# NOVAK'S GYNECOLOGIC AND OBSTETRIC PATHOLOGY

With Clinical and Endocrine Relations

Novak and Woodruff

# Eighth Edition

# Novak's

# Gynecologic and Obstetric Pathology

With Clinical and Endocrine Relations

# EDMUND R. NOVAK, A.B., M.D.

Associate Professor of Gynecology, The Johns Hopkins University School of Medicine; Gynecologist, Johns Hopkins and Union Memorial Hospitals, and Greater Baltimore Medical Center. Baltimore, Maryland

# J. DONALD WOODRUFF, B.S., M.D.

Professor, Obstetrics and Gynecology, Richard W. TeLinde Professor of Gynecologic Pathology, Johns Hopkins Hospital, Baltimore, Maryland

902 illustrations—18 in color

1979

W. B. SAUNDERS COMPANY / Philadelphia / London / Toronto

W. B. Saunders Company:

West Washington Square Philadelphia, PA 19105

1 St. Anne's Road

Eastbourne, East Sussex BN21, 3UN, England

1 Goldthorne Avenue

Toronto, Ontario M8Z 5T9, Canada

# **CONTRIBUTORS**

JOHN K. FROST, M.D. Professor of Pathology, Johns Hopkins University School of Medicine;
Pathologist and Head, Department of Cytopathology, Johns Hopkins Hospital, Baltimore, Maryland
Cytopathology

CARL J. PAUERSTEIN, M.D. Professor of Obstetrics and Gynecology and Professor of Physiology, The University of Texas Health Science Center at San Antonio; Chief, Gynecologic Service, Bexar County Hospital District, San Antonio, Texas Fertilization; Placental Abnormalities; Pathology of Abortion

Novak's Gynecologic and Obstetric Pathology with Clinical and Endocrine Relations

ISBN 0-7216-6869-0

© 1979 by W. B. Saunders Company. Copyright 1940, 1947, 1952, 1958, 1962, 1967 and 1974 by W. B. Saunders Company. Copyright under the International Copyright Union. All rights reserved. This book is protected by copyright. No part of it may be reproduced, stored in a retrieval system, or transmitted in any form or by any means, electronic, mechanical, photocopying, recording, or otherwise, without written permission from the publisher. Made in the United States of America. Press of W. B. Saunders Company. Library of Congress catalog card number 78-64719

Last digit is the print number: 9 8 7 6 5 4 3 2 1

# PREFACE TO THE EIGHTH EDITION

The eighth edition of our textbook has appeared somewhat later than new editions usually appear, but the last revision was so well received and has continued to have such a wide sale that we have procrastinated. Actually in a pathologic text there are usually relatively few truly revolutional concepts and new thoughts in any decade, and this seems especially true in the 1970s. There has been considerable experimentation in various methods of population control, and we have attempted a more detailed explanation and illustration as to how these may modify pelvic pathology. Newer concepts of oncology have likewise been reevaluated. In an effort to save space, we have on occasion abbreviated the titles of certain references, but the references themselves seem up to date. Various older ones may be found in earlier editions.

We trust that this eighth edition of our textbook will receive the same warm reception, and will continue to have the same extensive sale, as our previous edition. We have tried to improve it in every way possible. At the same time we have tried to streamline it somewhat to avoid making it too bulky and cumbersome. We had occasion to note that the seventh edition was approximately twice the size of the fourth edition, which was published in 1956 by Novak and Novak.

Obviously any pathology textbook will be concerned with oncology, and every attempt has been made to update the book by covering the newest and most recent concepts without becoming too involved with the treatment of these diseases, which seems permissible in a primarily pathologic textbook. As mentioned in earlier editions, it is our feeling that the TNM method of classification of the extent of pelvic tumors is extremely cumbersome and impractical, and consequently we have not utilized it extensively in this text.

Since this pathologic textbook is widely read over the entire world and appears in many different languages, we have enlarged discussion of various

tropical disorders that may afflict the female genital tract. This addition follows many suggestions by Middle and Far Eastern critics; although we have expanded our coverage, we still feel that these subjects belong primarily in various textbooks of tropical medicine, so coverage here is not extensive.

Adequate illustrative material is of extreme importance in any pathologic treatise, and we have continued to try to improve on these. In most instances black and white photographs seem satisfactory, but we have included a number of color plates, some of which are new in this textbook.

We should like to welcome Dr. Carl Pauerstein as a contributor to this text, and thank him for his outstanding chapters. We continue our appreciation to Dr. J.K. Frost for his well written chapter on cytopathology, which has received the highest approval in previous reviews of this textbook. Aside from these, we are the sole authors, and although each of us is responsible for certain chapters, there is frequent mutual discussion, so that the vast majority of the text represents a compendium of our ideas. This seems preferable to presenting isolated chapters by a variety of authors, who may discuss closely allied topics with a lack of consistency and coordination.

We wish to formally thank those who have been good enough to send material to our laboratory. In all cases we have attempted to give proper acknowledgment, but we may have inadvertently omitted credit in a few instances. On occasion we have utilized pictures that may not be of the highest technical quality, but this seems justifiable if it allows illustration of lesions only rarely encountered.

We are indebted to a number of individuals whose ideas or illustrations are incorporated. Thanks for invaluable help as always go to Miss Helen Clayton, as well as to Chester Reather and Raymond (Pete) Lund of our Photographic Department. We should also like to express our gratitude to the many nice people associated with the W. B. Saunders Company, who were so helpful and cognizant of the various difficulties associated with compiling this edition. Thanks again go to The Williams & Wilkins Company for permission to use various illustrations and figures, which we have done without specific notation.

We sincerely trust this edition of the text will represent continued improvement. We feel strongly that this is a legacy of Emil Novak and warrants our utmost dedicated efforts.

EDMUND R. NOVAK, M.D. J. DONALD WOODRUFF, M.D.

# CONTENTS

Chapter 1	
DISEASES OF THE VULVA	J
Normal Histology of Vulva	.]
Inflammatory Disease of Vulva	6
Systemic Disease of Vulva	Ç
Viral Disease of Vulva	Ç
Ulcerative Lesions of Vulva	13
Hyperplastic and "Atrophic" Changes	18
Benign Tumors of Vulva	25
Malignant Tumors of Vulva	37
Other Malignant Lesions of the Vulva	51
Diseases of the Urethra	53
Chapter 2	
DISEASES OF THE VAGINA	59
Embryology	59
Normal Histology of Vagina	59
Vaginal Histologic Cycle	59
Inflammatory Lesions	61
Ulcerative Lesions	64
Benign Tumors	64
Malignant Tumors	70
*	
Chapter 3	
HISTOLOGY OF THE CERVIX	82
Fistology	83
<del> </del>	хi

# xii / CONTENTS

0				
CI	111	1) t	er	4

BENIGN LESIONS OF THE CERVIX (CERVICITIS, THE METAPLASIAS, AND BENIGN TUMORS)	93
Cervicitis	93
Granulomatous Disease	96
Viral Diseases	97
Cervical Erosion	98
Cervical Polyp	104
Other Benign Lesions	107
•	
Chapter 5	
CERVICAL NEOPLASIA	111
Epidemiology	111
Precursors	112
Irritative Reactions Simulating Neoplasia	117
Carcinoma in Situ (Intraepithelial Carcinoma, Preinvasive	,
Carcinoma)	120
Microinvasive Carcinoma	123
Gross Characteristics of Epidermoid Cancer	123
Clinical Classification of Cervical Cancer	130
Diagnosis of Cervical Cancer	131
Histologic Grading of Epidermoid Cancer	137
Adenocarcinoma of Cervix	140
Adenosquamous and Adenoepidermoid Carcinoma	142
Extension and Metastasis of Cervical Cancer	146
Effects of Radiotherapy on Cervical Carcinoma	147
Recurrent Cancer	149
Other Malignancies	151
Metastatic Malignancy	153
Chapter 6	
ENDOCRINOLOGY OF THE ENDOMETRIUM AND OVARY	157
Ovarian Hormones	157
Histochemical Study of the Endometrium	160
Histochemical Study of the Ovary	160
The Hormones of Pregnancy	162

CONTENTS	xiii
The Anovulatory Cycle in Women	163
Induction of Ovulation	164
Progestogen Therapy	166
Postmenopausal Ovarian Function	166
Chapter 7	
HISTOLOGY OF THE ENDOMETRIUM	171
Menstruation	172
Histologic Phases of Menstrual Cycle	174
The Endometrium of Pregnancy	184
Chapter 8	
HYPERPLASIA OF THE ENDOMETRIUM	190
Genuine Hyperplasia of the Endometrium	191
Proliferative Pictures in Endometrium Associated with Functional Bleeding	195
Proliferative and Pseudomalignant Types	196
Chapter 9	
CARCINOMA OF THE ENDOMETRIUM	204
Treatment of Endometrial Adenocarcinoma	229
Salvage	236
Chapter 10	
ENDOMETRITIS AND OTHER BENIGN CONDITIONS OF THE ENDOMETRIUM	239
Endometritis	239
Myometritis	250
Myometrial Hypertrophy	251
Subinvolution of Uterus	251
Pyometra	
Endometrial Polyp	254
Chapter 11	
MYOMA AND OTHER BENIGN TUMORS OF THE UTERUS	260
Chapter 12	
ADENOMYOSIS (ADENOMYOMA) UTERI	280

### xiv / CONTENTS

Chapter 13

SARCOMA AND ALLIED LESIONS OF THE UTERUS	291
Leiomyosarcoma	291
Intravenous Leiomyomatosis	298
Metastasizing Myoma	299
Endometrial Sarcoma and Mixed Tumors (Mesenchymal)	299
Müllerian Adenosarcoma	300
Chapter 14	
HISTOLOGY OF FALLOPIAN TUBES	306
Pathophysiologic Changes in the Tubal Epithelium	311
Functions of Fallopian Tube	315
Heterotopic Alterations in the Tube	315
Chapter 15	
SALPINGITIS	319
Pelvic Inflammatory Disease (PID)	319
Acute and Subacute Salpingitis	320
Chronic Salpingitis	322
Granulomatous Salpingitis	328
Tuberculous Salpingitis	328
Other Granulomatous Diseases	330
Perisalpingitis,	332
Chapter 16	
TUMORS OF THE TUBE, PAROVARIUM, AND UTERINE LIGAMENTS	334
Benign Tumors	334
Carcinoma of the Tube	335
Other Tumors of the Tube	341
Parovarium Cysts	342
Tumors of the Para-Adnexal Structures (Including Peritoneum, Ligaments, and Embryonic Remnants)	346
Round Ligament	347
Tumors of the Broad Ligament	349

Chapter 17	
EMBRYOLOGY AND HISTOLOGY OF OVARIES	355
Embryology	355
Histology	357
Chapter 18	
INFLAMMATORY DISEASES OF THE OVARY	375
Acute Oophoritis	375
Abscess of the Ovary	375
Chronic Perioophoritis	377
Chronic Oophoritis	378
Chapter 19	
CLASSIFICATION OF OVARIAN TUMORS	380
Tumors of the Ovary	380
Classification of Ovarian Tumors Based on Pathophysiology and Embryology	381
FIGO Classification	382
WHO Classification	383
•	
Chapter 20	
	385
•	385
	385
Lutein Cysts (Luteinized Granulosa-Theca Cysts)	388
Chapter 21	
OVARIAN NEOPLASIA: MESOTHELIAL; STROMOEPITHELIAL LESIONS—PRIMARILY EPITHELIAL (BENIGN, OF LOW MALIGNANT POTENTIAL) AND MALIGNANT (SEROUS, MUCINOUS, ENDOMETRIOID, MESONEPHROID,	396
	397
	397
	409
- No. 1	416
	419
*	422

### xvi / CONTENTS

Mesothelial Reactions and the Mesothelioma	426
Unspecified Designations—Undifferentiated Carcinoma (Primary Solid Carcinoma of Ovary)	430
Extension and Metastasis of Ovarian Carcinoma	430
Treatment and Salvage of Ovarian Cancer	433
	•
Chapter 22	
OVARIAN NEOPLASIA: STROMOEPITHELIAL LESIONS —PRIMARILY STROMAL (BRENNER TUMORS, FIBROTHE- COMAS, SARCOMAS, ETC.)	437
Brenner Tumors of the Ovary	
Tumors with Functioning Matrix	
Tumors with Functioning Matrix	443
Chapter 23	٠
OVARIAN NEOPLASIA: PRIMARILY STROMAL LESIONS (FIBROMA, FIBROTHECOMA INCLUDING THE MIXED	451
MESODERMAL TUMORS)	451
Stromal Tumors	451
Fibroma (Fibrothecoma)	451
Other Solid Tumors	455
Ovarian Sarcomas	455
Lymphoma	459
Chapter 24	
OVARIAN NEOPLASIA: METASTATIC LESIONS	461
Tumors Arising in the Genital Canal	461
Metastatic Lesions with an Extragenital Primary Tumor	464
The Krukenberg Tumor of the Ovary	471
Primary Krukenberg Tumors	473
Chapter 25	
GERM CELL TUMORS	476
PART I. DYSGERMINOMA OF THE OVARY	
Germ Cell Tumors	476
Germ Cell Tumors and Precursory Conditions	476
Dysgerminoma (Germinoma)	477
Gonadoblastoma	485

PART II. OVARIAN TERATOMAS	ar a
Mature Teratomas	487
Immature Teratomas (Embryonal)	495
Extraembryonal Teratomas	496
Chapter 26	
GONADAL STROMAL TUMORS—FEMINIZING (GRANULOSA AND THECA CELL)	504
Histogenesis	504
Pathology of Granulosa Cell Tumor	505
Pathology of Thecoma	511
Diffuse Thecosis	512
Luteinization of Granulosa Cell Tumors and Thecomas	513
Effects of Granulosa Cell Tumors and Thecomas on Endometrium	513
Luteoma of Pregnancy	515
Association of Endometrial Adenocarcinoma with Feminizing Tumors	516
Experimental Production of Granulosal and Thecal Tumors	517
Clinical Characteristics	518
•	
Chapter 27	
GONADAL STROMAL TUMORS—VIRILIZING (ARRHENO-BLASTOMA, ADRENAL, AND HILUS CELL)	<b>52</b> 3
Arrhenoblastoma—Histogenesis and Types	523
Pathology	524
Clinical Characteristics	527
Gynandroblastoma	530
Adrenal Tumors of the Ovary	531
Hilus or Leydig Cell Tumors	534
Homology of Certain Ovarian and Testicular Tumors	535
Gonadal Stromal Tumors	536
Treatment	538
Chapter 28	
ECTOPIC PREGNANCY	541
Etiology of Tubal Pregnancy	544

# xviii / CONTENTS

Nidation in the Tube	545
Terminations of Tubal Pregnancy	549
Behavior of Uterine Mucosa in Cases of Tubal Pregnancy	550
Value of Diagnostic Curettage in Tubal Pregnancy	554
Source of External Bleeding in Tubal Pregnancy	.555
Pregnancy Tests in Tubal Pregnancy	556
Ovarian Pregnancy	556
Primary Abdominal (Peritoneal) Pregnancy	557
Cervical Pregnancy	558
Interstitial Pregnancy (Following Salpingectomy)	558
Combined Pregnancy (Intra- and Extrauterine)	558
Combined Pregnancy (Tubal)	558
Posthysterectomy Ectopic Pregnancy	558
Chapter 29	
PELVIC ENDOMETRIOSIS	561
Ovarian Endometriosis and Endometrial Cysts of the Ovary	561
Endometriosis of Uterosacral Ligaments	567
Endometriosis of Rectovaginal Septum	567
Endometriosis of Round Ligaments	569
Endometriosis of Umbilicus	570
Endometriosis in Laparotomy Scars	570
Other Sites of Endometriosis	570
Histogenesis of Endometriosis	573
Endometriosis as a Source of Ovarian Carcinoma	575
Endometrioid Carcinoma of the Ovary	578
Clinical Characteristics of Endometriosis	578
Endometrium Following Progestogen Therapy	580
Chapter 30	
FERTILIZATION, IMPLANTATION, AND PLACENTATION	585
Carl J. Pauerstein	
Fertilization	585
Implantation	587
Placentation	588

Chapter 31

ABNORMALITIES AND DISEASES OF THE PLACENTA AND APPENDAGES (OTHER THAN HYDATIDIFORM MOLE AN CHORIOCARCINOMA)	N <b>D</b> 594
Carl J. Pauerstein	
Placental Lesions in Toxemia of Pregnancy	612
Special Diseases of the Placenta	615
Abnormalities of Implantation and Separation	620
New Growths	624
Abnormalities of the Amnion	625
Abnormalities of the Umbilical Cord	627
Chapter 32	
PATHOLOGY OF ABORTION (IN UTERUS, PLACENTA, APPENDAGES, AND OVOFETUS)	633
Carl J. Pauerstein	
Endocrine Factors	635
Progestational Phase Endometrium	637
Mechanism of Abortion	638
Pathologic Characteristics of the Ovofetus	639
Classification of Incomplete Specimens	639
Classifications of Specimens in Special Clinical Groups	643
Changes in the Decidua and Uteroplacental Area	645
Changes in the Placenta	645
Changes in the Placental Appendages	647
Habitual Abortion	648
Chapter 33	
$ \   \textbf{HYDATIDIFORM MOLE AND CHORIOCARCINOMA~(TRD)}$	651
Hydatidiform Mole	652
Invasive Hydatidiform Mole (Chorioadenoma Destruens)	657
Choriocarcinoma	661
Syncytial Endometritis	666
Trophoblastic Pseudotumor	667

## XX / CONTENTS

	Diagnosis of Choriocarcinoma from Curettings	667
•	Gross Characteristics of Hydatidiform Mole and Choriocarcinoma	668
	Chief Clinical Features	671
	Extension and Metastasis in Choriocarcinoma	674
	Malignancy of Choriocarcinoma	675
	Vagaries of Benign Moles	676
	Ectopic Choriocarcinoma	676
	Tubal and Ovarian Hydatidiform Mole and Choriocarcinoma	677
	Ovarian Changes Associated with Hydatidiform Mole and Choriocarcinoma	677
	Biologic Tests in Hydatidiform Mole and Choriocarcinoma	680
	Treatment of Trophoblastic Disease	682
GY	npter 34  NECOLOGIC AND OBSTETRIC CLINICAL  TOPATHOLOGY	689
9	ohn K. Frost	000
,	Normal Squamous Cell Morphology	689
	Cytohormonal Evaluation	694
	Cytohormonal Patterns: Normal	
	Cytohormonal Patterns: Abnormal	709
	Sex Determination: Genetic and Endocrine Bases	714
	Normal Cell Morphology	719
	Macrophages	723
	Degeneration and Regeneration	725
	Nonspecific Inflammation, Chronic Irritation, and Senile Vaginitis	727
	Infections	729
	The "Shades of Gray" Lesions—Abnormalities of Epithelial Development, Cellular Orientation, and Nuclear and Cytoplasmic Relationships	736
	The Cancer Cell: Criteria of Malignancy	743
	Cancer Differentiation—Cellular Characteristics of Functional	
	Differentiation	748

CONTENTS	,,,,,,
Lesions of the Cervix Uteri	759
Lesions of the Vulva and Vagina	
Lesions of the Endometrium	771
Lesions of the Placenta, Tubes, and Ovaries	774
The Clinical Specimen	776
Cytopathologic Reports and Their Interpretations	
INDEX	

# DISEASES OF THE VULVA

### NORMAL HISTOLOGY OF VULVA

The external genitalia—specifically the labia majora and minora, the clitoris, and the vestibule with its associated glands—are of ectodermal origin (Fig. 1-1). The labia majora are longitudinal folds of fat whose lining epithelium is stratified squamous, with varying degrees of surface maturation and keratinization, and with an underlying laver of connective tissue corresponding to the dartos of the male scrotum. The labia majora are practically absent in the young child, their development—primarily the deposition of fat—being one of the secondary sex characteristics heralding puberty. The skin of the more prominent portions of the labia is pigmented. These folds are rich in hair follicles, sebaceous glands, and sudoriferous glands (Fig. 1-2). The latter include the unique apocrine glands found in special areas, e.g., the axilla, perianal region, and breast, and characterized by "decapitation" secretion, in contrast to the characteristic cellular loss of the sebaceous (holocrine) gland and the cytoplasmic secretory activity of the merocrine gland (Fig. 1-3). Because the onset of secretion occurs at puberty and the cyclic nature of the activity corresponds to that of the ovary, Way and Memmesheimer consider these apocrine glands to be "accessory sex glands." The knowledge of this cyclic activity is important in the diagnosis and treatment of certain vulvar diseases.

The mons pubis (mons veneris) is a cushion of fat covered by skin and its appendages, including the apocrine glands. The labia minora are firmer structures than the majora, and are composed primarily of vascular connective tissue. The surface stratified epithe-

lium is characterized by a relative absence of both the granular layer of the epithelium and hair follicles. The numerous sebaceous glands secrete directly onto the skin through epithelial tunnels (Fig. 1–4). Apocrine glands, although present, are infrequent.

The clitoris, like its male homologue, is made up of vascular erectile tissue, differing from the penis in that it lacks the corpus spongiosum. From an embryologic standpoint the two vestibulovaginal bulbs, which are congeries of veins situated beneath the anterior portion of the labial structures, correspond in the female to a divided corpus cavernosum, i.e., they are made up of two corpora with an intricate network of nerves (Fig. 1–5).

The female urethra, opening externally at the meatus urinarius, is lined by transitional epithelium with the stratified epithelium of the vaginal mucosa present at or near the orifice. At the lower border of the meatus are the openings of the Skene's ducts (paraurethral ducts), tiny tortuous canals coursing just beneath the urethra for a distance of about 1.5 cm.; they are lined by squamous epithelium, and may be the seat of infection inaccessible to treatment by local applications. Studies have shown that the canal is almost completely surrounded by a labyrinth of paraurethral glands entering the distal urethra from its posterior aspect. Huffman considered these structures to be the homologues of the male prostate.

Occlusion of one or more of these glands produces cyst formation and the subsequent infection may result in the development of a suburethral abscess, an occasional cause of urinary retention in the female. In cases of recurrent urinary tract infection, the paraurethral canals should be suspected as a

Text continued on page 6