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Series Editor

Molecular and Clinical Advances in Anticancer Drug Resistance

**edited by
Robert F. Ozols**

Kluwer Academic Publishers

Molecular And Clinical Advances In Anticancer Drug Resistance

Raphael E. Pollock



Molecular And Clinical Advances In Anticancer Drug Resistance:

Molecular and Clinical Advances in Anticancer Drug Resistance Robert F. Ozols, 2012-12-06 The importance of drug resistance in cancer chemotherapy cannot be over stated The 500 000 patients who die every year from cancer in the United States have in most cases been treated with chemotherapy Many of these patients responded initially to chemotherapy but death resulted from the development of drug resistant tumors In the first volume in the series Drug Resistance in Chemotherapy the results of comprehensive laboratory studies aimed at understanding the mechanisms for resistance to individual agents and to the development of broad cross resistance were described In the past 2 years there has been substantial progress in understanding the molecular biology associated with these mechanisms of drug resistance For the first time we are starting to understand which mechanisms are playing an important role in human tumors and even more importantly clinical trials have recently been initiated in an effort to reverse specific forms of drug resistance The purpose of this volume is to describe the new advances both at the molecular level and in the clinic regarding mechanisms of drug resistance and potential ways this resistance can be circumvented This volume is focused upon mechanisms of resistance associated with two major classes of anticancer drugs alkylating agents including cisplatin and the natural products e g adriamycin and vinblastine The first section of the book describes new insights into the genetic mechanisms associated with drug resistance

Molecular and Clinical Advances in Anticancer Drug Resistance Robert F Ozols, 1991-09-30 **Drug Resistance** William N. Hait, 2012-12-06 Resistance to treatment represents the final common outcome for far too many patients with cancer Even our most promising new drugs fall victim to drug resistance Hormones and newer biological therapies though safe and active also lose their activity over time In this volume of Drug Resistance leading investigators in the field have reviewed the most basic mechanisms of drug resistance and have proposed ways to modulate resistance This comprehensive volume should be of value for basic and clinical scientists who wish to delve more deeply into this intriguing problem in the laboratory and devastating problem in the clinic

Advances in Radiation Therapy Bharat B. Mittal, James A. Purdy, K.K. Ang, 2012-12-06 Recent advances in radiation oncology have depended upon and are intertwined with subsequent scientific discoveries and the development of new techniques in the fields of radiation and molecular biology physics electrical engineering surgery and medical oncology This volume describes how some of the recent discoveries in the radiological sciences have influenced the way radiation oncology is practised As there are many advances in this field the Editors have chosen to concentrate on selected topics in clinical radiotherapy radiation physics and biology and technical innovations that have had a major impact on radiation oncology in the past twenty years It is hoped that the techniques described in this volume will increase tumor control and prolong patient survival and at the same time decrease radiation induced side effects and complications

Advances in Allogeneic Hematopoietic Stem Cell Transplantation Richard K. Burt, Mary M. Brush, 2012-11-28 The field of hematopoietic stem cell transplantation is rapidly

evolving Realization that hematopoietic stem cells give rise to the immune compartment has resulted in clinical trials of hematopoietic stem cell transplantation for patients with autoimmune diseases Allogeneic hematopoietic transplants are a form of adoptive immunotherapy resulting in beneficial graft versus tumor effects Large numbers of hematopoietic cells can be collected with ease Therefore a renewable source of cells for ex vivo genetic manipulations is readily available Multiple trials combining hematopoietic transplants and gene therapy are in progress One such application is the infusion of allogeneic lymphocytes containing a suicide gene to abort graft versus host disease Hematopoietic stem cell transplantation is in reality the clinical and practical application of cellular therapy Hematopoietic transplant physicians are by design or by practical application evolving into cell and gene therapy specialists The excitement and enthusiasm in hematopoietic transplantation is that it offers a door to the future A future not of drugs or titrating poisonous chemotherapy but rather of cellular and gene therapy

1 ALLOGENEIC PERIPHERAL BLOOD STEM CELL TRANSPLANTATION FOR HEMATOLOGIC DISEASES

Martin Korbling University of Texas MD Anderson Cancer Center Houston Texas 77030

INTRODUCTION

Circulating hematopoietic stem cells have emerged as an alternative to bone marrow BM stem cells for allografting For many years the reconstitutive potential of circulating stem cells was questioned peripheral blood stem cells PBSC were even characterized a waste product

1 Cancer Policy: Research and Methods

C.L. Bennett,Tammy Stinson,2007-08-20

Policy and outcome studies represent a new area of research in oncology Understanding the factors that affect quality of life costs of care patterns of care and outcomes in oncology is important to providing comprehensive care Health services is meant to be both a productive area of research as well as a discipline that improves patient decision making In sum Cancer Policy highlights some of the important areas of health services research in oncology

Drug Resistance in Oncology

S. Bernal,1997-08-21 This timely new reference integrates the latest clinical results and laboratory studies on the resistance of specific cancers to chemotherapeutic drugs covering drug resistance in lung breast ovary and colon cancer as well as hematological malignancies

Diagnostic and Therapeutic Advances in Hematologic Malignancies

Martin S. Tallman,Leo I. Gordon,2007-11-23 There is no field of medicine in which advances in therapy have been so closely linked to a better understanding of molecular medicine than in the area of hematologic malignancies Diagnostic and Therapeutic Advances in Hematologic Malignancies will provide practitioners and academic clinicians with a better understanding of the constant evolving concepts in this area

Biological and Hormonal Therapies of Cancer

Kenneth A. Foon,Hyman B. Muss,2012-12-06 This volume Biological and Hormonal Therapies of Cancer which is part of the series Cancer Treatment and Research presents selected new information concerning biologic and hormonal therapy of cancer We have attempted to provide the reader with topics of major interest in a timely fashion There is renewed interest in biologic therapy of cancer Two chapters review the role of interferon in the hematologic malignancies and in solid tumors Vaccine therapies have come to the forefront of cancer therapy recently and two chapters approach different strategies of vaccine therapies one reviews

the cellular vaccine therapies and another the anti idotype approach The hormonal therapy chapters focus on current uses of endocrine therapy in endometrial breast and prostate cancer In addition hormonal strategies for the prevention of breast cancer and endometrial cancer including exciting information relating to phytochemicals are presented The effects of tamoxifen on endometrium is a topic of major interest and is discussed in detail Finally there is a chapter on estrogen receptor expression and regulation in human breast cancer These chapters are all written by experts in the field and contain timely and relevant information of interest to laboratory and clinical scientists and practitioners alike Biologic and endocrine therapies represent major areas of cancer research interest The advent of newer biologic therapies including new antibody targeted treatments and the use of biologics as tumor modulators to enhance the effects of other treatment regimens is an exploding avenue of research

Diagnostic and Therapeutic Advances in Pediatric Oncology David O. Walterhouse, Susan L. Cohn, 2012-12-06 The purpose of *Diagnostic and Therapeutic Advances in Pediatric Oncology* for the Cancer Treatment and Research Series is to provide an up to date summary of how recent advances in cancer research are being applied to the care of children with solid tumors The interface of cancer research with clinical practice in pediatric oncology has never been more intimate than today While researchers are identifying oncogenes and tumor suppressor genes and are studying their specific functions clinicians are using knowledge of oncogenes and tumor suppressor genes for diagnosing cancer in children for therapeutic decision making purposes and for prognostic purposes The first three chapters in this book describe models for understanding the causes of childhood cancer that were perhaps initially identified by clinicians and that are now being studied and understood by researchers These chapters will describe research evidence that supports roles for the involvement of normal developmental regulatory genes in childhood oncogenesis of abnormal immune regulation in childhood oncogenesis and of heredity in childhood oncogenesis The next eight chapters are devoted to descriptions of the application of new research developments to clinical practice with reference to the most common forms of solid tumors of childhood outside the central nervous system The final chapter will describe late effects of childhood cancer and its therapy and the impact research is having on understanding and perhaps preventing these late effects

Surgical Oncology Raphael E. Pollock, 2012-12-06 Among the standard oncology modalities surgical oncology is singular in that it lacks a separate board certification or even an added qualification mechanism Card carrying surgical oncologists are certified by the American Board of Surgery as are all other general surgeons What distinguishes the surgical oncologist is a set of cognitive skills rather than a specific armamentarium of surgical techniques This different conceptual framework is derived from extensive additional training that leads to an in depth understanding of the natural history and biologic behavior of the various solid tumor systems Equipped with this perspective the surgical oncologist is particularly well positioned to integrate the various available therapeutic modalities into a coherent care program for the solid tumor patient As a central theme the chapters of this book demonstrate that increasingly sophisticated diagnostic and staging approaches are helping to move chemotherapy

and radiotherapy into the preoperative neoadjuvant setting This fundamental alteration is based on the awareness that even early stage solid tumor disease is frequently systemic at the time of presentation at least on a subclinical level And although the primary tumor may be controllable by surgery with radiotherapy the uncontrolled and initially clinically unapparent distant disease ultimately determines patient survival The other perspective driving the neoadjuvant approach is an emerging awareness that for most solid tumor systems neoadjuvant treatment responses can facilitate less mutilating surgery with comparable levels of local disease control

Anticancer Drug Resistance Lori J. Goldstein, Robert F.

Ozols, 2012-12-06 Over the last 50 years drug development and clinical trials have resulted in successful complete responses in diseases such as childhood leukemia testicular cancer and Hodgkin's disease We are still however confronted with over 500 000 cancer related deaths per year Clearly the phenomenon of drug resistance is largely responsible for these failures and continues to be an area of active investigation Since the last volume in this series we have learned that the energy dependent drug efflux protein P-glycoprotein encoded by the MDR 1 gene is a member of a family of structurally related transport polypeptides thus allowing us to explore the relationship between structure and function In addition to ongoing well designed clinical trials aimed at reversing MDR mediated drug resistance the first gene therapy studies with the MDR 1 gene retrovirally transduced into human bone marrow cells are about to be initiated Although MDR is currently the most understood mechanism of drug resistance we are uncovering increasing knowledge of alternative molecular and biochemical mechanisms of drug resistance to antimetabolites cisplatin and alkylating agents and developing new strategies for circumventing such resistance It is clear that drug resistance is complex and many mechanisms exist by which cancer cells may overcome the cytotoxicity of our known chemotherapeutic agents As our understanding of each of these mechanisms expands well designed models will be necessary to test laboratory hypotheses and determine their relationship to drug resistance in humans It is this integration of basic science and clinical investigation that will both advance our scientific knowledge and result in the improvement of cancer therapy

Recent Advances in Clinical Trial Design and Analysis Peter

F. Thall, 2012-12-06 Clinical trials have two purposes to treat the patients in the trial and to obtain information which increases our understanding of the disease and especially how patients respond to treatment Statistical design provides a means to achieve both these aims while statistical data analysis provides methods for extracting useful information from the trial data Recent advances in statistical computing have enabled statisticians to implement very rapidly a broad array of methods which previously were either impractical or impossible Biostatisticians are now able to provide much greater support to medical researchers working in both clinical and laboratory settings As our collective toolkit of techniques for analyzing data has grown it has become increasingly difficult for biostatisticians to keep up with all the developments in our own field Recent Advances in Clinical Trial Design and Analysis brings together biostatisticians doing cutting edge research and explains some of the more recent developments in biostatistics to clinicians and scientists who work in clinical trials

Journal of the National Cancer Institute ,1995 **Current Catalog** National Library of Medicine (U.S.),1993 First multi year cumulation covers six years 1965 70 **Techniques in Aquatic Toxicology** Gary K. Ostrander,1996-08-07 This is a comprehensive gathering of measurement and assessment techniques for aquatic toxicants Covering everything from ASTM and similar standard methods to new and innovative techniques Techniques in Aquatic Toxicology provides necessary details on sampling testing and analysis in both saltwater and freshwater environments Research scientists and field and laboratory technicians will find help in testing for everything from assessing DNA damage to bioaccumulation of common toxins to assays of fish embryos and fish tissues **Adjuvant Therapy of Breast Cancer** I. Craig Henderson,2012-12-06 The results of randomized trials evaluating the use of early or adjuvant systemic treatment for patients with resectable breast cancer provide an eloquent rebuttal to those who would argue that we have made no progress in the treatment of cancer Many of the tumors that we have been most successful in curing with chemotherapy and other newer forms of treatment are relatively uncommon In contrast breast cancer continues to be the single most common malignancy among women in the western world is increasingly a cause of death throughout Asia and Third World countries and remains one of the most substantial causes of cancer mortality world wide The use of mammography as a means of early detection has been shown to reduce breast cancer mortality by 25 35% among those populations in which it is utilized The use of adjuvant systemic treatment in appropriate patients provides a similar and additional reduction in breast cancer mortality Few subjects have been so systematically studied in the history of medicine and it seems fair to conclude that the value to adjuvant systemic therapy in prolonging the lives of women with breast cancer is more firmly supported by empirical evidence than even the more conventional or primary treatments using various combinations of surgery and radiotherapy *Multidrug Resistance* Loretta D. Ulin,1992 Gastrointestinal Oncology A. B. Benson,2012-12-06 Gastrointestinal tract malignancies encompass a spectrum of diseases with marked differences in etiology incidence biological behavior and treatment outcome The incidence of new gastrointestinal cases recorded yearly for patients residing in the United States is approximately 230 000 representing nearly 20% of all cancer cases Worldwide gastrointestinal malignancies are responsible for the largest number of cancer deaths particularly because of the high incidence of hepatocellular and gastric cancer in other countries Gastrointestinal Oncology is directed to those most involved in the multidisciplinary approach to the gastrointestinal cancer patient including medical radiation and surgical oncologists gastroenterologists interventional radiologists pathologists oncology nurses and oncology physicians in training *Anticancer Research* ,2004

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