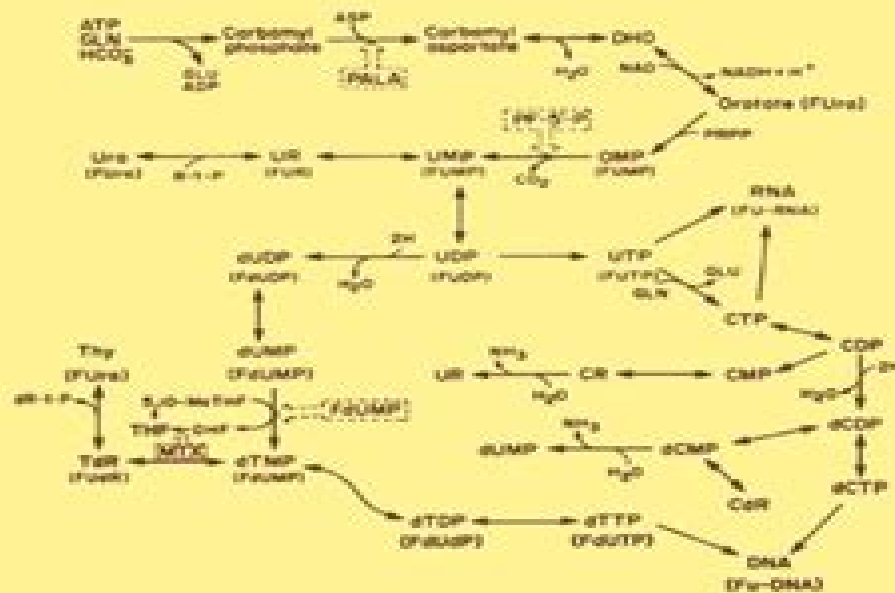


BIOCHEMICAL MODULATION OF ANTICANCER AGENTS: EXPERIMENTAL AND CLINICAL APPROACHES

Frederick A. Valeriote
Laurence H. Baker (editors)



Martinus Nijhoff Publishing

Biochemical Modulation Of Anticancer Agents

Lori J. Goldstein, Robert F. Ozols



Biochemical Modulation Of Anticancer Agents:

Biochemical Modulation of Anticancer Agents: Experimental and Clinical Approaches Frederick A. Valeriote, L.O. Baker, 2012-12-06 Biochemical Modulation at the present time defines an area of study in which the intracellular metabolism of a given anti cancer agent is modulated usually by a noncytotoxic agent or a cytotoxic agent at sufficiently low dosage to make it non cytotoxic in order to either increase the effectiveness of the particular agent against tumor cells or decrease its cytotoxicity against normal cells The major focus of modulation has been the agents 5 fluorouracil FUra arabinofuranosylcytosine ara C methotrexate MTX and a few alkylating agents The major thrust of the studies has been to increase the flow of the anticancer agent along the pathway responsible for the formation of the cytotoxic species for example FUra to FUTP or ara C to ara CTP While in most cases the application of research results to clinical trials does not require the subsequent expertise of the laboratory researchers application of biochemical modulatory schemes to clinical protocols necessitate a dramatic break with the past procedures As shown in the laboratory clinical loop below close collaboration between the laboratory and clinical investigator is essential While the laboratory REDEFINE TECHNOLOGY TESTS OR QUESTIONS FOR FURTHER THERAPEUTIC ADVANCE CLINICAL EXPERIMENTAL PROTOCOL LABORATORY RESEARCH STUDIES DEFINE AND TEST APPROPRIATE SCIENTIFIC PARAMETERS results define rationally based regimens it is essential that the clinical protocols contain the requirement that clinical material either tumor or normal tissues be sampled to determine whether the biochemical modulation being proposed is in fact being accomplished

Preclinical and Clinical Modulation of Anticancer Drugs Kenneth D. Tew, Peter J. Houghton, Janet A. Houghton, 1993-09-15 Preclinical and Clinical Modulation of Anticancer Drugs focuses on the theoretical and practical approaches to designing and enacting modulation principles Each class of anticancer drug and the different types of modulators used within each drug class are discussed within individual chapters The molecular and biochemical rationale for the use of specific modulators is discussed in detail and preclinical and clinical implications of the data are integrated into each chapter Mechanisms of drug resistance and the reasons behind circumventing the resistant phenotype are covered The book will interest cancer chemotherapists pharmacologists oncologists biochemists and experimental therapeutics researchers in addition to students studying the principles of drug discovery and protocol design **International Cancer Congress** , 1986-08-22

Biomedical Index to PHS-supported Research: pt. A. Subject access A-H , 1992 **Biomedical Index to PHS-supported Research** , Anticarcinogenesis and Radiation Protection Peter Cerutti, 2012-12-06 This book is based on the invited and contributed papers presented at the 2nd International Conference on Anticarcinogenesis and Radiation Protection held at the National Bureau of Standards Gaithersburg Maryland USA on March 8 12 1987 The conference documented developments that have taken place in areas that were addressed during the first conference in 1982 A number of new topics such as biological response modifiers were included because of their emerging relevance to anticarcinogenesis

and radiation protection The organization of the material in this book does not follow the conference program rather we have attempted to provide a different sequence for didactic reasons The aim of the conference which is reflected in this book was to promote further development of mechanistic approaches to cancer prevention and treatment based on recent progress in molecular biology and free radical chemistry At the basis of carcinogenesis lie changes in the dynamics of growth and differentiation of specific cell subpopulations in the target tissue These changes are brought about by selective toxicity and modulation of gene expression that are induced by xenobiotic carcinogens and affected by physiological and genetic factors The book deals with oxidative stress and molecular damage caused by radiation and chemical pro oxidants and their role in carcinogenesis and it discusses mechanisms of deregulation of the expression of oncogenes and other genes involved in carcinogenic initiation and promotion

Enzymes Dependent on Pyridoxal Phosphate and Other Carbonyl Compounds as Cofactors T. Fukui, 2013-10-22 This volume is a summary of the proceedings of the 8th International Symposium on Vitamin B6 and Carbonyl Catalysis held in Osaka Japan 15-19 October 1990 The development of molecular biology and x ray crystallography techniques have contributed greatly to the knowledge and understanding of PLP dependent enzymes The structural mechanistic and functional aspects of the PLP dependent enzymes have been studied for over 40 years by many active researchers Recent evidence indicates that the coenzyme is substituted by the pyruvoyl group which is a reactive carbonyl in some amino acid decarboxylases and the presence of pyrroloquinoline quinone PQQ another carbonyl coenzyme in other enzymes The contents of this volume attest to the exciting progress in elucidating structure mechanism of action and the function of enzymes dependent on coenzymes containing the carbonyl group

Cancer Treatment Reports, 1986

Novel Approaches to Selective Treatments of Human Solid Tumors Youcef M. Rustum, 2012-12-06

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

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