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Inflammation in the Pathogenesis of Chronic Diseases

The COX-2 Controversy

Edited by

Randall E. Harris



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Inflammation in the Pathogenesis of Chronic Diseases Randall E. Harris, 2007-05-09 In this book a worldwide panel of leading experts discuss the role of inflammation in the pathogenesis of major chronic diseases and the current controversy regarding risk versus benefit of selective cyclooxygenase 2 COX 2 inhibitors. The authors provide exciting and enlightening perspectives on COX 2 and related molecular targets in the future of medicine including historical perspectives Allergy Brian A. Baldo, Nghia H. Pham, 2020-12-08 The second edition of this book spans the broad range of modern therapeutic drugs from small molecules to biologic recombinant proteins It offers a comprehensive review of the classification and description of different drug induced systemic and cutaneous hypersensitivities an up to date coverage of individual culprit drugs in each group of therapeutics the diagnosis and mechanisms of reactions and important structure activity relationships New content expands to two areas of drug allergy that have recently experienced explosive growth biological therapies and new targeted chemotherapies Other new and expanded chapters address antimicrobials drugs used in anesthesia and surgery opioids non targeted anti cancer drugs vaccines and newly understood reaction mechanisms This new edition includes photographs of a wide variety of cutaneous manifestations that will be of use to other clinicians as well as allergists and dermatologists In addition to its wide clinical emphasis the book s mechanistic and structure activity detail will provide valuable background for researchers and investigators in universities medical research institutes drug companies and regulatory agencies The second edition of Drug Allergy is an essential reference for practitioners across the medical disciplines from specialist clinicians surgeons GPs residents and medical students to nurses pharmacists dentists and those taking undergraduate and graduate courses in the biomedical sciences Inflammation, Lifestyle and Chronic Diseases Bharat B. Aggarwal, Sunil Krishnan, Sushovan Guha, 2016-04-19 Oxidative stress and inflammation are among the most important factors of disease Chronic infections obesity alcohol and tobacco usage radiation environmental pollutants and high calorie diets have been recognized as major risk factors for a variety of chronic diseases from cancer to metabolic diseases All these risk factors are linked to ch *Improved Non-Steroid Anti-Inflammatory Drugs: COX-2 Enzyme Inhibitors* Sir John R. Vane, Jack H. Botting, R.M. Botting, 2012-12-06 In 1971 Vane proposed that the mechanism of action of the aspirin like drugs was through their inhibition of prostaglandin biosynthesis Since then there has been intense interest in the interaction between this diverse group of inhibitors and the enzyme known as cyclooxygenase COX It exists in two isoforms COX l and COX 2 discovered some 5 years ago Over the last two decades several new drugs have reached the market based on COX l enzyme screens Elucidation of the three dimensional structure of COX l has provided a new understanding for the actions of COX inhibitors The constitutive isoform of COX COX l has clear physiological functions Its activation leads for instance to the production of prostacyclin which when released by the endothelium is anti thrombogenic and anti atherosclerotic and in the gastric mucosa is cyto protective COX l also generates prostaglandins in the kidney where they

help to maintain blood flow and promote natriuresis The inducible isoform COX 2 was discovered through its activity being increased in a number of cells by pro inflammatory stimuli A year or so later COX 2 was identified as a distinct isoform encoded by a different gene from COX I COX 2 is induced by inflammatory stimuli and by cytokines in migratory and other cells Thus the anti inflammatory actions of non steroid anti inflammatory drugs NSAIDs may be due to the inhibition of COX 2 whereas the unwanted side effects such as irritation of the stomach lining and toxic effects on the kidney are due to Inflammation, Chronic Diseases and Cancer Mahin Khatami, 2012-03-09 inhibition of the constitutive enzyme COX I This book is a collection of excellent reviews and perspectives contributed by experts in the multidisciplinary field of basic science clinical studies and treatment options for a wide range of acute and chronic inflammatory diseases or cancer The goal has been to demonstrate that persistent or chronic unresolved or subclinical inflammation is a common denominator in the genesis progression and manifestation of many illnesses and or cancers particularly during the aging process Understanding the fundamental basis of shared and interrelated immunological features of unresolved inflammation in initiation and progression of chronic diseases or cancer are expected to hold real promises when the designs of cost effective strategies are considered for diagnosis prevention or treatment of a number of age associated illnesses such as autoimmune and neurodegenerative diseases as well as many cancers The Physiology of Inflammation - The Final Common Pathway to Disease Alexandrina Ferreira Mendes, Maria Teresa Cruz, Oreste Gualillo, 2019-02-20 Chronic diseases are increasingly recognized as involving low grade inflammation that is a self perpetuating tissue response to stress caused by exogenous or endogenous triggers that progressively evokes danger associated molecular pattern release ultimately driving tissue damage and loss of function This response is frequently unapparent clinically thus the designation low grade This eBook comprises nineteen reviews and original articles that provide the most updated knowledge on the causes and roles of this inflammatory response in a variety of diseases and conditions. The editorial that precedes these articles not only summarizes each one but provides a broader interpretation of the role of inflammation in health and a variety of disease conditions the underlying mechanisms and the targets more promising for therapy Finally it also highlights the most relevant and emerging research topics that are already shaping future directions for the development of more fine tuned and innovative therapies Regulation of Inflammation in Chronic Disease Jixin Zhong, Guixiu Shi, 2019-07-19 This eBook is a collection of articles from a Frontiers Research Topic Frontiers Research Topics are very popular trademarks of the Frontiers Journals Series they are collections of at least ten articles all centered on a particular subject With their unique mix of varied contributions from Original Research to Review Articles Frontiers Research Topics unify the most influential researchers the latest key findings and historical advances in a hot research area Find out more on how to host your own Frontiers Research Topic or contribute to one as an author by contacting the Frontiers Editorial Office frontiers in org about contact Biochemistry of Inflammation J. Whicher, S.W. Evans, 1992-02-29 Our understanding of inflammation has increased

rapidly in recent years due in large part to the impact of molecular biology and gene identification and cloning This book brings together ideas from a number of different biochemical disciplines which are frequently not integrated The first chapter gives a visual overview of the subject the remaining chapters are organized into three themes the affector molecules the regulatory components and the processes of inflammation itself This book is essential reading for the busy physician or pathologist who wants to be up to date with the latest developments in immunology as they affect the diagnosis and treatment of many conditions Chronic Inflammation Sashwati Roy, Debasis Bagchi, Siba P. Raychaudhuri, 2012-09-17 Although acute inflammation is a healthy physiological response indicative of wound healing chronic inflammation has been directly implicated in a wide range of degenerative human health disorders encompassing almost all present day non communicable diseases including autoimmune diseases obesity diabetes and atherosclerosis Chronic Inflammation Molecular Pathophysiology Nutritional and Therapeutic Interventions provides an exposition of the process of chronic inflammation in three parts Systems Biology of Inflammation and Regulatory Mechanisms describes the process of chronic inflammation including initiation progression and resolution Pathologies Associated with Inflammation gives a rigorous and critical treatment of specific human health disorders where chronic inflammation plays a major role Nutrition Therapeutics for Inflammatory Diseases details the protective abilities of structurally diverse antioxidants phytochemicals anti inflammatory diets omega 3 fatty acids NSAIDs disease modifying anti rheumatic drugs and novel regimens Designed for scientists as well as clinicians Chronic Inflammation provides critical understanding of the key checkpoints that regulate chronic inflammation Going beyond the epidemiology of chronic inflammation the text covers regulatory mechanisms controlling inflammation initiation progression and resolution The authors address pathologies associated with inflammation and provide various **New Targets in Inflammation** N. Bazan, Jack H. nutritional and therapeutic interventions for inflammatory diseases Botting, Sir John R. Vane, 2012-12-06 For the past 100 years the mainstay of therapy for rheumatoid arthritis RA has been aspirin or other drugs of the non steroid anti inflammatory group In 1971 Vane pro posed that both the beneficial and toxic actions of these drugs was through inhibition of prostaglandin synthesis The recent discovery that prostaglandins responsible for pain and other symptoms at inflammatory foci are synthesized by an inducible cyclooxygenase COX 2 that is encoded by a gene distinct from that of the constitutive enzyme COX I provided a new target for therapy of RA A drug that would selectively inhibit COX 2 would hopefully produce the symptomatic benefit provided by existing NSAIDs without the gastrointestinal and renal toxicity due to the inhibition of COX I Drugs selective for COX 2 are now available Experimental studies have shown them to be effective with minimal toxicity and in clinical trials gastric and renal toxicities are less Highly selective COX 2 inhibitors perhaps designed with knowledge of the crystal structures of COX I and COX 2 are also available Other experimental studies including those in animals lacking effective genes for COX lor COX 2 and in experimental carcinomas suggest there is still much to be learned of the pathophysiological functions of these enzymes. The inflammatory

response is a complex reaction involving many mediators that derive from white blood cells endothelial cells and other tissues Preliminary data have revealed that inhibitors of the cytokines and adhesion molecules that play a crucial role in the migration of white cells to inflammatory sites may be useful in RA Chronic Inflammation Masayuki Miyasaka, Kiyoshi Takatsu, 2016-11-01 This book provides readers with the most up to date information on cutting edge research concerning chronic inflammation We now know that when inflammation becomes chronic it acts as a strong disease promoting factor in a variety of disorders including arteriosclerosis obesity cancer and Alzheimer disease Chronic inflammation is hence called as the silent killer it upsets the body s homeostatic mechanism insidiously In spite of these developments we know very little about the mechanism underlying chronic inflammation Particularly we do not know precisely what induces chronic inflammation or what promotes its prolongation in a spatiotemporal framework Neither do we have clear knowledge about how chronic inflammation destroys various tissues or how it predisposes individuals to many different diseases To make the situation worse we have no effective treatment against chronic inflammation Since 2010 two major research programs CREST and PRESTO aimed at clarifying the mechanisms underlying chronic inflammation were launched in Japan and investigators of different research areas with a brilliant track record were selected by their research proposals Subsequently they have made their best efforts to answer the conundrum concerning chronic inflammation This book is a compendium of such research efforts In each chapter the CREST or PRESTO funded researchers summarize their original work concerning mechanisms of induction progression or resolution underlying chronic inflammation The most emphasized characteristic is the molecular aspect of chronic inflammation The book thus presents the most recent progress made in the molecular understanding of chronic inflammation Molecular and Cellular Basis of Inflammation Charles N. Serhan, Peter A. Ward,1998-12-09 The study of inflammation has captured the interest of scholars since the earliest recorded history Symbols identifying the cardinal signs of inflammation were uncovered in both Sanskrit and hieroglyphics 1 Since complete apprecia tion of the inflammatory process is underscored by the need for knowledge at both the cellular and molecular levels academic inquiry in the area of inflammation has led in many respects the foray of current biomedical research Molecular and Cellular Basis of Inflammation represents research from the cutting edge in the broad view of inflammation The chapters are written by experts with a multidisciplinary approach to the study of inflammatory and cellular processes and thus include contributions form the fields of molecular biology biochemistry pharmacology immunology and pathobiology Molecular and Cellular Basis of Inflammation was first conceived during a mini symposium sponsored by the American Society for Investigative Pathology held at FASEB in 1995 entitled The Role of Reactive Lipids Oxygen and Nitro gen Metabolites in Inflammation at which several of the contributing authors delivered lectures This present much extended volume includes leading front descriptions of both protein and lipid mediators The chapter devoted to the comple ment cascade by Ward and colleagues as well as Chapters 3 7 and 13 provide up to date descriptions of the biosynthesis molecular biology chemistry

and actions of both protein and lipid mediators Inflammation Jean-Marc Cavaillon, Mervyn Singer, 2017-10-02 The leading reference on this topic of increasing medical relevance is unique in offering unparalleled coverage. The editors are among the most respected researchers in inflammation worldwide and here have put together a prestigious team of contributors Starting with the molecular basis of inflammation from cytokines via the innate immune system to the different kinds of inflammatory cells they continue with the function of inflammation in infectious disease before devoting a large section to the relationship between inflammation and chronic diseases The book concludes with wound and tissue healing and options for therapeutic interventions A must have for clinicians and biomedical researchers alike The Molecular Mechanisms of Chronic Inflammation Development Masaaki Murakami, Toshio Hirano, Inflammation is critical for the development of many complex diseases and disorders including autoimmune diseases metabolic syndrome neurodegenerative diseases cancers and cardiovascular diseases Inflammation comes as two types chronic inflammation which can be defined as a dysregulated form of inflammation and acute inflammation which can defined as a regulated form Because of its special role in the aforementioned diseases establishing methods to control chronic inflammation is important for developing cures and treatments One challenge for this purpose has been the ability to distinguish chronic and acute inflammation based on molecular biology diagnostics Thus this Research Topic is focused on articles that can shed some new light on the molecular mechanisms responsible for the development of chronic inflammation and its related conditions COX-2 Inhibitors Michel Pairet, Joanne van Ryn, 2012-12-06 COX 2 inhibitors are important drugs with analgesic and anti inflammatory effects The discovery of COX 2 the evolution of drug development in this field and the implications of these developments in patient therapy are topics of this volume This book presents both pre clinical and clinical information and is important for clinicians interested in the latest information about this class of drugs for researchers and for students in the field Inhibitors Sir John R. Vane, Jack H. Botting, 2012-12-06 The mainstay of therapy for rheumatoid disease is the non steroid antiinflammatory drugs NSAIDs despite their inherent gastrointestinal toxicity and ability to cause renal damage in susceptible patients The theory that the beneficial and toxic effects of NSAIDs stem from a reduction in prostanoid production through inhibition of cyclooxygenase implied that particular toxicities were inevitable with NSAIDs and would always be correlated with efficacy However over the years it became apparent that at therapeutic doses some NSAIDs had greater toxic side effects than others a fact not explained by the general theory A significant clarification arose from the discovery that there are two distinct isoforms of COX a constitutive enzyme COX I responsible for the production of prostanoids necessary for platelet aggregation and protection of the gastric mucosa and kidney and an inducible enzyme COX 2 that is newly synthesized at sites of tissue damage and produces prostaglanding that manifest pathological effects It became clear that different NSAIDs had greater or lesser effects on COX I when used in therapeutic doses explaining the variation in side effects The elucidation of the crystal structure of these different enzymes and the skills of medicinal

chemists have led to the synthesis of new chemicals with a selectivity for the inducible enzyme and thus with therapeutic efficacy without those toxic effects result ing from inhibition of the constitutive enzyme Inflammation Protocols Paul Winyard, Derek A. Willoughby, 2010-11-10 Inflammation has been described as the basis of many pathologies of human disease When one considers the updated signs of inflammation they would be vasodilation cell migration and in the case of chronic inflam tion cell proliferation often with an underlying autoimmune basis Gen ally inflammation may be divided into acute chronic and autoimmune though the editors believe that most if not all chronic states are often the result of an autoimmune response to an endogenous antigen Thus a proper understanding of the inflammatory basis may provide clues to new therap tic targets not only in classical inflammatory diseases but atherosclerosis cancer and ischemic heart disease as well The lack of advances in classical inflammatory diseases such as rh matoid arthritis may in part arise from a failure to classify the disease into different forms That different forms exist is exemplified in patients with d fering responses to existing antiinflammatory drugs ranging from nonresponders to very positive responders for a particular nonsteroidal an inflammatory drug NSAID Though researchers have progressively unreled the mechanisms the story is far from complete It should also be noted that the inflammatory response is part of the innate immune response or to use John Hunter's words in 1795 inflammation is a salutary response That may be applied in particular to the defensive response to invading micro Inflammation Resolution and Chronic Diseases Anurag Tripathi, Ashish Dwivedi, Shailendra Gupta, Shiv ganisms Poojan, 2024-04-25 This book provides recent advances in the understanding of the central processes in the resolution of inflammation It also describes the active specialized pathways that bring the resolution of inflammation Further the book examines the key molecular and cellular events that are involved in the resolution of inflammation processes involved in tissue repair and regain of physiological function Importantly it sheds light on the cellular and molecular processes through which conventional inflammatory cytokines control mammalian tissue repair and regeneration It also explores the perturbation of processes that are involved in the resolution of inflammation with aging and reviews clinical trials of anti inflammatory and immunomodulatory therapies Towards the end the book discusses the role of inflammation in the development and progression of cancer This book is an invaluable resource for scientists academicians and research scholars interested in the latest cutting edge knowledge in the field of immunology inflammation inflammation resolution and COX-2 Inhibition Impaired Resolution of Chronic Inflammation in a Murine Model of Autoimmune inflammatory diseases Arthritis Andrea Rossi Moore.2010

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Introduction

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