

# **ENZYME-PRODRUG STRATEGIES FOR CANCER THERAPY**

**Edited by  
Roger G. Melton and  
Richard J. Knox**

# Enzyme Prodrug Strategies For Cancer Therapy

**Ajit S. Narang, Ram I. Mahato**



## **Enzyme Prodrug Strategies For Cancer Therapy:**

*Enzyme-Prodrug Strategies for Cancer Therapy* Roger G. Melton, Richard J. Knox, 2012-12-06 Antibody directed enzyme prodrug therapy ADEPT directly addresses the major problem in cancer chemotherapy its lack of selectivity Antibody delivery combined with the amplification provided by the enzymatic activation of prodrugs enables selection to be made between tumour and normal tissue ADEPT offers a novel field of opportunities in the therapy of systemic cancer and may be a major advance for the treatment of solid tumours This book is the first to describe ADEPT in detail Each chapter reviews an aspect of the immunology enzymology biochemistry chemistry and cancer chemotherapy which have been integrated into the ADEPT concept An additional chapter describes the related approach of gene directed enzyme prodrug therapy GDEPT This latter approach is still in its infancy but ADEPT has entered the clinic The initial clinical studies with ADEPT are included and discussed in detail

**Handbook of Oxidative Stress in Cancer: Therapeutic Aspects** Sajal Chakraborti, 2022-09-28 This reference book which is the second volume of Targeting Oxidative Stress in Cancer explores oxidative stress as the potential therapeutic target for cancer therapy The initial chapters discuss the molecular mechanisms of oxidative stress and its effects on different signaling pathways Subsequently the sections examine the impact of redox signaling on tumor cell proliferation and consider the therapeutic potential of dietary phytochemicals and nutraceuticals in reactive oxygen species ROS induced cancer In turn it examines the evidence supporting the use of Vitamin C in cancer management before presenting various synthetic and natural compounds that have therapeutic implications for oxidative stress induced cancer It also explores the correlation between non coding RNA and oxidative stress Furthermore the book summarizes the role of stem cells in ROS induced cancer therapy and reviews the therapeutic applications of nanoparticles to alter redox haemostasis in cancer cells Lastly it explores heat shock proteins ubiquitin ligases and probiotics as potential therapeutic agents in ROS mediated cancer This book is a useful resource for basic and translational scientists as well as clinicians interested in the field of oxidative stress and cancer therapy

**Molecular Biomarkers for Cancer Diagnosis and Therapy** Ranbir Chander Sobti, Haruhiko Sugimura, Aastha Sobti, 2024-06-29 This book presents recent advancements challenges and clinical implications of molecular biomarkers in various cancers This book highlights the clinical applications of biomarkers are extensive for cancer risk assessment screening and early detection of cancer accurate diagnosis patient prognosis prediction of response to therapy and cancer surveillance and monitoring response It also explores the progress of predictive biomarkers as an adjunctive tool to tumor immunotherapy in effectively identifying the efficacy of immune checkpoint inhibitors and discusses their future directions in achieving precision immuno oncology Further the book examines the combination of next generation sequencing and advanced computational data analysis approaches in the understanding of the genomic underpinnings of cancer development and progression Towards the end the chapter discusses the role of some commonly investigated phytochemicals and their epigenetic targets that are of particular interest in cancer

prevention and cancer therapy It is a must for researchers as well as advanced students and physicians in the field of cancer and clinical oncologists *Cancer Gene Therapy* David T. Curiel, Joanne T. Douglas, 2007-11-03 A complete introduction and guide to the latest developments in cancer gene therapy from bench to bedside The authors comprehensively review the anticancer genes and gene delivery methods currently available for cancer gene therapy including the transfer of genetic material into the cancer cells stimulation of the immune system to recognize and eliminate cancer cells and the targeting of the nonmalignant stromal cells that support their growth They also thoroughly examine the advantages and limitations of the different therapies and detail strategies to overcome obstacles to their clinical implementation Topics of special interest include vector targeting techniques the lessons learned to date from clinical trials of cancer gene therapy and the regulatory guidelines for future trials Noninvasive techniques to monitor the extent of gene transfer and disease regression during the course of treatment are also discussed

**Prodrugs and Targeted Delivery** Jarkko Rautio, 2011-01-11 This topical reference and handbook addresses the chemistry pharmacology toxicology and the patentability of prodrugs perfectly mirroring the integrated approach prevalent in today's drug design It summarizes current experiences and strategies for the rational design of prodrugs beginning at the early stages of the development process as well as discussing organ and site selective prodrugs Every company employing medicinal chemists will be interested in this practice oriented overview of a key strategy in modern drug discovery and development

Quinones and Quinone Enzymes, Part B, 2004-03-08 Quinones are members of a class of aromatic compounds with two oxygen atoms bonded to the ring as carbonyl groups This volume covers more clinical aspects of quinines such as anticancer properties as well as their role in nutrition and in age related diseases Mitochondrial Ubiquinone and Reductases Anticancer Quinones and Quinone Oxido Reductases Quinone Reductases Chemoprevention Nutrition Quinones and Age Related Diseases

*Suicide Gene Therapy* Caroline J. Springer, 2008-02-01 Gene therapy has expanded rapidly over the last decade The number of clinical trials reported by 2001 included 532 protocols and 3436 patients Phase I trials predominate with 359 trials of 1774 patients versus Phase II 57 trials with 507 patients and Phase III 3 trials of 251 patients The disease overwhelmingly targeted by gene therapy is cancer involving 331 trials with 2361 patients Despite the somewhat disappointing results of clinical trials to date gene therapy offers tremendous promise for the future of cancer therapy The area of gene therapy is vast and both malignant and nonmalignant cells can be targeted Suicide Gene Therapy Methods and Reviews covers gene therapy that targets malignant cells in a treatment that has become known as suicide gene therapy Basically this approach uses the transduction of cancer cells with a gene for a foreign enzyme that when expressed is able to activate a nontoxic prodrug into a highly cytotoxic drug able to kill the cancer cell population This is a major area in cancer gene therapy in 2001 this technique was represented by 52 clinical protocols with a total of 567 patients Additional trials used multiple gene therapy protocols that also involved suicide gene therapy 83 with 497 patients indicating that the interest in this area is considerable Suicide Gene Therapy Methods and Reviews aims to

cover comprehensively both in theoretical and practical terms the rapidly evolving area of suicide gene therapy for cancer

Pharmaceutical Biocatalysis Peter Grunwald, 2020-11-01 This volume of Pharmaceutical Biocatalysis starts with a discussion on the importance of biocatalytic synthesis approaches for a sustainable and environmentally friendly production of pharmaceuticals and active pharmaceutical ingredients Among the enzymes discussed in detail with respect to their pharmaceutical relevance are cyclic nucleotide phosphodiesterases playing an important role in modulating signal transduction in various cell types human DOPA decarboxylase related to Parkinson's disease and aromatic amino acid decarboxylase deficiency and phospholipase D enzymes as drug targets Isocitrate dehydrogenase 1 and 2 mutations are novel therapeutic targets in acute myeloid leukemia An additional chapter is devoted to the use of enzymes for prodrug activation in cancer therapy The other topics include small molecule inhibitors targeting receptor tyrosine kinases in cancer Lactams and related compounds as antibacterials non vitamin K oral anticoagulants for the treatment of thromboembolic diseases and the molecular mechanisms for statin pleiotropy and its clinical relevance in cardiovascular diseases The last chapter is a review of lysosomal storage disorders with an overview of approved drugs for treating these disorders by enzyme replacement therapy **Laser-Driven Particle Acceleration Towards Radiobiology and Medicine** Antonio

Giulietti, 2016-05-04 This book deals with the new method of laser driven acceleration for application to radiation biophysics and medicine It provides multidisciplinary contributions from world leading scientist in order to assess the state of the art of innovative tools for radiation biology research and medical applications of ionizing radiation The book contains insightful contributions on highly topical aspects of spatio temporal radiation biophysics evolving over several orders of magnitude typically from femtosecond and sub micrometer scales Particular attention is devoted to the emerging technology of laser driven particle accelerators and their application to spatio temporal radiation biology and medical physics customization of non conventional and selective radiotherapy and optimized radioprotection protocols *Targeted Delivery of Small and*

*Macromolecular Drugs* Ajit S. Narang, Ram I. Mahato, 2010-05-12 Site specific drug delivery and targeting attracts much research interest from both academia and industry but because of the many challenges faced in the development of these systems only a handful of targeted therapies have successfully made it into clinical practice Focusing on the delivery technologies that utilize both systemic and local routes Nanotechnologies for Drug Delivery and Biopharmaceutical Development Virat Khanna, Shweta Dang, Alka Ahuja, 2025-09-26 This book is a comprehensive coverage of the transformative

role of nanotechnology in the field of biopharmaceuticals The book covers the fundamental principles of nanotechnology and systematically explains how nanomaterials are revolutionizing drug delivery systems formulation techniques diagnostics and imaging modalities Through detailed discussions on lipid based nanocarriers polymeric nanoparticles dendrimers and nanogels readers gain insights into the diverse array of nanomaterials and their applications in enhancing drug efficacy targeting specific tissues and minimizing adverse effects Moreover the book addresses critical aspects such as

biocompatibility assessment regulatory considerations and ethical implications underscoring the importance of responsible innovation in the development and commercialization of nano biopharmaceuticals This book serves as an invaluable resource for researchers and practitioners to navigate the complex landscape of biopharmaceutical development Through its interdisciplinary approach and forward thinking perspectives on future trends and challenges it contributes to shaping the future of healthcare by harnessing the power of nanotechnology to deliver innovative and effective therapeutics The target audience for this book covers the fields of biopharmaceutical research development regulation and policymaking This includes scientists researchers and engineers seeking a deep understanding of nanotechnology s applications in drug delivery systems formulation techniques and diagnostic tools Additionally pharmaceutical industry professionals involved in drug development and manufacturing will find valuable insights into innovative nanoscale formulations and regulatory considerations Overall this book caters to a multidisciplinary audience seeking to leverage nanotechnologies for advancing biopharmaceutical research development and clinical practice

*Drug Delivery* Binghe Wang, Teruna J. Siahaan, Richard A. Soltero, 2005-04-29 An indispensable tool for those working at the front lines of new drug development Written for busy professionals at the forefront of new drug development *Drug Delivery* gets readers quickly up to speed on both the principles and latest applications in the increasingly important field of drug delivery Recent developments in such areas as combinatorial chemistry proteomics and genomics have revolutionized researchers ability to rapidly identify and synthesize new pharmacological compounds However delivery related properties remain a significant reason for clinical trial failures Bringing together contributions by leading international experts *Drug Delivery* covers the entire field in a systematic but concise way It begins with an in depth review of key fundamentals such as physiochemical and biological barriers drug delivery pathways metabolism drug formulation pharmacokinetic and pharmacodynamic issues and more The remainder of the book is devoted to the systematic examination including overviews timely examples and extensive references of a host of specific subjects including Receptor mediated drug delivery Prodrug delivery approaches Oral protein and peptide drug delivery Gene therapy and gene delivery Ultrasound mediated drug delivery Polycationic peptides and proteins in drug delivery Pulmonary drug delivery Antibody directed drug delivery Efflux transporters in drug excretion Intellectual property issues in drug delivery

*Novel Gene Therapy Approaches* Ming Wei, David Good, 2013-02-13 Gene therapy has the potential to be a tailor made therapeutic with increased specificity and decreased side effects that can offer a cure for many disorders The aim of this book is to provide up to date reviews of the rapidly growing field of gene therapy Chapters cover a large range of topics including methods of gene delivery and identification of targets with several papers on cancer gene therapy If more people become aware of the true nature and potential of gene therapy perhaps we can achieve the full benefit of such an innovative approach for the treatment of a range of diseases including cancer

**Tumor Targeting in Cancer Therapy** Michel Pagé, 2002-04-01 This volume introduces the principles and techniques of tumor targeting and critically surveys their

applications from laboratory to bedside By concisely synthesizing the many technical details the authors illuminate this innovative technique ranging from the fundamentals of drug targeting and in vivo and in vitro experimentation to such emerging therapeutic uses as radioimmunotherapy radioimmunodetection therapy with cytotoxic antibodies immunotoxins enzyme prodrug immunotherapy and immunotherapeutics with fusion proteins *Current Strategies in Cancer Gene Therapy* Wolfgang Walther,2017-01-17 This book describes important developments and emerging trends in experimental and clinical cancer gene therapy It reflects the tremendous advances made over recent years with respect to immunogenes suicide genes and gene correction therapies as well as in gene suppression and miRNA therapies Many of the described strategies focus on the generation of more efficient and specific means of attack at known and novel cellular targets associated with tumor development and progression The book also details parallel improvements in vector design vector delivery and therapeutic efficacy It offers readers a stimulating broad overview of advances in the field linking experimental strategies to their clinical applications Prodrugs Valentino Stella,Ronald Borchardt,Michael Hageman,Reza Oliyai,Hans Maag,Jefferson Tilley,2007-03-12 These volumes represent a comprehensive guide to prodrugs They guide the reader through the current status of the prodrug concept and its many applications and highlight its many successes in overcoming formulation and delivery of problematic drugs Replete with examples of approved and marketed prodrugs these volumes introduce the topic to the novice as well as professional in the design of prodrugs Recent Advancement in Prodrugs Kamal Shah,Durgesh Nandini Chauhan,Nagendra Singh Chauhan,Pradeep Mishra,2020-05-13 Recent Advancement in Prodrugs Drugs used as medicines have many limitations like low chemical stability aqueous solubility or oral absorption bioavailability rapid presystemic metabolism toxicity inadequate site specificity or poor patient acceptance compliance unwanted adverse effects unacceptable taste or odor irritation or pain Prodrugs design is an approach to overcome these limitations Key features Covers recent advancements in development of prodrugs Presents balanced synthesis and applications of prodrug chemistry Discusses broad spectrum of prodrug categories and outlines industrial applications Reviews prodrugs in cancer nanomedicine its therapy and treatment Elucidates mathematical models to study the kinetics of prodrugs This book covers recent advances in the design of prodrugs It contains all the significant recent examples of prodrug chemistry developments and will aid academics and researchers seeking to generate new projects in the field

**Innovative Approaches to Overcome Resistance and Toxicities of Anti-Cancer Drugs** Awanish Mishra,Dayanidhi Raman,Valeria Consoli,2025-08-18 Although diagnosis and treatment of various cancer types have made significant strides recently drug resistance is a major challenge faced in the cancer clinic Cancer cells evolve continuously through a combination of genetic mutations epigenetic changes support from cellular and acellular tumour microenvironment The chemoresistant tumour cells expand and become the dominant population and at this point it becomes difficult to treat The cancer cell heterogeneity is also a major contributing factor to chemoresistance The other challenge faced is the

development of adverse events due to drug toxicity which is overwhelming especially for immunocompromised patients Collectively these factors reduce the treatment response and overall survival Current neoadjuvant chemotherapy NACT and targeted therapies aim at drug efficacy with minimal toxicity along with employment of adjuvant immunotherapy Potential exploits include novel drug delivery platforms such as antibody drug conjugates combination therapies that target addicted signalling pathways transcription factors utilization of long noncoding RNAs including siRNA and miRNA using nanocarriers reprogramming the tumour immune microenvironment TIME employment of in silico approaches from docking drug like molecules to crystal structures of novel targets bioinformatics and machine learning approaches These approaches hold immense potential to enhance cancer therapeutics while minimizing toxicities These strategies aim to amplify therapy impact while minimizing toxicity leading to better patient outcome This research topic welcomes data and review articles on the following sub topics but are not limited to 1 Novel molecular targets targeting of signalling pathways transcription and epigenetic factors proteomic metabolomic and single cell analyses of therapy naïve and chemoresistant tumour cell populations 2 Role of non coding RNAs and microRNAs in chemoresistance 3 Advancements in tumour immune microenvironment TIME and therapies taking advantage of reprogramming the TIME 3 Novel synthetic and natural derived compounds for targeted therapy to improve anti cancer efficacy overcoming resistance and minimizing toxicities 4 Cancer stem or stem like cells in creation of minimal residual disease and induction of drug resistance cancer stemness factors that induce and orchestrate chemoresistance 5 Bioinformatics in silico studies and machine learning in design for the study of novel molecules to enhance efficacy and overcome resistance to anticancer drugs and toxicities In silico results should be validated through the exploitation of experimental methodologies

**Handbook of Oncobiology: From Basic to Clinical Sciences** R. C. Sobti, Nirmal K. Ganguly, Rakesh Kumar, 2024-05-14 This Handbook on cancer biology comprehensively reviews the current status of the oncobiology of major cancer types cancer detection and treatment strategies principles and processes of cancer drug development and nanomedicine and other emerging cancer medicine applications to cancer diagnosis and treatment The book also provides practical and implementable nutritional guidance in cancer prevention treatment and quality of life for cancer survivors It discusses pharmacogenetics strategies for predicting cancer prognosis and treatment exposure response and toxicity Further it presents bioinformatics approaches for predicting anti cancer drugs and drug combinations based on the multi omic data including transcriptomics toxicogenomics functional genomics and biological networks The Handbook also examines major factors and pathways that regulate cancer stem cells development and discusses potential targeted therapy for cancer stem cells The book explores the application of the CRISPR Cas9 based gene editing technique in basic cancer research diagnosis and treatment of cancer This Handbook is an invaluable source for oncologists researchers public health specialists epidemiologists and policy makers

*Drug Delivery in Oncology* Felix Kratz, Peter Senter, Henning Steinhagen, 2013-09-30 In this first authoritative overview on modern cancer chemotherapy 121



international specialists have contributed their experience and recent data for what is likely to become the gold standard in the field. The authors summarize knowledge gained over the past decade from basic concepts to successful applications in the clinic covering active and passive targeting strategies as well as tissue specific approaches. All current and future targeted delivery systems are discussed from ligand based to antibody based polymer based systems right up to micro and nanoparticulate systems. A special section covers the delivery of nucleic acid therapeutics such as siRNA, miRNA and antisense nucleotides. In each case a description of the basic technique is followed by a discussion of the latest preclinical and clinical developments in the field. By virtue of its clear and didactic structure, rich illustrative material and summary chapters, this handbook and ready reference enables the efficient transfer of knowledge between different disciplines from basic research to the clinician and vice versa. It is equally well suited for professionals, researchers and students in medical oncology and cancer biology and is also excellent for teaching medical students the foundations of 21st century cancer chemotherapy.

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