



Platt versus Pickering: what molecular insight to primary hyperaldosteronism tells us about hypertension

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DECLARATIONS

Competing interests

MJB cites no current conflicts of interest but is seeking collaboration with companies developing aldosterone synthase inhibitors, in response to the Medical Research Council call for proposals in Stratified Medicine

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Summary

Recent genome-wide analyses have found 50 loci associated with variation in blood pressure but failed to advance understanding of the molecular basis of hypertension. Whether hypertension is not after all due to multiple common variants or is simply an order of magnitude more complex than previously suspected remains unsettled – in part because only a minority of subjects in the analyses had true hypertension. A better starting point than normotensive subjects for explaining hypertension may be the most common distinct cause of hypertension, primary hyperaldosteronism (PHA). The findings that 40% of patients with an aldosterone-producing adenoma (APA) of the adrenal have somatic gain-of-function mutations in a single gene, *KCNJ5*, and that this gene is, less frequently, mutated in inherited cases of PHA, potentially transform the understanding and management of hypertension. Firstly, they illustrate how hypertension could be due to a multiplicity of uncommon variants. Mutations that present with abnormal electrolytes and anatomy are the easiest to detect but are likely the tip of the iceberg. Secondly, we found a genotype:phenotype pattern, with *KCNJ5* mutations inducing larger APAs in the cortisol-secreting zona fasciculata in young women. Smaller APAs without *KCNJ5* mutations usually present in older men with resistant hypertension, having been overlooked earlier because of their size. This reflects their compact zona glomerulosa cells. Routine measurement of plasma renin in hypertension and a new positron emission tomography/computerized tomography allow prompt diagnosis and management of PHA before resistant hypertension ensues. Wider recognition of distinct phenotypes should permit earlier, specific treatment and reduce life-time risk of complications.

Genetic analysis of complex disorders

The advent of genome-wide association studies (GWAS) has brought bitter-sweet results for complex disorders like hypertension. On the one hand, we can now finger – with the sort of statistical certainty previously reserved for forensic DNA

fingerprinting – dozens of sites in the genome that are contributing to inherited variation in blood pressure. On the other, the highest risk from any site is 1 mmHg of blood pressure – in most cases rather less; in no case can we be absolutely certain which specific gene is responsible, far less have any idea of where or how in the gene a variant is influencing blood pressure; and the

Platt Versus Pickering An Episode In Recent Medical History

**Ezra Susser M.D., Sharon
Schwartz, Alfredo Morabia M.D., Evelyn
J. Bromet**

Platt Versus Pickering An Episode In Recent Medical History:

, **Platt Versus Pickering** J. D. Swales, 1985 Histories of the Normal and the Abnormal Waltraud Ernst, 2006-09-27 This fascinating volume tackles the history of the terms normal and abnormal Originally meaning as occurring in nature normality has taken on significant cultural gravitas and this book recognizes and explores that fact The essays engage with the concepts of the normal and the abnormal from the perspectives of a variety of academic disciplines ranging from art history to social history of medicine literature and science studies to sociology and cultural anthropology The contributors use as their conceptual anchors the works of moral and political philosophers such as Canguilhem Foucault and Hacking as well as the ideas put forward by sociologists including Durkheim and Illich With contributions from a range of scholars across differing disciplines this book will have a broad appeal to students in many areas of history **National Library of Medicine Current Catalog** National Library of Medicine (U.S.), 1987 **The Rise and Fall of Modern Medicine** James Le Fanu, 2012-11-06 In the years following World War II medicine won major battles against smallpox diphtheria and polio In the same period it also produced treatments to control the progress of Parkinson s rheumatoid arthritis and schizophrenia It made realities of open heart surgery organ transplants test tube babies Unquestionably the medical accomplishments of the postwar years stand at the forefront of human endeavor yet progress in recent decades has slowed nearly to a halt In this judicious examination of medicine in our times which has won the Los Angeles Times Book Prize medical doctor and columnist James Le Fanu both surveys the glories of medicine in the postwar years and analyzes the factors that for the past twenty five years have increasingly widened the gulf between achievement and advancement the social theories of medicine ethical issues and political debates over health care that have hobbled the development of vaccines and discovery of new miracle cures While fully demonstrating the extraordinary progress effected by medical research in the latter half of the twentieth century Le Fanu also identifies the perils that confront medicine in the twenty first century From a respected science writer important information that has been overlooked or ignored by many physicians New Republic Provocative and engrossing and informative Houston Chronicle **Hypertension: New Frontiers** HK Chopra, 2021-03-31 Hypertension is another name for high blood pressure It can lead to severe complications and increases the risk of heart disease stroke and death Blood pressure is the force exerted by the blood against the walls of the blood vessels Comprising more than 1000 pages this book is a complete guide to the diagnosis and management of hypertension Divided into 19 sections the text begins with an introduction to hypertension its causes risk factors and how it presents in different sectors of the population The next chapters discuss diagnostic methods and echocardiography The following sections cover hypertension associated with other disorders including heart failure stroke renovascular diseases metabolic syndrome and many more Numerous clinical management strategies are discussed in depth as well as cardio oncology nutrition and ECMO Extracorporeal Membrane Oxygenation a procedure that uses a machine to take over the work of the

lungs and sometimes the heart A complete section is dedicated to paediatric cardiology and hypertension With an internationally recognised editor and author team the comprehensive text is highly illustrated with clinical photographs diagrams and tables The book also includes full online access Key points Comprehensive guide to diagnosis and management of hypertension comprising over 1000 pages Internationally recognised editor and author team Highly illustrated with clinical photographs diagrams and tables Includes full online access to the book

Biomedicine in the Twentieth Century: Practices, Policies, and Politics C. Hannaway, 2008-02-11 Biomedicine in the Twentieth Century Practices Policies and Politics is a testimony to the growing interest of scholars in the development of the biomedical sciences in the twentieth century and to the number of historians social scientists and health policy analysts now working on the subject The book is comprised of essays by noted historians and social scientists that offer insights on a range of subjects that should be a significant stimulus for further historical investigation It details the NIH s practices policies and politics on a variety of fronts including the development of the intramural program the National Institute of Mental Health and mental health policy the politics and funding of heart transplantation and the initial focus of the National Cancer Institute Comparisons can be made with the development of other American and British institutions involved in medical research such as the Rockefeller Institute and the Medical Research Council Discussions of the larger scientific and social context of United States federal support for research the role of lay institutions in federal funding of virus research the consequences of technology transfer and patenting the effects of vaccine and drug development and the environment of research discoveries all offer new insights and suggest questions for further exploration

General Practice Under the National Health Service 1948-1997 Irvine Loudon, John Horder, Charles Webster, 1998 This is a history of general practice under the National Health Service covering the whole of the first 50 years from 1948 to the present

Schizophrenia Bulletin, 2003

A Short History of Cardiology Peter Fleming, 2020-01-29 The story told in this book begins in about 1700 when the first attempts were made to study the diseased heart in life the subject matter of cardiology as distinct from its appearance after death it ends rather arbitrarily in 1970 The account of the development of knowledge of heart disease is mainly chronological with emphasis on the fruitful consequences of the cross fertilization of clinical practice with pathological anatomy at the beginning of the nineteenth century and with physiology at the end In addition shorter chapters deals with such topics as specific disease entities methods of investigation cardiac surgery and the work of two individuals Peter Latham an example of a physician practising with today s clinical skills but a very imperfect knowledge of the pathogenesis of heart disease and Etienne Marey an early exponent of the clinical physiology which would in time throw light on that pathogenesis

Public Expectations and Physicians' Responsibilities John K Crellin, 2016-07-12 This guide for practice managers in a question and answer format explains accounting procedures and describes GMS health authority and other sources of income The book covers payroll pension personnel and complaints matters and advises on optimizing income for the practice

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Library of Medicine (U.S.), First multi year cumulation covers six years 1965 70 **Comprehensive Hypertension E-Book** Gregory Y. H. Lip, John E. Hall, 2007-06-28 Here is today's most in depth reference for any cardiologist internist or nephrologist interested in hypertension Drawing from international experience in cardiology physiology and nephrology Drs Lip and Hall have assembled a group of section editors and contributors second to none You'll find the long term effects of primary and secondary hypertension and a lengthy section on hypertensions for special populations featured prominently Prevention and treatment of hypertension are covered in detail from lifestyle and diet issues to drug choice and delivery and the section on comparison of guidelines is unique to this book Find comprehensive coverage of hypertension including pathogenesis prevention and treatment all in one practical volume See the complete systemic problems of hypertension at a glance with detailed full color illustrations of cellular and clinical manifestations Simplify navigating the complexities of hypertension using algorithms for clinical exam and diagnosis Get specific insight into prevention and treatment of hypertension in special populations Go global with a comprehensive section on worldwide guidelines and the application of clinical material to local standards of practice **British Cardiology in the 20th Century** Mark E. Silverman, Peter R. Fleming, Arthur Hollman, Desmond G. Julian, Dennis M. Krikler, 2012-12-06 Cardiology as a medical specialty originated in the 20th century and Britain played an important role in its development British Cardiology in the 20th Century provides the first comprehensive account of the British contributions to this exciting field as well as the interesting story of many of the people and institutions who were involved Many of the key changes in the understanding of the physiology of the heart and their clinical implications were discovered by these individuals This book will be of great interest to clinicians students and medical historians who wish to gain a historical understanding and appreciation of this dynamic clinical discipline that has improved the health and prognosis for so many Public Health and the Risk Factor William G. Rothstein, 2003 A risk factor is anything that increases the risk of disease in an individual **The Risks of Medical Innovation** Thomas Schlich, Ulrich Tröhler, 2006 Presenting a new way of thinking about the risks of medical innovation this volume considers the issues from a social historical perspective and studies specific cases in their respective contexts **Psychiatric Epidemiology** Ezra Susser M.D., Sharon Schwartz, Alfredo Morabia M.D., Evelyn J. Bromet, 2006-06-01 Searching for the causes of mental disorders is as exciting as it is complex The relationship between pathophysiology and its overt manifestations is exceedingly intricate and often the causes of a disorder are elusive at best This book is an invaluable resource for anyone trying to track these causes whether they be clinical researchers public health practitioners or psychiatric epidemiologists in training Uniting theory and practice in very clear language it makes a wonderful contribution to both epidemiologic and psychiatric research Rather than attempting to review the descriptive epidemiology of mental disorders this book gives much more dynamic exposition of the thinking and techniques used to establish it Starting out by tracing the brief history of psychiatric epidemiology the book describes the study of risk factors as causes of mental disorders Subsequent sections discuss

approaches to investigation of biologic genetic or social causes and the statistical analysis of study results The book concludes by following some of the problems involved in the search for genetic causes of mental disorders and more complex casual relationships *Bibliography of the History of Medicine* , **Rutter's Child and Adolescent Psychiatry** Anita Thapar, Daniel S. Pine, James F. Leckman, Stephen Scott, Margaret J. Snowling, Eric A. Taylor, 2017-12-26 Rutter's Child and Adolescent Psychiatry is the leading textbook in its field Both interdisciplinary and international it provides a coherent appraisal of the current state of the field to help researchers trainees and practicing clinicians in their daily work Integrating science and clinical practice it is a comprehensive reference for all aspects of child and adolescent psychiatry New to this full color edition are expanded coverage on classification including the newly revised Diagnostic and Statistical Manual of Mental Disorders DSM 5 and new chapters on systems neuroscience relationship based treatments resilience global psychiatry and infant mental health From an international team of expert editors and contributors this sixth edition is essential reading for all professionals working and learning in the fields of child and adolescent mental health and developmental psychopathology as well as for clinicians working in primary care and pediatric settings Michael Rutter has contributed a number of new chapters and a Foreword for this edition I greatly welcome this new edition as providing both a continuity with the past and a substantial new look Professor Sir Michael Rutter extract from Foreword Reviews of previous editions This book is by far the best textbook of Child Adolescent Psychiatry written to date Dr Judith Rapoport NIH The editors and the authors are to be congratulated for providing us with such a high standard for a textbook on modern child psychiatry I strongly recommend this book to every child psychiatrist who wants a reliable up to date comprehensive informative and very useful textbook To my mind this is the best book of its kind available today *Journal of Child Psychology and Psychiatry* **Cumulated Index Medicus** ,1986

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