

Prenatal Diagnosis of Hereditary Disease and Congenital Defects

By MICHAEL M. KABACK, M.D.

Since 1966, more than 100,000 midtrimester amniocenteses have been performed throughout the world for the prenatal determination of fetal hereditary diseases or congenital defects. Recently, a growing number of pregnancies have been monitored by placental aspiration/fetoscopy or by serial ultrasonographic studies for the detection of other congenital/genetic disorders. A growing public awareness of the applicability of these technologies and their increasing availability portend a greatly increased utilization of such procedures in the coming decades.

Other, related advances also have increased the frequency with which fetal diagnostic studies are being conducted. Heterozygote-screening programs (i.e., for the identification of carriers of Tay-Sachs disease, sickle-cell anemia, and beta-thalassemia) and serum alpha-fetoprotein (AFP) screening of pregnant women (in several national and regional pilot programs) are also greatly increasing the numbers of indicated fetal diagnostic procedures that are being conducted.

The rationale for obtaining an amniotic-fluid sample is to enable study of the fluid and cellular constituents (which are of fetal origin) in order to determine whether or not the fetus in that pregnancy is afflicted with, or at high risk for, a specific hereditary disorder or developmental defect for which it is known to be at increased risk. Biochemical analyses of the fluid itself or, in most instances, cytologic (chromosomal) or biochemical determinations performed on cultivated populations of fetal cells, derived from the amniotic fluid cells, allow such determinations to be made. Results of such analyses are available generally within a few days (e.g., AFP measurement) or within two to four

weeks (chromosomal and biochemical determinations) of the procedure.

It is not the purpose of this report to review all of the sophisticated and extensive methods that have been developed for the amniocentesis-related detection of each of the hereditary diseases or developmental defects that can now be detected in the midtrimester fetus. Several hundred research articles have appeared, and pediatricians interested in learning more about these procedures are referred to the many extensive reviews that are available.¹⁻⁷

With current somatic cell tissue-culture methods, successful cultivation of amniotic fluid cells can be achieved in more than 98 percent of instances with a single sample of 10 ml. or more of amniotic fluid.⁸ Once a sufficient population of growing fetal cells is available, it is possible to thoroughly assess the chromosomal complement of the fetus and to identify virtually any fetal cytogenic abnormality. Fetal sex is established readily by karyotype assessment, nuclear chromatin-mass frequency, or Y-body analysis. A wide variety of genetically defined metabolic functions also can be assayed in such cells. In this way, the prenatal determination of about 75 serious inborn errors of metabolism, inherited as either X-linked or autosomal-recessive disorders, can be addressed in the midtrimester fetus (the metabolic function being absent, grossly deficient, or clearly abnormal in the cultivated fetal cells if the fetus is affected). Tables 1 through 5 list disorders that can now be detected in this way.

The liquid fraction of the amniotic-fluid sample, separated from the cellular component by centrifugation, also has important applications to the prenatal detection of hereditary diseases, particularly

(Continued)

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**Robert L. Brent, John E. Fogarty
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Genetic Disorders and the Fetus Aubrey Milunsky, 2012-12-06 Technological advances continue to expand the number of genetic disorders that can be diagnosed in utero Utilization of this new technology has demanded special expertise available in relatively few academic centers As these new applications have become more widespread so have the realities of the medicolegal implications Notwithstanding the laboratory challenges most legal action at least in the United States has arisen from the physician's failure to inform a patient about the risks of a genetic disorder or the opportunities presented by prenatal diagnosis Hence an extensive thorough reexamination of the subject seems appropriate and timely The steady escalation in the number of prenatal genetic studies now being done in the western world makes it imperative for the physician to have a thorough comprehension of the subject in its entirety I am therefore fortunate in having colleagues who as acknowledged experts have shared their knowledge and experience in order to make this volume a major critical repository of facts and guidance about prenatal genetic diagnosis The subject matter ranges from a consideration of required genetic counseling through the intricacies of establishing prenatal diagnoses Special attention is focused on new advances using ultrasound a fetoprotein fetoscopy and first trimester diagnosis Both ethical and legal implications are discussed in detail as is the development of public policy

Teratogenic Mechanisms T.V.N. Persaud, 2012-12-06 The study of birth defects has assumed an importance even greater now than in the past because mortality rates attributed to congenital anomalies have declined far less than those for other causes of death such as infectious and nutritional diseases It is estimated that as many as 50% of all pregnancies terminate as miscarriages In the majority of cases this is the result of faulty development Major congenital malformations are found in at least 2% of all liveborn infants and 22% of all stillbirths and infant deaths are associated with severe congenital anomalies Teratological studies of an experimental nature are neither ethical nor justifiable in humans Numerous investigations have been carried out in laboratory animals and other experimental models in order to improve our understanding of abnormal intra uterine development In less than two decades the field of experimental teratology has advanced phenomenally As a result of the wide range of information that is now accumulating it has become possible to obtain an insight into the causes mechanisms and prevention of birth defects However considerable work will be needed before these problems can be resolved This book brings together some of the more recent and important research findings related to the mechanisms and pathogenesis of abnormal development It is not only a documentation of the latest experimental work but it also points out future directions that seem productive and challenging

Creasy and Resnik's Maternal-Fetal Medicine: Principles and Practice E-Book Robert Resnik, Robert K. Creasy, Jay D. Iams, Charles J. Lockwood, Thomas Moore, Michael F Greene, 2008-11-25 In your practice you require advanced knowledge of the obstetrical medical genetic and surgical complications of pregnancy and their effects on the mother and fetus With both basic science and clinical information six new chapters and an updated color design you need

look no further than the 6th edition of this long time best seller Includes both basic science and clinical information to give you comprehensive knowledge of the biology of pregnancy Acts as an excellent resource for OB GYNs studying for their Maternal Fetal Medicine boards and for practitioners who need quick access to practical information Provides an updated and focused reference list to keep you up to date on the standards of care in maternal fetal medicine today Keeps you current with a new section Disorders at the Maternal Fetal Interface and 6 new chapters Biology of Parturition Developmental Origins of Health and Disease Intrapartum Assessment of Fetal Health Pathogenesis of Pre term Birth Maternal and Fetal Infectious Disorders and Benign Gynecological Conditions of Pregnancy Features over 50% new authorship with increased focus on international perspectives Includes the following hot topics in Maternal Fetal Medicine o Biology of Parturition o Fetal Growth o Prenatal Genetic Screening and Diagnosis o Fetal Cardiac Malformations and Arrhythmias o Thyroid Disease and Pregnancy o Management of Depression and Psychoses during Pregnancy and the Puerperium Focuses on evidence based medicine the current best practice in MFM for diagnosing and treating high risk pregnancies Includes new illustrations and an updated color design

Cumulated Index Medicus ,1974 **First Trimester Fetal Diagnosis**
 Marco Fraccaro,Giuseppe Simoni,Bruno Brambati,2012-12-06 An International Symposium Covento delle Clarisse Rapallo Italy October 25 27 1984 The Economic Ideal in British Government Phyllis Colvin,1985 **Prevention of Embryonic, Fetal, and Perinatal Disease** Robert L. Brent,John E. Fogarty International Center for Advanced Study in the Health Sciences,1976 **Congenital Malformations of the Brain and Skull** P. J. Vinken,G. W. Bruyn,Ntinou Cleovoulou Myrianthopoulos,1977 *Handbook of Hemophilia* Kenneth Merle Brinkhous,H. C. Hemker,1975 *Atlas of Inherited Metabolic Diseases* William L Nyhan,Georg F Hoffmann,2020-07-14 In a field where even experts may find that years have elapsed since they last encountered a child with a given disorder it is essential for the clinician to have a comprehensive source of practical and highly illustrated information covering the whole spectrum of metabolic disease to refer to The content is divided into sections of related disorders including disorders of amino acid metabolism lipid storage disorders and mitochondrial diseases for ease of reference with an introductory outline where appropriate summarizing the biochemical features and general management issues Within the sections each chapter deals with an individual disease opening with a useful summary of major phenotypic expression including clear and helpful biochemical pathways identifying for the reader exactly where the defect occurs Throughout the book plentiful photographs often showing extremely rare disorders are an invaluable aid to diagnosis Key Features Fully updated to incorporate all new developments in the field Brand new chapters cover methylmalonic aciduria of ACSF3 deficiency branched chain keto acid dehydrogenase deficiency serine deficiencies purine nucleoside phosphorylase deficiency antequitin deficiency and others Excellent and detailed clinical descriptions with numerous valuable hints and suggestions for management Helpful explanatory algorithms and decision trees and high quality illustrative material including biochemical pathways and an unrivaled photographic collection which enhance clinical

applicability The fourth edition of this highly regarded book authored by two of the foremost authorities in pediatric metabolic medicine continues to provide incomparable insight into the problems associated with metabolic diseases and remains invaluable to pediatricians geneticists and general clinicians worldwide **Pediatric Pathology** J. Thomas Stocker, Louis P. Dehner, 2001 This newly revised Second Edition has been reorganized to emphasize the practical aspects of pediatric pathology The two volume text is separated into two easy to follow sections General Pathology and Organ System Pathology Includes up to date coverage on AIDS new diagnostic techniques etiologic agents and drug reactions and tumors Nearly 1 200 illustrations supplement the text This one of a kind preeminent reference is a must for both practitioners and students who are connected with the practice of pediatric pathology Compatibility BlackBerry R OS 4 1 or Higher iPhone iPod Touch 2 0 or Higher Palm OS 3 5 or higher Palm Pre Classic Symbian S60 3rd edition Nokia Windows Mobile TM Pocket PC all versions Windows Mobile Smartphone Windows 98SE 2000 ME XP Vista Tablet PC **Chorion Villus Sampling** Mitchell S. Golbus, D. T. Y. Liu, E. M. Symonds, 2013-11-11 *Neuroethics and Neurodevelopment*, 2023-08-25 This volume focuses on topics at the intersection between neuroethics and neurodevelopment and brings together the perspectives of experts in both clinical assessment and intervention and researchers in child psychology neurosciences medicine health policy law and social work The goal is to review emerging issues related to the ethical ramifications of how variation in human neurodevelopment is described and the effects of these descriptions on those with lived experience clinical and intervention services and health and social policy Related topics are also explored including the effect of the Covid 19 pandemic the ethics of invasive neurotechnology interventions biomarkers machine learning precision medicine Provides novel and original research on the emerging field of the legal regulation of neuroscience Takes an interdisciplinary approach with chapters by global scholars from several disciplines including law philosophy and medicine Develops a global approach that will be useful in jurisdictions around the globe *Screening and Management of Potentially Treatable Genetic Metabolic Disorders* P.F. Benson, 1984-03-31 1 The success of early diagnosis and therefore of treatment of phenylketonuria inevitably suggests the possibility of early screening and treatment of other genetic metabolic abnormalities This volume contains a summary of papers delivered at a Workshop on Recent Advances in Screening and Management of Potentially Treatable Genetic Metabolic Disorders held under the auspices of the Commission of the European Communities in London U K on the 17th and 18th March 1983 to consider such possibilities The Workshop was not aimed at those disorders for which the principles of treatment and management have been soundly established or for which screening procedures are in general use The papers therefore do not form a comprehensive account of metabolic disorders The topics for discussion were selected mainly to highlight recent discoveries which might be exploited by concerted approaches between different centres especially when the immediate benefits were restricted to only a few centres Recent reports suggest that about 2% of infants with persistent hyperphenylalaninaemia do not respond to treatment by phenylalanine low diets and develop severe brain

damage Infants with such Malignant Hyperphenylalaninaemia due to one of several genetic causes of tetrahydrobiopterin deficiency should be rapidly identified since there is evidence that appropriate treatment will prevent brain damage The Workshop considered how appropriate screening could be made generally available

Pathology of the Human Placenta Kurt Benirschke, Graham J. Burton, Rebecca N Baergen, 2012-06-16 Pathology of the Human Placenta remains the most comprehensive and authoritative text in the field It provides extensive information on the normal placenta encompassing physiology metabolism and endocrinology and covers the full range of placental diseases in great detail Further chapters are devoted to abortions molar pregnancies multiple pregnancies and legal considerations This sixth edition of the book has been extensively revised and expanded to reflect the most recent progress in the field and a brand new chapter has been added on artificial reproductive technology Some 800 illustrations are included many of them in color The detailed index has been further improved and tables updated Pathology of the Human Placenta will be of enormous value to pathologists and obstetrician gynecologists alike

Histology Arthur Worth Ham, David H. Cormack, 1979

Monthly Bibliography of Medical Reviews, 1975

Placental Pharmacology B. V. Rama Sastry, 1996-05-06 Placenta plays a very significant role in the development of secretory and regulatory functions for the maintenance of pregnancy and in the nutrition of the embryo It shares many functions of the heart lung liver endocrines and other organs In Placental Pharmacology fourteen expert authors discuss how placenta and its component parts can be used effectively in drug development research and in the study of transfer mechanisms across membranes This reference begins by describing the use of modern biophysical methods to study placental function and prospects for their application in pharmacological research The next three chapters deal with placental hemodynamics use of placental tissues and cells in pharmacological assays and placental transfer of drugs used in pregnancy Three more chapters contain up to date information on molecular biological aspects of placenta currently receiving lots of attention phospholipase A2 isozymes imprinted genes and embryogenesis and placental growth factors The final chapter describes the clinically relevant topics of placental metabolism and toxicity of homocysteine in placenta Easy to understand thorough and complete Placental Pharmacology is the perfect resource for scientists interested in placenta and its uses in drug development and testing

Bibliography of Medical Reviews, 1971

The Kelalis--King--Belman Textbook of Clinical Pediatric Urology Steven G. Docimo, Douglas Canning, Antoine Khoury, Joao Luiz Pippi Salle, 2018-09-03 The sixth edition of The Kelalis King Belman Textbook of Clinical Pediatric Urology presents state of the art knowledge and common sense practice in all areas of clinical pediatric urology in a single easily accessible volume Clinical in orientation and practical in presentation it covers every disorder and disease diagnostic method and appropriate treatment in pediatric urology from the embryo onwards Widely acknowledged as the leading reference in the field the book is essential reading for all practitioners who deal with urological disorders of childhood whether trainees residents or experienced specialists The book begins with chapters on evaluating the patient and general principles of pediatric urology including the

use of advanced diagnostic imaging techniques and the increasing importance of minimally invasive surgical techniques It then proceeds through the entire urinary tract from adrenal gland to external genitalia discussing anatomy normal and abnormal development anomalies and diseases and their treatment

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