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In 1890 a case of myxedema was treated in Lisbon by the implantation of a sheep thyroid gland with the immediate improvement in the patient ' s condition. A few years later, medications for the then ill-explained condition of the menopause included tablets made from cow ovaries. In the first quarter of the 20th century the identification of vitamin D, and its sunlight driven production in skin, paved the way to the elimination of rickets as a major medical problem. Twenty years or so later, Sir Vincent Wigglesworth established the endocrine basis of developmental moulting in insects, arguably the most commonly performed animal behaviour on Planet Earth. A paradigm that would unify these disparate observations arose between 1985 and 1987 beginning with the identification of the glucocorticoid receptor and the nuclear receptor super-family. What follows is a timely and positive manifestation of the capacity, productivity and value of international human scientific endeavour. Based on intrigue, lively competition and cooperation a global effort has rapidly fostered a school of biology with widespread ramifications for the understanding of metazoan animals, the human condition and the state of the planet. This book is the first this century to try and capture the spirit of this endeavour, to depict where the field is now

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and to identify some of the challenges and opportunities for the future.

Acting principally to control patterns of gene expression, nuclear receptors play vital roles during embryonic development and in the regulation of metabolic and reproductive functions in adult life, which proves this superfamily of ligand-activated transcription factors to be a crucial part of biological life. In *The Nuclear Receptor Superfamily: Methods and Protocols*, expert researchers describe a range of molecular, structural and cell biological techniques currently used to investigate the structure-function of nuclear receptors, together with experimental approaches that may lead to new therapeutic strategies for treating nuclear receptor-associated diseases. Written in the highly successful *Methods in Molecular Biology*TM series format, the chapters in this volume contain brief introductions to the topics, lists of the necessary materials and reagents, step-by-step, readily reproducible laboratory protocols, as well as notes from the experts to highlight tips on troubleshooting and avoiding known pitfalls. Cutting-edge and easy to use, *The Nuclear Receptor Superfamily: Methods and Protocols* provides beneficial and time-saving guidance for all those undertaking research in this ever-growing field of study.

Nuclear receptors are ligand activated transcription factors that control numerous biological functions. Consequently, altering activity of these receptors is proposed, and indeed documented, to affect many physiological and pathological conditions in experimental animals and humans. Thus, nuclear receptors have become a major target in the effort to treat numerous diseases. This book will shed light on and emphasize intricate processes involved in designing as well as discovering physiological and pharmacological modulators of these important proteins. World-renowned scientists will share with the reader their professional expertise and extensive experience acquired through decades

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working with nuclear receptors. Chapters address the various means and consequences of modulating nuclear receptor activity will be presented and discussed. These modulators cover a wide span of moieties ranging from synthetic chemicals to natural products. In addition, the classification of these chemicals ranges from pan agonists to selective agonists and inverse agonists to antagonists. They also include proteolytic means to obliterate the receptor in the event that modulating its activity through canonical pharmacological agents becomes less effective and/or less desirable due to anticipated or experienced toxicities. Modulation of receptor activity may also take place in the absence of a ligand or through manipulating the structure of the receptor itself by controlling posttranslational events.

The publication of the extensive seven-volume work *Comprehensive Molecular Insect Science* provided a complete reference encompassing important developments and achievements in modern insect science. One of the most swiftly moving areas in entomological and comparative research is endocrinology, and this volume, *Insect Endocrinology*, is designed for those who desire a comprehensive yet concise work on important aspects of this topic. Because this area has moved quickly since the original publication, articles in this new volume are revised, highlighting developments in the related area since its original publication. *Insect Endocrinology* covers the mechanism of action of insect hormones during growth and metamorphosis as well as the role of insect hormones in reproduction, diapause and the regulation of metabolism. Contents include articles on the juvenile hormones, circadian organization of the endocrine system, ecdysteroid chemistry and biochemistry, as well as new chapters on insulin-like peptides and the peptide hormone Bursicon. This volume will be of great value to senior investigators, graduate students, post-doctoral fellows and advanced undergraduate research students. It can also be used as a reference for graduate courses and seminars on the topic. Chapters will also

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be valuable to the applied biologist or entomologist, providing the requisite understanding necessary for probing the more applied research areas. Articles selected by the known and respected editor-in-chief of the original major reference work, *Comprehensive Molecular Insect Science* Newly revised contributions bring together the latest research in the quickly moving field of insect endocrinology Review of the literature of the past five years is now included, as well as full use of data arising from the application of molecular technologies wherever appropriate

This book is a printed edition of the Special Issue "Molecular Science for Drug Development and Biomedicine" that was published in *IJMS*

Current Concepts in Cardiovascular Physiology examines seven different areas related to the field of cardiac physiology. In addition to the biochemistry and receptor pharmacology of the heart, this book explores coronary physiology, cardiovascular function, and neural and reflex control of the circulation. The electrophysiology and biophysics of cardiac excitation are also considered, along with humoral control of the circulation. This monograph consists of seven chapters and opens with an overview of the biochemistry of the heart, with emphasis on cardiac energy metabolism and the ways in which metabolism and the biochemical pathways are controlled. The mechanisms whereby physiological events influence biochemical activities and vice versa are also discussed. The following chapters look at the chemistry and physiology of myocardial receptors; the complex interplay between the nervous and cardiovascular systems; and the chemical and hormonal factors that regulate, modify, and modulate the cardiovascular system. The influence of humoral, neural, intrinsic, vascular, and myocardial factors on coronary blood flow is also examined, along with muscle mechanics; the biochemical basis of contraction; cardiac function; and the factors determining the heart's electrophysiologic behavior.

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This text is directed primarily at clinical cardiologists, cardiovascular surgeons, and trainees in their disciplines, as well as internists, medical students, and house officers.

Nuclear Receptors focuses on the structural analysis of nuclear receptors from the initial work using isolated protein domains to the more recent exciting developments investigating the conformational shape of full-length receptor complexes. The book also reviews the structure of key nuclear receptor co-regulatory proteins. It brings together, for the first time, a comprehensive review of nuclear receptor structure and the importance of receptor conformation underpinning allosteric regulation by different ligands (hormone, drugs, DNA response elements, protein-protein interactions) and receptor activity. The nuclear receptor superfamily, including receptors for steroid hormones and non-steroid ligands, are pivotal to normal physiology, regulating processes as diverse as reproduction, metabolism, the immune system and brain development. The first members of the family were cloned over 25 years ago, which heralded in the idea of a superfamily of intracellular receptor proteins that bound small molecule ligands: classical steroid hormones, vitamins, fatty acids and other products of metabolism. These signals are then transmitted through multiprotein receptor-DNA complexes, leading to the regulation of target genes, often in a cell-selective manner. The cloning of the receptor cDNAs also ushered in an era of unparalleled analysis of the mechanisms of action of these ligand-activated transcription factors.

This book covers a wide range of state-of-the-art methodologies and detailed protocols currently used to study the actions that lipid-activated nuclear receptors and their co-regulators have in tissues and immune cell types considered classic metabolic “ powerhouses ” . This includes the liver, adipose tissue, and monocytes/macrophages present in these and other metabolic

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tissues. While the main focus is on the oxysterol receptor or Liver X Receptor (LXR), the majority of the methods described can be easily applied to multiple nuclear receptors, as well as to other tissues or cell types. Written in the highly successful Methods in Molecular Biology series format, chapters include introductions to their respective topics, lists of the necessary materials and reagents, step-by-step, readily reproducible laboratory protocols, and tips on troubleshooting and avoiding known pitfalls. Authoritative and cutting-edge, *Lipid-Activated Nuclear Receptors: Methods and Protocols* serves as an ideal guide for researchers pursuing the vital study of nuclear receptor biology and beyond.

Edited by two experts working at the pioneering pharmaceutical company and major global player in hormone-derived drugs, this handbook and reference systematically treats the drug development aspects of all human nuclear receptors, including recently characterized receptors such as PPAR, FXR and LXR. Authors from leading pharmaceutical companies around the world present examples and real-life data from their own work.

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