Encyclopedic Reference of Cancer

Springer Berlin

Berlin Heidelberg New York Barcelona Hong Kong London Milan Paris Tokyo MANFRED SCHWAB (Ed.)

Encyclopedic Reference of Cancer

With 261 Figures and 64 Tables



MANFRED SCHWAB Professor for Genetics, Director, Division Cytogenetics - H-0400 German Cancer Research Center Im Neuenheimer Feld 280 69120 Heidelberg Germany

ISBN 3-540-66527-7 Springer-Verlag Berlin Heidelberg New York

Library of Congress Cataloging-in-Publication Data

Schwab, M. (Manfred), 1945-

Encyclopedic reference of cancer / Manfred Schwab. p. cm. ISBN 3540665277 1. Cancer-Encyclopedias. I. Title.

RC254.5 .S393 2001 616.99'4'003-dc21

2001049122

This work is subject to copyright. All rights are reserved, whether the whole or part of the material is concerned, specifically the rights of translation, reprinting, reuse of illustrations, recitation, broadcasting, reproduction on microfilms or in other ways, and storage in data banks. Duplication of this publication or parts thereof is only permitted under the provisions of the German Copyright Law of September 9, 1965, in its current version, and permission for use must always be obtained from Springer-Verlag. Violations are liable for prosecution under the German Copyright Law.

Springer-Verlag Berlin Heidelberg New York a member of BertelsmannSpringer Science+Business Media GmbH

http://www.springer.de

© Springer-Verlag Berlin Heidelberg 2001 Printed in Italy

The use of registered names, trademarks, etc. in this publication does not imply, even in the absence of a specific statement, that such names are exempt from the relevant protective laws and regulations and therefore free for general use.

Product liability: The publishers cannot guarantee the accuracy of any information about the application of operative techniques and medications contained in this book. In every individual case the user must check such information by consulting the relevant literature.

Production Editor: Andreas Gösling, Heidelberg Typesetting: Mitterweger & Partner Kommunikationsgesellschaft mbH, Plankstadt Cover design: Erich Kirchner, Heidelberg

Printed on acid-free paper SPIN: 10731831 14/3130/ag 5 4 3 2 1 0

Preface

Cancer, although a dreadful disease, is at the same time a fascinating biological phenotype. Around 1980, cancer was first attributed to malfunctioning genes and, subsequently, cancer research has become a major area of scientific research supporting the foundations of modern biology to a great extent. To unravel the human genome sequence was one of those extraordinary tasks, which has largely been fuelled by cancer research, and many of the fascinating insights into the genetic circuits that regulate developmental processes have also emerged from research on cancer. Diverse biological disciplines such as cytogenetics, virology, cell biology, classical and molecular genetics, epidemiology, biochemistry, together with the clinical sciences, have closed ranks in their search of how cancer develops and to find remedies to stop the abnormal growth that is characteristic of cancerous cells. In the attempt to establish how, why and when cancer occurs, a plethora of genetic pathways and regulatory circuits have been discovered that are necessary to maintain general cellular functions such as proliferation, differentiation and migration. Alterations of this fine-tuned network of cascades and interactions, due to endogenous failure or to exogenous challenges by environmental factors, may disable any member of such regulatory pathways. This could, for example, induce the death of the affected cell, may mark it for cancerous development or may immediately provide it with a growth advantage within a particular tissue.

Recent developments have seen the merger of basic and clinical science. Of the former, particularly genetics has provided instrumental and analytical tools with which to assess the role of environmental factors in cancer, to refine and enable diagnosis prior to the development of symptoms and to evaluate the prognosis of patients. Hopefully, even better strategies for causal therapy will become available in the future. Merging the basic and clinical science disciplines towards the common goal of fighting cancer, calls for a comprehensive reference source to serve both as a tool to close the language gap between clinical and basic science investigators and as an information platform for the student and the informed layperson alike. Obviously this was an extremely ambitious goal, and the immense progress in the field cannot always be portrayed in line with the latest developments. The aim of the Encyclopedia is to provide the reader with an entrance point to a particular topic. It should be of value to both basic and clinical scientists working in the field of cancer research. Additionally both students and lecturers in the life sciences should benefit highly from this database. I therefore hope that this Encyclopedia will become an essential complement to existing science resources.

The attempts to identify the mechanisms underlying cancer development and progression have produced a wealth of facts, and no single individual is capable of addressing the immense breadth of the field with undisputed authority. Hence, the 'Encyclopedic Reference of Cancer' is the work of many authors, all of whom are experts in their fields and reputable members of the international scientific community. Each author contributed a large number of keyword definitions and in-depth essays and in so doing it was possible to cover the broad field of cancerrelated topics within a single publication. Obviously this approach entails a form of presentation, in which the author has the freedom to set priorities and to promote an individual point of view. This is most obvious when it comes to nomenclature, particularly that of genes and proteins. Although the editorial intention was to apply the nomenclature of the Human Genome Organisation (HUGO), the more vigorous execution of this attempt has been left to future endeavours.

In the early phase of planning the Encyclopedia, exploratory contacts to potential authors produced an overwhelmingly positive response. The subsequent contact with almost 300 contributory authors was a marvellous experience, and I am extremely grateful for their excellent and constructive cooperation. An important element in the preparation of the Encyclopedia has been the competent secretarial assistance of Hiltrud Wilbertz of the Springer-Verlag and of Ingrid Cederlund and Cornelia Kirchner of the DKFZ. With great attention to detail they helped to keep track of the technical aspects in the preparation of the manuscript. It was a pleasure to work with the Springer crew, including Dr. Rolf Lange as the Editorial Director (Medicine) and Dr. Thomas Mager, Senior Editor for Encyclopedias and Dictionaries. In particular I wish to thank Dr. Walter Reuss, who untiringly has mastered all aspects and problems associated with the management of the numerous manuscripts that were received from authors of the international scientific community. It has been satisfying and at times comforting to see how he made illustration files come alive. Thanks also to Dr. Claudia Lange who, being herself a knowledgeable cell biologist, has worked as the scientific editor. Her commitment and interest have substantially improved this Encyclopedia.

As a final word, I would like to stress that although substantial efforts have been made to compose factually correct and well understandable presentations, there may be places where a definition is incomplete or a phrase in an essay is flawed. All contributors to this Encyclopedia will be extremely happy to receive possible corrections, or revisions, in order for them to be included in any future editions of the 'Encyclopedic Reference of Cancer'.

Heidelberg, September 2001

Manfred Schwab

List of Contributors

ABATE-SHEN, COry

Department of Neuroscience and Cell Biology, Center for Advanced Biotechnology and Medicine, UMDNJ-Robert Wood Johnson Medical School, 679 Hoes Lane, Piscataway, NJ 08854, USA

Ани, Natalie G.

Department of Chemistry and Biochemistry, Howard Huges Medical Institute, University of Colorado, Boulder, CO 80309, USA

Albini, Adriana

National Cancer Research Institute - Genova, c/o Advanced Biotechnology Center, Tumor Progression Session, Largo Rosanna Benzi 10, 16132 Genova, Italy

ANGEL, Peter

Division of Signal Transduction and Growth Control (B-0800), German Cancer Research Center (DKFZ) Heidelberg, Im Neuenheimer Feld 280, 69120 Heidelberg, Germany

Araki, Tsutomu

Department of Molecular Biology, Institute of Gerontology, Nippon Medical School, 1–396, Kosugi-cho, Nakahara-ku, Kawasaki 211–0063, Japan and Department of Obstetrics and Gynecology, Nippon Medical School, 1–1-5, Sendagi, Bunkyo-ku, Tokyo 113–0022, Japan

ATERINI, Stefano Department of Experimental Pathology and Oncology, University of Florence, Florence, Italy

AVET-LOISEAU, Herve Laboratory of Hematology, University Hospital, Nantes, France

BALE, Sherri GeneDx Inc., 9700 Great Seneca Highway, Suite 217, Rockville, MD 20850 USA

Bartsch, Helmut

Toxikologie und Krebsrisikofaktoren (C0200), German Cancer Research Center (DKFZ) Heidelberg, Im Neuenheimer Feld 280, 69120 Heidelberg, Germany

BECKER, Heinz Klinik für Allgemeinchirurgie, Universitätsklinikum Göttingen, Robert-Koch-Str. 40, 37075 Göttingen, Germany

BEN-DAVID, Yaacov

Division of Cancer Biology, Sunnybrook & Women's College Health Sciences Center, 2075 Bayview Avenue, S-218, Toronto, Ontario, M4N 3M5, Canada

BERG, Rob J.W. University Medical Center Utrecht, Plutohof 12, 3951 EG Maarn, The Netherlands

BERNARDS, Rene Division of Molecular Carcinogenesis, The Netherlands Cancer Institute, 121 Plesmanlaan, 1066 CX Amsterdam, The Netherlands

BHATTACHARYA-CHATTERJEE, Malaya The Barrett Cancer Center, Univeristy of Cincinnati, Cincinatti, OH 45267, USA

BIEGEL, Jaclyn A. Division of Human Genetics and Molecular Biology, Room 1002 Abramson Research Building, The Children's Hospital of Philadelphia, 3516 Civic Center Blvd., Philadelphia, PA 19104, USA

ВІGNAMI, Margherita Istituto Superiore di Sanita, Viale Regina Elena, 00161 Rome, Italy

BING, Chen Diabetes and Endocrinology Research Group, Department of Medicine, University of Liverpool, Liverpool L69 3GA, UK

BLOW, Julian CRC Chromosome Replication Research Group, Wellcome Trust Biocentre, University of Dundee, MSI/WTB Complex, Dow Street, Dundee DD1 5EH, Scotland, UK

BLUME-JENSEN, Peter Molecular and Cell Biology Laboratory, The Salk Institute, 10010 North Torrey Pines Road, La Jolla, CA 92037, USA

BOSCH, Valerie Angewandte Tumorvirologie (ATV) F0200, German Cancer Research Center (DKFZ) Heidelberg, Im Neuenheimer Feld 242, 69120 Heidelberg, Germany

Boshoff, Chris

The Cancer Research Campaign, Viral Oncology Group, Departments of Oncology and Molecular Pathology, Wolfson Institute for Biomedical Research, Cruciform Building, University College London, Gower Street, London WC1E 6BT, UK

BRAUCH, Hiltrud Dr. Margarete Fischer-Bosch-Institute of Clinical Pharmacology, Auerbachstr. 112, 70376 Stuttgart, Germany

BROWN, Vivette Ontario Cancer Center/Princess Margaret Hospital, Rm 9–204, 610 University Avenue, Toronto, Ontario, M5G 2M9 , Canada

BUDAY, Laszlo Department of Medical Chemistry, Semmelweis University Medical School, 9 Puskin Street, Budapest 1088, Hungary

BUENDIA, Marie-Annick Unité de Recombinaison et Expression Génétique, Institute Pasteur, Inserm U. 163, 28, Rue du Dr Roux, 75724 Paris Cédex 15, France

BÜRKLE, Alexander Department of Gerontology, IAH, University of Newcastle, Wolfson Research Centre, NGH, Newcastle upon Tyne NE4 6BE, UK

CALABRETTA, Bruno Kimmel Cancer Institute, Thomas Jefferson University, Philadelphia, USA CAPOBIANCO, Anthony J. Department of Molecular Genetics, University of Cincinnati College of Medicine, 231 Albert Sabine Way, Cincinnati, OH 45267–0524, USA

CARBONE, Michele Department of Pathology, Cardinal Bernardin Cancer Center, Room 205, Loyola University Medical Center, 2160 South First Ave, Maywood Il 60153, USA

CASELMANN, Wolfgang H. Medizinische Klinik und Poliklinik I, Rheinische Friedrich-Wilhelms-Universität, Sigmund-Freud-Str. 25, 53105 Bonn, Germany

CAVENEE, Webster K. Ludwig Institute for Cancer Research, University of California-San Diego (UCSD), 9500 Gilman Drive, La Jolla, CA 92093–0660, USA

CHAN, Homan Division of Biochemistry and Molecular Biology, Davidson Building, University of Glasgow, Glasgow G12 8QQ, UK

CHEN, Taosheng New Leads, Bristol-Myers Squibb, Wallingford, CT 06492, USA

CHENG, Jin Q. H.Lee Moffit Cancer Center, University of South Florida College of Medicine, Tampa, FL, USA

CHRISTOFORI, Gerhard Research Institute of Molecular Pathology, Dr. Bohr-Gasse 7, 1030 Vienna, Austria

CLAUDIO, Pier Paolo Dept. of Pathology, Anatomy and Cell Biology, Thomas Jefferson University, 1020 Locust Street, JAH 226, Philadelphia, PA 19107, USA

CLAUS, Elizabeth B. Department of Epidemiology and Public Health, Yale University, School of Medicine, New Haven, CT, USA

CLAYETTE, Pascal SPI-BIO, c/o CEA, DSV/DRM/SNV, 18 route du Panorama, B.P. 6, 92265 Fontenay aux Roses Cedex, France

COBLENTZ, Timothy Department of Urology, University of Virginia, Charlottesville, 22908 VA, USA

COLLINS, Andrew R. DNA Instability Group, Rowett Research Institute, Greenburn Road, Bucksburn, Aberdeen AB21 9SB, Scotland, UK

COLOMBO, Mario P. Immunotherapy & Gene Therapy Unit, Department of Experimental Oncology, Istituto Nazionale Tumori, Via Venezian, 1 20133 Milano, Italy

CONAWAY, Joan W. Stowers Institute for Medical Research, 1000 E. 50th Street, Kansas City, MO 64110, USA

CONAWAY, Ronald C. Stowers Institute for Medical Rearch, 1000 E. 50th Street, Kansas City, MO 64110, USA

DANESH, John Clinical Trial Service Unit and Epidemiological Studies Unit, University of Oxford, Oxford, UK DANILKOVITCH-MIAGKOVA, Alla Laboratory of Immunobiology, NCI-FCRDC, Building 560, Rm 12-46, Fort Detrick, Frederick, MD 21702, USA DANKS, Mary K. St. Jude Children's Research Hospital, Memphis, TN USA DE ALAVA, Enrique Departamento de Anatomía Patológica, Clínica Universitaria de Navarra, Apartado 4209, 31080 Pamplona, Spain DECKER, Jochen Bioscientia Institute, Konrad Adenauer Str. 17, 55218 Ingelheim, Germany DENNLER, Sylviane Division of Cellular Biochemistry (H3) The Netherlands Cancer Institute, Plesmanlaan 121, 1066 CX Amsterdam, The Netherlands DER, Channing J. Lineberger Comprehensive Cancer Center, CB# 7295, University of North Carolina at Chapel Hill, 102 Mason Farm Road, Chapel Hill, NC 27599-7295, USA DEVILEE, Peter Department of Human & Clinical Genetics, Leiden University Medical Center, Wassenaarseweg 72, 2333 Al Leiden, The Netherlands

DI FIORE, Pier Paolo Department of Experimental Oncology, European Institute of Oncology, Milan, Italy

DI LUCA, Dario Department of Experimental and Diagnostic Medicine, Section of Microbiology, University of Ferrara, Via Borsari 46, 44100 Ferrara, Italy

DIGWEED, Martin Institute of Human Genetics, Charité, Humboldt Universität zu Berlin, Campus Virchow Klinikum, Augustenburger Platz 1, 13353 Berlin, Germany

DITTMER, Jürgen Department of Molecular Biology, Institute of Cell Biology, University of Tübingen, Auf der Morgenstelle 15, 72076 Tübingen, Germany

DRAGANI, Tommaso A. Research Unit Polygenic Inheritance, Istituto Nazionale Tumori, Via G. Venezian 1, 20133 Milan, Italy

DuBois, Raymond N. Department of Medicine, Vanderbilt University, Medical Center, Nashville, TN, USA

EMI, Mitsuru Department of Molecular Biology, Institute of Gerontology, Nippon Medical School, 1–396, Kosugi-cho, Nakahara-ku, Kawasaki 211–0063, and Department of Obstetrics and Gynecology, Nippon Medical School, 1–1-5, Sendagi, Bunkyo-ku, Tokyo 113–0022, Japan

FAN, Saijun Long Island Jewish Medical Center, and Albert Einstein College of Medicine, Bronx, New York, NY, USA

FARRAR, William L. Cytokine Molecular Mechanisms Section, National Cancer Institute, P.O.Box B, Bldg.560, Rm.31–68, Frederick, MD21702, USA FEINBERG, Andrew P. Institute of Genetic Medicine, John Hopkins University School of Medicine, 1064 Ross, Johns Hopkins Medical School, 720 Rutland Ave. Baltimore, MD 21205, USA

FINLEY, Daniel Department of Cell Biology, Harvard Medical School, Boston, MA 02115, USA

FLETCHER, Jonathan Brigham and Women's Hospital and Harvard Medical School, Boston, MA, USA

FODDE, Riccardo Department of Human and Clinical Genetics, Leiden University Medical Center, Wassenaarseweg 72, PO Box 9503, 2300 RA Leiden, The Netherlands

FOON, Kenneth A. The Barrett Cancer Center, University of Cincinatti, 234 Goodman St., Suite 1091 45267–0502 Cincinatti, OH USA

FORNI, Guido Dipartimento di Scienze Clin. e Biologiche Ospedale San Luigi Gonzaga, 10043 Orbassano, Italy

FREI III, Emil Dana-Farber Cancer Institute, Boston, MA, USA

FRIEDBERG, Errol C. Department of Pathology, University of Texas, Southwestern Medical Center, Dallas, TX 75390–9072, USA

FRISCH, Steven M. The Burnham Institute, La Jolla, CA 92037, USA

FRÜHWALD, Michael C. Division of Pediatric Hematology and Oncology, The Ohio State University and the Comprehensive Cancer Center, Columbus, OH 43210, USA

FUJIMOTO, Jiro Department of Obstetrics and Gynecology, Gifu University School of Medicine, 40 Tsukasa-machi, Gifu City 500-8705, Japan

FURGE, Kyle Van Andel Research Institute, 333 Bostwick, NE, Grand Rapids, MI 49503, USA

GALLIE, Brenda University of Toronto, Head Cancer Informatics, Ontario Cancer Institute/Princess Margaret Hospital, University Health Network, RM 8-415, 610 University Avenue, Toronto M5G 2M9, Canada

GARCIA, ROY Molecular Oncology Program, H.Lee Moffitt Cancer Center and Research Institute, 12902 Magnolia Drive, MRC 4042, Tampa, FL 33612–9416, USA

GARROD, David School of Biological Sciences, University of Manchester, 3.239, Stopford Building, Oxford Road, Manchester M13 9PT, UK

GEISLER, Christian The Leukemia and Lymphoma Marker Laboratory, The Department of Hematology 4041, The Finsen Centre, Rigshospitalet, Copenhagen, Denmark GEORGATOS, Spyros D. Department of Basic Sciences, The University of Crete, School of Medicine, Heraklion, Crete, Greece

GERHARZ, Claus Dieter Institute of Pathology, Heinrich Heine-University, Moorenstr. 5, 40225 Düsseldorf, Germany

GHADIMI, B. Michael Klinik für Allgemeinchirurgie, Universitätsklinikum Göttingen, Robert-Koch-Str. 40, 37075 Göttingen, Germany

GILMORE, Thomas D. Biology Department, Boston University, Boston, MA, USA

GIMM, Oliver Department of General, Visceral, and Vascular Surgery, Martin-Luther-University Halle-Wittenberg, Ernst-Grube-Str. 40, 06097 Halle, Germany

GIORDANO, Antonio Department of Pathology, Anatomy and Cell Biology, Jefferson Medical College, 1020 Locust Street, Jeff Hall Rm. 226, Philadelphia, PA 19107, USA

GISSMANN, Lutz German Cancer Research Center (DKFZ) Heidelberg, Angewandte Tumorvirologie, Im Neuenheimer Feld 242, 69120 Heidelberg, Germany

GLOVER, Thomas W. Department of Human Genetics, 4909 Buhl, Box 0618, University of Michigan, Ann Arbor, MI 48109–0618, USA

GOLDBERG, Steven Long Island Jewish Medical Center, and Albert Einstein College of Medicine, Bronx, NY, USA

GOLDBERG, Itzhak D. Jake Gittlen Cancer Research Institute, The Pennsylvania State University College of Medicine, Hershey, PA, USA

GORES, Gregory J. Division of Gastroenterology and Hepatology, Department of Internal Medicine, Mayo Clinic, Medical School and Foundation, 200 First St SW, Rochester, MN 55905, USA

GRIFFIOEN, Arjan Tumor Angiogenesis Laboratory, University Hospital, Maastricht, The Netherlands,

GRIMM, Dirk German Cancer Research Center (DKFZ) Heidelberg, Im Neuenheimer Feld 280, 69120 Heidelberg, Germany

GROSS, Karen L. Department of Pathology, University of Vermont, Burlington, VT, USA

GRUNICKE, Hans H. Institut für medizinische Chemie und Biochemie der Universität Insbruck, Fritz-Pregl-Straße 3, 6020 Innsbruck, Austria

GUENGERICH, F. Peter Department of Biochemistry and Center in Molecular Toxicology, Vanderbilt University School of Medicine, 638 MRB I Robinson Research Bldg. (MRB I), Pierce Avenue, Nashville, TN 37232-0146, USA GULLICK, Bill Department of Biosciences, University of Kent at Canterbury, Canterbury, Kent CT2 7NJ, UK

GÜNTHERT, Ursula Institute for Medical Microbiology, University of Basel, Petersplatz 10, 4003 Basel, Switzerland

GUSELLA, James F. Molecular Neurogenetics Unit, Massachusetts General Hospital, Building 149, Rm 6214, 13th Street, Charlestown, MA 02129, USA

HAHN, Stephan A. Ruhr-Universität Bochum, Medizinische Klinik, Knappschaftskrankenhaus, Labor für Molekulare Onkologie, In der Schornau 23–25, 44892 Bochum, Germany

HAINAUT, Pierre International Agency for Research on Cancer (IARC), 150 Cours Albert Thomas, 69372 Lyon, France

HAMLIN, Joyce L. Department of Therapeutic Radiology and Oncology, Department of Biochemistry and Molecular Genetics, University of Virginia, School of Medicine, Charlotteville, VA, USA

HAMOUDA, Rasha S. National Institutes of Health (NIH), Bethesda, MD, USA

HAYES, John D. Biomedical Research Centre, Ninewells Hospital and Medical School, Level 5, University of Dundee, Dundee DD1 9SY, Scotland, UK

HAYWARD, Nicholas Queensland Institute of Medical Research, Brisbane, Australia

HE, Li-Zhen Memorial Sloan-Kettering Cancer Center, New York, USA

HECHT, Stephen S. University of Minnesota Cancer Center, Mayo Mail Code #806, 420 Delaware St., S.E., Minneapolis, MN 55455, USA

HELDIN, Carl-Henrik Ludwig Institute for Cancer Research, Box 595, Biomedical Center, 751 24 Uppsala, Sweden

HERZIG, Michaela Research Institute of Molecular Pathology, Dr. Bohr-Gasse 7, 1030 Vienna, Austria

HESLOP, Helen E. Baylor College of Medicine, Houston, TX, USA

HICKSON, Ian D. ICRF Medical Oncology Unit, Institute of Molecular Medicine, University of Oxford, John Radcliffe Hospital, Oxford OX3 9DS, UK

HIEBERT, Scott School of Medicine, Vanderbild University, Nashville, TN, USA

HIGGINS, Rachel R. Division of Cancer Biology, Sunnybrook & Women's College Health Sciences Center, 2075 Bayview Avenue, S-218, Toronto, Ontario, M4N 3M5, Canada

HUEBNER, Kav Kimmel Cancer Institute, Thomas Jefferson University, BLSB, Room 1008, 233 S. 10th Street, Philadelphia, PA 19107, USA HUNTER, TONY Molecular and Cell Biology Laboratory, The Salk Institute, 10010 North Torrey Pines Road, La Jolla, California 92037-1099, USA ISRAEL, Mark A. Norris Cotton Cancer Center, Dartmouth-Hitchcock Medical Center, One Medical Center Drive, Lebanon, NH 03756, USA Izzo, Paola Dipartimento di Biochimica e Biotechnologie, Medicine a Chirurgia, Facoltà di Medicina, Università di Napoli Federico II, Via S. Pansini 5, 80131 Napoli, Italy JACKMAN, Mark Wellcome CRC Institute, Tennis Court Road, Cambridge CB2 1QR, UK JANSEN-DÜRR, Pidder Institute for Biomedical Ageing Research, Austrian Academy of Sciences, Innsbruck, Austria JAY, Daniel G. Department of Physiology, Tufts University School of Medicine, 136 Harrison Avenue, Boston, MA 02111, USA JENSEN, Elwood V. PRE Branch, NICHD, National Institutes of Health (NIH), Rockledge 1, Suite 8000, 6705 Rockledge Drive - MSC 7970, Bethesda, MD 20817, USA JIN, Charlotte Department of Clinical Genetics, University Hospital, Lund, Sweden JOENJE, Hans Department of Clinical Genetics and Human Genetics, Free University Medical Center, Van der Boechorststraat 7, 1081 BT Amsterdam, The Netherlands JOVE, Richard Molecular Oncology Program, H.Lee Moffitt Cancer Center and Research Institute, 12902 Magnolia Drive, MRC 4042, Tampa, FL 33612-9416, USA JURLANDER, Jesper The Leukemia Laboratory, Department of Hematology L4041, The Finsen Centre, Rigshospitalet, Blegdamsvej 9, 2100 Copenhagen, Denmark KAELIN, William Howard Huges Medical Institute, Dana-Faber Cancer Institute and Harvard, Medical School, Boston, MA, USA KATAN, Matilda CRC Centre for Cell and Molecular Biology, Chester Beatty Laboratories, The Institute of Cancer Research, 237 Fulham Road, London SW3 6JB, UK KEFFORD, Richard Department of Medicine, University of Sydney at Westmead Hospital, Westmead, NSW 2145, Australia

KESKI-OJA, Jorma Departments of Virology and Pathology, Haartman Institute, University of Helsinki, Helsinki, Finland KIMCHI, Adi Weizmann Institute of Science, Department of Meolecular Genetics, Rehovot, Israel

KLEIHUES, Paul International Agency for Research on Cancer (IARC), 150 Cours Albert-Thomas, 69372 Lyon, France

KLEIN, George MTC, Karolinska Institute, Box 280, Nobelsväg 16, 171 77 Stockholm, Sweden

KOCHANEK, Stefan Center for Molecular Medicine (ZMMK), University of Cologne, Kerpener Str. 34, 50931 Cologne, Germany

KOVAR, Heinrich Children's Cancer Research Institute (CCRI), St.Anna Kinderspital, Kinderspitalgasse 6, 1090 Vienna, Austria

KUROSE, Keisuke Department of Molecular Biology, Institute of Gerontology, Nippon Medical School, 1–396, Kosugi-cho, Nakahara-ku, Kawasaki 211–0063, and Department of Obstetrics and Gynecology, Nippon Medical School, 1–1-5, Sendagi, Bunkyo-ku, Tokyo 113–0022, Japan

LA THANGUE, Nicholas B. Division of Biochemistry and Molecular Biology, Davidson Building, University of Glasgow, Glasgow, UK

LACAL, Juan Carlos Instituto de Investigaciones Biomédicas, Arturo Duperier 4, 28029 Madrid, Spain

LARNER, James M. Department of Therapeutic Radiology and Oncology, Department of Biochemistry and Molecular Genetics, Box 800383, University of Virginia - Health Science Center, 1300 Jefferson Park Ave, Carlottesville, VA. 22908, USA

LE ROITH, Derek Clinical Endocrinology Branch, National Institutes of Health, Bethesda, MD, USA

LEVY, Jay A. University of California, School of Medicine, San Francisco, CA, USA

LIPSICK, Joseph Stanford University, Stanford, CA, USA

LOEB, Lawrence A. Department of Pathology, Box 357705, University of Washington, Seattle, WA 98195–7705, USA

LOHMANN, Dietmar Institut für Humangenetik, Hufelandstr. 55, 45122 Essen, Germany

LOTHE, Ragnhild A. Institute for Cancer Research, Department of Genetics, The Norwegian Radium Hospital, 0310 Oslo, Norway

LUBENSKY, Irina A.

Molecular Pathogenesis Unit, Surgical Neurology Branch (SNB), National Institute of Neurological Disorders and Stroke (NINDS), National Institute of Health (NIH), Bldg 10, Rm 5D37, 10 Center Drive, Bethesda, MD 20892, USA LYNCH, Henry Department of Preventive Medicine and Public Health, Hereditary Cancer Prevention Clinic, Creighton University, 2500 California Plaza, Omaha, NE 68178, USA

MÄDGE, Britta Dept. of Cellular & Molecular Medicine, HHMI/CMM-West, UCSD School of Medicine, 9500 Gilman Drive, La Jolla, CA 92093–0686, USA

MANET, Evelyne Unite de Virologie humaine, INSERM U412, Ecole Normale Superieure de Lyon, 46 Allee d'Italie, 69364, Lyon cedex 07, France

MANNENS, Marcel DNA-Diagnostics Laboratory, Department of Clinical Genetics, Academic Medical Centre, University of Amsterdam, P.O. Box 22700, 1100 DE Amsterdam, The Netherlands

MANTANI, Akio Department of Psychiatry and Neuroscience, Hiroshima University of Medicine, 1–2-3 Kasumi, Minami-ku, Hiroshima 734–8551, Japan

MARMÉ, Dieter Institute of Molecular Oncology, Tumor Biology Center, Klinik fur Tumorbiologie, Breisacher Strasse 117, 79106 Freiburg, Germany

MARSH, Deborah J. Kolling Institute of Medical Research, Royal North Shore Hospital, St Leonards, Sydney, NSW 2065, Australia

MCCARTY, Joseph H. Massachusetts Institute of Technology (MIT), Center for Cancer Research, E17–230, 77 Massachusetts Avenue, Cambridge, MA 02139, USA

MCGREGOR, W. Glenn University of Louisville, 570 S. Preston Street Suite 221A, Louisville, KY 40202–1786, USA

MOESLEIN, Gabriela Klinik für Allgemeine und Unfallchirurgie, Düsseldorf, Germany

MOESTA, K.Thomas Klinik für Chirurgie und chirurgische Onkologie, Robert-Roessle-Klinik, Charité, Campus Buch, Lindenberger Weg 80, 13125 Berlin, Germany

MOLLENHAUER, Jan Department for Molecular Genome Analysis, German Cancer Research Center (DKFZ) Heidelberg, Im Neuenheimer Feld 280, 69120 Heidelberg, Germany

MONTESANO, Ruggero International Agency for Research on Cancer, 150 Cours Albert Thomas, 69372 Lyon Cedex 08, France

MORIN, Pat J. Laboratory of Cellular and Molecular Biology, National Institute on Aging, National Institutes of Health (NIH), 5600 Nathan Shock Drive, Baltimore, MD 21224, USA

MORRIS, Christine Cancer Genetics Research Group, Department of Pathology, Christchurch School of Medicine, PO Box 4385, Christchurch, New Zealand MORTON, Cynthia C. Department of Pathology, Brigham and Women's Hospital, 75 Francis Stren, Boston, MA 02115, USA

MOUSSES, Spyro Cancer Genetics Branch, National Human Genome Research Institute, National Institutes of Health (NIH), Room 4C20, MSC 4465, 49 Convent Drive, Bethesda, MD 20892–4465, USA

MÜLLER, Rolf Institute of Molecular Biology and Tumor Research (IMT), Philipps-University Marburg, Emil-Mannkopff-Strasse 2, 35033 Marburg, Germany

MÜLLER-HERMELINK, H.K. Pathologisches Institut der Universität Würzburg, Bau 21, Joseph-Schneider-Straße 2, 97080 Würzburg, Germany

MURPHY, Edward L. University of California, School of Medicine, San Francisco, CA, USA

MUSCHEL, Ruth J. Departments of Pathology and Laboratory Medicine, Rm 269 John Morgan Bld., University of Pennsylvania, Philadelphia, PA 19104, USA

NAGATA, Shigekazu Osaka University Medical School, Osaka, Japan

NAKAGAWARA, Akira Chiba Cancer Center, Research Institute, Chiba, Japan

NICOSIA, Santo V. H. Lee Moffitt Cancer Center, University of South Florida of Medicine, Tampa, FL, USA

NOONAN, Douglas National Cancer Research Institute - Genova, c/o Advanced Biotechnology Center, Tumor Progression Session, Largo Rosanna Benzi 10, 16132 Genova, Italy

NORDHEIM, Alfred Department of Molecular Biology, Institute for Cell Biology, Auf der Morgenstelle 15, University of Tübingen, 72076 Tübingen, Germany

OFFERMANNS, Stefan Pharmakologisches Institut, Universität Heidelberg, Im Neuenheimer Feld 366, 69120 Heidelberg, Germany

Онмо, Hitoshi The First Division, Department of Internal Medicine, Faculty of Medicine, Kyoto University, 54-Shogoin-Kawaramachi, Sakyo-ku, Kyoto 606–8507, Japan

OTT, G. Pathologisches Institut der Universität Würzburg, Josef-Schneider-Strasse 2, 97080 Würzburg, Germany

OTTENSMEIER, Christian CRC Wessex Oncology Unit, CF93, Southampton General Hospital, Tremona Road, Southampton SO16 6YD, UK

PANDOLFI, Pier Paolo

Sloan-Kettering Institute, Memorial Sloan-Kettering Cancer Center, Graduate School of Medical Sciences, Cornell University, 1275 York Avenue, Box 110, New York, NY 10021, USA PANTEL, Klaus Universitäts-Krankenhaus Eppendorf, Hamburg, Germany

PARSONS, Sarah J. Department of Microbiology, PO Box 800734, University of Virginia Health System, Charlottesville, VA 22908, USA

PELLICER, Angel Department of Pathology, University School of Medicine, 550 First Avenue, New York, NY 10028, USA

PFREUNDSCHUH, Michael Universitätsklinikum des Saarlandes, Medizinische Klinik 1, Kirrbergstr. 1, 66424 Homburg/Saar, Germany

PIEROTTI, Marco A. Department of Experimental Oncology, Istituto Nazionale Tumori, Via G. Venezin, 1, 20133 Milan, Italy

PIETSCH, Torsten Institut für Neuropathologie, Universitätskliniken Bonn, Sigmund-Freud-Strasse 25, 53105 Bonn, Germany

PIZER, Ellen S. National Institute on Aging, National Institutes of Health (NIH), Johns Hopkins University, School of Medicine, Baltimore, MD, USA

PLASS, Christoph Division of Human Cancer Genetics, The Ohio State University, Medical Research Facility 464A, 420 West 12th Ave., Columbus, Ohio 43210, USA

PLATT, Jeffrey L. Transplantation Biology and Departments of Surgery, Immunology and Pediatrics, Medical Sciences Building 2–66, Mayo Clinic, 200 1st Street S.W., Rochester, MN 55905, USA

POIRIER, Miriam C. Carcinogen-DNA Interactions Section, National Cancer Institute, Bldg. 37, Rm. 2A05, National Institutes of Health (NIH), 37 Convent Dr, MSC-4255, Bethesda, MD 20892–4255, USA

PONDER, Bruce Cambridge University, Cambridge Institute for Medical Research, Wellcome Center for Molecular Mechanisms in Disease, Addenbrooke's Hospital Site, Hills Road, Cambridge CB2 2XY, UK

POOL-ZOBEL, Beatrice L. Nutritional Toxicology, Institute for Nutritional Sciences, Friedrich Schiller University Jena, Dornburger Str. 25, 007743 Jena, Germany

POTTEN, Christopher S. CRC Department of Epithelial Biology, Paterson Institute of Cancer Research, Christie Hospital NHS Trust, Manchester, UK

POUSTKA, Annemarie Department for Molecular Genome Analysis, German Cancer Research Center (DKFZ) Heidelberg, Im Neuenheimer Feld 280, 69120 Heidelberg, Germany

PRALL, Richard T. Winona Memorial Hospital, Indianapolis, IN 46142, USA PRITCHARD-JONES, Kathy Institute of Cancer Research/Royal Marsden NHS Trust, Downs Road, Sutton, Surrey SM2 5PT, UK

PUI, Ching-Hon St. Jude Children's Research Hospital, 332 North Lauderdale, Memphis. TN 38105, USA

RABES, Hartmut M. Institute of Pathology, University of Munich, Munich, Germany

RADICH, Jerald P. Clinical Research Division, Program in Genetics and Genomics, Fred Hutchinson Cancer Research Center, 1100 Fairview Avenue N., D4–100, Seattle, WA 98109–1024, USA

RAMÓN Y CAJAL, Santiago Department of Pathology, Clinica Puerta de Hierro, c/ San Martin de Porres, 4, 28035 Madrid, Spain

REDDEL, Roger Children's Medical Research Institute, Locked Bag 23, Wentworthville, NSW 2145, Australia

REGO, Eduardo M. Memorial Sloan-Kettering Cancer Center, New York, NY, USA

RENEHAN, Andrew G. CRC Department of Epithelial Biology, Paterson Institute of Cancer Research, Christie Hospital NHS Trust, Manchester, UK

RIED, Thomas National Cancer Institute, National Institutes of Health (NIH), Building 9, Rm 1N-105, 9 Memorial Drive MSC 0913, Bethesda, MD 20892–0913, USA

RODRIGUEZ, Vilmarie Mayo Clinic, East 9, 200 First St SW, Rochester, MN 55905, USA

RONAI, Ze'ev Ruttenberg Cancer Center, Mount Sinai School of Medicine, 1425 Madison Ave, Rm 15-20, New York, NY 10029, USA

RONINSON, Igor B. Division of Molecular Oncology, Department of Molecular Genetics (M/C 669), University of Illinois at Chicago, 900 South Ashland Avenue, Chicago, IL 60607–7170, USA

ROSEN, Eliot M. Long Island Jewish Medical Center, and Albert Einstein College of Medicine, Bronx, NY, USA

Ross, Jeffrey S. Albany Medical College, Albany, NY, USA

RUGGIERO, Marco Dipartimento di Patologia e Oncologia Sperimentali, Viale Morgani 50, 50134 Firenze, Italy

RUIZ-CABELLO OSUNA, Francisco Servicio Analisis Clinicos, Hospital Virgen de las Nieves, Avda Fuerzas Armadas sn, Granada 18014, Spain

RUOSLAHTI, Erkki Cancer Research Center, The Burnham Institute, 10901 North Torrey Pines Road, La Jolla, CA 92037, USA RUSCIANO, Dario Medway Genetics, Via Cantonale, 6805 Mezzovico, Lugano, Switzerland

Russo, Giandomenico Instituto Dermopatico dell' Immacolata, Istituto die Ricerca e Cura a Carattere Scientifico, Rome, Italy

SAMANT, Rajeev S. Jake Gittlen Cancer Research Institute (H059), M. S. Hershey Medical Centre, The Pennsylvania State University College of Medicine, Hershey, PA 17033, USA

SCHÄFER, Reinhold Molekulare Tumorpathologie, Charité Campus Mitte, Institut für Pathologie, Schumannstr. 20/21, 10117 Berlin, Germany

SCHARTL, Manfred Biozentrum, Universität Würzburg, Am Hubland, 97074 Würzburg, Germany

SCHLAG, Peter Klinik für Chirurgie und chirurgische Onkologie, Robert-Roessle-Klinik, Charité, Campus Buch, Lindenberger Weg 80, 13125 Berlin, Germany

SCHMITT-GRÄFF, Annette Pathologisches Institut des Universitätsklinikums, Universität Freiburg, Albertstr. 19, 79104 Freiburg, Germany

SCHÖNTHAL, Axel H. University of Southern California, Keck School of Medicine, Los Angeles, CA, USA

SCHREIBER-AGUS, Nicole Department of Molecular Genetics, Albert Einstein College of Medicine, 1300 Morris Park Ave, Ullmann 809, Bronx, NY 10461, USA

SCHWAB, Manfred Division Cytogenetics - H-0400, German Cancer Research Center (DKFZ) Heidelberg, Im Neuenheimer Feld 280, 69120 Heidelberg, Germany

SELLERS, William R. Harvard Medical School, Dana-Farber Cancer Institute, D720C, 44 Binney Street, Boston, MA 02115, USA

SEN, Subrata M.D. Anderson Cancer Center, Houston, TX, USA

SHAY, Jerry W. Department of Cell Biology, University of Texas Southwestern Medical Center, Room K2–206, 5323 Harry Hines Boulevard, Dallas, TX 75390–9039, USA

SHEWACH, Donna 4713 Upjohn Center, Department of Pharmacology, University of Michigan Medical School, 1310 E. Catherine, Ann Arbor, MI 48109–0504, USA

SHILOH, Yosef Department of Human Genetics and Molecular Medicine, Sackler School of Medicine, Tel Aviv University, Tel Aviv 69978, Israel

SHIPLEY, Janet Institute of Cancer Research/Royal Marsden NHS Trust, Downs Road, Sutton, Surrey SM2 5PT, UK SOUSSI, Thierry Institut Curie, Laboratoire de Genotoxicologie des Tumeurs, 26 rue d'Ulm, 75005 Paris, France

SRIVASTAVA, Sudhir Division of Cancer Prevention, National Cancer Institute, National Institutes of Health (NIH), 6130 Executive Boulevard, Rm EPN 330F, Rockville, MD 20852, USA

STACK, M. Sharon Northwestern University Medical School, 303 E. Chicago Ave., Tarry 8–715, Chicago, IL 60611, USA

Stanbridge, Eric

Departments of Microbiology and Molecular Genetics, University of California, College of Medicine, 240 Medical Science, Building B, Irvine, CA 92697-4025, USA

STEEG, Patricia S Laboratory of Pathology, NCI, National Institutes of Health (NIH), Bethesda, MD, USA

STEIN, Stacey Department of Neuroscience and Cell Biology, Center for Advanced Biotechnology and Medicine, UMDMJ Robert Wood Johnson Medical School, Piscataway, NJ, USA

STEVENSON, Freda Molecular Immunology Group, Tenovus Laboratory, Southampton University Hospitals Trust, Southampton SO16 6YD, UK

SWANSTROM, Ron CB 7295 Lineberger Cancer Center, Room 22–006, University of North Carolina at Chapel Hill, Chapel Hill, NC 27599, USA

TEAN TEH, Bin Van Andel Research Institute, Grand Rapide, MI 49503, USA

TEN DIJKE, Peter Division of Cellular Biochemistry (H3), The Netherlands Cancer Institute, Plesmanlaan 121, 1066 CX Amsterdam, The Netherlands

TESTA, Joseph R. Fox Chase Cancer Center, 7701 Burholme Avenue, Philadelphia, PA 19111, USA

THACKER, John Medical Research Council, Radiation & Genome Stability Unit, Harwell, Oxfordshire OX11 0RD, UK

THEODORESCU, Dan Department of Urology, University of Virginia, Charlottesville, Virginia, 22908–0422, USA

THEODOROPOULOS, Panayiotis A. Department of Basic Sciences, The University of Crete, School of Medicine, Heraklion, Crete, Greece

THOMS, Sven Friedrich-Miescher-Laboratorium of the Max-Planck-Society, Spemannstr. 37, 72076 Tübingen, Germany

TIRELLI, Umberto National Cancer Institute, Aviano, Italy

TOMMASINO, Massimo Angewandte Tumorvirologie (ATV), German Cancer Research Center (DKFZ) Heidelberg, Im Neuenheimer feld 242, 69120 Heidelberg, Germany TORO, Jorge R. National Institutes of Health (NIH), Bethesda, MD, USA

TURENCHALK, Gregory S. Yale University, Yale, CT, USA

VANDE WOUDE, George F. Van Andel Research Institute, 333 Bostwick NE, Grand Rapids, MI 49503, USA

VAUPEL, Peter Institute of Physiology and Pathophysiology, University of Mainz, Duesbergweg 6, 55099 Mainz, Germany

VENKITARAMAN, Ashok R. Department of Oncology and The Wellcome Trust Centre for Molecular Mechanisms in Disease, The Cambridge Institute for Medical Research, Cancer Research Centre, University of Cambridge, Hills Road, Cambridge CB2 2XZ, UK

VERMA, Rakesh Antisoma Research Laboratories, St. George's Hospital Medical School, Cranmer Terrace, London, SW17 0QS, UK

VLEMINCKX, Kris Department of Molecular Biology, Ghent University - VIB, K.L. Ledeganckstraat 35, 9000 Ghent, Belgium

VLODAVSKY, Israel Department of Oncology, Hadassah-Hebrev University Hospital, POBox 12000, Jerusalem 91120, Israel

VOGEL, Charles L. Department of Surgery, Heinrich Heine Universität, Düsseldorf, Germany

VOGEL, Tilman Sylvester Cancer Center, School of Medicine, University of Miami, 600 S.Pine Island Rd. Ste 104, Plantation, FL 33324, USA

VON HOFF, Daniel D. Arizona Cancer Center, 1515 N. Campbell Ave, P.O. Box 245024, Tucson, AZ 85724, USA

VON SCHWEINITZ, Dietrich Kinderchirurgie, Universitäts-Kinderspital beider Basel (UKBB), Standort Bruderholz, Postfach, 4005 Basel, Switzerland

WEINSTEIN, I. Bernard Columbia University, New York, NY, USA

WEISS, Lawrence M. Division of Pathology, City of Hope National Medical Center, 1500 E. Duarte Road, Duarte, CA 91010, USA

WELCH, Danny R. Jake Gittlen Cancer Research Institute, H-059, Room C7850, The Pennsylvania State University College of Medicine, 500 University Drive, Hershey, PA 17033–2390, USA

WESTON, Ainsely National Institute of Occupational Safety and Health NIOSH-CDC, MS-L3014, 1095 Willowdale Road, Morgantown, WV 26505–2888, USA WICKING, Carol

Institute for Molecular Bioscience, The Centre for Molecular and Cellular Biology, The Centre for Drug Design and Development, ARC Special Research Centre for Functional and Applied Genomics, University of Queensland, St Lucia 4072, Queensland, Australia

WILLIAMS, Gareth Department of Medicine, Clinical Sciences Centre, University Hospital Aintree, Longmoor Lane, Liverpool L9 7AL, UK

WOLF, C. Roland University of Dundee Biomedical Research Centre, Level 5, Ninewells Hospital and Medical School, Dundee DD1 9SY, UK

Xu, Tian

Howard Hughes Medical Institute, Genetics Department, Yale University School of Medicine, 295 Congress Avenue BCMM 236, New Haven, CT 06536, USA

YANG, Xiao Yi National Cancer Institute-Frederick, Building 434, Room 4D, P.O.Box B, Frederick, MR 21702, USA

ZBAR, Berton Laboratory of Immunobiology, National Cancer Institute-Frederick, Building 560, Room 12–89, Frederick, MR 21702, USA

ZHOU, Hongyi Department of Molecular Pathology, M.D. Anderson Cancer Center, Box # 89, 1515 Holcombe BLVD, Houston TX 77030, USA

ZÖLLER, Margot German Cancer Research Center (DKFZ) Heidelberg, Abt. D0600, Im Neuenheimer Feld 280, 69120 Heidelberg, Germany

ZORI, Roberto T. Division of Genetics, Department of Pediatrics, PO Box 100296, University of Florida, Gainesville, FL 32610, USA