, Sakai T, Enomoto N.
Source: Cancer. 2001 March 1; 91(5): 949-56.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11251946&dopt=Abstract

- **Selection criteria for hepatectomy in patients with hepatocellular carcinoma and portal vein tumor thrombus.**

Author(s): Minagawa M, Makuuchi M, Takayama T, Ohtomo K.
Source: Annals of Surgery. 2001 March; 233(3): 379-84.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11224626&dopt=Abstract

- **Spontaneous gas-forming liver abscess caused by Salmonella within hepatocellular carcinoma: a case report and review of the literature.**

Author(s): Lee CC, Poon SK, Chen GH.
Source: Digestive Diseases and Sciences. 2002 March; 47(3): 586-9.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11911347&dopt=Abstract

- **Subphrenic abscess formation following superselective transcatheter chemoembolization for hepatocellular carcinoma.**

Author(s): Yokoi Y, Suzuki S, Sakaguchi T, Okumura T, Kurachi K, Konno H, Nakamura S.
Source: Radiat Med. 2002 January-February; 20(1): 45-9.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=12002604&dopt=Abstract

- **Surgical removal of a distinct subcutaneous metastasis of multilocular hepatocellular carcinoma 2 months after initial percutaneous ethanol injection therapy.**

Author(s): Braune C, Widjaja A, Bartels M, Bleck JS, Flemming P, Manns MP, Klempnauer J, Gebel M.
Source: Zeitschrift Fur Gastroenterologie. 2001 September; 39(9): 789-92.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11558070&dopt=Abstract

- **Surgical treatment of recurrent hepatocellular carcinoma based on the mode of recurrence: repeat hepatic resection or ablation are good choices for patients with recurrent multicentric cancer.**

Author(s): Matsuda M, Fujii H, Kono H, Matsumoto Y.

Source: Journal of Hepato-Biliary-Pancreatic Surgery. 2001; 8(4): 353-9.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11521181&dopt=Abstract

- **The mode of tumour progression in combined hepatocellular carcinoma and cholangiocarcinoma: an immunohistochemical analysis of E-cadherin, alpha-catenin and beta-catenin.**
 Author(s): Asayama Y, Taguchi Ki K, Aishima Si S, Nishi H, Masuda K, Tsuneyoshi M.
 Source: Liver. 2002 February; 22(1): 43-50.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11906618&dopt=Abstract

- **The potential role of bcl-2 mRNA and protein expression in hepatocellular carcinomas.**
 Author(s): Ravazoula P, Tsamandas AC, Kardamakis D, Gogos C, Karatza C, Thomopoulos K, Tepetes K, Kourelis T, Petsas T, Bonikos DS, Karavias D.
 Source: Anticancer Res. 2002 May-June; 22(3): 1799-805.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=12168872&dopt=Abstract

- **The role of diabetes in hepatocellular carcinoma: a case-control study among United States Veterans.**
 Author(s): El-Serag HB, Richardson PA, Everhart JE.
 Source: The American Journal of Gastroenterology. 2001 August; 96(8): 2462-7.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11513191&dopt=Abstract

- **The role of overexpression and gene amplification of cyclin D1 in intrahepatic cholangiocarcinoma.**
 Author(s): Sugimachi K, Aishima S, Taguchi K, Tanaka S, Shimada M, Kajiyama K, Sugimachi K, Tsuneyoshi M.
 Source: Journal of Hepatology. 2001 July; 35(1): 74-9.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11495045&dopt=Abstract

- **Three-dimensional conformal radiation therapy and periodic irradiation with the deep inspiration breath-hold technique for hepatocellular carcinoma.**

Author(s): Pattaranutaporn P, Chansilpa Y, Ieumwananonthachai N, Kakanaporn C, Onnomdee K, Mungkung N, Santisiri R.
Source: J Med Assoc Thai. 2001 December; 84(12): 1692-700.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11999815&dopt=Abstract

- **Thymidine phosphorylase activity in liver tissue and its correlation with multifocal occurrence of hepatocellular carcinomas.**

Author(s): Ikeguchi M, Sakatani T, Ueda T, Hirooka Y, Kaibara N.
Source: In Vivo. 2001 July-August; 15(4): 265-70.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11695216&dopt=Abstract

- **Treatment of adrenal metastases after hepatic resection of a hepatocellular carcinoma.**

Author(s): Shuto T, Hirohashi K, Kubo S, Tanaka H, Yamamoto T, Higaki I, Takemura S, Kinoshita H.
Source: Digestive Surgery. 2001; 18(4): 294-7.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11528139&dopt=Abstract

- **Treatment of ruptured hepatocellular carcinoma.**

Author(s): Tanaka A, Takeda R, Mukaihara S, Hayakawa K, Shibata T, Itoh K, Nishida N, Nakao K, Fukuda Y, Chiba T, Yamaoka Y.
Source: International Journal of Clinical Oncology / Japan Society of Clinical Oncology. 2001 December; 6(6): 291-5.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11828948&dopt=Abstract

- **Tumor heterogeneity in small hepatocellular carcinoma: analysis of tumor cell proliferation, expression and mutation of p53 AND beta-catenin.**

Author(s): An FQ, Matsuda M, Fujii H, Tang RF, Amemiya H, Dai YM, Matsumoto Y.
Source: International Journal of Cancer. Journal International Du Cancer. 2001 August 15; 93(4): 468-74.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11477549&dopt=Abstract

- **Viral serostatus and coexisting inflammatory activity affect metachronous carcinogenesis after hepatectomy for hepatocellular carcinoma. A further report.**

Author(s): Yamanaka N, Takada M, Tanaka T, Yamanaka J, Yasui C, Ando T, Maeda S, Matsushita K, Okamoto E.
Source: Journal of Gastroenterology. 2000; 35(3): 206-13.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=10755690&dopt=Abstract

Vocabulary Builder

Abscess: A localized collection of pus caused by suppuration buried in tissues, organs, or confined spaces. [EU]

Alimentary: Pertaining to food or nutritive material, or to the organs of digestion. [EU]

Alkaline: Having the reactions of an alkali. [EU]

Alkalosis: A pathologic condition resulting from accumulation of base, or from loss of acid without comparable loss of base in the body fluids, and characterized by decrease in hydrogen ion concentration (increase in pH). [EU]

Antigen: Any substance which is capable, under appropriate conditions, of inducing a specific immune response and of reacting with the products of that response, that is, with specific antibody or specifically sensitized T-lymphocytes, or both. Antigens may be soluble substances, such as toxins and foreign proteins, or particulate, such as bacteria and tissue cells; however, only the portion of the protein or polysaccharide molecule known as the antigenic determinant (q.v.) combines with antibody or a specific receptor on a lymphocyte. Abbreviated Ag. [EU]

Aspiration: Removal of fluid from a lump, often a cyst, with a needle and a syringe. [NIH]

Astrocytoma: A tumor that begins in the brain or spinal cord in small, star-shaped cells called astrocytes. [NIH]

Bacteremia: The presence of viable bacteria circulating in the blood. Fever, chills, tachycardia, and tachypnea are common acute manifestations of bacteremia. The majority of cases are seen in already hospitalized patients, most of whom have underlying diseases or procedures which render their bloodstreams susceptible to invasion. [NIH]

Carcinogen: Any substance that causes cancer. [NIH]

Carcinogenesis: The process by which normal cells are transformed into cancer cells. [NIH]

Carcinogenic: Producing carcinoma. [EU]

Cerebrospinal: Pertaining to the brain and spinal cord. [EU]

Cholangitis: Inflammation of a bile duct. [EU]

Chromosomal: Pertaining to chromosomes. [EU]

Chromosome: Part of a cell that contains genetic information. Except for sperm and eggs, all human cells contain 46 chromosomes. [NIH]

Coagulation: 1. the process of clot formation. 2. in colloid chemistry, the solidification of a sol into a gelatinous mass; an alteration of a disperse phase or of a dissolved solid which causes the separation of the system into a liquid phase and an insoluble mass called the clot or curd. Coagulation is usually irreversible. 3. in surgery, the disruption of tissue by physical means to form an amorphous residuum, as in electrocoagulation and photocoagulation. [EU]

Concomitant: Accompanying; accessory; joined with another. [EU]

Contamination: The soiling or pollution by inferior material, as by the introduction of organisms into a wound, or sewage into a stream. [EU]

Dentists: Individuals licensed to practice dentistry. [NIH]

Dysplasia: Cells that look abnormal under a microscope but are not cancer. [NIH]

Epidemiological: Relating to, or involving epidemiology. [EU]

Epidural: The space between the wall of the spinal canal and the covering of the spinal cord. An epidural injection is given into this space. [NIH]

Epigastralgia: Pain in the epigastrium. [EU]

Extravasation: A discharge or escape, as of blood, from a vessel into the tissues. [EU]

Fluorescence: The property of emitting radiation while being irradiated. The radiation emitted is usually of longer wavelength than that incident or absorbed, e.g., a substance can be irradiated with invisible radiation and emit visible light. X-ray fluorescence is used in diagnosis. [NIH]

Glioma: A cancer of the brain that comes from glial, or supportive, cells. [NIH]

Grade: The grade of a tumor depends on how abnormal the cancer cells look under a microscope and how quickly the tumor is likely to grow and spread. Grading systems are different for each type of cancer. [NIH]

Hematoma: An extravasation of blood localized in an organ, space, or tissue. [NIH]

Hemobilia: Hemorrhage in or through the biliary tract, due to trauma, inflammation, cholelithiasis, vascular disease, or neoplasms. [NIH]

Heparin: A drug that helps prevent blood clots from forming. It belongs to the family of drugs called anticoagulants (blood thinners). [NIH]

Hepatocyte: A liver cell. [NIH]

Hybridization: The genetic process of crossbreeding to produce a hybrid.

Hybrid nucleic acids can be formed by nucleic acid hybridization of DNA and RNA molecules. Protein hybridization allows for hybrid proteins to be formed from polypeptide chains. [NIH]

Hypertension: Abnormally high blood pressure. [NIH]

Immunization: The induction of immunity. [EU]

Immunology: The study of the body's immune system. [NIH]

Inflammation: A response of redness, swelling, pain, and a feeling of heat in certain areas which is meant to protect tissues affected by injury or disease. [NIH]

Intermittent: Occurring at separated intervals; having periods of cessation of activity. [EU]

Intracellular: Inside a cell. [NIH]

Intrahepatic: Within the liver. [NIH]

Intrathecal: Describes the fluid-filled space between the thin layers of tissue that cover the brain and spinal cord. Drugs can be injected into the fluid or a sample of the fluid can be removed for testing. [NIH]

Invasive: 1. having the quality of invasiveness. 2. involving puncture or incision of the skin or insertion of an instrument or foreign material into the body; said of diagnostic techniques. [EU]

Laparotomy: A surgical incision made in the wall of the abdomen. [NIH]

Lethal: Deadly, fatal. [EU]

LH: A small glycoprotein hormone secreted by the anterior pituitary. LH plays an important role in controlling ovulation and in controlling secretion of hormones by the ovaries and testes. [NIH]

Ligation: The process of tying off blood vessels so that blood cannot flow to a part of the body or to a tumor. [NIH]

Lobe: A portion of an organ such as the liver, lung, breast, or brain. [NIH]

Lymph: The almost colorless fluid that travels through the lymphatic system and carries cells that help fight infection and disease. [NIH]

Lymphocyte: A white blood cell. Lymphocytes have a number of roles in the immune system, including the production of antibodies and other substances that fight infection and diseases. [NIH]

Lymphoma: Cancer that arises in cells of the lymphatic system. [NIH]

Mediastinum: The area between the lungs. The organs in this area include the heart and its large blood vessels, the trachea, the esophagus, the bronchi, and lymph nodes. [NIH]

Molecular: Of, pertaining to, or composed of molecules : a very small mass of matter. [EU]

Myelosuppression: A condition in which bone marrow activity is decreased, resulting in fewer red blood cells, white blood cells, and platelets. Myelosuppression is a side effect of some cancer treatments. [NIH]

Neurology: A medical specialty concerned with the study of the structures, functions, and diseases of the nervous system. [NIH]

Neurosurgery: A surgical specialty concerned with the treatment of diseases and disorders of the brain, spinal cord, and peripheral and sympathetic nervous system. [NIH]

Oncogene: A gene that normally directs cell growth. If altered, an oncogene can promote or allow the uncontrolled growth of cancer. Alterations can be inherited or caused by an environmental exposure to carcinogens. [NIH]

Pancreatitis: Acute or chronic inflammation of the pancreas, which may be asymptomatic or symptomatic, and which is due to autodigestion of a pancreatic tissue by its own enzymes. It is caused most often by alcoholism or biliary tract disease; less commonly it may be associated with hyperlipaemia, hyperparathyroidism, abdominal trauma (accidental or operative injury), vasculitis, or uraemia. [EU]

Paraffin: A mixture of solid hydrocarbons obtained from petroleum. It has a wide range of uses including as a stiffening agent in ointments, as a lubricant, and as a topical anti-inflammatory. It is also commonly used as an embedding material in histology. [NIH]

Phosphorylase: An enzyme of the transferase class that catalyzes the phosphorylysis of a terminal alpha-1,4-glycosidic bond at the non-reducing end of a glycogen molecule, releasing a glucose 1-phosphate residue. Phosphorylase should be qualified by the natural substance acted upon. EC 2.4.1.1. [NIH]

Plasma: The clear, yellowish, fluid part of the blood that carries the blood cells. The proteins that form blood clots are in plasma. [NIH]

Polyposis: The development of numerous polyps (growths that protrude from a mucous membrane). [NIH]

Postoperative: After surgery. [NIH]

Postprandial: Occurring after dinner, or after a meal; postcibal. [EU]

Precancerous: A term used to describe a condition that may (or is likely to) become cancer. Also called premalignant. [NIH]

Preoperative: Preceding an operation. [EU]

Procarbazine: An anticancer drug that belongs to the family of drugs called alkylating agents. [NIH]

Radium: Radium. A radioactive element of the alkaline earth series of metals. It has the atomic symbol Ra, atomic number 88, and atomic weight

226. Radium is the product of the disintegration of uranium and is present in pitchblende and all ores containing uranium. It is used clinically as a source of beta and gamma-rays in radiotherapy, particularly brachytherapy. [NIH]

Receptor: A molecule inside or on the surface of a cell that binds to a specific substance and causes a specific physiologic effect in the cell. [NIH]

Refractory: Not readily yielding to treatment. [EU]

Salmonella: A genus of gram-negative, facultatively anaerobic, rod-shaped bacteria that utilizes citrate as a sole carbon source. It is pathogenic for humans, causing enteric fevers, gastroenteritis, and bacteremia. Food poisoning is the most common clinical manifestation. Organisms within this genus are separated on the basis of antigenic characteristics, sugar fermentation patterns, and bacteriophage susceptibility. [NIH]

Shunt: A surgically created diversion of fluid (e.g., blood or cerebrospinal fluid) from one area of the body to another area of the body. [NIH]

Skull: The skeleton of the head including the bones of the face and the bones enclosing the brain. [NIH]

Stool: The waste matter discharged in a bowel movement; feces. [NIH]

Subcutaneous: Beneath the skin. [NIH]

Substrate: A substance upon which an enzyme acts. [EU]

Thrombosis: The formation or presence of a blood clot inside a blood vessel. [NIH]

Thrombus: An aggregation of blood factors, primarily platelets and fibrin with entrapment of cellular elements, frequently causing vascular obstruction at the point of its formation. Some authorities thus differentiate thrombus formation from simple coagulation or clot formation. [EU]

Tomography: A series of detailed pictures of areas inside the body; the pictures are created by a computer linked to an x-ray machine. [NIH]

Tumour: 1. swelling, one of the cardinal signs of inflammations; morbid enlargement. 2. a new growth of tissue in which the multiplication of cells is uncontrolled and progressive; called also neoplasm. [EU]

Vaccination: Treatment with a vaccine. [NIH]

Veins: The vessels carrying blood toward the heart. [NIH]

Vincristine: An anticancer drug that belongs to the family of plant drugs called vinca alkaloids. [NIH]

Viruses: Minute infectious agents whose genomes are composed of DNA or RNA, but not both. They are characterized by a lack of independent metabolism and the inability to replicate outside living host cells. [NIH]

CHAPTER 5. BOOKS ON ADULT PRIMARY LIVER CANCER

Overview

This chapter provides bibliographic book references relating to adult primary liver cancer. You have many options to locate books on adult primary liver cancer. The simplest method is to go to your local bookseller and inquire about titles that they have in stock or can special order for you. Some patients, however, feel uncomfortable approaching their local booksellers and prefer online sources (e.g. **www.amazon.com** and **www.bn.com**). In addition to online booksellers, excellent sources for book titles on adult primary liver cancer include the Combined Health Information Database and the National Library of Medicine. Once you have found a title that interests you, visit your local public or medical library to see if it is available for loan.

The National Library of Medicine Book Index

The National Library of Medicine at the National Institutes of Health has a massive database of books published on healthcare and biomedicine. Go to the following Internet site, <http://locatorplus.gov/>, and then select "Search LOCATORplus." Once you are in the search area, simply type "adult primary liver cancer" (or synonyms) into the search box, and select "books only." From there, results can be sorted by publication date, author, or relevance. The following was recently catalogued by the National Library of Medicine:³¹

³¹ In addition to LOCATORPlus, in collaboration with authors and publishers, the National Center for Biotechnology Information (NCBI) is adapting biomedical books for the Web. The books may be accessed in two ways: (1) by searching directly using any search term or phrase (in the same way as the bibliographic database PubMed), or (2) by following the

- **Adult primary care.** Author: [edited by] Pamela Vesta Meredith, Nancy Mathes Horan; Year: 2000; Philadelphia: Saunders, c2000; ISBN: 0721660371
<http://www.amazon.com/exec/obidos/ASIN/0721660371/icongroupinterna>
- **Associated primary cancer of the larynx and bronchogenic carcinoma.** Author: Perez, Patricio Eugenio; Year: 1960; [Minneapolis] 1960
- **Clinical skills for adult primary care.** Author: editors, Mark E. Silverman, J. Willis Hurst; Year: 1996; Philadelphia: Lippincott-Raven, c1996; ISBN: 0781703271
<http://www.amazon.com/exec/obidos/ASIN/0781703271/icongroupinterna>
- **Diagnosis by ultrasound-guided fine-needle biopsy: studies on renal expansions, primary liver malignancies, enlarged lymph nodes and anterior mediastinal masses.** Author: Tapani Tikkakoski; Year: 1990; Oulu: University of Oulu, 1990; ISBN: 9514229673
- **Early detection and treatment of liver cancer.** Author: edited by Kunio Okuda, Takayoshi Tobe, Tomoyuki Kitagawa; Year: 1991; Tokyo: Japan Scientific Societies Press; London; Bristol, Pa.: Taylor ; Francis, 1991; ISBN: 4762246662 (Japan Scientific Societies Press)
<http://www.amazon.com/exec/obidos/ASIN/4762246662/icongroupinterna>
- **Handbook for the care of the older adult with cancer.** Author: edited by Ann Schmidt Luggen, Sue E. Meiner; Year: 2000; Pittsburgh, PA: Oncology Nursing Press, Inc., c2000; ISBN: 1890504165
<http://www.amazon.com/exec/obidos/ASIN/1890504165/icongroupinterna>
- **Handbook of adult primary care.** Author: Carla Greene, editor; Robert W. Singer, general consultant; illustrated by Masako Herman; Year: 1987; New York: Wiley, c1987; ISBN: 0471096210
<http://www.amazon.com/exec/obidos/ASIN/0471096210/icongroupinterna>
- **Hepatocellular carcinoma.** Author: edited by Kunio Okuda and Ian Mackay; Year: 1982; Geneva: International Union Against Cancer; Bern, Switzerland: Huber [distributor], 1982; ISBN: 9290180749 (pbk.)

links to PubMed abstracts. Each PubMed abstract has a "Books" button that displays a facsimile of the abstract in which some phrases are hypertext links. These phrases are also found in the books available at NCBI. Click on hyperlinked results in the list of books in which the phrase is found. Currently, the majority of the links are between the books and PubMed. In the future, more links will be created between the books and other types of information, such as gene and protein sequences and macromolecular structures. See <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=Books>.

<http://www.amazon.com/exec/obidos/ASIN/9290180749/icongroupinterna>

- **LEC rat: a new model for hepatitis and liver cancer.** Author: M. Mori ... [et al.]; [foreword by T. Sugimura]; Year: 1991; Tokyo; New York: Springer-Verlag, c1991; ISBN: 443170079X
<http://www.amazon.com/exec/obidos/ASIN/443170079X/icongroupinterna>
- **Liver and cancer; a new cancer theory.** Author: Blond, Kasper; Year: 1960; Bristol, Wright, 1960
- **Liver cancer risk from internally-deposited radionuclides: recommendation of the National Council on Radiation Protection and Measurements.** Author: National Council on Radiation Protection and Measurements; Year: 2000; Bethesda, Md.: The Council, 2000; ISBN: 0929600681 (alk. paper)
<http://www.amazon.com/exec/obidos/ASIN/0929600681/icongroupinterna>
- **Liver cancer.** Author: edited by Kunio Okuda, Edward Tabor; Year: 1997; New York: Churchill Livingstone, 1997; ISBN: 0443054819
<http://www.amazon.com/exec/obidos/ASIN/0443054819/icongroupinterna>
- **Liver cancer.** Author: Steven A. Curley, editor; Year: 1998; New York: Springer, c1998; ISBN: 0387983708 (alk. paper)
<http://www.amazon.com/exec/obidos/ASIN/0387983708/icongroupinterna>
- **Liver cancer.** Author: edited by Joseph C. Bottino, Richard W. Opfell, Franco M. Muggia; Year: 1985; Boston: Nijhoff; Hingham, MA: Distributors for North America, Kluwer Academic, c1985; ISBN: 0898387132
<http://www.amazon.com/exec/obidos/ASIN/0898387132/icongroupinterna>
- **Liver cell cancer.** Author: edited by H. M. Cameron, D.[i.e. C.] A. Linsell, and G. P. Warwick; Year: 1976; Amsterdam; New York: Elsevier Scientific Pub. Co., 1976; ISBN: 0444415424
<http://www.amazon.com/exec/obidos/ASIN/0444415424/icongroupinterna>
- **Management of gonadal toxicity resulting from the treatment of adult cancer: report of a working party of the Joint Council for Clinical Oncology.** Author: Joint Council for Clinical Oncology; Year: 1998; London: Royal College of Physicians: Royal College of Radiologist, c1998; ISBN: 1860160719

- **Multiple primary cancer; a clinical-statistical investigation based on 650 cases.** [Tr. by Eva Palmgren]. Author: Malmio, Kai; Year: 1959; Helsinki, 1959
- **National Health Services and Practice Patterns Survey first-year report on adult liver transplantaioin operating costs, Medicare payments, and utilization rates.** Author: prepared by Flora Chu, Dennis Cotter, Mari Anne T. Hamilton; Year: 1988; Washington, D.C. (2233 Wisconsin Ave. N.W., Washington): Medical Technology and Practice Patterns Institute, c1988
- **Prevention of liver cancer: report of a WHO meeting.** Author: Tikkakoski, Tapani; Year: 1983; Geneva: World Health Organization, 1983; ISBN: 9241206918 (pbk.)
- **Primary liver cancer: etiological and progression factors.** Author: edited by Christian Bréchet; Year: 1994; Boca Raton: CRC Press, c1994; ISBN: 0849349133 (alk. paper)
<http://www.amazon.com/exec/obidos/ASIN/0849349133/icongroupinterna>
- **Primary liver cancer.** Author: Tang Zhao-you, Wu Meng-chao, Xia Sui-sheng (eds.); associate editor, Ye Sheng-long; forewords by Li Bing and Qiu Fa-zu; Year: 1989; Beijing: China Academic Publishers; Berlin; New York: Springer-Verlag, c1989; ISBN: 0387502289 (SpringerVerlag: U.S.: alk. paper)
<http://www.amazon.com/exec/obidos/ASIN/0387502289/icongroupinterna>
- **Primary liver tumors: proceedings of the 25th Falk Symposium, on the occasion of the 5th centennial celebrations of the Eberhard-Karls-Universität Tübingen held at the Schwartzwaldhotel, Titisee, West Germany, October 3-5, 1977.** Author: edited by H. Remmer .; Year: 1978; Baltimore University Park Press c1978; ISBN: 0839112262
<http://www.amazon.com/exec/obidos/ASIN/0839112262/icongroupinterna>
- **Role of 239PU-induced chromosome alterations and mutated KI-V-RAS oncogene during liver-cancer induction in chinese hamsters and mice [microform].** Author: A.L. Brooks ... [et al.]; Year: 1994; [Washington, D.C.?]: U.S. Dept. of Energy, [1994]
- **Synthesis of a research on liver cancer in Thailand.** Author: by Paibul Suriyawongpaisal, Nilarat Premmanisakul; Year: 2002; [Bangkok?]: Thailand Research Fund, [2000?]
- **Therapeutic strategies in primary and metastatic liver cancer.** Author: edited by Ch. Herfarth, P. Schlag, and P. Hohenberger; Year: 1986; Berlin; New York: Springer-Verlag, c1986; ISBN: 0387160116 (U.S.)

<http://www.amazon.com/exec/obidos/ASIN/0387160116/icongroupinterna>

- **Towards the aggressive surgery of liver cancer based on the redox theory.** Author: Kazue Ozawa; Year: 1992; [Japan]: Nakayama Institute of Cancer Research, [1992]; ISBN: 4906225225
- **Unusual clinical features of cirrhosis and primary liver cell carcinoma, by Luis Leon-Sotomayor and Victor A. Moore.** Author: Leon-Sotomayor, Luis, 1931-; Year: 1967; Springfield, Ill., Thomas [c1967]
- **Update on regional treatment of liver cancer: the role of vascular occlusion.** Author: edited by N. Kemeny ... [et al.]; Year: 1992; Kent: Wells Medical Ltd., c1992; ISBN: 1869969634
- **Viruses and liver cancer.** Author: editor, Edward Tabor; Year: 2002; Amsterdam; Boston: Elsevier, 2002; ISBN: 0444505806 (alk. paper)
<http://www.amazon.com/exec/obidos/ASIN/0444505806/icongroupinterna>

Chapters on Adult Primary Liver Cancer

Frequently, adult primary liver cancer will be discussed within a book, perhaps within a specific chapter. In order to find chapters that are specifically dealing with adult primary liver cancer, an excellent source of abstracts is the Combined Health Information Database. You will need to limit your search to book chapters and adult primary liver cancer using the "Detailed Search" option. Go directly to the following hyperlink: <http://chid.nih.gov/detail/detail.html>. To find book chapters, use the drop boxes at the bottom of the search page where "You may refine your search by." Select the dates and language you prefer, and the format option "Book Chapter." By making these selections and typing in "adult primary liver cancer" (or synonyms) into the "For these words:" box, you will only receive results on chapters in books.

General Home References

In addition to references for adult primary liver cancer, you may want a general home medical guide that spans all aspects of home healthcare. The following list is a recent sample of such guides (sorted alphabetically by title; hyperlinks provide rankings, information, and reviews at Amazon.com):

- **Cancer: 50 Essential Things to Do** by Greg Anderson, O. Carl Simonton; Paperback - 184 pages; Revised & Updated edition (August 1999), Plume;

ISBN: 0452280745;

<http://www.amazon.com/exec/obidos/ASIN/0452280745/icongroupinterna>

- **Cancer Encyclopedia -- Collections of Anti-Cancer & Anti-Carcinogenic Agents, Chemicals, Drugs and Substances** by John C. Bartone; Paperback (January 2002), ABBE Publishers Association of Washington, DC; ISBN: 0788326791;
<http://www.amazon.com/exec/obidos/ASIN/0788326791/icongroupinterna>
- **Cancer Sourcebook: Basic Consumer Health Information About Major Forms and Stages of Cancer** by Edward J. Prucha (Editor); Library Binding - 1100 pages, 3rd edition (August 1, 2000), Omnigraphics, Inc.; ISBN: 0780802276;
<http://www.amazon.com/exec/obidos/ASIN/0780802276/icongroupinterna>
- **Cancer Supportive Care: A Comprehensive Guide for Patients and Their Families** by Ernest H. Rosenbaum, M.D., Isadora Rosenbaum, M.A.; Paperback - 472 pages (November 5, 1998), Somerville House Books Limited; ISBN: 1894042115;
<http://www.amazon.com/exec/obidos/ASIN/1894042115/icongroupinterna>
- **Cancer Symptom Management: Patient Self-Care Guides (Book with CD-ROM for Windows & Macintosh)** by Connie Henke Yarbro (Editor), et al; CD-ROM - 264 pages, 2nd Book & CD-Rom edition (January 15, 2000), Jones & Bartlett Publishing; ISBN: 0763711675;
<http://www.amazon.com/exec/obidos/ASIN/0763711675/icongroupinterna>
- **Diagnosis Cancer: Your Guide Through the First Few Months** by Wendy Schlessel Harpham, Ann Bliss Pilcher (Illustrator); Paperback: 230 pages; Revised & Updated edition (November 1997), .W. Norton & Company; ISBN: 0393316912;
<http://www.amazon.com/exec/obidos/ASIN/0393316912/icongroupinterna>
- **The Human Side of Cancer: Living with Hope, Coping with Uncertainty** by Jimmie C. Holland, M.D., Sheldon Lewis; Paperback - 368 pages (October 2, 2001), Quill; ISBN: 006093042X;
<http://www.amazon.com/exec/obidos/ASIN/006093042X/icongroupinterna>

Vocabulary Builder

Gonadal: Pertaining to a gonad. [EU]

Induction: The act or process of inducing or causing to occur, especially the production of a specific morphogenetic effect in the developing embryo through the influence of evocators or organizers, or the production of anaesthesia or unconsciousness by use of appropriate agents. [EU]

Larynx: The area of the throat containing the vocal cords and used for breathing, swallowing, and talking. Also called the voice box. [NIH]

Radiologist: A doctor who specializes in creating and interpreting pictures of areas inside the body. The pictures are produced with x-rays, sound waves, or other types of energy. [NIH]

CHAPTER 6. MULTIMEDIA ON ADULT PRIMARY LIVER CANCER

Overview

Information on adult primary liver cancer can come in a variety of formats. Among multimedia sources, video productions, slides, audiotapes, and computer databases are often available. In this chapter, we show you how to keep current on multimedia sources of information on adult primary liver cancer. We start with sources that have been summarized by federal agencies, and then show you how to find bibliographic information catalogued by the National Library of Medicine. If you see an interesting item, visit your local medical library to check on the availability of the title.

Bibliography: Multimedia on Adult Primary Liver Cancer

The National Library of Medicine is a rich source of information on healthcare-related multimedia productions including slides, computer software, and databases. To access the multimedia database, go to the following Web site: <http://locatorplus.gov/>. Select "Search LOCATORplus." Once in the search area, simply type in adult primary liver cancer (or synonyms). Then, in the option box provided below the search box, select "Audiovisuals and Computer Files." From there, you can choose to sort results by publication date, author, or relevance. The following multimedia has been indexed on adult primary liver cancer. For more information, follow the hyperlink indicated:

- **Adjuvant therapy of primary breast cancer, 1989.** Source: produced in the facilities of Instructional Media Services, University of Washington;

Year: 1989; Format: Videorecording; [Seattle, Wash.]: University of Washington, c1989

- **Adult primary care.** Source: a co-production of Multimedia Communications and Physical Education and Development; Year: 1998; Format: Videorecording; Oakland, CA: Kaiser Foundation Health Plan, c1998
- **Biliary and liver resection for cancer of the gallbladder or hilar cholangiocarcinoma.** Source: American College of Surgeons; from the Film Library and the Clinical Congress of ACS; Memorial Sloan Kettering Cancer Center; produced by the Hepat; Year: 2000; Format: Videorecording; [Woodbury, Conn.]: Ciné-Med, distributor, [2000]
- **Cancer of unknown primary site.** Source: Marshfield Clinic, Saint Joseph's Hospital; a presentation of the Marshfield Video Network; Year: 1995; Format: Videorecording; Marshfield, WI: Video Network, [1995]
- **Hepatic arterial chemotherapy for cancer in the liver.** Source: Biomedical Media Production Unit, the University of Michigan Medical Center, Office of Educational Resources & Research; Year: 1981; Format: Videorecording; Ann Arbor, Mich.: The University, c1981
- **Hepatobiliary cancer.** Source: Leslie H. Blumgart, Yuman Fong, and William R. Jarnagin; Year: 2000; Format: Edited by; Hamilton, Ont.: B C Decker: Lewiston, NY: Sales and distribution, US, B.C. Decker, 2001
- **International symposium: liver cancer, a global problem.** Source: ASCO 36th Annual Meeting, May 2000; Year: 2000; Format: Sound recording; [Alexandria, Va.]: American Society of Clinical Oncology, c2000
- **James Ewing lecture: one life, many livers.** Source: the Society of Surgical Oncology & the World Federation of Surgical Oncology Societies' Cancer Symposium, March 26-29, 1998, San Diego, CA; Year: 1998; Format: Videorecording; Chicago, Ill.: Teach 'em, [1998]
- **Medical informatics and medical management.** Source: director, Core Curriculum Committee, Michael K. Rees; Year: 2001; Format: Electronic resource; Nashville, TN: HealthStream, c2001
- **Mental health primary care in prison: a guide to mental ill health in adults and adolescents in prison and young offender institutions.** Source: edited by Jo Paton and Rachel Jenkins; Year: 2002; London: Royal Society of Medicine Press, c2002
- **Neoplasms of the pancreatobiliary system and liver.** Source: Walter Lawrence, Jr., William Gayle, Jose J. Terz; [made by Medical College of Virginia Visual Education Dept]; Year: 1971; Format: Slide; [Richmond: Medical College of Virginia Learning Resource Center; Chapel Hill, N.C.: for loan by Health Sciences Consortium, 1971]

- **Primary hepatobiliary tumors: an update.** Source: the Society of Surgical Oncology & the World Federation of Surgical Oncology Societies' Cancer Symposium, March 26-29, 1998, San Diego, CA; Year: 1998; Format: Videorecording; Chicago, Ill.: Teach'em, [1998]
- **Prostate cancer and primary care: detection, diagnosis, and tracking treatment response.** Source: Mitchell C. Benson; Year: 1998; Format: Videorecording; Clifton, N.J.: Network for Continuing Medical Education, c1998
- **Second primary gastric cancer of the reconstructed esophagus.** Source: from the Motion Picture Library of the American College of Surgeons; Year: 1986; Format: Videorecording; Danbury, Conn.: American College of Surgeons, Davis ; Geck Surgical Film-Video Library, [1986]
- **Surgical management of the patient with liver and pancreatic cancer.** Source: Memorial Sloan-Kettering Cancer Center; Year: 1976; Format: Sound recording; [New York]: The Center, c1976
- **Techniques of liver transplantation in children.** Source: [author, Thierry Yandza; collaborators, Hervé Blanchard, Jacques Corman, Catherine Ferell]; Year: 1996; Format: Electronic resource; Montreal, Quebec: SSB Multimedia, c1996

Vocabulary Builder

Esophagus: The muscular tube through which food passes from the throat to the stomach. [NIH]

Lobectomy: The removal of a lobe. [NIH]

Otolaryngology: A surgical specialty concerned with the study and treatment of disorders of the ear, nose, and throat. [NIH]

Pulmonary: Relating to the lungs. [NIH]

CHAPTER 7. PHYSICIAN GUIDELINES AND DATABASES

Overview

Doctors and medical researchers rely on a number of information sources to help patients with their conditions. Many will subscribe to journals or newsletters published by their professional associations or refer to specialized textbooks or clinical guides published for the medical profession. In this chapter, we focus on databases and Internet-based guidelines created or written for this professional audience.

NIH Guidelines

For the more common diseases, The National Institutes of Health publish guidelines that are frequently consulted by physicians. Publications are typically written by one or more of the various NIH Institutes. For physician guidelines, commonly referred to as “clinical” or “professional” guidelines, you can visit the following Institutes:

- Office of the Director (OD); guidelines consolidated across agencies available at <http://www.nih.gov/health/consumer/conkey.htm>
- National Institute of General Medical Sciences (NIGMS); fact sheets available at <http://www.nigms.nih.gov/news/facts/>
- National Library of Medicine (NLM); extensive encyclopedia (A.D.A.M., Inc.) with guidelines:
<http://www.nlm.nih.gov/medlineplus/healthtopics.html>
- National Cancer Institute (NCI); guidelines available at http://cancernet.nci.nih.gov/pdq/pdq_treatment.shtml

In this chapter, we begin by reproducing one such guideline for adult primary liver cancer:

What Is Adult Primary Liver Cancer?³²

Hepatocellular carcinoma is a tumor that is relatively uncommon in the United States, although its incidence is rising, principally in relation to the spread of hepatitis C infection.³³ It is the most common cancer in some other parts of the world. Hepatocellular carcinoma is potentially curable by surgical resection, but surgery is the treatment of choice for only the small fraction of patients with localized disease.³⁴ Prognosis depends on the degree of local tumor replacement and the extent of liver function impairment. Therapy other than surgical resection is best administered as part of a clinical trial. Such trials evaluate the efficacy of systemic or infusional chemotherapy, hepatic artery ligation or embolization, percutaneous ethanol injection, radiofrequency ablation, cryotherapy, and radiolabeled antibodies, often in conjunction with surgical resection and/or radiation therapy. In some studies of these approaches, long remissions have been reported.³⁵ Hepatocellular carcinoma should be distinguished from bile duct cancer (cholangiocarcinoma) as well as from metastatic cancer that originates in another organ.

Hepatocellular carcinoma is associated with cirrhosis in 50% to 80% of patients; 5% of cirrhotic patients eventually develop hepatocellular cancer, which is often multifocal.

Hepatitis B infection³⁶ and hepatitis C infection³⁷ appear to be the most significant causes of hepatocellular carcinoma worldwide, particularly in

³² The following guidelines appeared on the NCI website on Aug. 26, 2002. The text was last modified in February 2002. The text has been adapted for this sourcebook. Note: Some citations in the text of this section are followed by a level of evidence. The PDQ editorial boards use a formal ranking system to help the reader judge the strength of evidence linked to the reported results of a therapeutic strategy.

³³ El-Serag HB, Mason AC: Rising incidence of hepatocellular carcinoma in the United States. *New England Journal of Medicine* 340(10): 745-750, 1999.

³⁴ Mor E, Kasper RT, Sheiner P, et al.: Treatment of hepatocellular carcinoma associated with cirrhosis in the era of liver transplantation. *Annals of Internal Medicine* 129(8): 643-653, 1998.

³⁵ Mor E, Kasper RT, Sheiner P, et al.: Treatment of hepatocellular carcinoma associated with cirrhosis in the era of liver transplantation. *Annals of Internal Medicine* 129(8): 643-653, 1998.

³⁶ Mor E, Kasper RT, Sheiner P, et al.: Treatment of hepatocellular carcinoma associated with cirrhosis in the era of liver transplantation. *Annals of Internal Medicine* 129(8): 643-653, 1998.

patients with continuing antigenemia and in those who have chronic active hepatitis. A series found that male patients older than 50 years of age who have both hepatitis B and hepatitis C infection may be at particularly high risk for hepatocellular cancers.³⁸ [Level of evidence:3iii] There is evidence that patients with both hepatitis B and hepatitis C infection who consume more than 80 grams of alcohol per day have an increased risk of developing cancer (odds ratio of 7.3) when compared to patients who abstain from alcohol.³⁹ Additionally, having a first-degree relative with hepatitis B plus hepatocellular carcinoma is associated with an increased risk (odds ratio 2.41) for family members who are hepatitis B carriers.⁴⁰

Aflatoxin has also been implicated as a factor in the etiology of primary liver cancer in parts of the world where this mycotoxin occurs in high levels in ingested food.⁴¹ Workers who were exposed to vinyl chloride before controls on vinyl chloride dust were instituted developed sarcomas in the liver, most commonly angiosarcomas. Other sarcomas of smooth muscular and vascular origin are also found.

The primary symptoms of hepatocellular carcinoma are those of a hepatic mass. Among patients with underlying cirrhotic disease, a progressive increase in alpha-fetoprotein (AFP) and/or in alkaline phosphatase or a rapid deterioration of hepatic function may be the only clue to the presence of the neoplasm. Infrequently, patients with this disease have polycythemia, hypoglycemia, hypercalcemia, or dysfibrinogenemia.

The biologic marker AFP is useful for the diagnosis of this neoplasm. By a radioimmunoassay technique, 50% to 70% of patients in the United States who have hepatocellular carcinoma have elevated levels of AFP. However,

Blumberg BS, Larouze B, London WT, et al.: The relation of infection with hepatitis B agent to primary hepatic carcinoma. *American Journal of Pathology* 81(3): 669-682, 1975.

³⁷ Tsukuma H, Hiyama T, Tanaka S, et al.: Risk factors for hepatocellular carcinoma among patients with chronic liver disease. *New England Journal of Medicine* 328(25): 1797-1801, 1993.

³⁸ Chiamonte M, Stroffolini T, Vian A, et al.: Rate of incidence of hepatocellular carcinoma in patients with compensated viral cirrhosis. *Cancer* 85(10): 2132-2137, 1999.

³⁹ Tagger A, Donato F, Ribero ML, et al.: Case-control study on hepatitis C virus (HCV) as a risk factor for hepatocellular carcinoma: the role of HCV genotypes and the synergism with hepatitis B virus and alcohol. *International Journal of Cancer* 81(5): 695-699, 1999.

⁴⁰ Yu MW, Chang HC, Liaw YF, et al.: Familial risk of hepatocellular carcinoma among chronic hepatitis B carriers and their relatives. *Journal of the National Cancer Institute* 92(14): 1159-1164, 2000.

⁴¹ Blumberg BS, Larouze B, London WT, et al.: The relation of infection with hepatitis B agent to primary hepatic carcinoma. *American Journal of Pathology* 81(3): 669-682, 1975.

Alpert ME, Hutt MS, Wogan GN, et al.: Association between aflatoxin content of food and hepatoma frequency in Uganda. *Cancer* 28(1): 253-260, 1971.

patients with other malignancies (germ cell carcinoma and, rarely, pancreatic and gastric carcinoma) also demonstrate elevated serum levels of this protein. AFP levels have been shown to be prognostically important, with the median survival of AFP-negative patients significantly longer than that of AFP-positive patients.⁴² Other prognostic variables include performance status, liver functions,⁴³ and the presence or absence of cirrhosis and its severity in relation to the Child-Pugh classification.⁴⁴

Patients scheduled for possible resection require preoperative assessment with angiography in conjunction with helical computed tomographic (CT) scan or magnetic resonance imaging (MRI) with magnetic resonance angiography; these scans have obviated the need for angiography in most patients. Information on the arterial anatomy is helpful for the operating surgeon and may eliminate some patients from consideration for resection. Dynamic CT and MRI scans can document the relationship of the tumor to the hepatic and portal veins (and, on occasion, involvement of these structures), delineating tumors for which the chances for surgical cure are remote.⁴⁵ Laparoscopic evaluation may detect metastatic disease, bilobar disease, or inadequate liver remnant, and therefore obviate the need for open surgical exploration.⁴⁶

Cellular Classification

Malignant tumors of the liver are primarily adenocarcinomas, with 2 major cell types: hepatocellular and cholangiocarcinoma.

⁴² Stillwagon GB, Order SE, Guse C, et al.: Prognostic factors in unresectable hepatocellular cancer: Radiation Therapy Oncology Group study 83-01. *International Journal of Radiation Oncology, Biology, Physics* 20(1): 65-71, 1991.

Izumi R, Shimizu K, Kiriya M, et al.: Alpha-fetoprotein production by hepatocellular carcinoma is prognostic of poor patient survival. *Journal of Surgical Oncology* 49(3): 151-155, 1992.

⁴³ Yamashita Y, Takahashi M, Koga Y, et al.: Prognostic factors in the treatment of hepatocellular carcinoma with transcatheter arterial embolization and arterial infusion. *Cancer* 67(2): 385-391, 1991.

⁴⁴ Nakakura EK, Choti MA: Management of hepatocellular carcinoma. *Oncology (Huntington NY)* 14(7): 1085-1100, 2000.

⁴⁵ Karl RC, Morse SS, Halpert RD, et al.: Preoperative evaluation of patients for liver resection: appropriate CT imaging. *Annals of Surgery* 217(3): 226-232, 1993.

⁴⁶ Lo CM, Lai EC, Liu CL, et al.: Laparoscopy and laparoscopic ultrasonography avoid exploratory laparotomy in patients with hepatocellular carcinoma. *Annals of Surgery* 227(4): 527-532, 1998.

Histologic classification is as follows:

- Hepatocellular carcinoma (liver cell carcinoma)
- Hepatocellular carcinoma (fibrolamellar variant)
- Cholangiocarcinoma (intrahepatic bile duct carcinoma)
- Mixed hepatocellular cholangiocarcinoma
- Undifferentiated

Hepatoblastoma rarely occurs in adults.⁴⁷

Stage Information

The American Joint Committee on Cancer (AJCC) has designated TNM stages for liver cancer as follows:⁴⁸

TNM Definitions

Primary tumor (T):

- TX: Primary tumor cannot be assessed
- T0: No evidence of primary tumor
- T1: Solitary tumor 2 cm or less in greatest dimension without vascular invasion
- T2: Solitary tumor 2 cm or less in greatest dimension with vascular invasion; or multiple tumors limited to one lobe, none more than 2 cm in greatest dimension without vascular invasion; or a solitary tumor more than 2 cm in greatest dimension without vascular invasion
- T3: Solitary tumor more than 2 cm in greatest dimension with vascular invasion; or multiple tumors limited to one lobe, none more than 2 cm in greatest dimension, with vascular invasion; or multiple tumors limited to one lobe, any more than 2 cm in greatest dimension, with or without vascular invasion

⁴⁷ Note: The fibrolamellar variant is important because an increased proportion of these patients may be cured if the tumor can be resected. It is more frequent in young women. It also generally exhibits a slower clinical course than the more common hepatocellular carcinoma.

⁴⁸ Liver (including intrahepatic bile ducts). In: American Joint Committee on Cancer: AJCC Cancer Staging Manual. Philadelphia, Pa: Lippincott-Raven Publishers, 5th ed., 1997, pp 97-101.

- T4: Multiple tumors in more than one lobe or tumor(s) involving a major branch of portal or hepatic vein(s) or invasion of adjacent organs other than the gallbladder or perforation of the visceral peritoneum

Note: For classification, the plane projecting between the bed of the gallbladder and the inferior vena cava divides the liver into 2 lobes.

Regional lymph nodes (N):

- NX: Regional lymph nodes cannot be assessed
- N0: No regional lymph node metastasis
- N1: Regional lymph node metastasis

Note: The regional lymph nodes are the hilar (i.e., those in the hepatoduodenal ligament, hepatic and periportal nodes). Regional lymph nodes also include those along the inferior vena cava, hepatic artery, and portal vein. Any lymph node involvement beyond these nodes is considered distant metastasis and should be coded as M1. Involvement of the inferior phrenic lymph nodes should also be considered M1.

Distant metastasis (M):

- MX: Distant metastasis cannot be assessed
- M0: No distant metastasis
- M1: Distant metastasis

Note: Metastases occur most frequently in bones and lungs. Tumors may extend through the capsule to the diaphragm.

AJCC Stage Groupings

Stage I

- T1, N0, M0

Stage II

- T2, N0, M0

Stage IIIA

- T3, N0, M0

Stage IIIB

- T1, N1, M0
- T2, N1, M0
- T3, N1, M0

Stage IVA

- T4, Any N, M0

Stage IVB

- Any T, Any N, M1

For purposes of treatment, patients with liver cancer are grouped as localized resectable, localized unresectable, or advanced disease. These groups are described with the following TNM correlations:

Localized Resectable (T1, T2, T3, and Selected T4; N0; M0)

This type of liver cancer is confined to a solitary mass in a portion of the liver that allows the possibility of complete surgical removal of the tumor with a margin of normal liver. Liver function tests are usually normal or minimally abnormal, and there should be no evidence of cirrhosis or chronic hepatitis. Only a small percentage of liver cancer patients will prove to have such localized resectable disease. Preoperative assessment that includes computed tomography and/or MR scanning should be directed toward determining the presence of extension of tumor across interlobar planes, involvement of the hepatic hilus, or encroachment on the vena cava. A resected specimen should contain a 1- to 2-centimeter margin of normal liver. Patients with chronic hepatitis and cirrhosis are at high risk when surgical resection is performed.

Localized Unresectable (Selected T2, T3, and T4; N0; M0)

This type of cancer appears to be confined to the liver, but surgical resection of the entire tumor is not possible despite a localized mass because of location within the liver or concomitant medical conditions (such as

cirrhosis). Patients with locally unresectable fibrolamellar variant hepatomas may be considered for liver transplantation.⁴⁹ For other patients, chemoembolization may be an option.⁵⁰

Advanced (Any T, N1 or M1)

Advanced liver cancer is cancer that is present in both lobes of the liver or has metastasized to distant sites. Median survival is usually 2 to 4 months. The most common metastatic sites of hepatocellular cancer are the lungs and bone. Multifocal disease in the liver is common, particularly when cirrhosis or chronic hepatitis is present. Chemoembolization has been beneficial in selected patients who have no extrahepatic metastases.⁵¹

Treatment Option Overview

The designations in PDQ that treatments are “standard” or “under clinical evaluation” are not to be used as a basis for reimbursement determinations.

Localized Resectable Adult Primary Liver Cancer (T1, T2, T3, and Selected T4; N0; M0)⁵²

Standard treatment options:

- **Surgery:** Resection of the localized liver cancer varies from segmental resection to trisegmental (80%) resection. In series of carefully selected patients, partial hepatectomy has resulted in a 5-year survival of 10% to

⁴⁹ Farmer DG, Rosove MH, Shaked A, et al.: Current treatment modalities for hepatocellular carcinoma. *Annals of Surgery* 219(3): 236-247, 1994.

Ringe B, Wittekind C, Weimann A, et al.: Results of hepatic resection and transplantation for fibrolamellar carcinoma. *Surgery, Gynecology and Obstetrics* 175(4): 299-305, 1992.

Venook AP: Treatment of hepatocellular carcinoma: too many options? *Journal of Clinical Oncology* 12(6): 1323-1334, 1994.

Iwatsuki S, Starzl TE, Sheahan DG, et al.: Hepatic resection versus transplantation for hepatocellular carcinoma. *Annals of Surgery* 214(3):221-229, 1991.

⁵⁰ Tanaka K, Nakamura S, Numata K, et al.: The long term efficacy of combined transcatheter arterial embolization and percutaneous ethanol injection in the treatment of patients with large hepatocellular carcinoma and cirrhosis. *Cancer* 82(1): 78-85, 1998.

⁵¹ Tanaka K, Nakamura S, Numata K, et al.: The long term efficacy of combined transcatheter arterial embolization and percutaneous ethanol injection in the treatment of patients with large hepatocellular carcinoma and cirrhosis. *Cancer* 82(1): 78-85, 1998.

⁵² Note: Some citations in the text of this section are followed by a level of evidence. The PDQ editorial boards use a formal ranking system to help the reader judge the strength of evidence linked to the reported results of a therapeutic strategy.

30%. Hepatic carcinoma is frequently multifocal and may involve multiple sites throughout the liver at the time of exploration, even when a dominant mass is found on preoperative assessment. Preoperative assessment should also include a search for extrahepatic metastases, since this condition will also preclude the planned hepatic resection. Resection that involves more than a wedge of liver is poorly tolerated (high mortality rate) in patients with cirrhosis or chronic active hepatitis. These are generally contraindications to major hepatic resection but may not contraindicate hepatic transplantation.⁵³ Hepatic transplantation for hemangioendothelioma, fibrolamellar hepatocellular carcinoma, and small (<5 cm) hepatocellular carcinoma in patients with or without cirrhosis has been associated with 5-year survivals of 20% to 30%.⁵⁴ [Level of evidence: 3iiiA];⁵⁵

Treatment options under clinical evaluation:

- Because of the high proportion of patients who experience relapse following surgery for localized hepatic cancer, adjuvant approaches have been employed using regional arterial infusion of the liver or systemic therapy with chemotherapeutic agents. One randomized trial of 43 patients suggested improved survival with adjuvant injection of a single dose (1850 MBq) of I-131 lipiodol via the hepatic artery. Median disease-free survival in the treatment group was 57 months compared to 13.6 months in the group that did not receive treatment beyond resection (=0.037).⁵⁶ [Level of evidence: 1iiA,B] Lipiodol was nontoxic, but required thyroid suppression before and after surgery. Enrollment in this trial was prematurely terminated because of early differences in survival between the treatment and control arms. Therefore, the results must be considered preliminary and will require confirmation. Adoptive immunotherapy

⁵³ Starzl TE, Koep LJ, Weil R, et al.: Right trisegmentectomy for hepatic neoplasms. *Surgery, Gynecology and Obstetrics* 150(2): 208-214, 1980.

Nagorney DM, van Heerden JA, Ilstrup DM, et al.: Primary hepatic malignancy: surgical management and determinants of survival. *Surgery* 106(4): 740-749, 1989.

MacIntosh EL, Minuk GY.: Hepatic resection in patients with cirrhosis and hepatocellular carcinoma. *Surgery, Gynecology and Obstetrics* 174(3): 245-254, 1992.

Hemming AW, Cattral MS, Reed AI, et al.: Liver transplantation for hepatocellular carcinoma. *Annals of Surgery* 233(5): 652-659, 2001.

⁵⁴ Pichlmayr R, Weimann A, Oldhafer KJ, et al.: Appraisal of transplantation for malignant tumours of the liver with special reference to early stage hepatocellular carcinoma. *European Journal of Surgical Oncology* 24(1): 60-67, 1998.

⁵⁵ Yamamoto J, Iwatsuki S, Kosuge T, et al.: Should hepatomas be treated with hepatic resection or transplantation? *Cancer* 86(7): 1151-1158, 1999.

⁵⁶ Lau WY, Leung TW, Ho SK, et al.: Adjuvant intra-arterial iodine-131-labelled lipiodol for resectable hepatocellular carcinoma: a prospective randomised trial. *Lancet* 353(9155): 797-801, 1999.

with interleukin-2 and anti-CD3 activated autologous lymphocytes was found to have lengthened recurrence-free survival, but not overall survival, in 1 study.⁵⁷ [Level of evidence: 1iiD] Localized recurrences in the liver may occasionally be successfully treated by re-resection.⁵⁸

Localized Unresectable Adult Primary Liver Cancer (Selected T2, T3, and T4; N0; M0)

Patients whose tumors are localized but unresectable due to location in the liver, concomitant medical considerations (such as cirrhosis), or even limited bilateral tumors, may be candidates for chemoembolization, cryosurgery, percutaneous ethanol injection, or radiofrequency ablation for cancers smaller than 5 centimeters. Survivals equivalent to resection have been reported.⁵⁹

Clinical trials that use systemic chemotherapy, regional chemotherapy, and/or labeled or radiolabeled antibodies have demonstrated remission of unresectable hepatoma. Other approaches include embolization of the hepatic artery with gelfoam powder or muscle fragments and chemotherapy, usually adriamycin. These approaches often produce central tumor necrosis, reduction in tumor size, and relief of pain, but the benefits are usually transient. Any interference with arterial blood supply (including infusion chemotherapy) may be associated with significant morbidity and is contraindicated in the presence of portal hypertension, portal vein thrombosis, or clinical jaundice. A randomized study of chemoembolization versus conservative treatment found no survival advantage for chemoembolization.⁶⁰ This study was terminated early and was underpowered to detect any but large survival differences.

⁵⁷ Takayama T, Sekine T, Makuuchi M, et al.: Adoptive immunotherapy to lower postsurgical recurrence rates of hepatocellular carcinoma: a randomised trial. *Lancet* 356(9232): 802-807, 2000.

⁵⁸ Nakajima Y, Ko S, Kanamura T, et al.: Repeat liver resection for hepatocellular carcinoma. *Journal of the American College of Surgeons* 192(3): 339-344, 2001.
Neeleman N, Andersson R: Repeated liver resection for recurrent liver cancer. *British Journal of Surgery* 83(7): 893-901, 1996.

⁵⁹ Zhou XD, Tang ZY: Cryotherapy for primary liver cancer. *Seminars in Surgical Oncology* 14(2): 171-174, 1998.

⁶⁰ Groupe d'Etude et de Traitement du Carcinome Hépatocellulaire: A Comparison of lipiodol chemoembolization and conservative treatment for unresectable hepatocellular carcinoma. *New England Journal of Medicine* 332(19): 1256-1261, 1995.

Standard treatment options:

- Chemoembolization, cryosurgery, percutaneous ethanol injection, or radiofrequency ablation for small (<5 cm), localized, unresectable tumors.⁶¹
- For selected patients with localized unresectable hepatoma, particularly patients with fibrolamellar hepatomas, liver transplantation may offer a potentially curative treatment option.⁶²
- Chemotherapy (regional infusion of the liver): Chemotherapeutic agents may be infused with a subcutaneous portal or implantable pump via a catheter placed in the hepatic artery. Older studies that use standard agents have demonstrated responses in 15% to 30% of such cases, but newer agents and techniques (that is, biodegradable microspheres) have been evaluated in pilot trials,⁶³ as has regional chemotherapy with external-beam radiation therapy.⁶⁴ Many patients are not candidates for these approaches, which often require surgical intervention.
- Systemic chemotherapy: Durable remissions have rarely been reported, and no significant survival benefits have been conclusively demonstrated.
- Surgery, chemotherapy, and radiation therapy: These modalities may be combined in clinical trials for patients with a dominant hepatic mass and multifocal involvement with small amounts of tumor; surgical resection or cryosurgery of the mass may be followed by hepatic infusion of the remaining liver with chemotherapeutic agents alone or in combination

⁶¹ Zhou XD, Tang ZY: Cryotherapy for primary liver cancer. *Seminars in Surgical Oncology* 14(2): 171-174, 1998.

Livraghi T, Goldberg SN, Lazzaroni S, et al.: Small hepatocellular carcinoma: treatment with radio-frequency ablation versus ethanol injection. *Radiology* 210(3): 655-661, 1999.

Tanaka K, Nakamura S, Numata K, et al.: The long term efficacy of combined transcatheter arterial embolization and percutaneous ethanol injection in the treatment of patients with large hepatocellular carcinoma and cirrhosis. *Cancer* 82(1): 78-85, 1998.

Curley SA, Izzo F, Delrio P, et al.: Radiofrequency ablation of unresectable primary and metastatic hepatic malignancies: results in 123 patients. *Annals of Surgery* 230(1): 1-8, 1999.

⁶² Hemming AW, Cattral MS, Reed AI, et al.: Liver transplantation for hepatocellular carcinoma. *Annals of Surgery* 233(5): 652-659, 2001.

⁶³ Ensminger WD, Niederhuber JE, Dakhil J, et al.: Totally implanted drug delivery system for hepatic arterial chemotherapy. *Cancer Treatment Reports* 65(516): 393-400, 1981.

Dakhil S, Ensminger W, Cho K, et al.: Improved regional selectivity of hepatic arterial BCNU with degradable microspheres. *Cancer* 50(4): 631-635, 1982.

Choi BI, Kim HC, Han JK, et al.: Therapeutic effect of transcatheter oily chemoembolization therapy for encapsulated nodular hepatocellular carcinoma: CT and pathologic findings. *Radiology* 182(3): 709-713, 1992.

⁶⁴ Epstein B, Ettinger D, Leichner PK, et al.: Multimodality cisplatin treatment in nonresectable alpha-fetoprotein-positive hepatoma. *Cancer* 67(4): 896-900, 1991.

with hyperthermia, radiation, or radiation with radiosensitizers.⁶⁵ Chemotherapy plus radiation has also been used to shrink tumors prior to resection.⁶⁶

- Intratumoral injection of alcohol.⁶⁷
- Other approaches include the use of radiosensitizers and external-beam radiation therapy without chemotherapy. The relative radiosensitivity of normal liver tissue compared with tumor tissue must always be considered when radiation therapy is contemplated.⁶⁸
- Radiofrequency tissue ablation.⁶⁹

Advanced Adult Primary Liver Cancer (Any T, N1 or M1)

There is no standard therapy for patients with advanced metastatic liver cancer. Such patients should be considered candidates for clinical trials exploring the usefulness of new biologicals or antitumor drugs (phase I and II studies) or combinations of existing drugs, radiosensitizers, and radiation therapy. Palliation may sometimes be achieved in such studies. Randomized studies versus observation have shown a survival advantage for pravastatin in patients who have undergone chemoembolization;⁷⁰ [Level of evidence: 1iiA] and for megestrol acetate in patients with variant estrogen receptors.⁷¹ [Level of evidence: 1iiB]

⁶⁵ Zhou XD, Tang ZY: Cryotherapy for primary liver cancer. *Seminars in Surgical Oncology* 14(2): 171-174, 1998.

⁶⁶ Sitzmann JV, Abrams R: Improved survival for hepatocellular cancer with combination surgery and multimodality treatment. *Annals of Surgery* 217(2): 149-154, 1993.

⁶⁷ Livraghi T, Bolondi L, Lazzaroni S, et al.: Percutaneous ethanol injection in the treatment of hepatocellular carcinoma in cirrhosis: a study on 207 patients. *Cancer* 69(4): 925-929, 1992.

Livraghi T, Benedini V, Lazzaroni S, et al.: Long term results of single session percutaneous ethanol injection in patients with large hepatocellular carcinoma. *Cancer* 83(1): 48-57, 1998.

⁶⁸ Di Bisceglie AM, Rustgi VK, Hoofnagle JH, et al.: NIH conference: hepatocellular carcinoma. *Annals of Internal Medicine* 108(3): 390-401, 1988.

⁶⁹ Goldberg SN, Gazelle GS, Solbiati L, et al.: Ablation of liver tumors using percutaneous RF therapy. *American Journal of Roentgenology* 170(4): 1023-1028, 1998.

⁷⁰ Kawata S, Yamasaki E, Nagase T, et al.: Effect of pravastatin on survival in patients with advanced hepatocellular carcinoma. A randomized controlled trial. *British Journal of Cancer* 84(7): 886-891, 2001.

⁷¹ Villa E, Ferretti I, Grottola A, et al.: Hormonal therapy with megestrol in inoperable hepatocellular carcinoma characterized by variant oestrogen receptors. *British Journal of Cancer* 84(7): 881-885, 2001.

Recurrent Adult Primary Liver Cancer⁷²

The prognosis for any treated primary liver cancer patient with progressing, recurring, or relapsing disease is poor. The question and selection of further treatment depends on many factors, including prior treatment, site of recurrence, presence of cirrhosis, and hepatic function as well as individual patient considerations. Re-resection should be considered when feasible, but most patients experience recurrence, typically in the liver.⁷³ When re-resection is not possible, treatment options for patients with recurrent hepatocellular cancer may include the use of transarterial oily chemoembolization (TOCE), percutaneous ethanol injection therapy (PEIT), chemotherapy, or liver transplantation.⁷⁴ At a single institution in Hong Kong, 244 consecutive patients treated with curative resection were followed for intrahepatic recurrence. Of the 244 patients followed, 139 patients did not develop intrahepatic recurrence and had 1-, 3-, and 5-year survival rates of 87%, 79%, and 74%, respectively. Of the 105 patients who developed subsequent intrahepatic recurrences, 11 patients were treated with re-resection and had 1-, 3-, and 5-year survival rates of 81%, 70%, and 69%, respectively; 71 patients were treated with TOCE and had 1-, 3-, and 5-year survival rates of 72%, 38%, and 20%, respectively; 6 patients were treated with PEIT and had 1-, 3-, 5-year survival rates of 67%, 22%, 0%, respectively; the remaining 17 patients had either systemic chemotherapy or conservative treatment, and had no survivors at 3 years.⁷⁵ [Level of evidence: 3iiA] Clinical trials are appropriate and should be considered whenever possible.

NIH Databases

In addition to the various Institutes of Health that publish professional guidelines, the NIH has designed a number of databases for professionals.⁷⁶

⁷² Note: Some citations in the text of this section are followed by a level of evidence. The PDQ editorial boards use a formal ranking system to help the reader judge the strength of evidence linked to the reported results of a therapeutic strategy.

⁷³ Shimada M, Takenaka K, Taguchi K, et al.: Prognostic factors after repeat hepatectomy for recurrent hepatocellular carcinoma. *Annals of Surgery* 227(1): 80-85, 1998.

⁷⁴ Poon RT, Fan ST, Lo CM, et al.: Intrahepatic recurrence after curative resection of hepatocellular carcinoma: long-term results of treatment and prognostic factors. *Annals of Surgery* 229(2): 216-222, 1999.

⁷⁵ Poon RT, Fan ST, Lo CM, et al.: Intrahepatic recurrence after curative resection of hepatocellular carcinoma: long-term results of treatment and prognostic factors. *Annals of Surgery* 229(2): 216-222, 1999.

⁷⁶ Remember, for the general public, the National Library of Medicine recommends the databases referenced in MEDLINEplus (<http://medlineplus.gov/> or <http://www.nlm.nih.gov/medlineplus/databases.html>).

Physician-oriented resources provide a wide variety of information related to the biomedical and health sciences, both past and present. The format of these resources varies. Searchable databases, bibliographic citations, full text articles (when available), archival collections, and images are all available. The following are referenced by the National Library of Medicine:⁷⁷

- **Bioethics:** Access to published literature on the ethical, legal and public policy issues surrounding healthcare and biomedical research. This information is provided in conjunction with the Kennedy Institute of Ethics located at Georgetown University, Washington, D.C.:
http://www.nlm.nih.gov/databases/databases_bioethics.html
- **HIV/AIDS Resources:** Describes various links and databases dedicated to HIV/AIDS research:
<http://www.nlm.nih.gov/pubs/factsheets/aidsinfs.html>
- **NLM Online Exhibitions:** Describes “Exhibitions in the History of Medicine”: <http://www.nlm.nih.gov/exhibition/exhibition.html>.
Additional resources for historical scholarship in medicine:
<http://www.nlm.nih.gov/hmd/hmd.html>
- **Biotechnology Information:** Access to public databases. The National Center for Biotechnology Information conducts research in computational biology, develops software tools for analyzing genome data, and disseminates biomedical information for the better understanding of molecular processes affecting human health and disease: <http://www.ncbi.nlm.nih.gov/>
- **Population Information:** The National Library of Medicine provides access to worldwide coverage of population, family planning, and related health issues, including family planning technology and programs, fertility, and population law and policy:
http://www.nlm.nih.gov/databases/databases_population.html
- **Cancer Information:** Access to cancer-oriented databases:
http://www.nlm.nih.gov/databases/databases_cancer.html
- **Profiles in Science:** Offering the archival collections of prominent twentieth-century biomedical scientists to the public through modern digital technology: <http://www.profiles.nlm.nih.gov/>
- **Chemical Information:** Provides links to various chemical databases and references: <http://sis.nlm.nih.gov/Chem/ChemMain.html>
- **Clinical Alerts:** Reports the release of findings from the NIH-funded clinical trials where such release could significantly affect morbidity and mortality: http://www.nlm.nih.gov/databases/alerts/clinical_alerts.html

⁷⁷ See <http://www.nlm.nih.gov/databases/databases.html>.

- **Space Life Sciences:** Provides links and information to space-based research (including NASA):
http://www.nlm.nih.gov/databases/databases_space.html
- **MEDLINE:** Bibliographic database covering the fields of medicine, nursing, dentistry, veterinary medicine, the healthcare system, and the pre-clinical sciences:
http://www.nlm.nih.gov/databases/databases_medline.html
- **Toxicology and Environmental Health Information (TOXNET):** Databases covering toxicology and environmental health:
<http://sis.nlm.nih.gov/Tox/ToxMain.html>
- **Visible Human Interface:** Anatomically detailed, three-dimensional representations of normal male and female human bodies:
http://www.nlm.nih.gov/research/visible/visible_human.html

While all of the above references may be of interest to physicians who study and treat adult primary liver cancer, the following are particularly noteworthy.

The Combined Health Information Database

A comprehensive source of information on clinical guidelines written for professionals is the Combined Health Information Database. You will need to limit your search to "Brochure/Pamphlet," "Fact Sheet," or "Information Package" and adult primary liver cancer using the "Detailed Search" option. Go directly to the following hyperlink: <http://chid.nih.gov/detail/detail.html>. To find associations, use the drop boxes at the bottom of the search page where "You may refine your search by." For the publication date, select "All Years," select your preferred language, and the format option "Fact Sheet." By making these selections and typing "adult primary liver cancer" (or synonyms) into the "For these words:" box above, you will only receive results on fact sheets dealing with adult primary liver cancer. The following is a sample result:

- **Hepatocellular Carcinoma, Alcohol, and Cirrhosis: Facts and Hypotheses**

Source: Digestive Diseases and Sciences. 36(8): 1137-1142. August 1991.

Summary: Despite the epidemiological evidence of a correlation between ethanol abuse and hepatocellular carcinoma, some of the results of experimental and clinical studies remain controversial. This article presents the facts and hypotheses of the interrelationships among

hepatocellular carcinoma, alcohol, and cirrhosis. Topics covered include epidemiology, clinical studies, ethanol as a co-carcinogen, hepatocyte proliferation, and other mechanisms underlying ethanol's co-carcinogenic effect. The authors contend that apart from inducing cirrhosis, which may be viewed as a precancerous liver lesion, ethanol may act as a co-carcinogen. 81 references. (AA-M).

- **Successful Treatment of Bleeding Due to Ileal Varices in a Patient with Hepatocellular Carcinoma**

Source: *European Journal of Gastroenterology and Hepatology*. 13(1): 63-66. January 2001.

Contact: Available from Lippincott Williams and Wilkins. 241 Borough High Street, London SE1 1GB, UK 44(0)20-7940-7502. Fax: 44(0)20-7940-7574. Website: <http://www.eurojgh.com/>.

Summary: This case report concerns a 62 year old female who was known to have cirrhosis (liver scarring) and advanced hepatocellular carcinoma (HCC, liver cancer). An endoscopic examination showed no evidence of hemorrhaging (bleeding) due to either esophageal or gastric varices (twisted, dilated veins). Angiographic studies demonstrated extravasation (blood escaping into the tissues) from the ileal varices. There was a prominent arterio-portal shunt in the liver, and the shunt was considered to be a contributing factor to induce portal hypertension (high blood pressure in the blood supply of the liver) and variceal bleeding in the ileum. Therefore, transcatheter arterial embolization was performed, but was unsuccessful. As a result, the patient underwent a laparotomy, and a dilatating ileocecal vein and a communicating ovarian vein were selectively ligated. Following the procedure, the hemorrhaging stopped and she then recovered. The patient is doing well 21 months after surgery at the time of writing. 4 figures. 1 table. 39 references.

- **Refractory Pancreatitis Secondary to Ruptured Hepatocellular Carcinoma into the Common Bile Duct**

Source: *Digestive Diseases and Sciences*. 46(5): 1029-1033. May 2001.

Contact: Available from Kluwer Academic Publishers. Customer Service Department, P.O. Box 358, Accord Station, Hingham, MA 02018-0358. (781) 871-6600. Fax (781) 681-9045. E-mail: kluwer@wkap.com. Website: www.wkap.nl. Distribution Centre, P.O. Box 322, 3300 AH Dordrecht, The Netherlands. 31 78 6392392. Fax: 31 78 6546474. E-mail: orderdept@wkap.nl.

Summary: Hepatocellular carcinoma (HCC, liver cancer) is a common disease worldwide and continues to be the leading cause of death of

males in Taiwan (from where this article originates). Jaundice is present in 19 to 44 percent of cases of HCC at the time of diagnosis and is usually attributed to the preexisting liver cirrhosis (scarring) or extensive hepatic parenchymal (the liver body) destruction by tumor. Icteric hepatoma (a type of liver tumor) is characterized by intermittent obstructive jaundice with associated complications, such as cholangitis (inflammation of the bile ducts) and hemobilia. In this article, the authors report the first case of icteric hepatoma that initially presented as pancreatitis in addition to obstructive jaundice. The 59 year old man was admitted with a 2 week history of tea colored urine, intermittent tarry stool, vomiting, and postprandial epigastralgia (pain in the stomach after meals) with radiation to his back. He denied alcohol abuse and drugs consumption and he had never experienced pancreatitis. After 8 days of hospital treatment, the patient was released and able to eat a normal diet at an outpatient visit one month later. However, he was rehospitalized 8 weeks later with another episode of pain; surgical treatment was implemented. The patient died of multisystem organ failure on the 32nd postoperative day. For this case, the treatment was focused on two goals. First, the consequences of the biliary obstruction including the pancreatitis should be resolved by nonoperative methods, if available. Second, the origin of the migrating tumor should be eradicated either by transarterial chemoembolization or hepatic (liver) resection. The present case was not suitable for hepatic resection or hepatic artery ligation because of intrahepatic metastasis of both lobes and portal vein thrombosis (clotting) seen at exploration. Although palliation could be satisfactorily given, the prognosis continues to be dismal. 4 figures. 11 references.

- **Preventing Hepatitis B in People in Close Contact with Hepatocellular Carcinoma Patients**

Source: Public Health Reports. 112(1): 63-65. January-February 1997.

Contact: Available from Public Health Reports. Room 1855, JFK Federal Building, Boston, MA 02203. (617) 565-1440. E-mail: phr@nlm.nih.gov.

Summary: This article reports on a study to determine the prevalence of testing for hepatitis B virus (HBV) infection in the clinical management of primary liver cancer (hepatocellular carcinoma). The authors reviewed the records of 78 patients treated for hepatocellular carcinoma in hospitals in the Puget Sound area in 1988 and early 1989 and reviewed all 1990 U.S. death certificates on which primary liver cancer was listed. The records of 50 (64 percent) of 78 hepatocellular carcinoma patients contained no evidence that the patient's hepatitis B surface antigen (HBsAg) status had been determined. In addition, of 4,353 people who died in 1990 for whom the diagnosis of primary liver cancer was listed on

the death certificate, HBV infection was also listed for only 136 (3 percent), much less than expected based on case series. Many patients with hepatocellular carcinoma are not tested for HBV infection, suggesting that their close contacts are also not evaluated for HBV infection and the need for vaccination. Hepatitis B vaccination of close personal contacts of HBV-infected hepatocellular carcinoma patients is an important strategy for preventing HBV transmission. 14 references. (AA-M).

- **Worldwide Immunization Program Targets Hepatitis B and Liver Cancer**

Source: *Journal of the National Cancer Institute*. 83(10): 666-668. May 15, 1991.

Summary: This article reports on the worldwide immunization program that targets hepatitis B and liver cancer. Three doses of hepatitis B vaccine have been shown to be 95 percent effective in preventing chronic hepatitis B virus (HBV) infection, the cause of over four-fifths of the world's primary liver cancers. The article details the work of the International Task Force on Hepatitis B Immunization, an independent body of health professionals from eight countries that was created in 1986. A major accomplishment of the Task Force has been to foster more international acceptance of the hepatitis B vaccine and the concept of universal childhood immunization, even in areas with a relatively low prevalence for HBV, such as the United States. 1 figure.

- **Detection of Hepatitis B Virus (HBV) in HBsAG Negative Individuals with Primary Liver Cancer**

Source: *Digestive Diseases and Sciences*. 36(8): 1122-1129. August 1991.

Summary: The importance of chronic hepatitis B virus (HBV) infection in the development of primary liver cancer has been established by epidemiological studies. This article discusses the detection of HBV in HBsAG negative individuals with primary liver cancer. The authors report the use of the polymerase chain reaction to detect HBV DNA in the serum and liver of these patients. This technique allows both for the detection and cloning of HBV variants. The authors contend that the laboratory values obtained with this technique reinforce the role of HBV in the pathogenesis of this tumor. 5 figures. 39 references. (AA-M).

- **Impact Factors on Development of Cirrhosis and Subsequent Hepatocellular Carcinoma**

Source: *Compendium of Continuing Education in Dentistry*. 22(3): 19-33. July 2001.

Contact: Available from Dental Learning Systems. 241 Forsgate Drive, Jamesburg, NJ 08831. (800) 926-7636.

Summary: Hepatocellular carcinoma (HCC, liver cancer) on the rise in many countries, is of multifactorial etiology (caused by many factors). This article reviews the development of cirrhosis and subsequent HCC. The causes of liver cancer differ between populations at high and low risk. Africans and Chinese have the highest incidence of HCC, but other affected groups include African Americans, Japanese, and Native Americans. Chronic infection by hepatitis B and hepatitis C viruses are major risk factors worldwide, although mechanisms through which the infections cause liver cancer are yet to be explained. Other documented risk factors have been proposed and include dietary exposure, cigarette smoking, alcohol consumption, diabetes, oral infection, and oral contraceptive use. In addition, many naturally occurring and synthetic chemicals to which humans are exposed via accidental contamination of food or water have been shown to induce liver cancer in experimental animals. Consequently, assessment of possible human liver cancer risk associated with such exposures is complex. Early diagnosis and transplantation are the best treatments presently, although transplantation is not widely available due to donor shortage. The author stresses that every effort should be directed toward the prevention of HCC, through the treatment and prevention of hepatitis and oral infections, prevention of chronic hepatitis progressing to cirrhosis (liver scarring), and prevention of the cirrhotic liver from developing HCC through chemopreventive modalities. The article is designed as a continuing education curriculum for dentists and other oral health care professionals. 3 figures. 4 tables. 85 references.

The NLM Gateway⁷⁸

The NLM (National Library of Medicine) Gateway is a Web-based system that lets users search simultaneously in multiple retrieval systems at the U.S. National Library of Medicine (NLM). It allows users of NLM services to initiate searches from one Web interface, providing “one-stop searching” for many of NLM’s information resources or databases.⁷⁹ One target audience for the Gateway is the Internet user who is new to NLM’s online resources and does not know what information is available or how best to search for it.

⁷⁸ Adapted from NLM: <http://gateway.nlm.nih.gov/gw/Cmd?Overview.x>.

⁷⁹ The NLM Gateway is currently being developed by the Lister Hill National Center for Biomedical Communications (LHNCBC) at the National Library of Medicine (NLM) of the National Institutes of Health (NIH).

This audience may include physicians and other healthcare providers, researchers, librarians, students, and, increasingly, patients, their families, and the public.⁸⁰ To use the NLM Gateway, simply go to the search site at <http://gateway.nlm.nih.gov/gw/Cmd>. Type “adult primary liver cancer” (or synonyms) into the search box and click “Search.” The results will be presented in a tabular form, indicating the number of references in each database category.

Results Summary

Category	Items Found
Journal Articles	350750
Books / Periodicals / Audio Visual	2585
Consumer Health	294
Meeting Abstracts	2575
Other Collections	87
Total	356291

HSTAT⁸¹

HSTAT is a free, Web-based resource that provides access to full-text documents used in healthcare decision-making.⁸² HSTAT’s audience includes healthcare providers, health service researchers, policy makers, insurance companies, consumers, and the information professionals who serve these groups. HSTAT provides access to a wide variety of publications, including clinical practice guidelines, quick-reference guides for clinicians, consumer health brochures, evidence reports and technology assessments from the Agency for Healthcare Research and Quality (AHRQ), as well as AHRQ’s Put Prevention Into Practice.⁸³ Simply search by “adult primary

⁸⁰ Other users may find the Gateway useful for an overall search of NLM’s information resources. Some searchers may locate what they need immediately, while others will utilize the Gateway as an adjunct tool to other NLM search services such as PubMed® and MEDLINEplus®. The Gateway connects users with multiple NLM retrieval systems while also providing a search interface for its own collections. These collections include various types of information that do not logically belong in PubMed, LOCATORplus, or other established NLM retrieval systems (e.g., meeting announcements and pre-1966 journal citations). The Gateway will provide access to the information found in an increasing number of NLM retrieval systems in several phases.

⁸¹ Adapted from HSTAT: <http://www.nlm.nih.gov/pubs/factsheets/hstat.html>.

⁸² The HSTAT URL is <http://hstat.nlm.nih.gov/>.

⁸³ Other important documents in HSTAT include: the National Institutes of Health (NIH) Consensus Conference Reports and Technology Assessment Reports; the HIV/AIDS

liver cancer” (or synonyms) at the following Web site:
<http://text.nlm.nih.gov>.

Coffee Break: Tutorials for Biologists⁸⁴

Some patients may wish to have access to a general healthcare site that takes a scientific view of the news and covers recent breakthroughs in biology that may one day assist physicians in developing treatments. To this end, we recommend “Coffee Break,” a collection of short reports on recent biological discoveries. Each report incorporates interactive tutorials that demonstrate how bioinformatics tools are used as a part of the research process. Currently, all Coffee Breaks are written by NCBI staff.⁸⁵ Each report is about 400 words and is usually based on a discovery reported in one or more articles from recently published, peer-reviewed literature.⁸⁶ This site has new articles every few weeks, so it can be considered an online magazine of sorts, and intended for general background information. You can access the Coffee Break Web site at <http://www.ncbi.nlm.nih.gov/Coffeebreak/>.

Other Commercial Databases

In addition to resources maintained by official agencies, other databases exist that are commercial ventures addressing medical professionals. Here are a few examples that may interest you:

- **CliniWeb International:** Index and table of contents to selected clinical information on the Internet; see <http://www.ohsu.edu/clinweb/>.

Treatment Information Service (ATIS) resource documents; the Substance Abuse and Mental Health Services Administration’s Center for Substance Abuse Treatment (SAMHSA/CSAT) Treatment Improvement Protocols (TIP) and Center for Substance Abuse Prevention (SAMHSA/CSAP) Prevention Enhancement Protocols System (PEPS); the Public Health Service (PHS) Preventive Services Task Force’s *Guide to Clinical Preventive Services*; the independent, nonfederal Task Force on Community Services *Guide to Community Preventive Services*; and the Health Technology Advisory Committee (HTAC) of the Minnesota Health Care Commission (MHCC) health technology evaluations.

⁸⁴ Adapted from <http://www.ncbi.nlm.nih.gov/Coffeebreak/Archive/FAQ.html>.

⁸⁵ The figure that accompanies each article is frequently supplied by an expert external to NCBI, in which case the source of the figure is cited. The result is an interactive tutorial that tells a biological story.

⁸⁶ After a brief introduction that sets the work described into a broader context, the report focuses on how a molecular understanding can provide explanations of observed biology and lead to therapies for diseases. Each vignette is accompanied by a figure and hypertext links that lead to a series of pages that interactively show how NCBI tools and resources are used in the research process.

- **Image Engine:** Multimedia electronic medical record system that integrates a wide range of digitized clinical images with textual data stored in the University of Pittsburgh Medical Center's MARS electronic medical record system; see the following Web site: <http://www.cml.upmc.edu/cml/imageengine/imageEngine.html>.
- **Medical World Search:** Searches full text from thousands of selected medical sites on the Internet; see <http://www.mwsearch.com/>.
- **MedWeaver:** Prototype system that allows users to search differential diagnoses for any list of signs and symptoms, to search medical literature, and to explore relevant Web sites; see <http://www.med.virginia.edu/~wmd4n/medweaver.html>.
- **Metaphrase:** Middleware component intended for use by both caregivers and medical records personnel. It converts the informal language generally used by caregivers into terms from formal, controlled vocabularies; see the following Web site: <http://www.lexical.com/Metaphrase.html>.

The Genome Project and Adult Primary Liver Cancer

With all the discussion in the press about the Human Genome Project, it is only natural that physicians, researchers, and patients want to know about how human genes relate to adult primary liver cancer. In the following section, we will discuss databases and references used by physicians and scientists who work in this area.

Online Mendelian Inheritance in Man (OMIM)

The Online Mendelian Inheritance in Man (OMIM) database is a catalog of human genes and genetic disorders authored and edited by Dr. Victor A. McKusick and his colleagues at Johns Hopkins and elsewhere. OMIM was developed for the World Wide Web by the National Center for Biotechnology Information (NCBI).⁸⁷ The database contains textual information, pictures, and reference information. It also contains copious links to NCBI's Entrez database of MEDLINE articles and sequence information.

⁸⁷ Adapted from <http://www.ncbi.nlm.nih.gov/>. Established in 1988 as a national resource for molecular biology information, NCBI creates public databases, conducts research in computational biology, develops software tools for analyzing genome data, and disseminates biomedical information--all for the better understanding of molecular processes affecting human health and disease.

To search the database, go to <http://www.ncbi.nlm.nih.gov/Omim/searchomim.html>. Type "adult primary liver cancer" (or synonyms) in the search box, and click "Submit Search." If too many results appear, you can narrow the search by adding the word "clinical." Each report will have additional links to related research and databases. By following these links, especially the link titled "Database Links," you will be exposed to numerous specialized databases that are largely used by the scientific community. These databases are overly technical and seldom used by the general public, but offer an abundance of information. The following is an example of the results you can obtain from the OMIM for adult primary liver cancer:

- **Hepatocellular Carcinoma**
Web site: <http://www.ncbi.nlm.nih.gov/htbin-post/Omim/dispim?114550>

Genes and Disease (NCBI - Map)

The Genes and Disease database is produced by the National Center for Biotechnology Information of the National Library of Medicine at the National Institutes of Health. This Web site categorizes each disorder by the system of the body associated with it. Go to <http://www.ncbi.nlm.nih.gov/disease/>, and browse the system pages to have a full view of important conditions linked to human genes. Since this site is regularly updated, you may wish to re-visit it from time to time. The following systems and associated disorders are addressed:

- **Cancer:** Uncontrolled cell division.
Examples: Breast And Ovarian Cancer, Burkitt lymphoma, chronic myeloid leukemia, colon cancer, lung cancer, malignant melanoma, multiple endocrine neoplasia, neurofibromatosis, p53 tumor suppressor, pancreatic cancer, prostate cancer, Ras oncogene, RB: retinoblastoma, von Hippel-Lindau syndrome.
Web site: <http://www.ncbi.nlm.nih.gov/disease/Cancer.html>

Entrez

Entrez is a search and retrieval system that integrates several linked databases at the National Center for Biotechnology Information (NCBI). These databases include nucleotide sequences, protein sequences,

macromolecular structures, whole genomes, and MEDLINE through PubMed. Entrez provides access to the following databases:

- **PubMed:** Biomedical literature (PubMed),
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=PubMed>
- **Nucleotide Sequence Database (Genbank):**
Web site:
<http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=Nucleotide>
- **Protein Sequence Database:**
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=Protein>
- **Structure:** Three-dimensional macromolecular structures,
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=Structure>
- **Genome:** Complete genome assemblies,
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=Genome>
- **PopSet:** Population study data sets,
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=Popset>
- **OMIM:** Online Mendelian Inheritance in Man,
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=OMIM>
- **Taxonomy:** Organisms in GenBank,
Web site:
<http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=Taxonomy>
- **Books:** Online books,
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=books>
- **ProbeSet:** Gene Expression Omnibus (GEO),
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=geo>
- **3D Domains:** Domains from Entrez Structure,
Web site: <http://www.ncbi.nlm.nih.gov/entrez/query.fcgi?db=geo>
- **NCBI's Protein Sequence Information Survey Results:**
Web site: <http://www.ncbi.nlm.nih.gov/About/proteinsurvey/>

To access the Entrez system at the National Center for Biotechnology Information, go to <http://www.ncbi.nlm.nih.gov/entrez/>, and then select the database that you would like to search. The databases available are listed in the drop box next to "Search." In the box next to "for," enter "adult primary liver cancer" (or synonyms) and click "Go."

Jablonski's Multiple Congenital Anomaly/Mental Retardation (MCA/MR) Syndromes Database⁸⁸

This online resource can be quite useful. It has been developed to facilitate the identification and differentiation of syndromic entities. Special attention is given to the type of information that is usually limited or completely omitted in existing reference sources due to space limitations of the printed form.

At the following Web site you can also search across syndromes using an index: http://www.nlm.nih.gov/mesh/jablonski/syndrome_toc/toc_a.html. You can search by keywords at this Web site: http://www.nlm.nih.gov/mesh/jablonski/syndrome_db.html.

The Genome Database⁸⁹

Established at Johns Hopkins University in Baltimore, Maryland in 1990, the Genome Database (GDB) is the official central repository for genomic mapping data resulting from the Human Genome Initiative. In the spring of 1999, the Bioinformatics Supercomputing Centre (BiSC) at the Hospital for Sick Children in Toronto, Ontario assumed the management of GDB. The Human Genome Initiative is a worldwide research effort focusing on structural analysis of human DNA to determine the location and sequence of the estimated 100,000 human genes. In support of this project, GDB stores and curates data generated by researchers worldwide who are engaged in the mapping effort of the Human Genome Project (HGP). GDB's mission is to provide scientists with an encyclopedia of the human genome which is continually revised and updated to reflect the current state of scientific knowledge. Although GDB has historically focused on gene mapping, its focus will broaden as the Genome Project moves from mapping to sequence, and finally, to functional analysis.

To access the GDB, simply go to the following hyperlink: <http://www.gdb.org/>. Search "All Biological Data" by "Keyword." Type "adult primary liver cancer" (or synonyms) into the search box, and review the results. If more than one word is used in the search box, then separate each one with the word "and" or "or" (using "or" might be useful when using synonyms). This database is extremely technical as it was created for

⁸⁸ Adapted from the National Library of Medicine:

http://www.nlm.nih.gov/mesh/jablonski/about_syndrome.html.

⁸⁹ Adapted from the Genome Database:

<http://gdbwww.gdb.org/gdb/aboutGDB.html#mission>.

specialists. The articles are the results which are the most accessible to non-professionals and often listed under the heading "Citations." The contact names are also accessible to non-professionals.

Specialized References

The following books are specialized references written for professionals interested in adult primary liver cancer (sorted alphabetically by title, hyperlinks provide rankings, information, and reviews at Amazon.com):

- **Advanced and Critical Care Oncology Nursing: Managing Primary Complications** by Cynthia C. Chernecky (Editor), et al; Paperback - 736 pages (September 18, 1997), W B Saunders Co; ISBN: 0721668607;
<http://www.amazon.com/exec/obidos/ASIN/0721668607/icongroupinterna>
- **Cancer: Etiology, Diagnosis, and Treatment** by Walter J. Burdette; Paperback - 287 pages, 1st edition (January 15, 1998), McGraw Hill Text; ISBN: 0070089922;
<http://www.amazon.com/exec/obidos/ASIN/0070089922/icongroupinterna>
- **Cancer Management: A Multidisciplinary Approach: Medical, Surgical & Radiation** by Richard Pazdur (Editor), et al; Paperback - 982 pages, 5th edition (June 15, 2001), Publisher Research & Representation, Inc.; ISBN: 1891483080;
<http://www.amazon.com/exec/obidos/ASIN/1891483080/icongroupinterna>
- **Familial Cancer and Prevention: Molecular Epidemiology: A New Strategy Toward Cancer Control** by Joji Utsunomiya (Editor), et al; Hardcover (April 1999), Wiley-Liss; ISBN: 0471249378;
<http://www.amazon.com/exec/obidos/ASIN/0471249378/icongroupinterna>
- **Fundamentals of Cancer Epidemiology** by Philip C. Nasca, Ph.D. (Editor), Pastides Harris, Ph.D., MPH (Editor); Hardcover - 368 pages, 1st edition (February 15, 2001), Aspen Publishers, Inc.; ISBN: 0834217767;
<http://www.amazon.com/exec/obidos/ASIN/0834217767/icongroupinterna>
- **Helping Cancer Patients Cope: A Problem-Solving Approach** by Arthur M. Nezu (Editor), et al; Hardcover - 314 pages (December 15, 1998), American Psychological Association (APA); ISBN: 1557985332;
<http://www.amazon.com/exec/obidos/ASIN/1557985332/icongroupinterna>
- **Quantitative Estimation and Prediction of Human Cancer Risks (Iarc Scientific Publications, 131)** by Suresh H. Moolgavkar (Editor), et al; Paperback (September 1999), Oxford University Press; ISBN: 9283221311;
<http://www.amazon.com/exec/obidos/ASIN/9283221311/icongroupinterna>

- **Textbook of Cancer Epidemiology** by ADAMI, et al; Hardcover - 385 pages, 1st edition (July 15, 2002), Oxford University Press; ISBN: 0195109694;
<http://www.amazon.com/exec/obidos/ASIN/0195109694/icongroupinterna>

Vocabulary Builder

Angiosarcoma: A type of cancer that begins in the lining of blood vessels.

[NIH]

Autologous: Taken from an individual's own tissues, cells, or DNA. [NIH]

Cryotherapy: Any method that uses cold temperature to treat disease. [NIH]

Diaphragm: The thin muscle below the lungs and heart that separates the chest from the abdomen. [NIH]

Hypercalcemia: Abnormally high blood calcium. [NIH]

Hypoglycemia: Abnormally low blood sugar [NIH]

Ligament: A band of fibrous tissue that connects bones or cartilages, serving to support and strengthen joints. [EU]

Microspheres: Small uniformly-sized spherical particles frequently labeled with radioisotopes or various reagents acting as tags or markers. [NIH]

Necrosis: Refers to the death of living tissues. [NIH]

Perforation: 1. the act of boring or piercing through a part. 2. a hole made through a part or substance. [EU]

Peritoneum: The tissue that lines the abdominal wall and covers most of the organs in the abdomen. [NIH]

Radiosensitizers: Drugs that make tumor cells more sensitive to radiation. [NIH]

Resected: Surgical removal of part of an organ. [NIH]

Visceral: , from viscus a viscus) pertaining to a viscus. [EU]

PART III. APPENDICES

ABOUT PART III

Part III is a collection of appendices on general medical topics which may be of interest to patients with adult primary liver cancer and related conditions.

APPENDIX A. RESEARCHING YOUR MEDICATIONS

Overview

There are a number of sources available on new or existing medications which could be prescribed to patients with adult primary liver cancer. While a number of hard copy or CD-Rom resources are available to patients and physicians for research purposes, a more flexible method is to use Internet-based databases. In this chapter, we will begin with a general overview of medications. We will then proceed to outline official recommendations on how you should view your medications. You may also want to research medications that you are currently taking for other conditions as they may interact with medications for adult primary liver cancer. Research can give you information on the side effects, interactions, and limitations of prescription drugs used in the treatment of adult primary liver cancer. Broadly speaking, there are two sources of information on approved medications: public sources and private sources. We will emphasize free-to-use public sources.

Your Medications: The Basics⁹⁰

The Agency for Health Care Research and Quality has published extremely useful guidelines on how you can best participate in the medication aspects of adult primary liver cancer. Taking medicines is not always as simple as swallowing a pill. It can involve many steps and decisions each day. The AHCRCQ recommends that patients with adult primary liver cancer take part in treatment decisions. Do not be afraid to ask questions and talk about your concerns. By taking a moment to ask questions early, you may avoid problems later. Here are some points to cover each time a new medicine is prescribed:

- Ask about all parts of your treatment, including diet changes, exercise, and medicines.
- Ask about the risks and benefits of each medicine or other treatment you might receive.
- Ask how often you or your doctor will check for side effects from a given medication.

Do not hesitate to ask what is important to you about your medicines. You may want a medicine with the fewest side effects, or the fewest doses to take each day. You may care most about cost, or how the medicine might affect how you live or work. Or, you may want the medicine your doctor believes will work the best. Telling your doctor will help him or her select the best treatment for you.

Do not be afraid to “bother” your doctor with your concerns and questions about medications for adult primary liver cancer. You can also talk to a nurse or a pharmacist. They can help you better understand your treatment plan. Feel free to bring a friend or family member with you when you visit your doctor. Talking over your options with someone you trust can help you make better choices, especially if you are not feeling well. Specifically, ask your doctor the following:

- The name of the medicine and what it is supposed to do.
- How and when to take the medicine, how much to take, and for how long.
- What food, drinks, other medicines, or activities you should avoid while taking the medicine.
- What side effects the medicine may have, and what to do if they occur.

⁹⁰ This section is adapted from AHCRCQ: <http://www.ahcpr.gov/consumer/ncpiebro.htm>.

- If you can get a refill, and how often.
- About any terms or directions you do not understand.
- What to do if you miss a dose.
- If there is written information you can take home (most pharmacies have information sheets on your prescription medicines; some even offer large-print or Spanish versions).

Do not forget to tell your doctor about all the medicines you are currently taking (not just those for adult primary liver cancer). This includes prescription medicines and the medicines that you buy over the counter. Then your doctor can avoid giving you a new medicine that may not work well with the medications you take now. When talking to your doctor, you may wish to prepare a list of medicines you currently take, the reason you take them, and how you take them. Be sure to include the following information for each:

- Name of medicine
- Reason taken
- Dosage
- Time(s) of day

Also include any over-the-counter medicines, such as:

- Laxatives
- Diet pills
- Vitamins
- Cold medicine
- Aspirin or other pain, headache, or fever medicine
- Cough medicine
- Allergy relief medicine
- Antacids
- Sleeping pills
- Others (include names)

Learning More about Your Medications

Because of historical investments by various organizations and the emergence of the Internet, it has become rather simple to learn about the medications your doctor has recommended for adult primary liver cancer. One such source is the United States Pharmacopeia. In 1820, eleven physicians met in Washington, D.C. to establish the first compendium of standard drugs for the United States. They called this compendium the “U.S. Pharmacopeia (USP).” Today, the USP is a non-profit organization consisting of 800 volunteer scientists, eleven elected officials, and 400 representatives of state associations and colleges of medicine and pharmacy. The USP is located in Rockville, Maryland, and its home page is located at www.usp.org. The USP currently provides standards for over 3,700 medications. The resulting USP DI® Advice for the Patient® can be accessed through the National Library of Medicine of the National Institutes of Health. The database is partially derived from lists of federally approved medications in the Food and Drug Administration’s (FDA) Drug Approvals database.⁹¹

While the FDA database is rather large and difficult to navigate, the Pharmacopeia is both user-friendly and free to use. It covers more than 9,000 prescription and over-the-counter medications. To access this database, simply type the following hyperlink into your Web browser: <http://www.nlm.nih.gov/medlineplus/druginformation.html>. To view examples of a given medication (brand names, category, description, preparation, proper use, precautions, side effects, etc.), simply follow the hyperlinks indicated within the United States Pharmacopoeia (USP). It is important to read the disclaimer by the USP (<http://www.nlm.nih.gov/medlineplus/drugdisclaimer.html>) before using the information provided.

Of course, we as editors cannot be certain as to what medications you are taking. Therefore, we have compiled a list of medications associated with the treatment of adult primary liver cancer. Once again, due to space limitations, we only list a sample of medications and provide hyperlinks to ample documentation (e.g. typical dosage, side effects, drug-interaction risks, etc.). The following drugs have been mentioned in the Pharmacopeia and other sources as being potentially applicable to adult primary liver cancer:

⁹¹ Though cumbersome, the FDA database can be freely browsed at the following site: www.fda.gov/cder/da/da.htm.

Cisplatin

- **Systemic - U.S. Brands:** Platinol; Platinol-AQ
<http://www.nlm.nih.gov/medlineplus/druginfo/cisplatinsystemic202143.html>

Doxorubicin

- **Systemic - U.S. Brands:** Rubex
<http://www.nlm.nih.gov/medlineplus/druginfo/doxorubicinsystemic202209.html>

Etoposide

- **Systemic - U.S. Brands:** Etopophos; Toposar; VePesid
<http://www.nlm.nih.gov/medlineplus/druginfo/etoposidesystemic202234.html>

Fluorouracil

- **Systemic - U.S. Brands:** Adrucil
<http://www.nlm.nih.gov/medlineplus/druginfo/fluorouracilsystemic202245.html>

Hepatitis B Vaccine Recombinant

- **Systemic - U.S. Brands:** Engerix-B
<http://www.nlm.nih.gov/medlineplus/druginfo/hepatitisbvaccinerecombinantsy202281.html>

Commercial Databases

In addition to the medications listed in the USP above, a number of commercial sites are available by subscription to physicians and their institutions. You may be able to access these sources from your local medical library or your doctor's office.

Reuters Health Drug Database

The Reuters Health Drug Database can be searched by keyword at the hyperlink: <http://www.reutershealth.com/frame2/drug.html>.

Mosby's GenRx

Mosby's GenRx database (also available on CD-Rom and book format) covers 45,000 drug products including generics and international brands. It provides prescribing information, drug interactions, and patient information. Information can be obtained at the following hyperlink: <http://www.genrx.com/Mosby/PhyGenRx/group.html>.

Physicians Desk Reference

The Physicians Desk Reference database (also available in CD-Rom and book format) is a full-text drug database. The database is searchable by brand name, generic name or by indication. It features multiple drug interactions reports. Information can be obtained at the following hyperlink: http://physician.pdr.net/physician/templates/en/acl/psuser_t.htm.

Other Web Sites

A number of additional Web sites discuss drug information. As an example, you may like to look at www.drugs.com which reproduces the information in the Pharmacopeia as well as commercial information. You may also want to consider the Web site of the Medical Letter, Inc. which allows users to download articles on various drugs and therapeutics for a nominal fee: <http://www.medletter.com/>.

Drug Development and Approval

The following Web sites can be valuable resources when conducting research on the development and approval of new cancer drugs:

- FDA Home Page: Search for drugs currently in development or those which have been recently approved by the FDA.
<http://redir.nci.nih.gov/cgi-bin/redir.pl?section=Cancerinfo&destURI=http://www.fda.gov/>
- Cancer Liaison Program: Answers questions from the public about drug approval processes, cancer clinical trials, and access to investigational therapies.
<http://redir.nci.nih.gov/cgi-bin/redir.pl?section=Cancerinfo&destURI=http://www.fda.gov/oashi/cancer/cancer.html>

- Center for Drug Evaluation and Research
<http://redir.nci.nih.gov/cgi-bin/redir.pl?section=Cancerinfo&destURI=http://www.fda.gov/cder/>
- Drug Approvals by Cancer Indications (Alphabetical List)
<http://redir.nci.nih.gov/cgi-bin/redir.pl?section=Cancerinfo&destURI=http://www.fda.gov/oashi/cancer/cdrugalpha.html>
- Drug Approvals by Cancer Indications (Cancer Type)
<http://redir.nci.nih.gov/cgi-bin/redir.pl?section=Cancerinfo&destURI=http://www.fda.gov/oashi/cancer/cdrugind.html>
- Electronic Orange Book of Approved Drug Products
<http://redir.nci.nih.gov/cgi-bin/redir.pl?section=Cancerinfo&destURI=http://www.fda.gov/cder/ob/default.htm>
- Guidance Documents for Industry: Contains an archive of documents describing FDA policies on specific topics.
<http://redir.nci.nih.gov/cgi-bin/redir.pl?section=Cancerinfo&destURI=http://www.fda.gov/cder/guidance/index.htm>
- Industry Collaboration: Provides information to industry on the process for getting new drugs into clinical trials.
<http://ctep.cancer.gov/industry/index.html>
- Investigator's Handbook: Provides information to investigators on specific procedures related to clinical trial development.
<http://ctep.cancer.gov/handbook/index.html>
- Questions and Answers About NCI's Natural Products Branch: A fact sheet that describes the functions of this branch, which collects and analyzes specimens of plant, marine, and microbial origin for possible anticancer properties.
http://cis.nci.nih.gov/fact/7_33.htm

Understanding the Approval Process for New Cancer Drugs⁹²

Since June 1996, about 80 new cancer-related drugs, or new uses for drugs already on the market, have been approved by the U.S. Food and Drug

⁹² Adapted from the NCI:

http://www.cancer.gov/clinical_trials/doc_header.aspx?viewid=d94cbfac-e478-4704-9052-d8e8a3372b56.

Administration (FDA), the division of the U.S. Department of Health and Human Services charged with ensuring the safety and effectiveness of new drugs before they can go on the market. (The FDA maintains an annotated online list of drugs approved for use with cancer since 1996.) Some of these drugs treat cancer, some alleviate pain and other symptoms, and, in one case, reduce the risk of invasive cancer in people who are considered high-risk. The FDA relied on the results of clinical trials in making every one of these approvals. Without reliable information about a drug's effects on humans, it would be impossible to approve any drug for widespread use.

When considering a new drug, the FDA faces two challenges:

- First, making sure that the drug is safe and effective before it is made widely available;
- Second, ensuring that drugs which show promise are made available as quickly as possible to the people they can help.

To deal with these challenges, the FDA maintains a rigorous review process but also has measures in place to make some drugs available in special cases. This aim of this section is to acquaint you with the drug approval process and point you to other resources for learning more about it.

The Role of the Federal Drug Administration (FDA)

Approval is only one step in the drug development process. In fact, the FDA estimates that, on average, it takes eight and a half years to study and test a new drug before it can be approved for the general public. That includes early laboratory and animal testing, as well as the clinical trials that evaluate the drugs in humans. The FDA plays a key role at three main points in this process:

- Determining whether or not a new drug shows enough promise to be given to people in clinical trials
- Once clinical trials begin, deciding whether or not they should continue, based on reports of efficacy and adverse reactions
- When clinical trials are completed, deciding whether or not the drug can be sold to the public and what its label should say about directions for use, side effects, warnings, and the like.

To make these decisions, the FDA must review studies submitted by the drug's sponsor (usually the manufacturer), evaluate any adverse reports from preclinical studies and clinical trials (that is, reports of side effects or

complications), and review the adequacy of the chemistry and manufacturing. This process is lengthy, but it is meant to ensure that only beneficial drugs with acceptable side effects will make their way into the hands of the public. At the same time, recent legislative mandates and streamlined procedures within the FDA have accelerated the approval of effective drugs, especially for serious illnesses such as cancer. In addition, specific provisions make some drugs available to patients with special needs even before the approval process is complete.

From Lab to Patient Care

By law, the Food and Drug Administration (FDA) must review all test results for new drugs to ensure that products are safe and effective for specific uses. “Safe” does not mean that the drug is free of possible adverse side effects; rather, it means that the potential benefits have been determined to outweigh any risks. The testing process begins long before the first person takes the drug, with preliminary research and animal testing.

If a drug proves promising in the lab, the drug company or sponsor must apply for FDA approval to test it in clinical trials involving people. For drugs, the application, called an Investigational New Drug (IND) Application, is sent through the Center for Drug Evaluation and Research’s (CDER) IND Review Process; for biological agents, the IND is sent to the Center for Biologics Evaluation and Research (CBER). Once the IND is approved by CDER or CBER, clinical trials can begin.

If the drug makes it through the clinical trials process—that is, the studies show that it is superior to current drugs—the manufacturer must submit a New Drug Application (NDA) or (for biological agents) a Biologics License Application (BLA) to the FDA. (Biological agents, such as serums, vaccines, and cloned proteins, are manufactured from substances taken from living humans or animals.) This application must include:

- The exact chemical makeup of the drug or biologic and the mechanisms by which it is effective
- Results of animal studies
- Results of clinical trials
- How the drug or biologic is manufactured, processed, and packaged
- Quality control standards
- Samples of the product in the form(s) in which it is to be administered.

Once the FDA receives the NDA or BLA from the manufacturer or developer, the formal New Drug Application Review Process or Biologics/Product License Application Review Process begins.

For an overview of the entire process from start to finish, see the CDER's visual representation of The New Drug Development Process: Steps from Test Tube to New Drug Application Review, which is available for public viewing at the following Web address: <http://www.fda.gov/cder/handbook/develop.htm>.

Speed versus Safety in the Approval Process

The FDA's current goal is that no more than ten months will pass between the time that a complete application is submitted and the FDA takes action on it. But the process is not always smooth. Sometimes FDA's external advisory panels call for additional research or data. In other cases, the FDA staff asks for more information or revised studies. Some new drug approvals have taken as little as 42 days; other more difficult NDAs have spent years in the approval process.

Setting Priorities

The order in which NDAs are assessed by the FDA is determined by a classification system designed to give priority to drugs with the greatest potential benefits. All drugs that offer significant medical advances over existing therapies for any disease are considered "priority" drugs in the approval process. NDAs for cancer treatment drugs are reviewed for this status primarily by the Division of Oncology Drug Products in the FDA's Center for Drug Evaluation and Research (CDER). For Biologic License Applications (vaccines, blood products, and medicines made from animal products), the Center for Biologics Evaluation and Research (CBER) provides additional regulation and oversight.

Expert Advice

The FDA relies on a system of independent advisory committees, made up of professionals from outside the agency, for expert advice and guidance in making sound decisions about drug approval. Each committee meets as needed to weigh available evidence and assess the safety, effectiveness, and appropriate use of products considered for approval. In addition, these

committees provide advice about general criteria for evaluation and scientific issues not related to specific products. The Oncologic Drugs Advisory Committee (ODAC) meets regularly to provide expert advice on cancer-related treatments and preventive drugs.

Each committee is composed of representatives from the research science and medical fields. At least one member on every advisory committee must represent the consumer perspective.

Final Approval

As the FDA looks at all the data submitted and the results of its own review, it applies two benchmark questions to each application for drug approval:

- Do the results of well-controlled studies provide substantial evidence of effectiveness?
- Do the results show the product is safe under the conditions of use in the proposed labeling? In this context, “safe” means that potential benefits have been determined to outweigh any risks.

Continued Vigilance

The FDA’s responsibility for new drug treatments does not stop with final approval. The Office of Compliance in the Center for Drug Evaluation and Research (CDER) implements and tracks programs to make sure manufacturers comply with current standards and practice regulations. CDER’s Office of Drug Marketing, Advertising, and Communication monitors new drug advertising to make sure it is truthful and complete. At the Center for Biologic Evaluation and Research, biologics are followed with the same vigilance after approval. And through a system called MedWatch, the FDA gets feedback from health professionals and consumers on how the new drugs are working, any adverse reactions, and potential problems in labeling and dosage.

Online FDA Resources

The following information from the FDA should help you better understand the drug approval process:

- Center for Drug Evaluation and Research:
<http://www.fda.gov/cder/handbook>

- From Test Tube to Patient: New Drug Development in the U.S. – a special January 1995 issue of the magazine FDA Consumer:
http://www.fda.gov/fdac/special/newdrug/ndd_toc.html
- Milestones in U.S. Food and Drug Law History:
<http://www.fda.gov/opacom/backgrounders/miles.html>
- Drug Approvals for Cancer Indications:
<http://www.fda.gov/oashi/cancer/cdrug.html>

Getting Drugs to Patients Who Need Them

Clinical trials provide the most important information used by the FDA in determining whether a new drug shows “substantial evidence of effectiveness,” or whether an already-approved drug can be used effectively in new ways (for example, to treat or prevent other types of cancer, or at a different dosage). The FDA must certify that a drug has shown promise in laboratory and animal trials before human testing can begin. The trials process includes three main stages and involves continuous review, which ensures that the sponsor can stop the study early if major problems develop or unexpected levels of treatment benefit are found. As with all clinical trials, benefits and risks must be carefully weighed by the researchers conducting the study and the patients who decide to participate.

Not everyone is eligible to participate in a clinical trial. Some patients do not fit the exact requirements for studies, some have rare forms of cancer for which only a limited number of studies are underway, and others are too ill to participate. Working with the NCI and other sponsors, the FDA has established special conditions under which a patient and his or her physician can apply to receive cancer drugs that have not yet been through the approval process. In the past, these special case applications for new drugs were grouped under the name “compassionate uses.” More recently, such uses have expanded to include more patients and more categories of investigational drugs.

Access to Investigational Drugs

The process of new drug development has many parts. In the United States, until a drug has been approved by the FDA, it can generally be obtained only through several mechanisms: enrollment in a clinical trial studying the drug, an expanded access program or special exemption/compassionate use programs. For more information about investigational drugs, see “Questions

and Answers: Access to Investigational Drugs” at http://www.cancer.gov/cancer_information/doc_img.aspx?viewid=74b62d84-e135-451f-9bc9-d54358ede947.

“Group C” Drugs

In the 1970s, researchers from the NCI became concerned about the lag between the date when an investigational drug was found to have anti-tumor activity and the time that drug became available on the market. Working with the FDA, the NCI established the “Group C” classification to allow access to drugs with reproducible activity. Group C drugs are provided to properly trained physicians who have registered using a special form to assure that their patient qualifies under guideline protocols for the drug. Each Group C drug protocol specifies patient eligibility, reporting methodology, and drug use. Not only does Group C designation (now called Group C/Treatment INDs) speed new drugs to patients who need them most, but the process also allows the NCI to gather important information on the safety as well as activity of the drugs in the settings in which they will be most used after final FDA approval. Drugs are placed in the Group C category by agreement between the FDA and the NCI. Group C drugs are always provided free of charge, and the Health Care Financing Administration provides coverage for care associated with Group C therapy.

Treatment INDs

In 1987, the FDA began authorizing the use of new drugs still in the development process to treat certain seriously ill patients. In these cases, the process is referred to as a treatment investigational new drug application (Treatment IND). Clinical trials of the new drug must already be underway and have demonstrated positive results that are reproducible. The FDA sets guidelines about what constitutes serious and life-threatening illnesses, how much must already be known about a drug’s side effects and benefits, and where physicians can obtain the drug for treatment. For many seriously ill patients, the risks associated with taking a not-yet-completely proven drug are outweighed by the possible benefits.

Accelerated Approval

“Accelerated approval” is the short-hand term for the FDA’s new review system which, in the 1990s, has been used to ensure rapid approval while at

the same time putting new safeguards into place. Accelerated approval is based on “surrogate endpoint” judgments: FDA can grant marketing approval to drugs and treatments that, according to certain indicators, prove they are likely to have beneficial effects on a disease or condition, even before such direct benefits have been shown clinically. Accelerated approval does NOT mean that additional clinical trials are not needed or that FDA stops gathering information about the effects of the drug; a follow-up study is required to demonstrate activity by more conventional endpoints.

Contraindications and Interactions (Hidden Dangers)

Some of the medications mentioned in the previous discussions can be problematic for patients with adult primary liver cancer--not because they are used in the treatment process, but because of contraindications, or side effects. Medications with contraindications are those that could react with drugs used to treat adult primary liver cancer or potentially create deleterious side effects in patients with adult primary liver cancer. You should ask your physician about any contraindications, especially as these might apply to other medications that you may be taking for common ailments.

Drug-drug interactions occur when two or more drugs react with each other. This drug-drug interaction may cause you to experience an unexpected side effect. Drug interactions may make your medications less effective, cause unexpected side effects, or increase the action of a particular drug. Some drug interactions can even be harmful to you.

Be sure to read the label every time you use a nonprescription or prescription drug, and take the time to learn about drug interactions. These precautions may be critical to your health. You can reduce the risk of potentially harmful drug interactions and side effects with a little bit of knowledge and common sense.

Drug labels contain important information about ingredients, uses, warnings, and directions which you should take the time to read and understand. Labels also include warnings about possible drug interactions. Further, drug labels may change as new information becomes available. This is why it's especially important to read the label every time you use a medication. When your doctor prescribes a new drug, discuss all over-the-counter and prescription medications, dietary supplements, vitamins, botanicals, minerals and herbals you take as well as the foods you eat. Ask your pharmacist for the package insert for each prescription drug you take.

The package insert provides more information about potential drug interactions.

A Final Warning

At some point, you may hear of alternative medications from friends, relatives, or in the news media. Advertisements may suggest that certain alternative drugs can produce positive results for patients with adult primary liver cancer. Exercise caution--some of these drugs may have fraudulent claims, and others may actually hurt you. The Food and Drug Administration (FDA) is the official U.S. agency charged with discovering which medications are likely to improve the health of patients with adult primary liver cancer. The FDA warns patients to watch out for⁹³:

- Secret formulas (real scientists share what they know)
- Amazing breakthroughs or miracle cures (real breakthroughs don't happen very often; when they do, real scientists do not call them amazing or miracles)
- Quick, painless, or guaranteed cures
- If it sounds too good to be true, it probably isn't true.

If you have any questions about any kind of medical treatment, the FDA may have an office near you. Look for their number in the blue pages of the phone book. You can also contact the FDA through its toll-free number, 1-888-INFO-FDA (1-888-463-6332), or on the World Wide Web at www.fda.gov.

General References

In addition to the resources provided earlier in this chapter, the following general references describe medications (sorted alphabetically by title; hyperlinks provide rankings, information and reviews at Amazon.com):

- **Antifolate Drugs in Cancer Therapy (Cancer Drug Discovery and Development)** by Ann L. Jackman (Editor); Hardcover: 480 pages; (March 1999), Humana Press; ISBN: 0896035964;
<http://www.amazon.com/exec/obidos/ASIN/0896035964/icongroupinterna>
- **Consumers Guide to Cancer Drugs** by Gail M. Wilkes, et al; Paperback - 448 pages, 1st edition (January 15, 2000), Jones & Bartlett Publishing; ISBN:

⁹³ This section has been adapted from <http://www.fda.gov/opacom/lowlit/medfraud.html>.

0763711705;

<http://www.amazon.com/exec/obidos/ASIN/0763711705/icongroupinterna>

- **Patient Education Guide to Oncology Drugs (Book with CD-ROM)** by Gail M. Wilkes, et al; CD-ROM - 447 pages, 1st edition (January 15, 2000), Jones & Bartlett Publishing; ISBN: 076371173X;
<http://www.amazon.com/exec/obidos/ASIN/076371173X/icongroupinterna>
- **The Role of Multiple Intensification in Medical Oncology** by M. S. Apro (Editor), D. Maraninchi (Editor); Hardcover (June 1998), Springer Verlag; ISBN: 3540635432;
<http://www.amazon.com/exec/obidos/ASIN/3540635432/icongroupinterna>

Vocabulary Builder

The following vocabulary builder gives definitions of words used in this chapter that have not been defined in previous chapters:

Aspirin: A drug that reduces pain, fever, inflammation, and blood clotting. Aspirin belongs to the family of drugs called nonsteroidal anti-inflammatory agents. It is also being studied in cancer prevention. [NIH]

Etoposide: An anticancer drug that is a podophyllotoxin derivative and belongs to the family of drugs called mitotic inhibitors. [NIH]

Preclinical: Before a disease becomes clinically recognizable. [EU]

APPENDIX B. RESEARCHING ALTERNATIVE MEDICINE

Overview⁹⁴

Research indicates that the use of complementary and alternative therapies is increasing. A large-scale study published in the November 11, 1998, issue of the *Journal of the American Medical Association* found that CAM use among the general public increased from 34 percent in 1990 to 42 percent in 1997.

Several surveys of CAM use by cancer patients have been conducted with small numbers of patients. One study published in the February 2000 issue of the journal *Cancer* reported that 37 percent of 46 patients with prostate cancer used one or more CAM therapies as part of their cancer treatment. These therapies included herbal remedies, old-time remedies, vitamins, and special diets. A larger study of CAM use in patients with different types of cancer was published in the July 2000 issue of the *Journal of Clinical Oncology*. That study found that 83 percent of 453 cancer patients had used at least one CAM therapy as part of their cancer treatment. The study included CAM therapies such as special diets, psychotherapy, spiritual practices, and vitamin supplements. When psychotherapy and spiritual practices were excluded, 69 percent of patients had used at least one CAM therapy in their cancer treatment.

In this chapter, we will begin by giving you a broad perspective on complementary and alternative therapies. Next, we will introduce you to official information sources on CAM relating to adult primary liver cancer. Finally, at the conclusion of this chapter, we will provide a list of readings on adult primary liver cancer from various authors. We will begin, however,

⁹⁴Adapted from the NCI: http://cis.nci.nih.gov/fact/9_14.htm.

with the National Center for Complementary and Alternative Medicine's (NCCAM) overview of complementary and alternative medicine.

What Is CAM?⁹⁵

Complementary and alternative medicine (CAM) covers a broad range of healing philosophies, approaches, and therapies. Generally, it is defined as those treatments and healthcare practices which are not taught in medical schools, used in hospitals, or reimbursed by medical insurance companies. Many CAM therapies are termed "holistic," which generally means that the healthcare practitioner considers the whole person, including physical, mental, emotional, and spiritual health. Some of these therapies are also known as "preventive," which means that the practitioner educates and treats the person to prevent health problems from arising, rather than treating symptoms after problems have occurred.

People use CAM treatments and therapies in a variety of ways. Therapies are used alone (often referred to as alternative), in combination with other alternative therapies, or in addition to conventional treatment (sometimes referred to as complementary). Complementary and alternative medicine, or "integrative medicine," includes a broad range of healing philosophies, approaches, and therapies. Some approaches are consistent with physiological principles of Western medicine, while others constitute healing systems with non-Western origins. While some therapies are far outside the realm of accepted Western medical theory and practice, others are becoming established in mainstream medicine.

Complementary and alternative therapies are used in an effort to prevent illness, reduce stress, prevent or reduce side effects and symptoms, or control or cure disease. Some commonly used methods of complementary or alternative therapy include mind/body control interventions such as visualization and relaxation, manual healing including acupuncture and massage, homeopathy, vitamins or herbal products, and acupuncture.

Should you wish to explore non-traditional types of treatment, be sure to discuss all issues concerning treatments and therapies with your healthcare provider, whether a physician or practitioner of complementary and alternative medicine. Competent healthcare management requires knowledge of both conventional and alternative therapies you are taking for the practitioner to have a complete picture of your treatment plan.

⁹⁵ Adapted from the NCCAM: <http://nccam.nih.gov/nccam/fcp/faq/index.html#what-is>.

The decision to use complementary and alternative treatments is an important one. Consider before selecting an alternative therapy, the safety and effectiveness of the therapy or treatment, the expertise and qualifications of the healthcare practitioner, and the quality of delivery. These topics should be considered when selecting any practitioner or therapy.

What Are the Domains of Alternative Medicine?⁹⁶

The list of CAM practices changes continually. The reason being is that these new practices and therapies are often proved to be safe and effective, and therefore become generally accepted as “mainstream” healthcare practices. Today, CAM practices may be grouped within five major domains: (1) alternative medical systems, (2) mind-body interventions, (3) biologically-based treatments, (4) manipulative and body-based methods, and (5) energy therapies. The individual systems and treatments comprising these categories are too numerous to list in this sourcebook. Thus, only limited examples are provided within each.

Alternative Medical Systems

Alternative medical systems involve complete systems of theory and practice that have evolved independent of, and often prior to, conventional biomedical approaches. Many are traditional systems of medicine that are practiced by individual cultures throughout the world, including a number of venerable Asian approaches.

Traditional oriental medicine emphasizes the balance or disturbances of qi (pronounced chi) or vital energy in health and disease, respectively. Traditional oriental medicine consists of a group of techniques and methods including acupuncture, herbal medicine, oriental massage, and qi gong (a form of energy therapy). Acupuncture involves stimulating specific anatomic points in the body for therapeutic purposes, usually by puncturing the skin with a thin needle.

Ayurveda is India’s traditional system of medicine. Ayurvedic medicine (meaning “science of life”) is a comprehensive system of medicine that places equal emphasis on body, mind, and spirit. Ayurveda strives to restore the innate harmony of the individual. Some of the primary Ayurvedic

⁹⁶ Adapted from the NCCAM: <http://nccam.nih.gov/nccam/fcp/classify/index.html>.

treatments include diet, exercise, meditation, herbs, massage, exposure to sunlight, and controlled breathing.

Other traditional healing systems have been developed by the world's indigenous populations. These populations include Native American, Aboriginal, African, Middle Eastern, Tibetan, and Central and South American cultures. Homeopathy and naturopathy are also examples of complete alternative medicine systems.

Homeopathic medicine is an unconventional Western system that is based on the principle that "like cures like," i.e., that the same substance that in large doses produces the symptoms of an illness, in very minute doses cures it. Homeopathic health practitioners believe that the more dilute the remedy, the greater its potency. Therefore, they use small doses of specially prepared plant extracts and minerals to stimulate the body's defense mechanisms and healing processes in order to treat illness.

Naturopathic medicine is based on the theory that disease is a manifestation of alterations in the processes by which the body naturally heals itself and emphasizes health restoration rather than disease treatment. Naturopathic physicians employ an array of healing practices, including the following: diet and clinical nutrition, homeopathy, acupuncture, herbal medicine, hydrotherapy (the use of water in a range of temperatures and methods of applications), spinal and soft-tissue manipulation, physical therapies (such as those involving electrical currents, ultrasound, and light), therapeutic counseling, and pharmacology.

Mind-Body Interventions

Mind-body interventions employ a variety of techniques designed to facilitate the mind's capacity to affect bodily function and symptoms. Only a select group of mind-body interventions having well-documented theoretical foundations are considered CAM. For example, patient education and cognitive-behavioral approaches are now considered "mainstream." On the other hand, complementary and alternative medicine includes meditation, certain uses of hypnosis, dance, music, and art therapy, as well as prayer and mental healing.

Biological-Based Therapies

This category of CAM includes natural and biological-based practices, interventions, and products, many of which overlap with conventional medicine's use of dietary supplements. This category includes herbal, special dietary, orthomolecular, and individual biological therapies.

Herbal therapy employs an individual herb or a mixture of herbs for healing purposes. An herb is a plant or plant part that produces and contains chemical substances that act upon the body. Special diet therapies, such as those proposed by Drs. Atkins, Ornish, Pritikin, and Weil, are believed to prevent and/or control illness as well as promote health. Orthomolecular therapies aim to treat disease with varying concentrations of chemicals such as magnesium, melatonin, and mega-doses of vitamins. Biological therapies include, for example, the use of laetrile and shark cartilage to treat cancer and the use of bee pollen to treat autoimmune and inflammatory diseases.

Manipulative and Body-Based Methods

This category includes methods that are based on manipulation and/or movement of the body. For example, chiropractors focus on the relationship between structure and function, primarily pertaining to the spine, and how that relationship affects the preservation and restoration of health. Chiropractors use manipulative therapy as an integral treatment tool.

In contrast, osteopaths place particular emphasis on the musculoskeletal system and practice osteopathic manipulation. Osteopaths believe that all of the body's systems work together and that disturbances in one system may have an impact upon function elsewhere in the body. Massage therapists manipulate the soft tissues of the body to normalize those tissues.

Energy Therapies

Energy therapies focus on energy fields originating within the body (biofields) or those from other sources (electromagnetic fields). Biofield therapies are intended to affect energy fields (the existence of which is not yet experimentally proven) that surround and penetrate the human body. Some forms of energy therapy manipulate biofields by applying pressure and/or manipulating the body by placing the hands in or through these fields. Examples include Qi gong, Reiki and Therapeutic Touch.

Qi gong is a component of traditional oriental medicine that combines movement, meditation, and regulation of breathing to enhance the flow of vital energy (qi) in the body, improve blood circulation, and enhance immune function. Reiki, the Japanese word representing Universal Life Energy, is based on the belief that, by channeling spiritual energy through the practitioner, the spirit is healed and, in turn, heals the physical body. Therapeutic Touch is derived from the ancient technique of "laying-on of hands." It is based on the premises that the therapist's healing force affects the patient's recovery and that healing is promoted when the body's energies are in balance. By passing their hands over the patient, these healers identify energy imbalances.

Bioelectromagnetic-based therapies involve the unconventional use of electromagnetic fields to treat illnesses or manage pain. These therapies are often used to treat asthma, cancer, and migraine headaches. Types of electromagnetic fields which are manipulated in these therapies include pulsed fields, magnetic fields, and alternating current or direct current fields.

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Several surveys of CAM use by cancer patients have been conducted with small numbers of patients. One study published in the February 2000 issue of the journal *Cancer* reported that 37 percent of 46 patients with prostate cancer used one or more CAM therapies as part of their cancer treatment. These therapies included herbal remedies, old-time remedies, vitamins, and special diets. A larger study of CAM use in patients with different types of cancer was published in the July 2000 issue of the *Journal of Clinical Oncology*. That study found that 83 percent of 453 cancer patients had used at least one CAM therapy as part of their cancer treatment. The study included CAM therapies such as special diets, psychotherapy, spiritual practices, and vitamin supplements. When psychotherapy and spiritual practices were excluded, 69 percent of patients had used at least one CAM therapy in their cancer treatment.

How Are Complementary and Alternative Approaches Evaluated?⁹⁷

It is important that the same scientific evaluation which is used to assess conventional approaches be used to evaluate complementary and alternative therapies. A number of medical centers are evaluating complementary and alternative therapies by developing clinical trials (research studies with people) to test them.

Conventional approaches to cancer treatment have generally been studied for safety and effectiveness through a rigorous scientific process, including clinical trials with large numbers of patients. Often, less is known about the safety and effectiveness of complementary and alternative methods. Some of these complementary and alternative therapies have not undergone rigorous evaluation. Others, once considered unorthodox, are finding a place in cancer treatment—not as cures, but as complementary therapies that may help patients feel better and recover faster. One example is acupuncture. According to a panel of experts at a National Institutes of Health (NIH) Consensus Conference in November 1997, acupuncture has been found to be effective in the management of chemotherapy-associated nausea and vomiting and in controlling pain associated with surgery. Some approaches, such as laetrile, have been studied and found ineffective or potentially harmful.

NCI-Sponsored Clinical Trials in Complementary and Alternative Medicine

The NCI is currently sponsoring several clinical trials (research studies with patients) that study complementary and alternative treatments for cancer. Current trials include enzyme therapy with nutritional support for the treatment of inoperable pancreatic cancer, shark cartilage therapy for the treatment of non-small cell lung cancer, and studies of the effects of diet on prostate and breast cancers. Some of these trials compare alternative therapies with conventional treatments, while others study the effects of complementary approaches used in addition to conventional treatments. Patients who are interested in taking part in these or any clinical trials should talk with their doctor.

More information about clinical trials sponsored by the NCI can be obtained from NCCAM (<http://nccam.nih.gov>, 1-888-644-6226), OCCAM

⁹⁷Adapted from the NCI: http://cis.nci.nih.gov/fact/9_14.htm

(<http://occam.nci.nih.gov>), and the NCI's Cancer Information Service (CIS) (<http://cis.nci.nih.gov>, 1-800-4-CANCER).

Questions to Ask Your Healthcare Provider about CAM

When considering complementary and alternative therapies, ask your healthcare provider the following questions:

- What benefits can be expected from this therapy?
- What are the risks associated with this therapy?
- Do the known benefits outweigh the risks?
- What side effects can be expected?
- Will the therapy interfere with conventional treatment?
- Is this therapy part of a clinical trial? If so, who is sponsoring the trial?
- Will the therapy be covered by health insurance?
- How can patients and their health care providers learn more about complementary and alternative therapies?

Finding CAM References on Adult Primary Liver Cancer

Having read the previous discussion, you may be wondering which complementary or alternative treatments might be appropriate for adult primary liver cancer. For the remainder of this chapter, we will direct you to a number of official sources which can assist you in researching studies and publications. Some of these articles are rather technical, so some patience may be required.

National Center for Complementary and Alternative Medicine

The National Center for Complementary and Alternative Medicine (NCCAM) of the National Institutes of Health (<http://nccam.nih.gov>) has created a link to the National Library of Medicine's databases to allow patients to search for articles that specifically relate to adult primary liver cancer and complementary medicine. To search the database, go to the following Web site: www.nlm.nih.gov/nccam/camonpubmed.html. Select "CAM on PubMed." Enter "adult primary liver cancer" (or synonyms) into the search box. Click "Go." The following references provide information on

particular aspects of complementary and alternative medicine (CAM) that are related to adult primary liver cancer:

- **5-Fluorouracil plus high dose levofolinic acid and oral hydroxyurea for the treatment of primary hepatocellular carcinomas: results of a phase II multicenter study of the Southern Italy Oncology Group (G.O.I.M.).**
 Author(s): Gebbia V, Maiello E, Serravezza G, Giotta F, Testa A, Borsellino N, Pezzella G, Colucci G.
 Source: *Anticancer Res.* 1999 March-April; 19(2B): 1407-10.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=10365114&dopt=Abstract
- **A preliminary report on the intervention trials of primary liver cancer in high-risk populations with nutritional supplementation of selenium in China.**
 Author(s): Yu SY, Zhu YJ, Li WG, Huang QS, Huang CZ, Zhang QN, Hou C.
 Source: *Biological Trace Element Research.* 1991 June; 29(3): 289-94.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=1726411&dopt=Abstract
- **Adriamycin in the treatment of resectible and irresectible primary hepatocellular carcinoma.**
 Author(s): Oon CJ, Chua EJ, Foong WC, Tan LK, Yo SL, Chang CH, Ho ST, Seah CS.
 Source: *Ann Acad Med Singapore.* 1980 April; 9(2): 256-9.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=6252816&dopt=Abstract
- **Analytical study of the clinical response to two distinct adoptive immunotherapies for advanced hepatocellular carcinoma: comparison between LAK cell and CTL therapy.**
 Author(s): Haruta I, Yamauchi K, Aruga A, Komatsu T, Takasaki K, Hayashi N, Hanyu F.
 Source: *J Immunother Emphasis Tumor Immunol.* 1996 May; 19(3): 218-23.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=8811496&dopt=Abstract
- **Clinical and chemotherapeutic study of hepatocellular carcinoma in Malaysia: a comparison with African and American patients.**
 Author(s): Joishy SK, Bennett JM, Balasegaram M, MacIntyre JM, Falkson G, Moertel C, Carbone PP.

Source: *Cancer*. 1982 September 15; 50(6): 1065-9.

http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=6286085&dopt=Abstract

- **Combined effects of hepatic arterial embolization using degradable starch microspheres (DSM) in hyperthermia for liver cancer.**

Author(s): Akuta K, Abe M, Kondo M, Yoshikawa T, Tanaka Y, Yoshida M, Miura T, Nakao N, Onoyama Y, Yamada T, et al.

Source: *International Journal of Hyperthermia : the Official Journal of European Society for Hyperthermic Oncology, North American Hyperthermia Group*. 1991 March-April; 7(2): 231-42.

http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=1715374&dopt=Abstract

- **Combined traditional Chinese medicine and Western medicine. Relieving effects of Chinese herbs, ear-acupuncture and epidural morphine on postoperative pain in liver cancer.**

Author(s): Li QS, Cao SH, Xie GM, Gan YH, Ma HJ, Lu JZ, Zhang ZH.

Source: *Chin Med J (Engl)*. 1994 April; 107(4): 289-94.

http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=8088198&dopt=Abstract

- **Complete remission of unresectable hepatocellular carcinoma on healthy liver by the combination of aggressive surgery and high-dose-intensity chemotherapy by CPT-11.**

Author(s): Gornet JM, Azoulay D, Duclos-Vallee JC, Goldwasser F.

Source: *Anti-Cancer Drugs*. 2000 September; 11(8): 649-52.

http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11081458&dopt=Abstract

- **Hepatitis B antigen, alpha-fetoglobulins and primary hepatocellular carcinoma in Ethiopia.**

Author(s): Tsega E, Gold P, Shuster J, Whittemore B, Lester FT.

Source: *J Trop Med Hyg*. 1976 October; 79(10): 230-4. No Abstract Available.

http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=64619&dopt=Abstract

- **Hepatocellular carcinoma, hepatic cirrhosis, and hepatitis B virus infection in Nigeria.**

Author(s): Otu AA.

Source: *Cancer*. 1987 November 15; 60(10): 2581-5.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=2822223&dopt=Abstract

- **Primary hepatocellular carcinoma in Ile-Ife, Nigeria: a prospective study of 154 cases.**
 Author(s): Ndububa DA, Ojo OS, Adeodu OO, Adetiloye VA, Olasode BJ, Famurewa OC, Durosinmi MA, Agbakwuru AE.
 Source: *Niger J Med*. 2001 April-June; 10(2): 59-63.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=11705059&dopt=Abstract

- **Primary liver cancer in Hong Kong.**
 Author(s): Shiu WC.
 Source: *Cancer Chemotherapy and Pharmacology*. 1992; 31 Suppl: S143-5.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=1458565&dopt=Abstract

- **Primary liver cell carcinoma associated with infective liver disease.**
 Author(s): Atoba MA, Otulana BA, Adebajo AO.
 Source: *Trop Geogr Med*. 1988 July; 40(3): 244-6.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=2847379&dopt=Abstract

- **Quadruple chemotherapy versus radiotherapy in treatment of primary hepatocellular carcinoma.**
 Author(s): Cochrane AM, Murray-Lyon IM, Brinkley DM, Williams R.
 Source: *Cancer*. 1977 August; 40(2): 609-14. No Abstract Available.
http://www.ncbi.nlm.nih.gov:80/entrez/query.fcgi?cmd=Retrieve&db=PubMed&list_uids=196734&dopt=Abstract

Additional Web Resources

A number of additional Web sites offer encyclopedic information covering CAM and related topics. The following is a representative sample:

- Alternative Medicine Foundation, Inc.: <http://www.herbmed.org/>
- AOL: <http://search.aol.com/cat.adp?id=169&layer=&from=subcats>
- Chinese Medicine: <http://www.newcenturynutrition.com/>

- drkoop.com[®]:
<http://www.drkoop.com/InteractiveMedicine/IndexC.html>
- Family Village: http://www.familyvillage.wisc.edu/med_altn.htm
- Google: <http://directory.google.com/Top/Health/Alternative/>
- Healthnotes: <http://www.thedacare.org/healthnotes/>
- Open Directory Project: <http://dmoz.org/Health/Alternative/>
- TPN.com: <http://www.tnp.com/>
- Yahoo.com: http://dir.yahoo.com/Health/Alternative_Medicine/
- WebMD[®]Health: http://my.webmd.com/drugs_and_herbs
- WellNet: <http://www.wellnet.ca/herbsa-c.htm>
- WholeHealthMD.com:
<http://www.wholehealthmd.com/reflib/0,1529,,00.html>

The following is a specific Web list relating to adult primary liver cancer; please note that any particular subject below may indicate either a therapeutic use, or a contraindication (potential danger), and does not reflect an official recommendation:

- **General Overview**

- **Breast Cancer**

- Source: Healthnotes, Inc.; www.healthnotes.com

- Hyperlink:

- http://www.thedacare.org/healthnotes/Concern/Cancer_Breast.htm

- **Cancer Prevention (Reducing the Risk)**

- Source: Prima Communications, Inc.

- Hyperlink: <http://www.personalhealthzone.com/pg000272.html>

- **Colon Cancer**

- Source: Healthnotes, Inc.; www.healthnotes.com

- Hyperlink:

- http://www.thedacare.org/healthnotes/Concern/Cancer_Colon.htm

Hepatitis

Source: Healthnotes, Inc.; www.healthnotes.com

Hyperlink:

<http://www.thedacare.org/healthnotes/Concern/Hepatitis.htm>

Liver cancer

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/InteractiveMedicine/ConsLookups/Uses/livercancer.html>

Liver Cirrhosis

Source: Healthnotes, Inc.; www.healthnotes.com

Hyperlink:

http://www.thedacare.org/healthnotes/Concern/Liver_Cirrhosis.htm

Lung Cancer

Source: Healthnotes, Inc.; www.healthnotes.com

Hyperlink:

http://www.thedacare.org/healthnotes/Concern/Cancer_Lung.htm

Prostate Cancer

Source: Healthnotes, Inc.; www.healthnotes.com

Hyperlink:

http://www.thedacare.org/healthnotes/Concern/Cancer_Prostate.htm

Viral Hepatitis

Source: Prima Communications, Inc.

Hyperlink: <http://www.personalhealthzone.com/pg000255.html>

- **Herbs and Supplements**

Acorus

Alternative names: Sweet Flag; *Acorus calamus* L.

Source: Alternative Medicine Foundation, Inc.;

www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Aloe

Alternative names: Aloe vera L.

Source: Alternative Medicine Foundation, Inc.;

www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Androstenedione

Source: Prima Communications, Inc.

Hyperlink: <http://www.personalhealthzone.com/pg000095.html>

Arctium

Alternative names: Burdock, Gobo; Arctium lappa L.

Source: Alternative Medicine Foundation, Inc.;

www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Aristolochia

Alternative names: Snakeroot, Guaco; Aristolochia sp

Source: Alternative Medicine Foundation, Inc.;

www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Asian Ginseng

Alternative names: Panax ginseng

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsHerbs/GinsengAsiach.html>

Bupleurum

Alternative names: Bupleurum chinense, Bupleurum falcatum

Source: Healthnotes, Inc.; www.healthnotes.com

Hyperlink:

<http://www.thedacare.org/healthnotes/Herb/Bupleurum.htm>

DHEA

Source: WholeHealthMD.com, LLC.; www.wholehealthmd.com

Hyperlink:

http://www.wholehealthmd.com/refshelf/substances_view/0,1525,10022,00.html

DHEA (Dehydroepiandrosterone)

Source: Prima Communications, Inc.

Hyperlink: <http://www.personalhealthzone.com/pg000146.html>

Echinacea

Source: Alternative Medicine Foundation, Inc.;

www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Echinacea

Alternative names: Echinacea angustifolia, Echinacea pallida,
Echinacea purpurea, Purple Coneflower

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsHerbs/Echinaceach.html>

Echinacea angustifolia

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsHerbs/Echinaceach.html>

Echinacea pallida

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsHerbs/Echinaceach.html>

Echinacea purpurea

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsHerbs/Echinaceach.html>

Ginseng, Asian

Alternative names: Panax ginseng

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsHerbs/GinsengAsianch.html>

Glutathione

Source: Healthnotes, Inc.; www.healthnotes.com

Hyperlink:

<http://www.thedacare.org/healthnotes/Supp/Glutathione.htm>

Glycyrrhiza1

Alternative names: Licorice; Glycyrrhiza glabra L.

Source: Alternative Medicine Foundation, Inc.;

www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

IP-6

Source: Healthnotes, Inc.; www.healthnotes.com

Hyperlink: <http://www.thedacare.org/healthnotes/Supp/IP-6.htm>

Lavandula

Alternative names: Lavender; Lavandula sp.

Source: Alternative Medicine Foundation, Inc.;

www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Menadione

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsSupplements/VitamKcs.html>

Menaphthone

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsSupplements/VitamKcs.html>

Menaquinone

Source: Integrative Medicine Communications;
www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsSupplements/VitaminKcs.html>

Ornithine Alpha-Ketoglutarate

Source: Prima Communications, Inc.

Hyperlink: <http://www.personalhealthzone.com/pg000215.html>

Panax

Alternative names: Ginseng; Panax ginseng

Source: Alternative Medicine Foundation, Inc.;
www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Panax ginseng

Source: Integrative Medicine Communications;
www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsHerbs/GinsengAsiach.html>

Phylloquinone

Source: Integrative Medicine Communications;
www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsSupplements/VitaminKcs.html>

Pimpinella

Alternative names: Anise; Pimpinella anisum (L)

Source: Alternative Medicine Foundation, Inc.;
www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Piper nigrum

Alternative names: Black Pepper

Source: Alternative Medicine Foundation, Inc.;
www.amfoundation.org

Hyperlink: <http://www.herbmed.org/>

Purple Coneflower

Source: Integrative Medicine Communications;

www.onemedicine.com

Hyperlink:

<http://www.drkoop.com/interactivemedicine/ConsHerbs/Echinacea.html>

General References

A good place to find general background information on CAM is the National Library of Medicine. It has prepared within the MEDLINEplus system an information topic page dedicated to complementary and alternative medicine. To access this page, go to the MEDLINEplus site at: www.nlm.nih.gov/medlineplus/alternativemedicine.html. This Web site provides a general overview of various topics and can lead to a number of general sources. The following additional references describe, in broad terms, alternative and complementary medicine (sorted alphabetically by title; hyperlinks provide rankings, information, and reviews at Amazon.com):

- **Alternative Medicine Definitive Guide to Cancer** by W. John Diamond, et al; Hardcover - 1120 pages Package edition (March 18, 1997), Alternativemedicine.Com Books; ISBN: 1887299017; <http://www.amazon.com/exec/obidos/ASIN/1887299017/icongroupinterna>
- **Beating Cancer With Nutrition - Revised** by Patrick Quillin, Noreen Quillin (Contributor); Paperback - 352 pages; Book & CD edition (January 1, 2001), Bookworld Services; ISBN: 0963837281; <http://www.amazon.com/exec/obidos/ASIN/0963837281/icongroupinterna>
- **Cancer: Increasing Your Odds for Survival - A Resource Guide for Integrating Mainstream, Alternative and Complementary Therapies** by David Bognar, Walter Cronkite; Paperback (August 1998), Hunter House; ISBN: 0897932471; <http://www.amazon.com/exec/obidos/ASIN/0897932471/icongroupinterna>
- **Choices in Healing** by Michael Lerner; Paperback - 696 pages; (February 28, 1996), MIT Press; ISBN: 0262621045; <http://www.amazon.com/exec/obidos/ASIN/0262621045/icongroupinterna>
- **The Gerson Therapy: The Amazing Nutritional Program for Cancer and Other Illnesses** by Charlotte Gerson, Morton Walker, D.P.M.; Paperback - 448 pages (October 2001), Kensington Publishing Corp.; ISBN: 1575666286; <http://www.amazon.com/exec/obidos/ASIN/1575666286/icongroupinterna>

- **Natural Compounds in Cancer Therapy** by John C. Boik; Paperback - 520 pages (March 2001), Oregon Medical Press; ISBN: 0964828014;
<http://www.amazon.com/exec/obidos/ASIN/0964828014/icongroupinterna>
- **There's No Place Like Hope: A Guide to Beating Cancer in Mind-Sized Bites** by Vickie Girard, Dan Zadra (Editor); Hardcover - 161 pages (April 2001), Compendium Inc.; ISBN: 1888387416;
<http://www.amazon.com/exec/obidos/ASIN/1888387416/icongroupinterna>
- **Your Life in Your Hands** by Jane A. Plant, Ph.D; Hardcover - 272 pages (December 13, 2000), St. Martins Press (Trade); ISBN: 0312275617;
<http://www.amazon.com/exec/obidos/ASIN/0312275617/icongroupinterna>

For additional information on complementary and alternative medicine, ask your doctor or write to:

National Institutes of Health
National Center for Complementary and Alternative Medicine
Clearinghouse
P. O. Box 8218
Silver Spring, MD 20907-8218

Vocabulary Builder

The following vocabulary builder gives definitions of words used in this chapter that have not been defined in previous chapters:

Androstenedione: A steroid with androgenic properties that is produced in the testis, ovary, and adrenal cortex. It is a precursor to testosterone and other androgenic hormones. [NIH]

Dehydroepiandrosterone: DHEA. A substance that is being studied as a cancer prevention drug. It belongs to the family of drugs called steroids. [NIH]

DHEA: Dehydroepiandrosterone. A substance that is being studied as a cancer prevention drug. It belongs to the family of drugs called steroids. [NIH]

Echinacea: A genus of perennial herbs used topically and internally. It contains echinacoside, glycosides, inulin, isobutyl amides, resin, and sesquiterpenes. [NIH]

Ginseng: An herb with a root that has been used in some cultures to treat certain medical problems. It may have anticancer effects. [NIH]

Hydroxyurea: An anticancer drug that belongs to the family of drugs called antimetabolites. [NIH]

Morphine: A narcotic drug used in the treatment of pain. [NIH]

Nausea: An unpleasant sensation, vaguely referred to the epigastrium and abdomen, and often culminating in vomiting. [EU]

Non-small cell lung cancer: A group of lung cancers that includes squamous cell carcinoma, adenocarcinoma, and large cell carcinoma. [NIH]

Ornithine: An amino acid produced in the urea cycle by the splitting off of urea from arginine. [NIH]

Psychotherapy: A generic term for the treatment of mental illness or emotional disturbances primarily by verbal or nonverbal communication. [NIH]

Radiotherapy: The treatment of disease by ionizing radiation. [EU]

Selenium: An essential dietary mineral. [NIH]

Supplementation: Adding nutrients to the diet. [NIH]

APPENDIX C. RESEARCHING NUTRITION

Overview

Since the time of Hippocrates, doctors have understood the importance of diet and nutrition to patients' health and well-being. Since then, they have accumulated an impressive archive of studies and knowledge dedicated to this subject. Based on their experience, doctors and healthcare providers may recommend particular dietary supplements to patients with adult primary liver cancer. Any dietary recommendation is based on a patient's age, body mass, gender, lifestyle, eating habits, food preferences, and health condition. It is therefore likely that different patients with adult primary liver cancer may be given different recommendations. Some recommendations may be directly related to adult primary liver cancer, while others may be more related to the patient's general health. These recommendations, themselves, may differ from what official sources recommend for the average person.

In this chapter we will begin by briefly reviewing the essentials of diet and nutrition that will broadly frame more detailed discussions of adult primary liver cancer. We will then show you how to find studies dedicated specifically to nutrition and adult primary liver cancer.

Food and Nutrition: General Principles

What Are Essential Foods?

Food is generally viewed by official sources as consisting of six basic elements: (1) fluids, (2) carbohydrates, (3) protein, (4) fats, (5) vitamins, and (6) minerals. Consuming a combination of these elements is considered to be a healthy diet:

- **Fluids** are essential to human life as 80-percent of the body is composed of water. Water is lost via urination, sweating, diarrhea, vomiting, diuretics (drugs that increase urination), caffeine, and physical exertion.
- **Carbohydrates** are the main source for human energy (thermoregulation) and the bulk of typical diets. They are mostly classified as being either simple or complex. Simple carbohydrates include sugars which are often consumed in the form of cookies, candies, or cakes. Complex carbohydrates consist of starches and dietary fibers. Starches are consumed in the form of pastas, breads, potatoes, rice, and other foods. Soluble fibers can be eaten in the form of certain vegetables, fruits, oats, and legumes. Insoluble fibers include brown rice, whole grains, certain fruits, wheat bran and legumes.
- **Proteins** are eaten to build and repair human tissues. Some foods that are high in protein are also high in fat and calories. Food sources for protein include nuts, meat, fish, cheese, and other dairy products.
- **Fats** are consumed for both energy and the absorption of certain vitamins. There are many types of fats, with many general publications recommending the intake of unsaturated fats or those low in cholesterol.

Vitamins and minerals are fundamental to human health, growth, and, in some cases, disease prevention. Most are consumed in your diet (exceptions being vitamins K and D which are produced by intestinal bacteria and sunlight on the skin, respectively). Each vitamin and mineral plays a different role in health. The following outlines essential vitamins:

- **Vitamin A** is important to the health of your eyes, hair, bones, and skin; sources of vitamin A include foods such as eggs, carrots, and cantaloupe.
- **Vitamin B¹**, also known as thiamine, is important for your nervous system and energy production; food sources for thiamine include meat, peas, fortified cereals, bread, and whole grains.
- **Vitamin B²**, also known as riboflavin, is important for your nervous system and muscles, but is also involved in the release of proteins from

nutrients; food sources for riboflavin include dairy products, leafy vegetables, meat, and eggs.

- **Vitamin B³**, also known as niacin, is important for healthy skin and helps the body use energy; food sources for niacin include peas, peanuts, fish, and whole grains
- **Vitamin B⁶**, also known as pyridoxine, is important for the regulation of cells in the nervous system and is vital for blood formation; food sources for pyridoxine include bananas, whole grains, meat, and fish.
- **Vitamin B¹²** is vital for a healthy nervous system and for the growth of red blood cells in bone marrow; food sources for vitamin B¹² include yeast, milk, fish, eggs, and meat.
- **Vitamin C** allows the body's immune system to fight various diseases, strengthens body tissue, and improves the body's use of iron; food sources for vitamin C include a wide variety of fruits and vegetables.
- **Vitamin D** helps the body absorb calcium which strengthens bones and teeth; food sources for vitamin D include oily fish and dairy products.
- **Vitamin E** can help protect certain organs and tissues from various degenerative diseases; food sources for vitamin E include margarine, vegetables, eggs, and fish.
- **Vitamin K** is essential for bone formation and blood clotting; common food sources for vitamin K include leafy green vegetables.
- **Folic Acid** maintains healthy cells and blood and, when taken by a pregnant woman, can prevent her fetus from developing neural tube defects; food sources for folic acid include nuts, fortified breads, leafy green vegetables, and whole grains.

It should be noted that one can overdose on certain vitamins which become toxic if consumed in excess (e.g. vitamin A, D, E and K).

Like vitamins, minerals are chemicals that are required by the body to remain in good health. Because the human body does not manufacture these chemicals internally, we obtain them from food and other dietary sources. The more important minerals include:

- **Calcium** is needed for healthy bones, teeth, and muscles, but also helps the nervous system function; food sources for calcium include dry beans, peas, eggs, and dairy products.

- **Chromium** is helpful in regulating sugar levels in blood; food sources for chromium include egg yolks, raw sugar, cheese, nuts, beets, whole grains, and meat.
- **Fluoride** is used by the body to help prevent tooth decay and to reinforce bone strength; sources of fluoride include drinking water and certain brands of toothpaste.
- **Iodine** helps regulate the body's use of energy by synthesizing into the hormone thyroxine; food sources include leafy green vegetables, nuts, egg yolks, and red meat.
- **Iron** helps maintain muscles and the formation of red blood cells and certain proteins; food sources for iron include meat, dairy products, eggs, and leafy green vegetables.
- **Magnesium** is important for the production of DNA, as well as for healthy teeth, bones, muscles, and nerves; food sources for magnesium include dried fruit, dark green vegetables, nuts, and seafood.
- **Phosphorous** is used by the body to work with calcium to form bones and teeth; food sources for phosphorous include eggs, meat, cereals, and dairy products.
- **Selenium** primarily helps maintain normal heart and liver functions; food sources for selenium include wholegrain cereals, fish, meat, and dairy products.
- **Zinc** helps wounds heal, the formation of sperm, and encourage rapid growth and energy; food sources include dried beans, shellfish, eggs, and nuts.

The United States government periodically publishes recommended diets and consumption levels of the various elements of food. Again, your doctor may encourage deviations from the average official recommendation based on your specific condition. To learn more about basic dietary guidelines, visit the Web site: <http://www.health.gov/dietaryguidelines/>. Based on these guidelines, many foods are required to list the nutrition levels on the food's packaging. Labeling Requirements are listed at the following site maintained by the Food and Drug Administration: <http://www.cfsan.fda.gov/~dms/lab-cons.html>. When interpreting these requirements, the government recommends that consumers become familiar with the following abbreviations before reading FDA literature:⁹⁸

- **DVs (Daily Values):** A new dietary reference term that will appear on the food label. It is made up of two sets of references, DRVs and RDIs.

⁹⁸ Adapted from the FDA: <http://www.fda.gov/fdac/special/foodlabel/dvs.html>.

- **DRVs (Daily Reference Values):** A set of dietary references that applies to fat, saturated fat, cholesterol, carbohydrate, protein, fiber, sodium, and potassium.
- **RDIs (Reference Daily Intakes):** A set of dietary references based on the Recommended Dietary Allowances for essential vitamins and minerals and, in selected groups, protein. The name “RDI” replaces the term “U.S. RDA.”
- **RDAs (Recommended Dietary Allowances):** A set of estimated nutrient allowances established by the National Academy of Sciences. It is updated periodically to reflect current scientific knowledge.

What Are Dietary Supplements?⁹⁹

Dietary supplements are widely available through many commercial sources, including health food stores, grocery stores, pharmacies, and by mail. Dietary supplements are provided in many forms including tablets, capsules, powders, gel-tabs, extracts, and liquids. Historically in the United States, the most prevalent type of dietary supplement was a multivitamin/mineral tablet or capsule that was available in pharmacies, either by prescription or “over the counter.” Supplements containing strictly herbal preparations were less widely available. Currently in the United States, a wide array of supplement products are available, including vitamin, mineral, other nutrients, and botanical supplements as well as ingredients and extracts of animal and plant origin.

The Office of Dietary Supplements (ODS) of the National Institutes of Health is the official agency of the United States which has the expressed goal of acquiring “new knowledge to help prevent, detect, diagnose, and treat disease and disability, from the rarest genetic disorder to the common cold.”¹⁰⁰ According to the ODS, dietary supplements can have an important impact on the prevention and management of disease and on the maintenance of health.¹⁰¹ The ODS notes that considerable research on the

⁹⁹ This discussion has been adapted from the NIH:

<http://ods.od.nih.gov/whatare/whatare.html>.

¹⁰⁰ Contact: The Office of Dietary Supplements, National Institutes of Health, Building 31, Room 1B29, 31 Center Drive, MSC 2086, Bethesda, Maryland 20892-2086, Tel: (301) 435-2920, Fax: (301) 480-1845, E-mail: ods@nih.gov.

¹⁰¹ Adapted from <http://ods.od.nih.gov/about/about.html>. The Dietary Supplement Health and Education Act defines dietary supplements as “a product (other than tobacco) intended to supplement the diet that bears or contains one or more of the following dietary ingredients: a vitamin, mineral, amino acid, herb or other botanical; or a dietary substance for use to supplement the diet by increasing the total dietary intake; or a concentrate,

effects of dietary supplements has been conducted in Asia and Europe where the use of plant products, in particular, has a long tradition. However, the overwhelming majority of supplements have not been studied scientifically. To explore the role of dietary supplements in the improvement of health care, the ODS plans, organizes, and supports conferences, workshops, and symposia on scientific topics related to dietary supplements. The ODS often works in conjunction with other NIH Institutes and Centers, other government agencies, professional organizations, and public advocacy groups.

To learn more about official information on dietary supplements, visit the ODS site at <http://ods.od.nih.gov/whatare/whatare.html>. Or contact:

The Office of Dietary Supplements
National Institutes of Health
Building 31, Room 1B29
31 Center Drive, MSC 2086
Bethesda, Maryland 20892-2086
Tel: (301) 435-2920
Fax: (301) 480-1845
E-mail: ods@nih.gov

Finding Studies on Adult Primary Liver Cancer

The NIH maintains an office dedicated to patient nutrition and diet. The National Institutes of Health's Office of Dietary Supplements (ODS) offers a searchable bibliographic database called the IBIDS (International Bibliographic Information on Dietary Supplements). The IBIDS contains over 460,000 scientific citations and summaries about dietary supplements and nutrition as well as references to published international, scientific literature on dietary supplements such as vitamins, minerals, and botanicals.¹⁰² IBIDS is available to the public free of charge through the ODS Internet page: <http://ods.od.nih.gov/databases/ibids.html>.

metabolite, constituent, extract, or combination of any ingredient described above; and intended for ingestion in the form of a capsule, powder, softgel, or gelcap, and not represented as a conventional food or as a sole item of a meal or the diet."

¹⁰² Adapted from <http://ods.od.nih.gov>. IBIDS is produced by the Office of Dietary Supplements (ODS) at the National Institutes of Health to assist the public, healthcare providers, educators, and researchers in locating credible, scientific information on dietary supplements. IBIDS was developed and will be maintained through an interagency partnership with the Food and Nutrition Information Center of the National Agricultural Library, U.S. Department of Agriculture.

After entering the search area, you have three choices: (1) IBIDS Consumer Database, (2) Full IBIDS Database, or (3) Peer Reviewed Citations Only. We recommend that you start with the Consumer Database. While you may not find references for the topics that are of most interest to you, check back periodically as this database is frequently updated. More studies can be found by searching the Full IBIDS Database. Healthcare professionals and researchers generally use the third option, which lists peer-reviewed citations. In all cases, we suggest that you take advantage of the “Advanced Search” option that allows you to retrieve up to 100 fully explained references in a comprehensive format. Type “adult primary liver cancer” (or synonyms) into the search box. To narrow the search, you can also select the “Title” field.

The following information is typical of that found when using the “Full IBIDS Database” when searching using “adult primary liver cancer” (or a synonym):

- **Evaluation of antiandrogen therapy in unresectable hepatocellular carcinoma: results of a European Organization for Research and Treatment of Cancer multicentric double-blind trial.**
 Author(s): Hopital de Cimiez, Nice, France.
 Source: Grimaldi, C Bleiberg, H Gay, F Messner, M Rougier, P Kok, T C Cirera, L Cervantes, A De Greve, J Paillot, B Buset, M Nitti, D Sahmoud, T Duez, N Wils, J J-Clin-Oncol. 1998 February; 16(2): 411-7 0732-183X
- **Sodium butyrate enhances STAT 1 expression in PLC/PRE/5 hepatoma cells and augments their responsiveness to interferon-alpha.**
 Author(s): School of Technology for Medical Sciences, Kaohsiung Medical College, Taiwan, Republic of China.
 Source: Hung, W C Chuang, L Y Br-J-Cancer. 1999 May; 80(5-6): 705-10 0007-0920

Federal Resources on Nutrition

In addition to the IBIDS, the United States Department of Health and Human Services (HHS) and the United States Department of Agriculture (USDA) provide many sources of information on general nutrition and health. Recommended resources include:

- healthfinder®, HHS’s gateway to health information, including diet and nutrition:
<http://www.healthfinder.gov/scripts/SearchContext.asp?topic=238&page=0>

- The United States Department of Agriculture's Web site dedicated to nutrition information: www.nutrition.gov
- The Food and Drug Administration's Web site for federal food safety information: www.foodsafety.gov
- The National Action Plan on Overweight and Obesity sponsored by the United States Surgeon General:
<http://www.surgeongeneral.gov/topics/obesity/>
- The Center for Food Safety and Applied Nutrition has an Internet site sponsored by the Food and Drug Administration and the Department of Health and Human Services: <http://vm.cfsan.fda.gov/>
- Center for Nutrition Policy and Promotion sponsored by the United States Department of Agriculture: <http://www.usda.gov/cnpp/>
- Food and Nutrition Information Center, National Agricultural Library sponsored by the United States Department of Agriculture: <http://www.nal.usda.gov/fnic/>
- Food and Nutrition Service sponsored by the United States Department of Agriculture: <http://www.fns.usda.gov/fns/>

Additional Web Resources

A number of additional Web sites offer encyclopedic information covering food and nutrition. The following is a representative sample:

- AOL: <http://search.aol.com/cat.adp?id=174&layer=&from=subcats>
- Family Village: http://www.familyvillage.wisc.edu/med_nutrition.html
- Google: <http://directory.google.com/Top/Health/Nutrition/>
- Healthnotes: <http://www.thedacare.org/healthnotes/>
- Open Directory Project: <http://dmoz.org/Health/Nutrition/>
- Yahoo.com: <http://dir.yahoo.com/Health/Nutrition/>
- WebMD[®]Health: <http://my.webmd.com/nutrition>
- WholeHealthMD.com:
<http://www.wholehealthmd.com/reflib/0,1529,,00.html>

The following is a specific Web list relating to adult primary liver cancer; please note that any particular subject below may indicate either a

therapeutic use, or a contraindication (potential danger), and does not reflect an official recommendation:

- **Vitamins**

- **Vitamin K**

- Alternative names: Menadione, Menaphthone, Menaquinone, Phylloquinone

- Source: Integrative Medicine Communications; www.onemedicine.com

- Hyperlink:

- <http://www.drkoop.com/interactivemedicine/ConsSupplements/VitaminKcs.html>

- **Minerals**

- **Selenium**

- Source: Prima Communications, Inc.

- Hyperlink: <http://www.personalhealthzone.com/pg000233.html>

- **Food and Diet**

- **Cancer Prevention and Diet**

- Source: Healthnotes, Inc.; www.healthnotes.com

- Hyperlink:

- http://www.thedacare.org/healthnotes/Concern/Cancer_Diet.htm

Vocabulary Builder

The following vocabulary builder defines words used in the references in this chapter that have not been defined in previous chapters:

Bacteria: A large group of single-cell microorganisms. Some cause infections and disease in animals and humans. The singular of bacteria is bacterium. [NIH]

Calcium: A mineral found in teeth, bones, and other body tissues. [NIH]

Capsules: Hard or soft soluble containers used for the oral administration of medicine. [NIH]

Carbohydrate: An aldehyde or ketone derivative of a polyhydric alcohol, particularly of the pentahydric and hexahydric alcohols. They are so named

because the hydrogen and oxygen are usually in the proportion to form water, $(CH_2O)_n$. The most important carbohydrates are the starches, sugars, celluloses, and gums. They are classified into mono-, di-, tri-, poly- and heterosaccharides. [EU]

Cholesterol: The principal sterol of all higher animals, distributed in body tissues, especially the brain and spinal cord, and in animal fats and oils. [NIH]

Degenerative: Undergoing degeneration : tending to degenerate; having the character of or involving degeneration; causing or tending to cause degeneration. [EU]

Diarrhea: Passage of excessively liquid or excessively frequent stools. [NIH]

Fetus: The developing offspring from 7 to 8 weeks after conception until birth. [NIH]

Neural: 1. pertaining to a nerve or to the nerves. 2. situated in the region of the spinal axis, as the neutral arch. [EU]

Niacin: Water-soluble vitamin of the B complex occurring in various animal and plant tissues. Required by the body for the formation of coenzymes NAD and NADP. Has pellagra-curative, vasodilating, and antilipemic properties. [NIH]

Overdose: 1. to administer an excessive dose. 2. an excessive dose. [EU]

Phosphorous: Having to do with or containing the element phosphorus. [NIH]

Potassium: A metallic element that is important in body functions such as regulation of blood pressure and of water content in cells, transmission of nerve impulses, digestion, muscle contraction, and heart beat. [NIH]

Riboflavin: Nutritional factor found in milk, eggs, malted barley, liver, kidney, heart, and leafy vegetables. The richest natural source is yeast. It occurs in the free form only in the retina of the eye, in whey, and in urine; its principal forms in tissues and cells are as FMN and FAD. [NIH]

Thermoregulation: Heat regulation. [EU]

Thyroxine: An amino acid of the thyroid gland which exerts a stimulating effect on thyroid metabolism. [NIH]

APPENDIX D. FINDING MEDICAL LIBRARIES

Overview

At a medical library you can find medical texts and reference books, consumer health publications, specialty newspapers and magazines, as well as medical journals. In this appendix, we show you how to quickly find a medical library in your area.

Preparation

Before going to the library, highlight the references mentioned in this sourcebook that you find interesting. Focus on those items that are not available via the Internet, and ask the reference librarian for help with your search. He or she may know of additional resources that could be helpful to you. Most importantly, your local public library and medical libraries have Interlibrary Loan programs with the National Library of Medicine (NLM), one of the largest medical collections in the world. According to the NLM, most of the literature in the general and historical collections of the National Library of Medicine is available on interlibrary loan to any library. NLM's interlibrary loan services are only available to libraries. If you would like to access NLM medical literature, then visit a library in your area that can request the publications for you.¹⁰³

¹⁰³ Adapted from the NLM: <http://www.nlm.nih.gov/psd/cas/interlibrary.html>.

Finding a Local Medical Library

The quickest method to locate medical libraries is to use the Internet-based directory published by the National Network of Libraries of Medicine (NN/LM). This network includes 4626 members and affiliates that provide many services to librarians, health professionals, and the public. To find a library in your area, simply visit <http://nnlm.gov/members/adv.html> or call 1-800-338-7657.

Medical Libraries Open to the Public

In addition to the NN/LM, the National Library of Medicine (NLM) lists a number of libraries that are generally open to the public and have reference facilities. The following is the NLM's list plus hyperlinks to each library Web site. These Web pages can provide information on hours of operation and other restrictions. The list below is a small sample of libraries recommended by the National Library of Medicine (sorted alphabetically by name of the U.S. state or Canadian province where the library is located):¹⁰⁴

- **Alabama:** Health InfoNet of Jefferson County (Jefferson County Library Cooperative, Lister Hill Library of the Health Sciences), <http://www.uab.edu/infonet/>
- **Alabama:** Richard M. Scrushy Library (American Sports Medicine Institute), <http://www.asmi.org/LIBRARY.HTM>
- **Arizona:** Samaritan Regional Medical Center: The Learning Center (Samaritan Health System, Phoenix, Arizona), <http://www.samaritan.edu/library/bannerlibs.htm>
- **California:** Kris Kelly Health Information Center (St. Joseph Health System), <http://www.humboldt1.com/~kkhic/index.html>
- **California:** Community Health Library of Los Gatos (Community Health Library of Los Gatos), <http://www.healthlib.org/orgresources.html>
- **California:** Consumer Health Program and Services (CHIPS) (County of Los Angeles Public Library, Los Angeles County Harbor-UCLA Medical Center Library) - Carson, CA, <http://www.colapublib.org/services/chips.html>
- **California:** Gateway Health Library (Sutter Gould Medical Foundation)
- **California:** Health Library (Stanford University Medical Center), <http://www-med.stanford.edu/healthlibrary/>

¹⁰⁴ Abstracted from <http://www.nlm.nih.gov/medlineplus/libraries.html>.

- **California:** Patient Education Resource Center - Health Information and Resources (University of California, San Francisco), <http://sfghdean.ucsf.edu/barnett/PERC/default.asp>
- **California:** Redwood Health Library (Petaluma Health Care District), <http://www.phcd.org/rdwplib.html>
- **California:** San José PlaneTree Health Library, <http://planetreesanjose.org/>
- **California:** Sutter Resource Library (Sutter Hospitals Foundation), <http://go.sutterhealth.org/comm/resc-library/sac-resources.html>
- **California:** University of California, Davis. Health Sciences Libraries
- **California:** ValleyCare Health Library & Ryan Comer Cancer Resource Center (ValleyCare Health System), <http://www.valleycare.com/library.html>
- **California:** Washington Community Health Resource Library (Washington Community Health Resource Library), <http://www.healthlibrary.org/>
- **Colorado:** William V. Gervasini Memorial Library (Exempla Healthcare), <http://www.exempla.org/conslib.htm>
- **Connecticut:** Hartford Hospital Health Science Libraries (Hartford Hospital), <http://www.harthosp.org/library/>
- **Connecticut:** Healthnet: Connecticut Consumer Health Information Center (University of Connecticut Health Center, Lyman Maynard Stowe Library), <http://library.uchc.edu/departm/hnet/>
- **Connecticut:** Waterbury Hospital Health Center Library (Waterbury Hospital), <http://www.waterburyhospital.com/library/consumer.shtml>
- **Delaware:** Consumer Health Library (Christiana Care Health System, Eugene du Pont Preventive Medicine & Rehabilitation Institute), http://www.christianacare.org/health_guide/health_guide_pmri_health_info.cfm
- **Delaware:** Lewis B. Flinn Library (Delaware Academy of Medicine), <http://www.delamed.org/chls.html>
- **Georgia:** Family Resource Library (Medical College of Georgia), http://cmc.mcg.edu/kids_families/fam_resources/fam_res_lib/frl.htm
- **Georgia:** Health Resource Center (Medical Center of Central Georgia), <http://www.mccg.org/hrc/hrchome.asp>
- **Hawaii:** Hawaii Medical Library: Consumer Health Information Service (Hawaii Medical Library), <http://hml.org/CHIS/>

- **Idaho:** DeArmond Consumer Health Library (Kootenai Medical Center), <http://www.nicon.org/DeArmond/index.htm>
- **Illinois:** Health Learning Center of Northwestern Memorial Hospital (Northwestern Memorial Hospital, Health Learning Center), http://www.nmh.org/health_info/hlc.html
- **Illinois:** Medical Library (OSF Saint Francis Medical Center), <http://www.osfsaintfrancis.org/general/library/>
- **Kentucky:** Medical Library - Services for Patients, Families, Students & the Public (Central Baptist Hospital), <http://www.centralbap.com/education/community/library.htm>
- **Kentucky:** University of Kentucky - Health Information Library (University of Kentucky, Chandler Medical Center, Health Information Library), <http://www.mc.uky.edu/PatientEd/>
- **Louisiana:** Alton Ochsner Medical Foundation Library (Alton Ochsner Medical Foundation), <http://www.ochsner.org/library/>
- **Louisiana:** Louisiana State University Health Sciences Center Medical Library-Shreveport, <http://lib-sh.lsuhscc.edu/>
- **Maine:** Franklin Memorial Hospital Medical Library (Franklin Memorial Hospital), <http://www.fchn.org/fmh/lib.htm>
- **Maine:** Gerrish-True Health Sciences Library (Central Maine Medical Center), <http://www.cmmc.org/library/library.html>
- **Maine:** Hadley Parrot Health Science Library (Eastern Maine Healthcare), <http://www.emh.org/hll/hpl/guide.htm>
- **Maine:** Maine Medical Center Library (Maine Medical Center), <http://www.mmc.org/library/>
- **Maine:** Parkview Hospital, <http://www.parkviewhospital.org/communit.htm#Library>
- **Maine:** Southern Maine Medical Center Health Sciences Library (Southern Maine Medical Center), <http://www.smmc.org/services/service.php3?choice=10>
- **Maine:** Stephens Memorial Hospital Health Information Library (Western Maine Health), http://www.wmhcc.com/hil_frame.html
- **Manitoba, Canada:** Consumer & Patient Health Information Service (University of Manitoba Libraries), <http://www.umanitoba.ca/libraries/units/health/reference/chis.html>
- **Manitoba, Canada:** J.W. Crane Memorial Library (Deer Lodge Centre), <http://www.deerlodge.mb.ca/library/libraryservices.shtml>

- **Maryland:** Health Information Center at the Wheaton Regional Library (Montgomery County, Md., Dept. of Public Libraries, Wheaton Regional Library), <http://www.mont.lib.md.us/healthinfo/hic.asp>
- **Massachusetts:** Baystate Medical Center Library (Baystate Health System), <http://www.baystatehealth.com/1024/>
- **Massachusetts:** Boston University Medical Center Alumni Medical Library (Boston University Medical Center), <http://med-libwww.bu.edu/library/lib.html>
- **Massachusetts:** Lowell General Hospital Health Sciences Library (Lowell General Hospital), <http://www.lowellgeneral.org/library/HomePageLinks/WWW.htm>
- **Massachusetts:** Paul E. Woodard Health Sciences Library (New England Baptist Hospital), http://www.nebh.org/health_lib.asp
- **Massachusetts:** St. Luke's Hospital Health Sciences Library (St. Luke's Hospital), <http://www.southcoast.org/library/>
- **Massachusetts:** Treadwell Library Consumer Health Reference Center (Massachusetts General Hospital), <http://www.mgh.harvard.edu/library/chrcindex.html>
- **Massachusetts:** UMass HealthNet (University of Massachusetts Medical School), <http://healthnet.umassmed.edu/>
- **Michigan:** Botsford General Hospital Library - Consumer Health (Botsford General Hospital, Library & Internet Services), <http://www.botsfordlibrary.org/consumer.htm>
- **Michigan:** Helen DeRoy Medical Library (Providence Hospital and Medical Centers), <http://www.providence-hospital.org/library/>
- **Michigan:** Marquette General Hospital - Consumer Health Library (Marquette General Hospital, Health Information Center), <http://www.mgh.org/center.html>
- **Michigan:** Patient Education Resource Center - University of Michigan Cancer Center (University of Michigan Comprehensive Cancer Center), <http://www.cancer.med.umich.edu/learn/leares.htm>
- **Michigan:** Sladen Library & Center for Health Information Resources - Consumer Health Information, <http://www.sladen.hfhs.org/library/consumer/index.html>
- **Montana:** Center for Health Information (St. Patrick Hospital and Health Sciences Center), <http://www.saintpatrick.org/chi/librarydetail.php3?ID=41>

- **National:** Consumer Health Library Directory (Medical Library Association, Consumer and Patient Health Information Section), <http://caphis.mlanet.org/directory/index.html>
- **National:** National Network of Libraries of Medicine (National Library of Medicine) - provides library services for health professionals in the United States who do not have access to a medical library, <http://nnlm.gov/>
- **National:** NN/LM List of Libraries Serving the Public (National Network of Libraries of Medicine), <http://nnlm.gov/members/>
- **Nevada:** Health Science Library, West Charleston Library (Las Vegas Clark County Library District), http://www.lvccld.org/special_collections/medical/index.htm
- **New Hampshire:** Dartmouth Biomedical Libraries (Dartmouth College Library), http://www.dartmouth.edu/~biomed/resources.html#conshealth.html#
- **New Jersey:** Consumer Health Library (Rahway Hospital), <http://www.rahwayhospital.com/library.htm>
- **New Jersey:** Dr. Walter Phillips Health Sciences Library (Englewood Hospital and Medical Center), <http://www.englewoodhospital.com/links/index.htm>
- **New Jersey:** Meland Foundation (Englewood Hospital and Medical Center), <http://www.geocities.com/ResearchTriangle/9360/>
- **New York:** Choices in Health Information (New York Public Library) - NLM Consumer Pilot Project participant, <http://www.nypl.org/branch/health/links.html>
- **New York:** Health Information Center (Upstate Medical University, State University of New York), <http://www.upstate.edu/library/hic/>
- **New York:** Health Sciences Library (Long Island Jewish Medical Center), <http://www.lij.edu/library/library.html>
- **New York:** ViaHealth Medical Library (Rochester General Hospital), <http://www.nyam.org/library/>
- **Ohio:** Consumer Health Library (Akron General Medical Center, Medical & Consumer Health Library), <http://www.akrongeneral.org/hwlibrary.htm>
- **Oklahoma:** Saint Francis Health System Patient/Family Resource Center (Saint Francis Health System), <http://www.sfh-tulsa.com/patientfamilycenter/default.asp>

- **Oregon:** Planetree Health Resource Center (Mid-Columbia Medical Center), <http://www.mcmc.net/phrc/>
- **Pennsylvania:** Community Health Information Library (Milton S. Hershey Medical Center), <http://www.hmc.psu.edu/commhealth/>
- **Pennsylvania:** Community Health Resource Library (Geisinger Medical Center), <http://www.geisinger.edu/education/commlib.shtml>
- **Pennsylvania:** HealthInfo Library (Moses Taylor Hospital), <http://www.mth.org/healthwellness.html>
- **Pennsylvania:** Hopwood Library (University of Pittsburgh, Health Sciences Library System), <http://www.hsls.pitt.edu/chi/hhrcinfo.html>
- **Pennsylvania:** Koop Community Health Information Center (College of Physicians of Philadelphia), <http://www.collphyphil.org/koopp1.shtml>
- **Pennsylvania:** Learning Resources Center - Medical Library (Susquehanna Health System), <http://www.shscare.org/services/lrc/index.asp>
- **Pennsylvania:** Medical Library (UPMC Health System), <http://www.upmc.edu/passavant/library.htm>
- **Quebec, Canada:** Medical Library (Montreal General Hospital), <http://ww2.mcgill.ca/mghlib/>
- **South Dakota:** Rapid City Regional Hospital - Health Information Center (Rapid City Regional Hospital, Health Information Center), <http://www.rcrh.org/education/LibraryResourcesConsumers.htm>
- **Texas:** Houston HealthWays (Houston Academy of Medicine-Texas Medical Center Library), <http://hwh.library.tmc.edu/>
- **Texas:** Matustik Family Resource Center (Cook Children's Health Care System), http://www.cookchildrens.com/Matustik_Library.html
- **Washington:** Community Health Library (Kittitas Valley Community Hospital), <http://www.kvch.com/>
- **Washington:** Southwest Washington Medical Center Library (Southwest Washington Medical Center), <http://www.swmedctr.com/Home/>

APPENDIX E. YOUR RIGHTS AND INSURANCE

Overview

Any patient with adult primary liver cancer faces a series of issues related more to the healthcare industry than to the medical condition itself. This appendix covers two important topics in this regard: your rights and responsibilities as a patient, and how to get the most out of your medical insurance plan.

Your Rights as a Patient

The President's Advisory Commission on Consumer Protection and Quality in the Healthcare Industry has created the following summary of your rights as a patient.¹⁰⁵

Information Disclosure

Consumers have the right to receive accurate, easily understood information. Some consumers require assistance in making informed decisions about health plans, health professionals, and healthcare facilities. Such information includes:

- *Health plans.* Covered benefits, cost-sharing, and procedures for resolving complaints, licensure, certification, and accreditation status, comparable measures of quality and consumer satisfaction, provider

¹⁰⁵Adapted from Consumer Bill of Rights and Responsibilities:
<http://www.hcqualitycommission.gov/press/cbor.html#head1>.

network composition, the procedures that govern access to specialists and emergency services, and care management information.

- **Health professionals.** Education, board certification, and recertification, years of practice, experience performing certain procedures, and comparable measures of quality and consumer satisfaction.
- **Healthcare facilities.** Experience in performing certain procedures and services, accreditation status, comparable measures of quality, worker, and consumer satisfaction, and procedures for resolving complaints.
- **Consumer assistance programs.** Programs must be carefully structured to promote consumer confidence and to work cooperatively with health plans, providers, payers, and regulators. Desirable characteristics of such programs are sponsorship that ensures accountability to the interests of consumers and stable, adequate funding.

Choice of Providers and Plans

Consumers have the right to a choice of healthcare providers that is sufficient to ensure access to appropriate high-quality healthcare. To ensure such choice, the Commission recommends the following:

- **Provider network adequacy.** All health plan networks should provide access to sufficient numbers and types of providers to assure that all covered services will be accessible without unreasonable delay -- including access to emergency services 24 hours a day and 7 days a week. If a health plan has an insufficient number or type of providers to provide a covered benefit with the appropriate degree of specialization, the plan should ensure that the consumer obtains the benefit outside the network at no greater cost than if the benefit were obtained from participating providers.
- **Women's health services.** Women should be able to choose a qualified provider offered by a plan -- such as gynecologists, certified nurse midwives, and other qualified healthcare providers -- for the provision of covered care necessary to provide routine and preventative women's healthcare services.
- **Access to specialists.** Consumers with complex or serious medical conditions who require frequent specialty care should have direct access to a qualified specialist of their choice within a plan's network of providers. Authorizations, when required, should be for an adequate number of direct access visits under an approved treatment plan.

- **Transitional care.** Consumers who are undergoing a course of treatment for a chronic or disabling condition (or who are in the second or third trimester of a pregnancy) at the time they involuntarily change health plans or at a time when a provider is terminated by a plan for other than cause should be able to continue seeing their current specialty providers for up to 90 days (or through completion of postpartum care) to allow for transition of care.
- **Choice of health plans.** Public and private group purchasers should, wherever feasible, offer consumers a choice of high-quality health insurance plans.

Access to Emergency Services

Consumers have the right to access emergency healthcare services when and where the need arises. Health plans should provide payment when a consumer presents to an emergency department with acute symptoms of sufficient severity--including severe pain--such that a "prudent layperson" could reasonably expect the absence of medical attention to result in placing that consumer's health in serious jeopardy, serious impairment to bodily functions, or serious dysfunction of any bodily organ or part.

Participation in Treatment Decisions

Consumers have the right and responsibility to fully participate in all decisions related to their healthcare. Consumers who are unable to fully participate in treatment decisions have the right to be represented by parents, guardians, family members, or other conservators. Physicians and other health professionals should:

- Provide patients with sufficient information and opportunity to decide among treatment options consistent with the informed consent process.
- Discuss all treatment options with a patient in a culturally competent manner, including the option of no treatment at all.
- Ensure that persons with disabilities have effective communications with members of the health system in making such decisions.
- Discuss all current treatments a consumer may be undergoing.
- Discuss all risks, benefits, and consequences to treatment or nontreatment.

- Give patients the opportunity to refuse treatment and to express preferences about future treatment decisions.
- Discuss the use of advance directives -- both living wills and durable powers of attorney for healthcare -- with patients and their designated family members.
- Abide by the decisions made by their patients and/or their designated representatives consistent with the informed consent process.

Health plans, health providers, and healthcare facilities should:

- Disclose to consumers factors -- such as methods of compensation, ownership of or interest in healthcare facilities, or matters of conscience -- that could influence advice or treatment decisions.
- Assure that provider contracts do not contain any so-called "gag clauses" or other contractual mechanisms that restrict healthcare providers' ability to communicate with and advise patients about medically necessary treatment options.
- Be prohibited from penalizing or seeking retribution against healthcare professionals or other health workers for advocating on behalf of their patients.

Respect and Nondiscrimination

Consumers have the right to considerate, respectful care from all members of the healthcare industry at all times and under all circumstances. An environment of mutual respect is essential to maintain a quality healthcare system. To assure that right, the Commission recommends the following:

- Consumers must not be discriminated against in the delivery of healthcare services consistent with the benefits covered in their policy, or as required by law, based on race, ethnicity, national origin, religion, sex, age, mental or physical disability, sexual orientation, genetic information, or source of payment.
- Consumers eligible for coverage under the terms and conditions of a health plan or program, or as required by law, must not be discriminated against in marketing and enrollment practices based on race, ethnicity, national origin, religion, sex, age, mental or physical disability, sexual orientation, genetic information, or source of payment.

Confidentiality of Health Information

Consumers have the right to communicate with healthcare providers in confidence and to have the confidentiality of their individually identifiable healthcare information protected. Consumers also have the right to review and copy their own medical records and request amendments to their records.

Complaints and Appeals

Consumers have the right to a fair and efficient process for resolving differences with their health plans, healthcare providers, and the institutions that serve them, including a rigorous system of internal review and an independent system of external review. A free copy of the Patient's Bill of Rights is available from the American Hospital Association.¹⁰⁶

Patient Responsibilities

Treatment is a two-way street between you and your healthcare providers. To underscore the importance of finance in modern healthcare as well as your responsibility for the financial aspects of your care, the President's Advisory Commission on Consumer Protection and Quality in the Healthcare Industry has proposed that patients understand the following "Consumer Responsibilities."¹⁰⁷ In a healthcare system that protects consumers' rights, it is reasonable to expect and encourage consumers to assume certain responsibilities. Greater individual involvement by the consumer in his or her care increases the likelihood of achieving the best outcome and helps support a quality-oriented, cost-conscious environment. Such responsibilities include:

- Take responsibility for maximizing healthy habits such as exercising, not smoking, and eating a healthy diet.
- Work collaboratively with healthcare providers in developing and carrying out agreed-upon treatment plans.
- Disclose relevant information and clearly communicate wants and needs.

¹⁰⁶ To order your free copy of the Patient's Bill of Rights, telephone 312-422-3000 or visit the American Hospital Association's Web site: <http://www.aha.org>. Click on "Resource Center," go to "Search" at bottom of page, and then type in "Patient's Bill of Rights." The Patient's Bill of Rights is also available from Fax on Demand, at 312-422-2020, document number 471124.

¹⁰⁷ Adapted from <http://www.hcqualitycommission.gov/press/cbor.html#head1>.

- Use your health insurance plan's internal complaint and appeal processes to address your concerns.
- Avoid knowingly spreading disease.
- Recognize the reality of risks, the limits of the medical science, and the human fallibility of the healthcare professional.
- Be aware of a healthcare provider's obligation to be reasonably efficient and equitable in providing care to other patients and the community.
- Become knowledgeable about your health plan's coverage and options (when available) including all covered benefits, limitations, and exclusions, rules regarding use of network providers, coverage and referral rules, appropriate processes to secure additional information, and the process to appeal coverage decisions.
- Show respect for other patients and health workers.
- Make a good-faith effort to meet financial obligations.
- Abide by administrative and operational procedures of health plans, healthcare providers, and Government health benefit programs.

Choosing an Insurance Plan

There are a number of official government agencies that help consumers understand their healthcare insurance choices.¹⁰⁸ The U.S. Department of Labor, in particular, recommends ten ways to make your health benefits choices work best for you.¹⁰⁹

1. Your options are important. There are many different types of health benefit plans. Find out which one your employer offers, then check out the plan, or plans, offered. Your employer's human resource office, the health plan administrator, or your union can provide information to help you match your needs and preferences with the available plans. The more information you have, the better your healthcare decisions will be.

2. Reviewing the benefits available. Do the plans offered cover preventive care, well-baby care, vision or dental care? Are there deductibles? Answers to these questions can help determine the out-of-pocket expenses you may

¹⁰⁸ More information about quality across programs is provided at the following AHRQ Web site:

<http://www.ahrq.gov/consumer/qntascii/qnthplan.htm>.

¹⁰⁹ Adapted from the Department of Labor:

<http://www.dol.gov/dol/pwba/public/pubs/health/top10-text.html>.

face. Matching your needs and those of your family members will result in the best possible benefits. Cheapest may not always be best. Your goal is high quality health benefits.

3. Look for quality. The quality of healthcare services varies, but quality can be measured. You should consider the quality of healthcare in deciding among the healthcare plans or options available to you. Not all health plans, doctors, hospitals and other providers give the highest quality care. Fortunately, there is quality information you can use right now to help you compare your healthcare choices. Find out how you can measure quality. Consult the U.S. Department of Health and Human Services publication “Your Guide to Choosing Quality Health Care” on the Internet at www.ahcpr.gov/consumer.

4. Your plan’s summary plan description (SPD) provides a wealth of information. Your health plan administrator can provide you with a copy of your plan’s SPD. It outlines your benefits and your legal rights under the Employee Retirement Income Security Act (ERISA), the federal law that protects your health benefits. It should contain information about the coverage of dependents, what services will require a co-pay, and the circumstances under which your employer can change or terminate a health benefits plan. Save the SPD and all other health plan brochures and documents, along with memos or correspondence from your employer relating to health benefits.

5. Assess your benefit coverage as your family status changes. Marriage, divorce, childbirth or adoption, and the death of a spouse are all life events that may signal a need to change your health benefits. You, your spouse and dependent children may be eligible for a special enrollment period under provisions of the Health Insurance Portability and Accountability Act (HIPAA). Even without life-changing events, the information provided by your employer should tell you how you can change benefits or switch plans, if more than one plan is offered. If your spouse’s employer also offers a health benefits package, consider coordinating both plans for maximum coverage.

6. Changing jobs and other life events can affect your health benefits. Under the Consolidated Omnibus Budget Reconciliation Act (COBRA), you, your covered spouse, and your dependent children may be eligible to purchase extended health coverage under your employer’s plan if you lose your job, change employers, get divorced, or upon occurrence of certain other events. Coverage can range from 18 to 36 months depending on your situation. COBRA applies to most employers with 20 or more workers and

requires your plan to notify you of your rights. Most plans require eligible individuals to make their COBRA election within 60 days of the plan's notice. Be sure to follow up with your plan sponsor if you don't receive notice, and make sure you respond within the allotted time.

7. HIPAA can also help if you are changing jobs, particularly if you have a medical condition. HIPAA generally limits pre-existing condition exclusions to a maximum of 12 months (18 months for late enrollees). HIPAA also requires this maximum period to be reduced by the length of time you had prior "creditable coverage." You should receive a certificate documenting your prior creditable coverage from your old plan when coverage ends.

8. Plan for retirement. Before you retire, find out what health benefits, if any, extend to you and your spouse during your retirement years. Consult with your employer's human resources office, your union, the plan administrator, and check your SPD. Make sure there is no conflicting information among these sources about the benefits you will receive or the circumstances under which they can change or be eliminated. With this information in hand, you can make other important choices, like finding out if you are eligible for Medicare and Medigap insurance coverage.

9. Know how to file an appeal if your health benefits claim is denied. Understand how your plan handles grievances and where to make appeals of the plan's decisions. Keep records and copies of correspondence. Check your health benefits package and your SPD to determine who is responsible for handling problems with benefit claims. Contact PWBA for customer service assistance if you are unable to obtain a response to your complaint.

10. You can take steps to improve the quality of the healthcare and the health benefits you receive. Look for and use things like Quality Reports and Accreditation Reports whenever you can. Quality reports may contain consumer ratings -- how satisfied consumers are with the doctors in their plan, for instance-- and clinical performance measures -- how well a healthcare organization prevents and treats illness. Accreditation reports provide information on how accredited organizations meet national standards, and often include clinical performance measures. Look for these quality measures whenever possible. Consult "Your Guide to Choosing Quality Health Care" on the Internet at www.ahcpr.gov/consumer.

Medicare and Medicaid

Illness strikes both rich and poor families. For low-income families, Medicaid is available to defer the costs of treatment. The Health Care Financing Administration (HCFA) administers Medicare, the nation's largest health insurance program, which covers 39 million Americans. In the following pages, you will learn the basics about Medicare insurance as well as useful contact information on how to find more in-depth information about Medicaid.¹¹⁰

Who is Eligible for Medicare?

Generally, you are eligible for Medicare if you or your spouse worked for at least 10 years in Medicare-covered employment and you are 65 years old and a citizen or permanent resident of the United States. You might also qualify for coverage if you are under age 65 but have a disability or End-Stage Renal disease (permanent kidney failure requiring dialysis or transplant). Here are some simple guidelines:

You can get Part A at age 65 without having to pay premiums if:

- You are already receiving retirement benefits from Social Security or the Railroad Retirement Board.
- You are eligible to receive Social Security or Railroad benefits but have not yet filed for them.
- You or your spouse had Medicare-covered government employment.

If you are under 65, you can get Part A without having to pay premiums if:

- You have received Social Security or Railroad Retirement Board disability benefit for 24 months.
- You are a kidney dialysis or kidney transplant patient.

Medicare has two parts:

- Part A (Hospital Insurance). Most people do not have to pay for Part A.
- Part B (Medical Insurance). Most people pay monthly for Part B.

¹¹⁰ This section has been adapted from the Official U.S. Site for Medicare Information: <http://www.medicare.gov/Basics/Overview.asp>.

Part A (Hospital Insurance)

Helps Pay For: Inpatient hospital care, care in critical access hospitals (small facilities that give limited outpatient and inpatient services to people in rural areas) and skilled nursing facilities, hospice care, and some home healthcare.

Cost: Most people get Part A automatically when they turn age 65. You do not have to pay a monthly payment called a premium for Part A because you or a spouse paid Medicare taxes while you were working.

If you (or your spouse) did not pay Medicare taxes while you were working and you are age 65 or older, you still may be able to buy Part A. If you are not sure you have Part A, look on your red, white, and blue Medicare card. It will show "Hospital Part A" on the lower left corner of the card. You can also call the Social Security Administration toll free at 1-800-772-1213 or call your local Social Security office for more information about buying Part A. If you get benefits from the Railroad Retirement Board, call your local RRB office or 1-800-808-0772. For more information, call your Fiscal Intermediary about Part A bills and services. The phone number for the Fiscal Intermediary office in your area can be obtained from the following Web site: <http://www.medicare.gov/Contacts/home.asp>.

Part B (Medical Insurance)

Helps Pay For: Doctors, services, outpatient hospital care, and some other medical services that Part A does not cover, such as the services of physical and occupational therapists, and some home healthcare. Part B helps pay for covered services and supplies when they are medically necessary.

Cost: As of 2001, you pay the Medicare Part B premium of \$50.00 per month. In some cases this amount may be higher if you did not choose Part B when you first became eligible at age 65. The cost of Part B may go up 10% for each 12-month period that you were eligible for Part B but declined coverage, except in special cases. You will have to pay the extra 10% cost for the rest of your life.

Enrolling in Part B is your choice. You can sign up for Part B anytime during a 7-month period that begins 3 months before you turn 65. Visit your local Social Security office, or call the Social Security Administration at 1-800-772-1213 to sign up. If you choose to enroll in Part B, the premium is usually taken out of your monthly Social Security, Railroad Retirement, or Civil Service Retirement payment. If you do not receive any of the above

payments, Medicare sends you a bill for your part B premium every 3 months. You should receive your Medicare premium bill in the mail by the 10th of the month. If you do not, call the Social Security Administration at 1-800-772-1213, or your local Social Security office. If you get benefits from the Railroad Retirement Board, call your local RRB office or 1-800-808-0772. For more information, call your Medicare carrier about bills and services. The phone number for the Medicare carrier in your area can be found at the following Web site: <http://www.medicare.gov/Contacts/home.asp>. You may have choices in how you get your healthcare including the Original Medicare Plan, Medicare Managed Care Plans (like HMOs), and Medicare Private Fee-for-Service Plans.

Medicaid

Medicaid is a joint federal and state program that helps pay medical costs for some people with low incomes and limited resources. Medicaid programs vary from state to state. People on Medicaid may also get coverage for nursing home care and outpatient prescription drugs which are not covered by Medicare. You can find more information about Medicaid on the HCFA.gov Web site at <http://www.hcfa.gov/medicaid/medicaid.htm>.

States also have programs that pay some or all of Medicare's premiums and may also pay Medicare deductibles and coinsurance for certain people who have Medicare and a low income. To qualify, you must have:

- Part A (Hospital Insurance),
- Assets, such as bank accounts, stocks, and bonds that are not more than \$4,000 for a single person, or \$6,000 for a couple, and
- A monthly income that is below certain limits.

For more information on these programs, look at the Medicare Savings Programs brochure, <http://www.medicare.gov/Library/PDFNavigation/PDFInterim.asp?Language=English&Type=Pub&PubID=10126>. There are also Prescription Drug Assistance Programs available. Find information on these programs which offer discounts or free medications to individuals in need at <http://www.medicare.gov/Prescription/Home.asp>.

Financial Assistance for Cancer Care¹¹¹

Cancer imposes heavy economic burdens on both patients and their families. For many people, a portion of medical expenses is paid by their health insurance plan. For individuals who do not have health insurance or who need financial assistance to cover health care costs, resources are available, including government-sponsored programs and services supported by voluntary organizations.

Cancer patients and their families should discuss any concerns they may have about health care costs with their physician, medical social worker, or the business office of their hospital or clinic.

The organizations and resources listed below may offer financial assistance. Organizations that provide publications in Spanish or have Spanish-speaking staff have been identified.

- The American Cancer Society (ACS) office can provide the telephone number of the local ACS office serving your area. The local ACS office may offer reimbursement for expenses related to cancer treatment including transportation, medicine, and medical supplies. The ACS also offers programs that help cancer patients, family members, and friends cope with the emotional challenges they face. Some publications are available in Spanish. Spanish-speaking staff are available. Telephone: 1-800-ACS-2345 (1-800-227-2345). Web site: <http://www.cancer.org>
- The *AVONCares* Program for Medically Underserved Women provides financial assistance and relevant education and support to low income, under- and uninsured, underserved women throughout the country in need of diagnostic and/or related services (transportation, child care, and social support) for the treatment of breast, cervical, and ovarian cancers. Telephone: 1-800-813-HOPE (1-800-813-4673). Web site: <http://www.cancercare.org>.

Community voluntary agencies and service organizations such as the Salvation Army, Lutheran Social Services, Jewish Social Services, Catholic Charities, and the Lions Club may offer help. These organizations are listed in your local phone directory. Some churches and synagogues may provide financial help or services to their members.

Fundraising is another mechanism to consider. Some patients find that friends, family, and community members are willing to contribute

¹¹¹ Adapted from the NCI: http://cis.nci.nih.gov/fact/8_3.htm.

financially if they are aware of a difficult situation. Contact your local library for information about how to organize fundraising efforts.

General assistance programs provide food, housing, prescription drugs, and other medical expenses for those who are not eligible for other programs. Funds are often limited. Information can be obtained by contacting your state or local Department of Social Services; this number is found in the local telephone directory.

Hill-Burton is a program through which hospitals receive construction funds from the Federal Government. Hospitals that receive Hill-Burton funds are required by law to provide some services to people who cannot afford to pay for their hospitalization. Information about which facilities are part of this program is available by calling the toll-free number or visiting the Web site shown below. A brochure about the program is available in Spanish. Telephone: 1-800-638-0742. Web site: <http://www.hrsa.gov/osp/dfcr/obtain/consfaq.htm>.

Income Tax Deductions

Medical costs that are not covered by insurance policies sometimes can be deducted from annual income before taxes. Examples of tax deductible expenses might include mileage for trips to and from medical appointments, out-of-pocket costs for treatment, prescription drugs or equipment, and the cost of meals during lengthy medical visits. The local Internal Revenue Service office, tax consultants, or certified public accountants can determine medical costs that are tax deductible. These telephone numbers are available in the local telephone directory. Web site: <http://www.irs.ustreas.gov>.

The Patient Advocate Foundation

The Patient Advocate Foundation (PAF) is a national nonprofit organization that provides education, legal counseling, and referrals to cancer patients and survivors concerning managed care, insurance, financial issues, job discrimination, and debt crisis matters. Telephone: 1-800-532-5274. **Web site:** <http://www.patientadvocate.org>.

Patient Assistance Programs are offered by some pharmaceutical manufacturers to help pay for medications. To learn whether a specific drug might be available at reduced cost through such a program, talk with a physician or a medical social worker.

Transportation

There are nonprofit organizations that arrange free or reduced cost air transportation for cancer patients going to or from cancer treatment centers. Financial need is not always a requirement. To find out about these programs, talk with a medical social worker. Ground transportation services may be offered or mileage reimbursed through the local ACS or your state or local Department of Social Services.

Veterans Benefits

Eligible veterans and their dependents may receive cancer treatment at a Veterans Administration Medical Center. Treatment for service-connected conditions is provided, and treatment for other conditions may be available based on the veteran's financial need. Some publications are available in Spanish. Spanish-speaking staff are available in some offices. Telephone: 1-877-222-VETS. Web site: <http://www.va.gov/vbs/health>.

NORD's Medication Assistance Programs

Finally, the National Organization for Rare Disorders, Inc. (NORD) administers medication programs sponsored by humanitarian-minded pharmaceutical and biotechnology companies to help uninsured or underinsured individuals secure life-saving or life-sustaining drugs.¹¹² NORD programs ensure that certain vital drugs are available "to those individuals whose income is too high to qualify for Medicaid but too low to pay for their prescribed medications." The program has standards for fairness, equity, and unbiased eligibility. It currently covers some 14 programs for nine pharmaceutical companies. NORD also offers early access programs for investigational new drugs (IND) under the approved "Treatment INDs" programs of the Food and Drug Administration (FDA). In these programs, a limited number of individuals can receive investigational drugs that have yet to be approved by the FDA. These programs are generally designed for rare diseases or disorders. For more information, visit www.rarediseases.org.

¹¹² Adapted from NORD: http://www.rarediseases.org/cgi-bin/nord/progserv#patient?id=rPIzL9oD&mv_pc=30.

Additional Resources

In addition to the references already listed in this chapter, you may need more information on health insurance, hospitals, or the healthcare system in general. The NIH has set up an excellent guidance Web site that addresses these and other issues. Topics include:¹¹³

- Health Insurance:
<http://www.nlm.nih.gov/medlineplus/healthinsurance.html>
- Health Statistics:
<http://www.nlm.nih.gov/medlineplus/healthstatistics.html>
- HMO and Managed Care:
<http://www.nlm.nih.gov/medlineplus/managedcare.html>
- Hospice Care: <http://www.nlm.nih.gov/medlineplus/hospicecare.html>
- Medicaid: <http://www.nlm.nih.gov/medlineplus/medicaid.html>
- Medicare: <http://www.nlm.nih.gov/medlineplus/medicare.html>
- Nursing Homes and Long-term Care:
<http://www.nlm.nih.gov/medlineplus/nursinghomes.html>
- Patient's Rights, Confidentiality, Informed Consent, Ombudsman Programs, Privacy and Patient Issues:
<http://www.nlm.nih.gov/medlineplus/patientissues.html>
- Veteran's Health, Persian Gulf War, Gulf War Syndrome, Agent Orange:
<http://www.nlm.nih.gov/medlineplus/veteranshealth.html>

Vocabulary Builder

Bilirubin: A bile pigment that is a degradation product of HEME. [NIH]

Endoscopic retrograde cholangiopancreatography: ERCP. A procedure to x-ray the pancreatic duct, hepatic duct, common bile duct, duodenal papilla, and gallbladder. In this procedure, a thin, lighted tube (endoscope) is passed through the mouth and down into the first part of the small intestine (duodenum). A smaller tube (catheter) is then inserted through the endoscope into the bile and pancreatic ducts. A dye is injected through the catheter into the ducts, and an x-ray is taken. [NIH]

ERCP: Endoscopic retrograde cholangiopancreatography (en-do-SKAH-pik

¹¹³ You can access this information at:

<http://www.nlm.nih.gov/medlineplus/healthsystem.html>.

RET-ro-grade ko-LAN-jee-o-PAN-kree-a-TAW-gra-fee). A procedure to x-ray the bile and pancreatic ducts. In this procedure, a thin, lighted tube (endoscope) is passed through the mouth and down into the first part of the small intestine (duodenum). A smaller tube (catheter) is then inserted through the endoscope into the bile and pancreatic ducts. A dye is injected through the catheter into the ducts, and an x-ray is taken. [NIH]

Retrograde: 1. moving backward or against the usual direction of flow. 2. degenerating, deteriorating, or catabolic. [EU]

Toxins: Poisons produced by certain animals, plants, or bacteria. [NIH]

ONLINE GLOSSARIES

The Internet provides access to a number of free-to-use medical dictionaries and glossaries. The National Library of Medicine has compiled the following list of online dictionaries:

- ADAM Medical Encyclopedia (A.D.A.M., Inc.), comprehensive medical reference: <http://www.nlm.nih.gov/medlineplus/encyclopedia.html>
- MedicineNet.com Medical Dictionary (MedicineNet, Inc.):
<http://www.medterms.com/Script/Main/hp.asp>
- Merriam-Webster Medical Dictionary (Inteli-Health, Inc.):
<http://www.intelihealth.com/IH/>
- Multilingual Glossary of Technical and Popular Medical Terms in Eight European Languages (European Commission) - Danish, Dutch, English, French, German, Italian, Portuguese, and Spanish:
<http://allserv.rug.ac.be/~rvdstich/eugloss/welcome.html>
- On-line Medical Dictionary (CancerWEB):
<http://www.graylab.ac.uk/omd/>
- Technology Glossary (National Library of Medicine) - Health Care Technology: <http://www.nlm.nih.gov/nichsr/ta101/ta10108.htm>
- Terms and Definitions (Office of Rare Diseases):
http://rarediseases.info.nih.gov/ord/glossary_a-e.html

Beyond these, MEDLINEplus contains a very user-friendly encyclopedia covering every aspect of medicine (licensed from A.D.A.M., Inc.). The ADAM Medical Encyclopedia Web site address is <http://www.nlm.nih.gov/medlineplus/encyclopedia.html>. ADAM is also available on commercial Web sites such as [drkoop.com](http://www.drkoop.com/) (<http://www.drkoop.com/>) and Web MD (http://my.webmd.com/adam/asset/adam_disease_articles/a_to_z/a). Topics of interest can be researched by using keywords before continuing elsewhere, as these basic definitions and concepts will be useful in more advanced areas of research. You may choose to print various pages specifically relating to adult primary liver cancer and keep them on file. The NIH, in particular, suggests that patients with adult primary liver cancer visit the following Web sites in the ADAM Medical Encyclopedia:

- **Basic Guidelines for Adult Primary Liver Cancer**

Cholangiocarcinoma

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/000291.htm>

Hepatocellular carcinoma

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/000280.htm>

- **Signs & Symptoms for Adult Primary Liver Cancer**

Abdominal pain

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003120.htm>

Fever

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003090.htm>

Itching

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003217.htm>

Jaundice

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003243.htm>

Loss of appetite

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003121.htm>

Stress

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003211.htm>

Weight loss

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003107.htm>

- **Diagnostics and Tests for Adult Primary Liver Cancer**

5'-N'Tase

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003575.htm>

Abdominal CT scan

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003789.htm>

Abdominal ultrasound

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003777.htm>

Alpha-1 antitrypsin

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003715.htm>

ALT

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003473.htm>

AST

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003472.htm>

Bilirubin

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003479.htm>

Bilirubin; urine

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003595.htm>

Biopsy

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003894.htm>

CT

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003330.htm>

Cysts

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003240.htm>

Delta-ALA

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003457.htm>

Endoscopic retrograde cholangiopancreatography

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003893.htm>

ERCP

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003893.htm>

Gall bladder radionuclide scan

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003826.htm>

Gamma-glutamyl transpeptidase

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003458.htm>

Leucine aminopeptidase - serum

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003559.htm>

Leucine aminopeptidase - urine

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003617.htm>

Liver biopsy

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003895.htm>

Liver function tests

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003436.htm>

Liver scan

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003825.htm>

PBG

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003596.htm>

Percutaneous transhepatic cholangiogram (PTCA)

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003820.htm>

Porphyryns; urine

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003614.htm>

Serum alpha-fetoprotein

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/003573.htm>

- **Background Topics for Adult Primary Liver Cancer**

Bile

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002237.htm>

Cancer - support group

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002166.htm>

Chemotherapy

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002324.htm>

Chronic

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002312.htm>

Cytology

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002323.htm>

Endoscopic

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002360.htm>

Hepatic

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002378.htm>

Incidence

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002387.htm>

Liver disease - support group

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002182.htm>

Metastasis

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002260.htm>

Physical examination

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002274.htm>

Primary

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002312.htm>

Radiation therapy

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/001918.htm>

Support group

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002150.htm>

Toxins

Web site:

<http://www.nlm.nih.gov/medlineplus/ency/article/002331.htm>

Online Dictionary Directories

The following are additional online directories compiled by the National Library of Medicine, including a number of specialized medical dictionaries and glossaries:

- Medical Dictionaries: Medical & Biological (World Health Organization):
<http://www.who.int/hlt/virtuallibrary/English/diction.htm#Medical>
- MEL-Michigan Electronic Library List of Online Health and Medical Dictionaries (Michigan Electronic Library):
<http://mel.lib.mi.us/health/health-dictionaries.html>
- Patient Education: Glossaries (DMOZ Open Directory Project):
http://dmoz.org/Health/Education/Patient_Education/Glossaries/
- Web of Online Dictionaries (Bucknell University):
<http://www.yourdictionary.com/diction5.html#medicine>

ADULT PRIMARY LIVER CANCER GLOSSARY

The following is a complete glossary of terms used in this sourcebook. The definitions are derived from official public sources including the National Institutes of Health [NIH] and the European Union [EU]. After this glossary, we list a number of additional hardbound and electronic glossaries and dictionaries that you may wish to consult.

Abdomen: The part of the body that contains the pancreas, stomach, intestines, liver, gallbladder, and other organs. [NIH]

Abdominal: Having to do with the abdomen, which is the part of the body between the chest and the hips that contains the pancreas, stomach, intestines, liver, gallbladder, and other organs. [NIH]

Abscess: A localized collection of pus caused by suppuration buried in tissues, organs, or confined spaces. [EU]

Adenocarcinoma: Cancer that begins in cells that line certain internal organs and that have glandular (secretory) properties. [NIH]

Adjuvant: A substance which aids another, such as an auxiliary remedy; in immunology, nonspecific stimulator (e.g., BCG vaccine) of the immune response. [EU]

AFP: Alpha-fetoprotein. A protein normally produced by a developing fetus. AFP levels are usually undetectable in the blood of healthy nonpregnant adults. An elevated level of AFP suggests the presence of either a primary liver cancer or germ cell tumor. [NIH]

Alanine: A non-essential amino acid that occurs in high levels in its free state in plasma. It is produced from pyruvate by transamination. It is involved in sugar and acid metabolism, increases immunity, and provides energy for muscle tissue, brain, and the central nervous system. [NIH]

Alimentary: Pertaining to food or nutritive material, or to the organs of digestion. [EU]

Alkaline: Having the reactions of an alkali. [EU]

Alkalosis: A pathologic condition resulting from accumulation of base, or from loss of acid without comparable loss of base in the body fluids, and characterized by decrease in hydrogen ion concentration (increase in pH). [EU]

American Cancer Society: A voluntary organization concerned with the prevention and treatment of cancer through education and research. [NIH]

Androstenedione: A steroid with androgenic properties that is produced in the testis, ovary, and adrenal cortex. It is a precursor to testosterone and

other androgenic hormones. [NIH]

Anesthesia: Loss of feeling or awareness. Local anesthetics cause loss of feeling in a part of the body. General anesthetics put the person to sleep. [NIH]

Angiogenesis: Blood vessel formation. Tumor angiogenesis is the growth of blood vessels from surrounding tissue to a solid tumor. This is caused by the release of chemicals by the tumor. [NIH]

Angiography: A procedure to x-ray blood vessels. The blood vessels can be seen because of an injection of a dye that shows up in the x-ray pictures. [NIH]

Angiosarcoma: A type of cancer that begins in the lining of blood vessels. [NIH]

Antiangiogenesis: Prevention of the growth of new blood vessels. [NIH]

Antibody: A type of protein made by certain white blood cells in response to a foreign substance (antigen). Each antibody can bind to only a specific antigen. The purpose of this binding is to help destroy the antigen. Antibodies can work in several ways, depending on the nature of the antigen. Some antibodies destroy antigens directly. Others make it easier for white blood cells to destroy the antigen. [NIH]

Antiemetic: An agent that prevents or alleviates nausea and vomiting. Also antinauseant. [EU]

Antigen: Any substance which is capable, under appropriate conditions, of inducing a specific immune response and of reacting with the products of that response, that is, with specific antibody or specifically sensitized T-lymphocytes, or both. Antigens may be soluble substances, such as toxins and foreign proteins, or particulate, such as bacteria and tissue cells; however, only the portion of the protein or polysaccharide molecule known as the antigenic determinant (q.v.) combines with antibody or a specific receptor on a lymphocyte. Abbreviated Ag. [EU]

Antineoplastons: Substances isolated from normal human blood and urine being tested as a type of treatment for some tumors and AIDS. [NIH]

Arterial: Pertaining to an artery or to the arteries. [EU]

Arteries: The vessels carrying blood away from the heart. [NIH]

Aspiration: Removal of fluid from a lump, often a cyst, with a needle and a syringe. [NIH]

Aspirin: A drug that reduces pain, fever, inflammation, and blood clotting. Aspirin belongs to the family of drugs called nonsteroidal anti-inflammatory agents. It is also being studied in cancer prevention. [NIH]

Astrocytoma: A tumor that begins in the brain or spinal cord in small, star-shaped cells called astrocytes. [NIH]

Autologous: Taken from an individual's own tissues, cells, or DNA. [NIH]

Bacteremia: The presence of viable bacteria circulating in the blood. Fever, chills, tachycardia, and tachypnea are common acute manifestations of bacteremia. The majority of cases are seen in already hospitalized patients, most of whom have underlying diseases or procedures which render their bloodstreams susceptible to invasion. [NIH]

Bacteria: A large group of single-cell microorganisms. Some cause infections and disease in animals and humans. The singular of bacteria is bacterium. [NIH]

Bereavement: Refers to the whole process of grieving and mourning and is associated with a deep sense of loss and sadness. [NIH]

Bile: A fluid made by the liver and stored in the gallbladder. Bile is excreted into the small intestine where it helps digest fat. [NIH]

Bilirubin: A bile pigment that is a degradation product of heme. [NIH]

Biopsy: The removal of cells or tissues for examination under a microscope. When only a sample of tissue is removed, the procedure is called an incisional biopsy or core biopsy. When an entire tumor or lesion is removed, the procedure is called an excisional biopsy. When a sample of tissue or fluid is removed with a needle, the procedure is called a needle biopsy or fine-needle aspiration. [NIH]

Bypass: A surgical procedure in which the doctor creates a new pathway for the flow of body fluids. [NIH]

Calcium: A mineral found in teeth, bones, and other body tissues. [NIH]

Capecitabine: An anticancer drug that belongs to the family of drugs called antimetabolites. [NIH]

Capsules: Hard or soft soluble containers used for the oral administration of medicine. [NIH]

Carbohydrate: An aldehyde or ketone derivative of a polyhydric alcohol, particularly of the pentahydric and hexahydric alcohols. They are so named because the hydrogen and oxygen are usually in the proportion to form water, (CH₂O)_n. The most important carbohydrates are the starches, sugars, celluloses, and gums. They are classified into mono-, di-, tri-, poly- and heterosaccharides. [EU]

Carboplatin: An anticancer drug that belongs to the family of drugs called platinum compounds. [NIH]

Carcinogen: Any substance that causes cancer. [NIH]

Carcinogenesis: The process by which normal cells are transformed into cancer cells. [NIH]

Carcinogenic: Producing carcinoma. [EU]

Carcinoma: Cancer that begins in the skin or in tissues that line or cover

internal organs. [NIH]

Cardiology: The study of the heart, its physiology, and its functions. [NIH]

Catheter: A flexible tube used to deliver fluids into or withdraw fluids from the body. [NIH]

Cell: The individual unit that makes up all of the tissues of the body. All living things are made up of one or more cells. [NIH]

Cerebrospinal: Pertaining to the brain and spinal cord. [EU]

Cervical: Relating to the neck, or to the neck of any organ or structure. Cervical lymph nodes are located in the neck; cervical cancer refers to cancer of the uterine cervix, which is the lower, narrow end (the "neck") of the uterus. [NIH]

Charities: Social welfare organizations with programs designed to assist individuals in times of need. [NIH]

Chemoembolization: A procedure in which the blood supply to the tumor is blocked surgically or mechanically, and anticancer drugs are administered directly into the tumor. This permits a higher concentration of drug to be in contact with the tumor for a longer period of time. [NIH]

Chemotherapy: Treatment with anticancer drugs. [NIH]

Cholangitis: Inflammation of a bile duct. [EU]

Cholesterol: The principal sterol of all higher animals, distributed in body tissues, especially the brain and spinal cord, and in animal fats and oils. [NIH]

Chromosomal: Pertaining to chromosomes. [EU]

Chromosome: Part of a cell that contains genetic information. Except for sperm and eggs, all human cells contain 46 chromosomes. [NIH]

Chronic: A disease or condition that persists or progresses over a long period of time. [NIH]

Chronotherapy: The adaptation of the administration of drugs to circadian rhythms. The concept is based on the response of biological functions to time-related events, such as the low point in epinephrine levels between 10 p.m. and 4 a.m. or the elevated histamine levels between midnight and 4 a.m. The treatment is aimed at supporting normal rhythms or modifying therapy based on known variations in body rhythms. While chronotherapy is commonly used in cancer chemotherapy, it is not restricted to cancer therapy or to chemotherapy. [NIH]

Cirrhosis: A type of chronic, progressive liver disease. [NIH]

Cisplatin: An anticancer drug that belongs to the family of drugs called platinum compounds. [NIH]

Coagulation: 1. the process of clot formation. 2. in colloid chemistry, the

solidification of a sol into a gelatinous mass; an alteration of a disperse phase or of a dissolved solid which causes the separation of the system into a liquid phase and an insoluble mass called the clot or curd. Coagulation is usually irreversible. 3. in surgery, the disruption of tissue by physical means to form an amorphous residuum, as in electrocoagulation and photocoagulation. [EU]

Collagen: A fibrous protein found in cartilage and other connective tissue. [NIH]

Colon: The long, coiled, tubelike organ that removes water from digested food. The remaining material, solid waste called stool, moves through the colon to the rectum and leaves the body through the anus. [NIH]

Colorectal: Having to do with the colon or the rectum. [NIH]

Concomitant: Accompanying; accessory; joined with another. [EU]

Contamination: The soiling or pollution by inferior material, as by the introduction of organisms into a wound, or sewage into a stream. [EU]

Contraceptive: An agent that diminishes the likelihood of or prevents conception. [EU]

Cryosurgery: Treatment performed with an instrument that freezes and destroys abnormal tissues. This procedure is a form of cryotherapy. [NIH]

Cryotherapy: Any method that uses cold temperature to treat disease. [NIH]

Curative: Tending to overcome disease and promote recovery. [EU]

Cytokines: A class of substances that are produced by cells of the immune system and can affect the immune response. Cytokines can also be produced in the laboratory by recombinant DNA technology and given to people to affect immune responses. [NIH]

Degenerative: Undergoing degeneration : tending to degenerate; having the character of or involving degeneration; causing or tending to cause degeneration. [EU]

Dehydroepiandrosterone: DHEA. A substance that is being studied as a cancer prevention drug. It belongs to the family of drugs called steroids. [NIH]

Dentists: Individuals licensed to practice dentistry. [NIH]

Dexamethasone: A synthetic steroid (similar to steroid hormones produced naturally in the adrenal gland). Dexamethasone is used to treat leukemia and lymphoma and may be used to treat some of the problems caused by other cancers and their treatment. [NIH]

DHEA: Dehydroepiandrosterone. A substance that is being studied as a cancer prevention drug. It belongs to the family of drugs called steroids. [NIH]

Diaphragm: The thin muscle below the lungs and heart that separates the chest from the abdomen. [NIH]

Diarrhea: Passage of excessively liquid or excessively frequent stools. [NIH]

Doxorubicin: An anticancer drug that belongs to the family of drugs called antitumor antibiotics. It is an anthracycline. [NIH]

Dysplasia: Cells that look abnormal under a microscope but are not cancer. [NIH]

Echinacea: A genus of perennial herbs used topically and internally. It contains echinacoside, glycosides, inulin, isobutyl amides, resin, and sesquiterpenes. [NIH]

Endocrinologist: A doctor that specializes in diagnosing and treating hormone disorders. [NIH]

Endocrinology: A subspecialty of internal medicine concerned with the metabolism, physiology, and disorders of the endocrine system. [NIH]

Endometrial: Having to do with the endometrium (the layer of tissue that lines the uterus). [NIH]

Endoscopic retrograde cholangiopancreatography: ERCP. A procedure to x-ray the pancreatic duct, hepatic duct, common bile duct, duodenal papilla, and gallbladder. In this procedure, a thin, lighted tube (endoscope) is passed through the mouth and down into the first part of the small intestine (duodenum). A smaller tube (catheter) is then inserted through the endoscope into the bile and pancreatic ducts. A dye is injected through the catheter into the ducts, and an x-ray is taken. [NIH]

Enzyme: A protein that speeds up chemical reactions in the body. [NIH]

Epidemiological: Relating to, or involving epidemiology. [EU]

Epidural: The space between the wall of the spinal canal and the covering of the spinal cord. An epidural injection is given into this space. [NIH]

Epigastralgia: Pain in the epigastrium. [EU]

Epithelial: Refers to the cells that line the internal and external surfaces of the body. [NIH]

ERCP: Endoscopic retrograde cholangiopancreatography (en-do-SKAH-pik RET-ro-grade ko-LAN-jee-o-PAN-kree-a-TAW-gra-fee). A procedure to x-ray the bile and pancreatic ducts. In this procedure, a thin, lighted tube (endoscope) is passed through the mouth and down into the first part of the small intestine (duodenum). A smaller tube (catheter) is then inserted through the endoscope into the bile and pancreatic ducts. A dye is injected through the catheter into the ducts, and an x-ray is taken. [NIH]

Esophageal: Having to do with the esophagus, the muscular tube through which food passes from the throat to the stomach. [NIH]

Esophagus: The muscular tube through which food passes from the throat to the stomach. [NIH]

Ethanol: A clear, colorless liquid rapidly absorbed from the gastrointestinal

tract and distributed throughout the body. It has bactericidal activity and is used often as a topical disinfectant. It is widely used as a solvent and preservative in pharmaceutical preparations as well as serving as the primary ingredient in alcoholic beverages. [NIH]

Etoposide: An anticancer drug that is a podophyllotoxin derivative and belongs to the family of drugs called mitotic inhibitors. [NIH]

Extravasation: A discharge or escape, as of blood, from a vessel into the tissues. [EU]

Extremity: A limb; an arm or leg (membrum); sometimes applied specifically to a hand or foot. [EU]

Fetus: The developing offspring from 7 to 8 weeks after conception until birth. [NIH]

Fibrosis: The growth of fibrous tissue. [NIH]

Fluorescence: The property of emitting radiation while being irradiated. The radiation emitted is usually of longer wavelength than that incident or absorbed, e.g., a substance can be irradiated with invisible radiation and emit visible light. X-ray fluorescence is used in diagnosis. [NIH]

Fluorouracil: An anticancer drug that belongs to the family of drugs called antimetabolites. [NIH]

Gallbladder: The pear-shaped organ that sits below the liver. Bile is concentrated and stored in the gallbladder. [NIH]

Gastric: Having to do with the stomach. [NIH]

Gastrointestinal: Refers to the stomach and intestines. [NIH]

Gemcitabine: An anticancer drug that belongs to the family of drugs called antimetabolites. [NIH]

Genitourinary: Pertaining to the genital and urinary organs; urogenital; urinosexual. [EU]

Ginseng: An herb with a root that has been used in some cultures to treat certain medical problems. It may have anticancer effects. [NIH]

Glioma: A cancer of the brain that comes from glial, or supportive, cells. [NIH]

Gonadal: Pertaining to a gonad. [EU]

Grade: The grade of a tumor depends on how abnormal the cancer cells look under a microscope and how quickly the tumor is likely to grow and spread. Grading systems are different for each type of cancer. [NIH]

Groin: The area where the thigh meets the abdomen. [NIH]

Hematoma: An extravasation of blood localized in an organ, space, or tissue. [NIH]

Hemobilia: Hemorrhage in or through the biliary tract, due to trauma, inflammation, cholelithiasis, vascular disease, or neoplasms. [NIH]

Heparin: A drug that helps prevent blood clots from forming. It belongs to the family of drugs called anticoagulants (blood thinners). [NIH]

Hepatic: Refers to the liver. [NIH]

Hepatitis: Inflammation of the liver. [NIH]

Hepatobiliary: Pertaining to the liver and the bile or the biliary ducts. [EU]

Hepatoblastoma: A type of liver tumor that occurs in infants and children. [NIH]

Hepatocyte: A liver cell. [NIH]

Hepatoma: A liver tumor. [NIH]

Hybridization: The genetic process of crossbreeding to produce a hybrid. Hybrid nucleic acids can be formed by nucleic acid hybridization of DNA and RNA molecules. Protein hybridization allows for hybrid proteins to be formed from polypeptide chains. [NIH]

Hydroxyurea: An anticancer drug that belongs to the family of drugs called antimetabolites. [NIH]

Hypercalcemia: Abnormally high blood calcium. [NIH]

Hypertension: Abnormally high blood pressure. [NIH]

Hyperthermia: A type of treatment in which body tissue is exposed to high temperatures to damage and kill cancer cells or to make cancer cells more sensitive to the effects of radiation and certain anticancer drugs. [NIH]

Hypervascular: Having a large number of blood vessels. [NIH]

Hypoglycemia: Abnormally low blood sugar [NIH]

Immunization: The induction of immunity. [EU]

Immunodeficiency: The decreased ability of the body to fight infection and disease. [NIH]

Immunology: The study of the body's immune system. [NIH]

Immunotherapy: Treatment to stimulate or restore the ability of the immune system to fight infection and disease. Also used to lessen side effects that may be caused by some cancer treatments. Also called biological therapy or biological response modifier (BRM) therapy. [NIH]

Induction: The act or process of inducing or causing to occur, especially the production of a specific morphogenetic effect in the developing embryo through the influence of evocators or organizers, or the production of anaesthesia or unconsciousness by use of appropriate agents. [EU]

Infarction: 1. the formation of an infarct. 2. an infarct. [EU]

Inflammation: A response of redness, swelling, pain, and a feeling of heat

in certain areas which is meant to protect tissues affected by injury or disease. [NIH]

Infusion: A method of putting fluids, including drugs, into the bloodstream. Also called intravenous infusion. [NIH]

Inoperable: Not suitable to be operated upon. [EU]

Interferon: A biological response modifier (a substance that can improve the body's natural response to disease). Interferons interfere with the division of cancer cells and can slow tumor growth. There are several types of interferons, including interferon-alpha, -beta, and -gamma. These substances are normally produced by the body. They are also made in the laboratory for use in treating cancer and other diseases. [NIH]

Intermittent: Occurring at separated intervals; having periods of cessation of activity. [EU]

Intracellular: Inside a cell. [NIH]

Intrahepatic: Within the liver. [NIH]

Intrathecal: Describes the fluid-filled space between the thin layers of tissue that cover the brain and spinal cord. Drugs can be injected into the fluid or a sample of the fluid can be removed for testing. [NIH]

Intravenous: IV. Into a vein. [NIH]

Invasive: 1. having the quality of invasiveness. 2. involving puncture or incision of the skin or insertion of an instrument or foreign material into the body; said of diagnostic techniques. [EU]

Iodine: A nonmetallic element of the halogen group that is represented by the atomic symbol I, atomic number 53, and atomic weight of 126.90. It is a nutritionally essential element, especially important in thyroid hormone synthesis. In solution, it has anti-infective properties and is used topically. [NIH]

Irinotecan: An anticancer drug that belongs to a family of anticancer drugs called topoisomerase inhibitors. It is a camptothecin analogue. Also called CPT 11. [NIH]

Jaundice: A condition in which the skin and the whites of the eyes become yellow, urine darkens, and stool becomes clay colored. Jaundice occurs when the liver is not working properly or when a bile duct is blocked. [NIH]

Laparoscopy: The insertion of a thin, lighted tube (called a laparoscope) through the abdominal wall to inspect the inside of the abdomen and remove tissue samples. [NIH]

Laparotomy: A surgical incision made in the wall of the abdomen. [NIH]

Larynx: The area of the throat containing the vocal cords and used for breathing, swallowing, and talking. Also called the voice box. [NIH]

Lesion: An area of abnormal tissue change. [NIH]

Lethal: Deadly, fatal. [EU]

LH: A small glycoprotein hormone secreted by the anterior pituitary. LH plays an important role in controlling ovulation and in controlling secretion of hormones by the ovaries and testes. [NIH]

Ligament: A band of fibrous tissue that connects bones or cartilages, serving to support and strengthen joints. [EU]

Ligation: The process of tying off blood vessels so that blood cannot flow to a part of the body or to a tumor. [NIH]

Liver: A large, glandular organ located in the upper abdomen. The liver cleanses the blood and aids in digestion by secreting bile. [NIH]

Lobe: A portion of an organ such as the liver, lung, breast, or brain. [NIH]

Lobectomy: The removal of a lobe. [NIH]

Lymph: The almost colorless fluid that travels through the lymphatic system and carries cells that help fight infection and disease. [NIH]

Lymphocyte: A white blood cell. Lymphocytes have a number of roles in the immune system, including the production of antibodies and other substances that fight infection and diseases. [NIH]

Lymphoma: Cancer that arises in cells of the lymphatic system. [NIH]

Malignant: Cancerous; a growth with a tendency to invade and destroy nearby tissue and spread to other parts of the body. [NIH]

Mammogram: An x-ray of the breast. [NIH]

Mammography: The use of x-rays to create a picture of the breast. [NIH]

Mediastinum: The area between the lungs. The organs in this area include the heart and its large blood vessels, the trachea, the esophagus, the bronchi, and lymph nodes. [NIH]

Megestrol: A drug that belongs to the group of hormones called progestins, used as hormone therapy to block estrogen and to suppress the effects of estrogen and androgens. It is also used to stimulate the appetite in people with cancer. [NIH]

Melanoma: A form of skin cancer that arises in melanocytes, the cells that produce pigment. Melanoma usually begins in a mole. [NIH]

Melphalan: An anticancer drug that belongs to the family of drugs called alkylating agents. [NIH]

Metaplasia: A change of cells to a form that does not normally occur in the tissue in which it is found. [NIH]

Metastasis: The spread of cancer from one part of the body to another. Tumors formed from cells that have spread are called "secondary tumors"

and contain cells that are like those in the original (primary) tumor. The plural is metastases. [NIH]

Metastatic: Having to do with metastasis, which is the spread of cancer from one part of the body to another. [NIH]

Microspheres: Small uniformly-sized spherical particles frequently labeled with radioisotopes or various reagents acting as tags or markers. [NIH]

Molecular: Of, pertaining to, or composed of molecules : a very small mass of matter. [EU]

Morphine: A narcotic drug used in the treatment of pain. [NIH]

MRI: Magnetic resonance imaging (mag-NET-ik REZ-o-nans IM-a-jing). A procedure in which a magnet linked to a computer is used to create detailed pictures of areas inside the body. [NIH]

Myelosuppression: A condition in which bone marrow activity is decreased, resulting in fewer red blood cells, white blood cells, and platelets. Myelosuppression is a side effect of some cancer treatments. [NIH]

Nausea: An unpleasant sensation, vaguely referred to the epigastrium and abdomen, and often culminating in vomiting. [EU]

Necrosis: Refers to the death of living tissues. [NIH]

Neoplasm: A new growth of benign or malignant tissue. [NIH]

Neural: 1. pertaining to a nerve or to the nerves. 2. situated in the region of the spinal axis, as the neutral arch. [EU]

Neurology: A medical specialty concerned with the study of the structures, functions, and diseases of the nervous system. [NIH]

Neurosurgery: A surgical specialty concerned with the treatment of diseases and disorders of the brain, spinal cord, and peripheral and sympathetic nervous system. [NIH]

Niacin: Water-soluble vitamin of the B complex occurring in various animal and plant tissues. Required by the body for the formation of coenzymes NAD and NADP. Has pellagra-curative, vasodilating, and antilipemic properties. [NIH]

Non-small cell lung cancer: A group of lung cancers that includes squamous cell carcinoma, adenocarcinoma, and large cell carcinoma. [NIH]

Oncogene: A gene that normally directs cell growth. If altered, an oncogene can promote or allow the uncontrolled growth of cancer. Alterations can be inherited or caused by an environmental exposure to carcinogens. [NIH]

Oncologist: A doctor who specializes in treating cancer. Some oncologists specialize in a particular type of cancer treatment. For example, a radiation oncologist specializes in treating cancer with radiation. [NIH]

Oncology: The study of cancer. [NIH]

Ondansetron: A drug that prevents or reduces nausea and vomiting. [NIH]

Ophthalmology: A surgical specialty concerned with the structure and function of the eye and the medical and surgical treatment of its defects and diseases. [NIH]

Ornithine: An amino acid produced in the urea cycle by the splitting off of urea from arginine. [NIH]

Otolaryngology: A surgical specialty concerned with the study and treatment of disorders of the ear, nose, and throat. [NIH]

Overdose: 1. to administer an excessive dose. 2. an excessive dose. [EU]

Palliative: 1. affording relief, but not cure. 2. an alleviating medicine. [EU]

Pancreas: A glandular organ located in the abdomen. It makes pancreatic juices, which contain enzymes that aid in digestion, and it produces several hormones, including insulin. The pancreas is surrounded by the stomach, intestines, and other organs. [NIH]

Pancreatic: Having to do with the pancreas. [NIH]

Pancreatitis: Acute or chronic inflammation of the pancreas, which may be asymptomatic or symptomatic, and which is due to autodigestion of a pancreatic tissue by its own enzymes. It is caused most often by alcoholism or biliary tract disease; less commonly it may be associated with hyperlipaemia, hyperparathyroidism, abdominal trauma (accidental or operative injury), vasculitis, or uraemia. [EU]

Pap test: The collection of cells from the cervix for examination under a microscope. It is used to detect changes that may be cancer or may lead to cancer, and can show noncancerous conditions, such as infection or inflammation. Also called a Pap smear. [NIH]

Paraffin: A mixture of solid hydrocarbons obtained from petroleum. It has a wide range of uses including as a stiffening agent in ointments, as a lubricant, and as a topical anti-inflammatory. It is also commonly used as an embedding material in histology. [NIH]

Particle: A tiny mass of material. [EU]

Pelvis: The lower part of the abdomen, located between the hip bones. [NIH]

Percutaneous: Performed through the skin, as injection of radiopaque material in radiological examination, or the removal of tissue for biopsy accomplished by a needle. [EU]

Perforation: 1. the act of boring or piercing through a part. 2. a hole made through a part or substance. [EU]

Perfusion: Bathing an organ or tissue with a fluid. In regional perfusion, a specific area of the body (usually an arm or a leg) receives high doses of

anticancer drugs through a blood vessel. Such a procedure is performed to treat cancer that has not spread. [NIH]

Peritoneum: The tissue that lines the abdominal wall and covers most of the organs in the abdomen. [NIH]

Phosphorous: Having to do with or containing the element phosphorus. [NIH]

Phosphorylase: An enzyme of the transferase class that catalyzes the phosphorylysis of a terminal alpha-1,4-glycosidic bond at the non-reducing end of a glycogen molecule, releasing a glucose 1-phosphate residue. Phosphorylase should be qualified by the natural substance acted upon. EC 2.4.1.1. [NIH]

Plasma: The clear, yellowish, fluid part of the blood that carries the blood cells. The proteins that form blood clots are in plasma. [NIH]

Polyposis: The development of numerous polyps (growths that protrude from a mucous membrane). [NIH]

Postoperative: After surgery. [NIH]

Postprandial: Occurring after dinner, or after a meal; postcibal. [EU]

Potassium: A metallic element that is important in body functions such as regulation of blood pressure and of water content in cells, transmission of nerve impulses, digestion, muscle contraction, and heart beat. [NIH]

Precancerous: A term used to describe a condition that may (or is likely to) become cancer. Also called premalignant. [NIH]

Preclinical: Before a disease becomes clinically recognizable. [EU]

Preoperative: Preceding an operation. [EU]

Prevalence: The total number of cases of a given disease in a specified population at a designated time. It is differentiated from incidence, which refers to the number of new cases in the population at a given time. [NIH]

Procarbazine: An anticancer drug that belongs to the family of drugs called alkylating agents. [NIH]

Progression: Increase in the size of a tumor or spread of cancer in the body. [NIH]

Prostate: A gland in males that surrounds the neck of the bladder and the urethra. It secretes a substance that liquifies coagulated semen. It is situated in the pelvic cavity behind the lower part of the pubic symphysis, above the deep layer of the triangular ligament, and rests upon the rectum. [NIH]

Proteins: Polymers of amino acids linked by peptide bonds. The specific sequence of amino acids determines the shape and function of the protein. [NIH]

Psychotherapy: A generic term for the treatment of mental illness or

emotional disturbances primarily by verbal or nonverbal communication. [NIH]

Pulmonary: Relating to the lungs. [NIH]

Radioactive: Giving off radiation. [NIH]

Radiofrequency ablation: The use of electrical current to destroy tissue. [NIH]

Radiolabeled: Any compound that has been joined with a radioactive substance. [NIH]

Radiologist: A doctor who specializes in creating and interpreting pictures of areas inside the body. The pictures are produced with x-rays, sound waves, or other types of energy. [NIH]

Radiology: The use of radiation (such as x-rays) or other imaging technologies (such as ultrasound and magnetic resonance imaging) to diagnose or treat disease. [NIH]

Radiosensitization: The use of a drug that makes tumor cells more sensitive to radiation therapy. [NIH]

Radiosensitizers: Drugs that make tumor cells more sensitive to radiation. [NIH]

Radiotherapy: The treatment of disease by ionizing radiation. [EU]

Radium: Radium. A radioactive element of the alkaline earth series of metals. It has the atomic symbol Ra, atomic number 88, and atomic weight 226. Radium is the product of the disintegration of uranium and is present in pitchblende and all ores containing uranium. It is used clinically as a source of beta and gamma-rays in radiotherapy, particularly brachytherapy. [NIH]

Randomized: Describes an experiment or clinical trial in which animal or human subjects are assigned by chance to separate groups that compare different treatments. [NIH]

Rebexamycin: An anticancer drug that belongs to the family of drugs called antineoplastic antibiotics. [NIH]

Receptor: A molecule inside or on the surface of a cell that binds to a specific substance and causes a specific physiologic effect in the cell. [NIH]

Recombinant: 1. a cell or an individual with a new combination of genes not found together in either parent; usually applied to linked genes. [EU]

Recurrence: The return of cancer, at the same site as the original (primary) tumor or in another location, after the tumor had disappeared. [NIH]

Refractory: Not readily yielding to treatment. [EU]

Regimen: A treatment plan that specifies the dosage, the schedule, and the duration of treatment. [NIH]

Remission: A decrease in or disappearance of signs and symptoms of cancer. In partial remission, some, but not all, signs and symptoms of cancer

have disappeared. In complete remission, all signs and symptoms of cancer have disappeared, although there still may be cancer in the body. [NIH]

Resected: Surgical removal of part of an organ. [NIH]

Resection: Removal of tissue or part or all of an organ by surgery. [NIH]

Retrograde: 1. moving backward or against the usual direction of flow. 2. degenerating, deteriorating, or catabolic. [EU]

Ribavirin: A drug used to treat respiratory syncytial virus (RSV) infection in the lungs. [NIH]

Riboflavin: Nutritional factor found in milk, eggs, malted barley, liver, kidney, heart, and leafy vegetables. The richest natural source is yeast. It occurs in the free form only in the retina of the eye, in whey, and in urine; its principal forms in tissues and cells are as FMN and FAD. [NIH]

Salmonella: A genus of gram-negative, facultatively anaerobic, rod-shaped bacteria that utilizes citrate as a sole carbon source. It is pathogenic for humans, causing enteric fevers, gastroenteritis, and bacteremia. Food poisoning is the most common clinical manifestation. Organisms within this genus are separated on the basis of antigenic characteristics, sugar fermentation patterns, and bacteriophage susceptibility. [NIH]

Sarcoma: A cancer of the bone, cartilage, fat, muscle, blood vessels or other connective or supportive tissue. [NIH]

Sargramostim: A colony-stimulating factor that stimulates the production of blood cells, especially platelets, during chemotherapy. It is a cytokine that belongs to the family of drugs called hematopoietic (blood-forming) agents. Also called GM-CSF. [NIH]

Screening: Checking for disease when there are no symptoms. [NIH]

Selenium: An essential dietary mineral. [NIH]

Serum: The clear liquid part of the blood that remains after blood cells and clotting proteins have been removed. [NIH]

Shunt: A surgically created diversion of fluid (e.g., blood or cerebrospinal fluid) from one area of the body to another area of the body. [NIH]

Skull: The skeleton of the head including the bones of the face and the bones enclosing the brain. [NIH]

Squamous: Scaly, or platelike. [EU]

Staging: Performing exams and tests to learn the extent of the cancer within the body, especially whether the disease has spread from the original site to other parts of the body. [NIH]

Stomach: An organ that is part of the digestive system. It helps in the digestion of food by mixing it with digestive juices and churning it into a thin liquid. [NIH]

Stool: The waste matter discharged in a bowel movement; feces. [NIH]

Subcutaneous: Beneath the skin. [NIH]

Substrate: A substance upon which an enzyme acts. [EU]

Supplementation: Adding nutrients to the diet. [NIH]

Symptomatic: Having to do with symptoms, which are signs of a condition or disease. [NIH]

Systemic: Affecting the entire body. [NIH]

Tamoxifen: An anticancer drug that belongs to the family of drugs called antiestrogens. Tamoxifen blocks the effects of the hormone estrogen in the body. It is used to prevent or delay the return of breast cancer or to control its spread. [NIH]

Testicular: Pertaining to a testis. [EU]

Thalidomide: A drug that belongs to the family of drugs called angiogenesis inhibitors. It prevents the growth of new blood vessels into a solid tumor. [NIH]

Thermoregulation: Heat regulation. [EU]

Thrombosis: The formation or presence of a blood clot inside a blood vessel. [NIH]

Thrombus: An aggregation of blood factors, primarily platelets and fibrin with entrapment of cellular elements, frequently causing vascular obstruction at the point of its formation. Some authorities thus differentiate thrombus formation from simple coagulation or clot formation. [EU]

Thyroid: A gland located near the windpipe (trachea) that produces thyroid hormone, which helps regulate growth and metabolism. [NIH]

Thyroxine: An amino acid of the thyroid gland which exerts a stimulating effect on thyroid metabolism. [NIH]

Tolerance: 1. the ability to endure unusually large doses of a drug or toxin. 2. acquired drug tolerance; a decreasing response to repeated constant doses of a drug or the need for increasing doses to maintain a constant response. [EU]

Tomography: A series of detailed pictures of areas inside the body; the pictures are created by a computer linked to an x-ray machine. [NIH]

Toxicity: The quality of being poisonous, especially the degree of virulence of a toxic microbe or of a poison. [EU]

Toxins: Poisons produced by certain animals, plants, or bacteria. [NIH]

Transplantation: The replacement of an organ with one from another person. [NIH]

Tumour: 1. swelling, one of the cardinal signs of inflammations; morbid enlargement. 2. a new growth of tissue in which the multiplication of cells is

uncontrolled and progressive; called also neoplasm. [EU]

Unresectable: Unable to be surgically removed. [NIH]

Urine: Fluid containing water and waste products. Urine is made by the kidneys, stored in the bladder, and leaves the body through the urethra. [NIH]

Vaccination: Treatment with a vaccine. [NIH]

Vaccine: A substance or group of substances meant to cause the immune system to respond to a tumor or to microorganisms, such as bacteria or viruses. [NIH]

Veins: The vessels carrying blood toward the heart. [NIH]

Venereal: Pertaining or related to or transmitted by sexual contact. [EU]

Vincristine: An anticancer drug that belongs to the family of plant drugs called vinca alkaloids. [NIH]

Viral: Pertaining to, caused by, or of the nature of virus. [EU]

Virus: Submicroscopic organism that causes infectious disease. In cancer therapy, some viruses may be made into vaccines that help the body build an immune response to, and kill, tumor cells. [NIH]

General Dictionaries and Glossaries

While the above glossary is essentially complete, the dictionaries listed here cover virtually all aspects of medicine, from basic words and phrases to more advanced terms (sorted alphabetically by title; hyperlinks provide rankings, information and reviews at Amazon.com):

- **The Cancer Dictionary** by Roberta Altman, Michael J., Md Sarg; Paperback - 368 pages, 2nd Revised edition (November 1999), Checkmark Books; ISBN: 0816039542;
<http://www.amazon.com/exec/obidos/ASIN/0816039542/icongroupinterna>
- **Dictionary of Medical Acronymns & Abbreviations** by Stanley Jablonski (Editor), Paperback, 4th edition (2001), Lippincott Williams & Wilkins Publishers, ISBN: 1560534605,
<http://www.amazon.com/exec/obidos/ASIN/1560534605/icongroupinterna>
- **Dictionary of Medical Terms : For the Nonmedical Person (Dictionary of Medical Terms for the Nonmedical Person, Ed 4)** by Mikel A. Rothenberg, M.D, et al, Paperback - 544 pages, 4th edition (2000), Barrons Educational Series, ISBN: 0764112015,
<http://www.amazon.com/exec/obidos/ASIN/0764112015/icongroupinterna>

- **A Dictionary of the History of Medicine** by A. Sebastian, CD-Rom edition (2001), CRC Press-Parthenon Publishers, ISBN: 185070368X,
<http://www.amazon.com/exec/obidos/ASIN/185070368X/iconegroupinterna>
- **Dorland's Illustrated Medical Dictionary (Standard Version)** by Dorland, et al, Hardcover - 2088 pages, 29th edition (2000), W B Saunders Co, ISBN: 0721662544,
<http://www.amazon.com/exec/obidos/ASIN/0721662544/iconegroupinterna>
- **Dorland's Electronic Medical Dictionary** by Dorland, et al, Software, 29th Book & CD-Rom edition (2000), Harcourt Health Sciences, ISBN: 0721694934,
<http://www.amazon.com/exec/obidos/ASIN/0721694934/iconegroupinterna>
- **Dorland's Pocket Medical Dictionary (Dorland's Pocket Medical Dictionary, 26th Ed)** Hardcover - 912 pages, 26th edition (2001), W B Saunders Co, ISBN: 0721682812,
<http://www.amazon.com/exec/obidos/ASIN/0721682812/iconegroupinterna/103-4193558-7304618>
- **Melloni's Illustrated Medical Dictionary (Melloni's Illustrated Medical Dictionary, 4th Ed)** by Melloni, Hardcover, 4th edition (2001), CRC Press-Parthenon Publishers, ISBN: 85070094X,
<http://www.amazon.com/exec/obidos/ASIN/85070094X/iconegroupinterna>
- **Stedman's Electronic Medical Dictionary Version 5.0 (CD-ROM for Windows and Macintosh, Individual)** by Stedmans, CD-ROM edition (2000), Lippincott Williams & Wilkins Publishers, ISBN: 0781726328,
<http://www.amazon.com/exec/obidos/ASIN/0781726328/iconegroupinterna>
- **Stedman's Medical Dictionary** by Thomas Lathrop Stedman, Hardcover - 2098 pages, 27th edition (2000), Lippincott, Williams & Wilkins, ISBN: 068340007X,
<http://www.amazon.com/exec/obidos/ASIN/068340007X/iconegroupinterna>
- **Stedman's Oncology Words** by Beverly J. Wolpert (Editor), Stedmans; Paperback - 502 pages, 3rd edition (June 15, 2000), Lippincott, Williams & Wilkins; ISBN: 0781726549;
<http://www.amazon.com/exec/obidos/ASIN/0781726549/iconegroupinterna>
- **Tabers Cyclopedic Medical Dictionary (Thumb Index)** by Donald Venes (Editor), et al, Hardcover - 2439 pages, 19th edition (2001), F A Davis Co., ISBN: 0803606540,
<http://www.amazon.com/exec/obidos/ASIN/0803606540/iconegroupinterna>

INDEX

- A**
 Abdomen 11, 12, 20, 24, 26, 59, 64, 93,
 94, 125, 167, 206, 249, 253, 255, 257,
 258, 259, 260, 261
 Abdominal.....26, 126, 167, 257, 260, 261
 Abscess 120
 Adenocarcinoma.....75, 206, 259
 Adjuvant..... 14, 15, 149
 Alkaline 102, 126, 143, 262
 Alkalosis 102
 Anesthesia.....61
 Angiogenesis71, 91, 95, 250, 264
 Angiography 12, 105, 144
 Antiangiogenesis71
 Antibody..... 14, 25, 123, 250
 Antigen25, 157, 196, 250
 Arterial 58, 119, 138, 144, 148, 149,
 150, 151, 156, 196
 Arteries 14, 91, 250
 Aspiration.....25, 112, 251
 Autologous..... 150
- B**
 Bacteremia 112, 123, 127, 251, 263
 Bacteria 96, 123, 127, 208, 215, 240,
 250, 251, 263, 264, 265
 Bereavement36
 Bile26, 54, 64, 66, 75, 124, 142, 145,
 157, 239, 240, 251, 252, 254, 256, 257,
 258
 Biopsy 12, 25, 94, 130, 244, 251, 260
 Bypass.....59
- C**
 Calcium..... 167, 209, 210, 256
 Capecitabine.....54, 75
 Carbohydrate.....211
 Carboplatin55
 Carcinogen 156
 Carcinogenesis..... 122
 Carcinogenic..... 156
 Catheter 12, 52, 58, 151, 239, 240, 254
 Cerebrospinal 103, 127, 263
 Cervical..... 35, 47, 64, 72, 236, 252
 Chemoembolization.. 16, 52, 69, 119, 120,
 148, 150, 151, 152, 153, 157
 Cholangitis..... 107, 157
 Cholesterol208, 211
 Chromosomal 110
 Chromosome109, 132
 Chronic25, 53, 62, 74, 100, 114, 126,
 143, 142, 143, 147, 148, 149, 158, 159,
 163, 227, 252, 260
- Chronotherapy 92, 252
 Cisplatin 54, 55, 56, 151
 Coagulation..... 118, 119, 127, 264
 Collagen..... 52
 Colon.. 25, 58, 64, 68, 71, 72, 73, 92, 163,
 253
 Colorectal..... 58, 64, 71, 72, 73, 109
 Concomitant..... 112, 147, 150
 Contamination..... 100, 159
 Contraceptive..... 100, 159
 Cryosurgery 16, 150, 151
 Cryotherapy 26, 142, 253
 Curative..... 37, 70, 151, 153, 216, 259
 Cytokines 53
- D**
 Degenerative 209
 Dexamethasone..... 64
 Diaphragm 146
 Diarrhea 208
 Doxorubicin 52, 55, 56, 76
 Dysplasia 107
- E**
 Endometrial..... 72
 Enzyme 126, 127, 193, 261, 264
 Epidemiological..... 155, 158
 Epidural..... 117, 124, 196, 254
 Epigastralgia 157
 Epithelial 58, 64, 71, 72, 73
 Esophageal 54, 58, 68, 71, 72, 156
 Esophagus 92, 125, 139, 254, 258
 Ethanol..... 16, 120, 142, 148, 150, 151,
 152, 153, 155
 Extravasation 124, 156, 255
 Extremity..... 61
- F**
 Fetus 25, 209, 249
 Fluorescence 109, 124, 255
 Fluorouracil 64
- G**
 Gallbladder.... 24, 54, 65, 75, 91, 93, 138,
 146, 239, 249, 251, 254, 255
 Gastric..... 54, 58, 64, 68, 71, 72, 73, 139,
 144, 156
 Gastrointestinal..... 26, 254
 Gemcitabine..... 75
 Ginseng..... 200, 202, 203
 Glioma..... 103
 Gonadal 131
 Groin 59
- H**
 Hematoma 117

- Hemobilia..... 157
 Heparin 107
 Hepatic ... 14, 16, 52, 58, 60, 67, 120, 122,
 142, 143, 142, 143, 144, 146, 147, 148,
 149, 150, 151, 153, 157, 196, 239, 254
 Hepatitis.... 11, 70, 74, 100, 109, 113, 114,
 115, 131, 142, 142, 143, 147, 148, 149,
 157, 158, 159, 196
 Hepatobiliary..... 139
 Hepatoblastoma 55
 Hepatocyte 156
 Hepatoma 68, 143, 150, 151, 157, 213
 Hybridization..... 109, 110, 118
 Hydroxyurea 195
 Hypercalcemia..... 143
 Hypertension 150, 156
 Hyperthermia 152, 196
 Hypervascular..... 71
 Hypoglycemia 143
I
 Immunization 112, 158
 Immunology 24, 249
 Immunotherapy..... 15, 149, 150
 Induction 125, 132, 256
 Inflammation 47, 124, 126, 157, 186,
 250, 256, 260
 Infusion ... 58, 93, 103, 144, 149, 150, 151,
 257
 Inoperable..... 52, 60, 104, 152, 193
 Interferon 61, 70, 74, 93, 119, 213, 257
 Intermittent..... 157
 Intracellular 102
 Intrahepatic..... 105, 114, 121, 145, 153,
 157
 Intrathecal..... 103
 Intravenous..... 74, 83, 93, 257
 Invasive 178
 Iodine 65, 149
 Irinotecan 72
L
 Laparoscopy 12
 Laparotomy..... 144, 156
 Larynx 130
 Lesion 25, 70, 156, 251
 Lethal 102
 Ligament..... 47, 146, 261
 Ligation 142, 157
 Lobe..... 139, 145, 146, 258
 Lymph ... 47, 108, 125, 130, 146, 252, 258
 Lymphocyte 123, 250
 Lymphoma..... 92, 109, 163, 253
M
 Malignant 4, 11, 68, 94, 102, 149, 163,
 259
 Mammography..... 34
 Mediastinum 112
 Megestrol 63, 152
 Melanoma 58, 61, 163
 Melphalan 59, 60
 Metaplasia..... 75
 Metastasis.... 94, 108, 113, 114, 120, 146,
 157, 259
 Metastatic..... 54, 68, 71, 72, 74, 75, 76,
 112, 114, 119, 132, 142, 144, 148, 151,
 152
 Microspheres 151, 196
 Molecular 107, 117, 154, 161, 162
 Morphine 196
 Myelosuppression..... 101
N
 Nausea..... 91, 94, 193, 250, 260
 Necrosis 150
 Neoplasm..... 4, 127, 143, 265
 Neural 209
 Non-small cell lung cancer..... 193
O
 Oncogene 126, 132, 163, 259
 Oncologist..... 47, 259
 Oncology..... 18
 Ondansetron 64
 Overdose 209
P
 Palliative..... 37, 70
 Pancreas..... 24, 68, 94, 126, 249, 260
 Pancreatic..... 54, 58, 64, 71, 72, 73, 94,
 126, 139, 144, 163, 193, 239, 240, 254,
 260
 Pancreatitis 157
 Paraffin..... 109
 Particle 62, 66
 Pelvis 59
 Percutaneous..... 70, 119, 120, 142, 148,
 150, 151, 152, 153
 Perforation 146
 Perfusion..... 60, 94, 260
 Peritoneum..... 146
 Phosphorous..... 210
 Phosphorylase 122
 Plasma 25, 117, 126, 249, 261
 Polyposis..... 107
 Postoperative 105, 111, 157, 196
 Postprandial 157
 Potassium 211
 Precancerous..... 156
 Preclinical..... 178
 Preoperative 105, 144, 149
 Prevalence 70, 157, 158
 Procarbazine..... 102
 Progression.... 62, 71, 111, 119, 121, 132
 Progressive 25, 70, 74, 127, 143, 252,
 265

- Proteins ..73, 95, 103, 123, 125, 126, 179,
 208, 210, 250, 256, 261, 263
 Psychotherapy..... 187, 192
R
 Radioactive..... 14, 27, 58, 65, 126, 262
 Radiofrequency ablation..... 118, 119, 142,
 150, 151
 Radiolabeled..... 14, 142, 150
 Radiosensitization 14
 Radiosensitizers 152
 Radiotherapy 127, 197, 262
 Randomized 61, 110, 149, 150, 152
 Rebeccamycin 66
 Receptor 106, 123, 250
 Recombinant 74, 92, 101, 253
 Recurrence ...65, 106, 108, 110, 115, 116,
 119, 120, 150, 153
 Regimen55, 56
 Remission.....74, 95, 150, 196, 262
 Resected 145, 147
 Resection..... 15, 70, 105, 107, 116, 119,
 120, 122, 138, 142, 144, 147, 148, 149,
 150, 151, 153, 157
 Retrograde.....239, 244, 254
 Ribavirin.....62
 Riboflavin.....208
S
 Sarcoma56, 58, 68, 72
 Sargramostim 73
 Screening37, 79, 81, 84
 Selenium..... 195, 210
 Serum 144, 158, 244
 Shunt 156
 Skull..... 113
 Squamous75, 206, 259
 Staging 12, 105, 115
 Stool26, 92, 157, 253, 257
 Subcutaneous..... 120, 151
 Substrate..... 102
 Supplementation 195
 Symptomatic 74, 106, 126, 260
 Systemic 14, 15, 142, 149, 150, 153
T
 Tamoxifen 56
 Testicular 64, 73
 Thalidomide 52, 69, 70
 Thermoregulation..... 208
 Thrombosis 150, 157
 Thrombus..... 120, 127, 264
 Thyroid..... 27, 56, 73, 93, 149, 216, 257,
 264
 Thyroxine 210
 Tolerance 67, 95, 264
 Tomography..... 147
 Toxicity..... 53, 102, 131
 Toxins 123, 250
 Transplantation 15, 16, 59, 62, 68, 70,
 100, 107, 112, 113, 116, 139, 142, 148,
 149, 151, 153, 159
 Tumour..... 119, 121
U
 Unresectable.... 53, 54, 55, 56, 57, 58, 63,
 65, 67, 69, 70, 72, 73, 75, 106, 144,
 147, 148, 150, 151, 196, 213
 Urine 26, 53, 91, 157, 216, 243, 244,
 245, 250, 257, 263
V
 Vaccination 158
 Vaccine 24, 73, 127, 158, 249, 265
 Veins 144, 156
 Vincristine 102
 Viral..... 11, 74, 143
 Virus..... 27, 62, 95, 109, 113, 114, 115,
 143, 157, 158, 196, 263, 265
 Visceral 146

