

# Textbook of Pharmacology, Pathology & Genetics for NURSES-II

As per the Revised INC Syllabus for BSc Nursing

Suresh Sharma





# Textbook of Pharmacology, Pathology and Genetics for Nurses–II

As per the Revised INC Syllabus for BSc Nursing

**SECOND EDITION** 

Suresh Sharma MSc (N) PhD FNRS RN (USA) Professor and Principal College of Nursing All India Institute of Medical Sciences (AIIMS) Jodhpur, Rajasthan, India



JAYPEE BROTHERS MEDICAL PUBLISHERS

The Health Sciences Publisher New Delhi | London

# Contents

# SECTION 1: PHARMACOLOGY-II

# 1. Drugs Used in Disorders of Ear, Nose, Throat and Eye

Hemlata, Nipin Kalal, Suresh Sharma

Antihistamines 3 Topical Application for Eye, Ear, Nose and Buccal Cavity 5

#### 2. Drugs Used on Urinary System

Farhanul Huda, Hemlata, Suresh Sharma Renin-Angiotensin Aldosterone System 11 Diuretics and Antidiuretics 12 Alkalinizers 28

#### 3. Drugs Acting on Nervous System 33

Raj Kumar, Suresh Sharma, Rajneesh Arora, Poonam Arora, Hemlata

Basis and Applied Pharmacology of Commonly Used Drugs 33 Analgesics and Anesthetics 33 Non-Selective COX Inhibitors 33 Preferential COX-2 Inhibitors 37 Selective COX Inhibitors 38 Analgesic-Antipyretics with Poor Anti-inflammatory Action 38 **Opioids and Other Central** Analgesics 39 Anesthesia 42 General Anesthesia 42 Classification 43 Hypnotics and Sedatives 52 Skeletal Muscle Relaxants 55 Peripherally Acting Muscle Relaxants 55 Centrally Acting Muscle Relaxants 62 Antipsychotics 63 Antidepressants 74 Antianxiety Drugs 81 Anticonvulsants 86

Drugs for Neurodegenerative Disorders and Miscellaneous Drugs 92 Stimulants, Ethyl Alcohol and Treatment of Methyl Alcohol Poisoning 94

4. Drugs Used for Hormonal Disorders and Supplementation, Contraception and Medical Termination of Pregnancy 112

Prasuna Jelly, Hemlata, Suresh Sharma

Estrogen and Progesterone 112 Oral Contraceptive and Hormone Replacement Therapy 119 Vaginal Contraceptives 125 Drugs for Infertility and Medical Termination of Pregnancy 126 Uterine Stimulants and Relaxants 129 Uterine Relaxants (Tocolytics) 131

#### 5. Drugs Used for Pregnant Women During Antenatal, Labor and Postnatal Period 135

Hemlata, Suresh Sharma

Tetanus Prophylaxis 135 Iron and Vitamin K1 Supplementation 136 Oxytocin, Misoprostol 136 Ergometrine 140 Methyl Prostaglandin E2 Alpha 141 Magnesium Sulfate 144 Calcium Gluconate 145

6. Miscellaneous Drugs 148

Vasantha Kalyani, Hemlata, Suresh Sharma Drugs Used for Deaddiction 148 Drugs Used in CPR and Emergency 148 IV Fluids and Electrolyte Replacement 153 Common Poisons, Drugs Used for Treatment of Poisoning 153

3

#### Contents

xii

Vitamins and Minerals Supplementations 161 Vaccines and Sera (Universal Immunization Program Schedule) 167 Anticancer Drugs: Chemotherapeutic Drugs Commonly Used 171 Notes on Individual Drugs 171 Nursing Responsibilities 176 Patient Education 176 Causes for Failure of Chemotherapy 177 Superinfection 177 Chemoprophylaxis 178 Immunosuppressants and Immunostimulants 178

#### 7. Introduction to Drugs Used in Alternative Systems of Medicine 183

Raj Kumar, Hemlata, Suresh Sharma Ayurveda, Homeopathy,

Unani, and Siddha, etc. *183* Department of AYUSH Under the Ministry of Health and Family Welfare *190* Drugs Used in Common Ailments *190* 

8. Fundamental Principles of Prescribing

Suresh Sharma

Prescriptive Role of Nurse Practitioners: Introduction 193 Legal and Ethical Issues Related to Prescribing 194 Principles of Prescribing 198 Steps of Prescribing 198 Prescribing Competencies 202

# SECTION 2: PATHOLOGY-II

#### . Kidney and Urinary System

Aminder Singh, Nancy Kurien, Suresh Sharma Glomerulonephritis 209 Pyelonephritis 216 Renal Calculi 219 Cystitis 221 Renal Cell Carcinoma 223 Renal Failure 225

#### 10. Male Genital System

Aminder Singh, Nancy Kurien, Suresh Sharma Cryptorchidism 229 Testicular Atrophy 231 Prostatic Hyperplasia 231 Carcinoma of Penis 233 Carcinoma of Prostate 234

11. Female Genital System

Neena Sood, Nancy Kurien, Suresh Sharma Carcinoma Cervix 237 Carcinoma Endometrium 238 Uterine Fibroids 240 Vesicular Mole (Hydatidiform Mole) 242 Choriocarcinoma 243 Ovarian Cysts 244 Ovarian Tumors 245

#### 12. Breast

193

209

Aminder Singh, Joyce Joseph, Suresh Sharma Fibrocystic Changes 251 Fibroadenoma 251 Breast Cancer 252

#### 13. Central Nervous System

Neena Sood, Nancy Kurien, Suresh Sharma Meningitis 256 Encephalitis 257 Stroke 258 CNS Tumors 264 Metastatic CNS Tumors 266

#### **SECTION 3: CLINICAL PATHOLOGY**

14. Examination of Body Cavity Fluids

271

229

237

251

256

Neena Sood, Nancy Kurien, Suresh Sharma Cerebrospinal Fluid Analysis 271 Examination of Sputum 273 Analysis of Gastric Constituents 275 Analysis of Duodenal Contents 276 Analysis of Peritoneal Fluid 277 Analysis of Pericardial Fluid 278 Analysis of Semen 279 Urine Examination 282 Fecal Examination 287

#### **SECTION 4: GENETICS**

| 15. Introduction to Genetics 29 |
|---------------------------------|
|---------------------------------|

Suresh Sharma, Sohinder Kaur Concept of Genetics 293 Basic Genetic Terms 294 **Practical Applications** of Genetics in Nursing 295 Review of Cellular Division 302 Genes 316 Protein Biosynthesis 325 Genetic Code 327 Naming Genes 328 Chromosome 329 Chromosomal Aberrations/ Chromosomal Mutation 332 Patterns of Inheritance 336 Sex-linked Inheritance 341 Mendelian Theory of Inheritance 348 Multiple Allots and Blood Groups 348 Facts about Transmission of Genetic Disorders 351 Mechanism of Inheritance 352 Gene Mutation (Errors of Transmission) 353

#### 16. Emerging Paradigm of Genetics in Nursing

Suresh Sharma

Genetic Nursing Practice Milestones 363 Roles of Nurses in Genetics 364 Importance of Genetics in Nursing Curriculum 364 Barriers and Approaches for Implementation of Genetics in Nursing 367 Assumptions Regarding Genetics and Health Care in Future 368

17. Maternal and Prenatal Genetics 371

Suresh Sharma, Vandana Mehta Genetics and Infection 371 Consanguinity Atopy 375 Prenatal Nutrition and Food Allergies 378 Role of Prenatal Nutrition in Prevention of Genetic Disorders 380 Maternal Age 381 Maternal Drug Therapy 384 Effects of Radiation, Drugs and Chemicals 389 Prenatal Testing and Diagnosis 396 Noninvasive Tests 398 Invasive Tests 400 Preimplantation Prenatal Diagnosis 408 Infertility 410 Spontaneous Abortions 417 Neural Tube Defects 420 Down Syndrome 426

#### 18. Neonatal and Children Testing or Screening

434

xiii

Suresh Sharma, Jyoti Arora Meaning and Purpose 434 Newborn Screening 435 Genetic Testing and Screening in Children 437 Congenital Abnormalities 441 Developmental Delay 442 Dysmorphism 446 Role of Nurses in Genetic Testing for Neonates and Children 447

# 19. Genetic Conditions of<br/>Adolescents and Adults450

Suresh Sharma, Kumar Satish Ravi Genetic Statistics 450 Naming Genetic Conditions 450 Common Genetic Conditions 451 Therapeutic Approaches for Genetic Disorders 484 Nursing Management in Genetic Disorders 486

#### 20. Services Related to Genetics 496

Suresh Sharma, Mohd Salahuddin Ansari Genetic Testing 496 Gene Therapy 503 Genetic Counseling 510

#### Index

362

# Drugs Used for Hormonal Disorders and Supplementation, Contraception and Medical Termination of Pregnancy

Prasuna Jelly, Hemlata, Suresh Sharma

# **ESTROGEN AND PROGESTERONE**

Hypothalamus releases gonadotropin-releasing hormone (GnRH). This stimulates the anterior pituitary to release FSH and LH. FSH stimulates maturation of primary oocyte in an immature follicle. Follicle produces estrogen. Estrogen builds the uterine wall (the endometrium) and inhibits secretion of FSH. High levels of estrogen further stimulate secretion of LH by anterior pituitary. This plus FSH also causes ovulation of the secondary oocyte—leaving follicle without egg (the corpus luteum).

Corpus luteum secretes estrogen and progesterone. This maintains the endometrium for 15–16 days and inhibits LH (if oocyte is not fertilized and implanted in the uterine wall). The corpus degenerates (to corpus albicans) and stops producing estrogen and progesterone. Without estrogen and progesterone, endometrium breaks down—menstruation occurs. Menstruation is the sloughing off of the enlarged endometrial wall along with blood and mucus. This decrease in progesterone and LH. Low LH causes secretion of FSH by pituitary again. The cycle repeats.

# Estrogen

CHAPTER

A group of steroid hormones/female sex hormone that readily diffuse across the cell membrane. Inside the cell, they interact with estrogen receptors.

# Natural Estrogens

Estradiol is the major estrogen secreted by ovary. It is synthesized in Graafian follicle, corpus luteum and placenta from cholesterol. Oxidized in the liver to weak estrogens like estrone, estriol.

# Synthetic Estrogens

- To overcome the shortcomings of natural estrogens
- Steroidal-ethinyl estradiol, mestranol, tibolone
- Nonsteroidal—diethylstilbestrol, hexestrol, dienestrol

# Estrogen Synthesis (Fig. 4.1)

- Estrogen is produced primarily by developing follicles in the ovaries, the corpus luteum, and the placenta
- The most abundant estrogen secreted by ovaries is estradiol
- The FSH and LH stimulate the production of estrogen in the ovaries
- Some estrogens are also produced in smaller amounts by other tissues such as the liver, adrenal glands and the breasts.

113



Fig. 4.1: Synthesis of estrogen and progesterone.

## Effects of estrogen at various sites in the body

- Female reproductive system:
  - Fallopian tubes, uterus, vagina—pubertal growth and development. Thickening of vaginal epithelium
  - Mammary glands—proliferations of ducts and stroma
  - Uterine endometrium—proliferation
  - Cervix—watery secretion to facilitate sperm penetration
- CNS:
  - FSH/LH secretion—feedback control
  - Nausea and vomiting
- Blood:
  - Coagulation factors—decreased antithrombin III. Increasing circulating levels of factors II, VII, IX and X
  - Lipid profile—increase in HDL, decrease in LDL, increase in triglycerides
- Metabolic effects:
  - Anabolic effects
    - Glucose intolerance
    - Sodium and water retention
    - Maintain bone mass and decrease in bone resorption.
- Other effects:
  - Growth of hair, fat deposition, pigmentation on nipples
  - Gallbladder stones, cholestatic jaundice
  - Increased circulating level of proteins, hepatic adenoma on prolonged use.

#### Pharmacokinetics

#### Absorption

114

Natural estrogens are orally inactive; synthetics estrogens are well absorbed orally and transdermally, estrogen ester in the form of IM injections are slowly absorbed and have prolonged effect.

#### Metabolism

Estradiol is converted reversibly to estrone and both can be converted to estriol, which is the major urinary metabolite. Estrogens also undergo enterohepatic recirculation via sulfate and glucuronide conjugation in the liver, biliary secretion of conjugates into the intestine, and hydrolysis in the gut followed by reabsorption.

#### Excretion

Estradiol, estrone and estriol are excreted in the urine along with glucuronide and sulfate conjugates.

#### Preparations and Doses

The preparations and doses of estrogen along with their routes of administration has been presented in **Table 4.1**.

#### Indications

- **Hormonal replacement therapy:** It includes administrations of estrogen and progesterone combinations. Due to cessation of ovarian function at menopause women suffer a number of physical, psychological and emotional consequences..
- Acne and hirsutism—estrogen benefits by suppressing ovarian production of androgen by inhibiting gonadotropins released from pituitary.

 Table 4.1: Estrogen preparations with doses and routes of administration.

| Drug   | Dose                              | Route   |  |
|--|-----------------------------------|---|--|
| Natural steroidal estrogen: Estradiol benzoate, cypionate, enanthate | 2.5–10 mg                         | IM  |  |
| Conjugated estrogen  | 0.625–1.25 mg/day                 | Oral  |  |
| Ethinylestradiol   | 0.02–0.2 mg/day                   | Oral  |  |
| Mestranol  | 0.1–0.2 mg/day                    | Oral  |  |
| Estriol succinate  | 4–8 mg/tds                        | Oral, cream tds   |  |
| Dienestrol   | 0.01%                             | Topically in vagina   |  |
| Transdermal estradiol: Estraderm-MX                                  | 25, 50,100 μg/24 h for 3–4<br>day | Apply to non-hairy skin below<br>the waist, oral progestin is<br>added        |  |
| Gel formulations: Oestrogel  | 3 mg/5 g in 80 g tube             | Applied over the arms<br>once daily for hormonal<br>replacement therapy (HRT) |  |

**CHAPTER 4:** Drugs Used for Hormonal Disorders and Supplementation...

- Dysmenorrhea—estrogen therapy benefits by inhibiting ovulation (anovulatory cycles) and decreasing prostaglandins secretions in endometrium.
- Carcinoma prostate—acts by suppressing the androgen production through pituitary.
- Senile vaginitis—effective in preventing as well as treating atrophic vaginitis that occurs in elderly women. An antibacterial may be combine.
- Delayed puberty in girls—Turner's syndrome or hypopituitarism.
- Dysfunctional uterine bleeding—this type of bleeding can result from the following three reasons:
  - i. Estrogen withdrawal bleeding occurs when the estrogen given to postmenopausal women
  - ii. Estrogen break through bleeding that occurs when there is a continuous stimulation of endometrium not interrupted by progesterone secretion, e.g. polycystic ovarian syndrome
  - iii. Progesterone break through bleeding that occurs in presence of abnormally high progesterone, e.g. in women using low dose oral contraceptives.

# Contraindications

- Pregnancy
- Thromboembolic disorders
- Diabetes
- Hepatic failure
- Estrogen dependent carcinoma of breast
- Endometrial carcinoma
- Endometriosis and undiagnosed genital bleeding.

# Side Effects

- In males— gynecomastia, feminisation and decreased libido
- In females—breast tenderness, migraine, nausea, withdrawal bleeding, amenorrhea, endometrial hyperplasia, increased risk of vaginal and cervical adenocarcinoma
- In both sexes—gallstones and gallbladder, hepatic dysfunction, predisposition to thromboembolic disorders, precipitation of diabetes and fluid retention
- Fusion of epiphyses and reduction of adult stature when given to children.

# **Drug Interactions**

- Need to increase the dose of warfarin, oral hypoglycemic or insulin
- Barbiturates, phenytoin, carbamazepine and rifampicin may decrease the effectiveness of estrogen
- Most antibiotics reduce the microbial flora of GIT, which retards the enterohepatic circulation of estrogens.

# Nursing Responsibilities

- Assess BP prior to and periodically during use
- Monthly breast self-exam and annual mammograms are recommended
- In clients with breast cancer and bone metastasis, severe hypercalcemia may be caused by this therapy
- Nausea frequently occurs in morning, but disappears after 1–2 weeks of treatment.

# Patient Education

- Teach client the correct dosage and amount and route of administration
- Advise client to take medicine with food, if nausea occurs
- Inform client that cigarette smoking can increase the risk of thrombus formation
- Advise client to report the signs of fluid retention
- Inform client that when cyclically taken vaginal bleeding will occur during the week each month when estrogen is withheld
- Instruct client to remain in lying down position for 30 minutes after administration of vaginal creams
- Instruct client to report positive Homan's sign
- Instruct client to take medication exactly as prescribed.

# Antiestrogens and Selective Estrogen Receptor Modulators (SERM)

- Antiestrogens:
  - Pure antagonists
  - Clomiphene is for treatment of infertility in anovulatory women 50 mg OD for 5 days starting from 5th day of menstrual cycle.
  - Systemic inquiry (S/E) ovarian tumor, polycystic ovaries and gastric upset
  - Fulvestrant is used for the treatment of breast cancer
- Selective estrogen receptor modulators (SERMs):
  - Compounds with tissue-selective actions
  - The goal of these drugs is to produce beneficial estrogenic actions in certain tissues (e.g. brain, bone, liver) during postmenopausal hormone therapy
  - Antiestrogenic action is due to inhibition of human breast cancer cells
  - Tamoxifen(10-20 mg bd), raloxifene, toremifene.

# Progesterone

- Progesterone is one of the steroid hormones
- It is secreted by the corpus luteum and by the placenta, and is responsible for preparing the body for pregnancy and, if pregnancy occurs, maintaining it until birth
- C21 steroid hormone—involved in the female menstrual cycle, pregnancy and embryogenesis
- Progesterone belongs to a class of hormones called progestogens and is the major naturally occurring human progestogen
- Progesterone should not be confused with progestins, which are synthetically produced progestogens
- **Synthetic progestogens:** Two classes of progestins are derivatives of either C21 or C19 steroid compounds:
  - The C21 derivatives are almost pure progestins, have weaker antiovulatory action and are used primarily as an adjuvant to estrogens for HRT, e.g. medroxyprogesterone, dydrogesterone
  - The C19 nortestosterone derivatives are having most potent antiovulatory action and more androgenic activity used in combined contraceptive pills, e.g. norethisterone, levonorgestrel, desogestrel

# Synthesis

- Synthesized from pregnenolone, a derivative of cholesterol
- The precursor of the mineralocorticoid aldosterone
- Conversion to 17-hydroxyprogesterone of cortisol and androstenedione
- Androstenedione can be converted to testosterone, estrone and estradiol.

#### Sources

- Progesterone is produced in the adrenal glands, gonads, brain, and during pregnancy, in the placenta
- Increasing amounts are produced during pregnancy:
  - Initially, the source is the corpus luteum
  - After the 8th week production of progesterone shifts over to the placenta
  - The placenta utilizes maternal cholesterol as the initial substrate
  - Most of the produced progesterone enters the maternal circulation
  - Some is picked up by the fetal circulation and is used as substrate for fetal corticosteroids
  - At term the placenta produces about 250 mg progesterone per day

## **Mechanism of Action**

- The progesterone receptor (PR) has a limited distribution in the body; confined mainly to female genital tract, breast, CNS and pituitary
- Since the ligand-binding domains of the two PR isoforms are identical, there is no difference in ligand binding
- However, the biological activities of PR-A and PR-B are distinct, and depend on the target gene in question:
  - PR-B mediates the stimulatory activities of progesterone
  - PR-A strongly inhibits this action of PR-B.
- Upon binding progesterone, the heat shock proteins dissociate, and the receptors are phosphorylated and that bind with high selectivity to progesterone response elements located on target genes and regulates transcription.

# Pharmacokinetics

- **Absorption:** Progesterone undergoes high first pass metabolism. Therefore synthetic preparations are more commonly used
- Progesterone esters in oily solution for IM administration
- $t\frac{1}{2}$  is 5–7 minutes
- t<sup>1</sup>/<sub>2</sub> of synthetic progestins is 8–24 hours
- Metabolism: By liver enzymes
- Excretion by urine after conjugation.

# **Effects of Progesterone**

#### **Reproductive System**

- Fallopian tubes—growth and development, inhibition of uterine contraction during pregnancy and of immunologic rejection of fetus
- Uterine endometrium—induction of secretory phase in estrogen primed endometrium
- Mammary glands-development of secretory system for lactation

#### SECTION 1: Pharmacology-II

- Cervix—viscous, scanty mucus secretion as barrier to sperm penetration.
- CNS—body temperature increases, depressant effect and feedback control.
- **Metabolic effects**—increases basal insulin levels, increased appetite, fat deposition increases, decrease in HDL and catabolic action.

#### Preparations and Doses

The preparations and doses of progesterone along with their routes of administration has been presented in **Table 4.2**.

#### Indications

118

- As contraceptive
- Hormone replacement therapy
- Dysfunctional uterine bleeding
- To treat endometriosis—long-term therapy with progestins may be used as the progestin can inhibit estrogen-dependent growth of ectopic endometrial tissue
- Premenstrual syndrome/tension
- Threatened abortion—occurs in progesterone deficiency; pure progestins are used
- Endometrial carcinoma—repress the metastatic endometrial mass.

#### Contraindications

- In patients with hepatic disease or dysfunction
- Incomplete abortion, suspected pregnancy or undiagnosed vaginal bleeding
- Relatively contraindicated in patients with hyperlipidemia, thromboembolic disease (especially in smokers)
- Caution is appropriate in patients with heart disease due to increased fluid retention and edema.

#### Side Effects

- Breast engorgement, headache, rise in body temperature, edema, acne and mood swings may occur
- Irregular bleeding and amenorrhea can occur

| 1 5                  |  |
|----------------------|--|
| Dose                 | Route  |
| 10–100 mg,100–400 mg | IM, oral   |
| 250–500 mg           | IM   |
| 5–20 mg, 50–150 mg   | Oral, IM   |
| 5–10 mg              | Oral   |
| 5–10 mg              | Oral   |
| 0.1–0.5 mg/day       | Oral   |
| 5–10 mg              | Oral   |
| 10–40 mg/day         | Oral   |
| 150 µg               | Oral   |
|                      | Dose         10–100 mg,100–400 mg         250–500 mg         5–20 mg, 50–150 mg         5–10 mg         5–10 mg         0.1–0.5 mg/day         5–10 mg         10–40 mg/day         150 µg |

#### Table 4.2: Preparation and doses of progesterone.

- The 19-nortestosterone derivatives can lower plasma HDL levels—may promote atherogenesis
- Long-term use may increase the risk of breast cancer
- Blood sugar may rise and diabetes can be precipitated.

#### **Nursing Responsibilities**

- Prior to administration in clients with current H/o depression, make a plan to deal with worsening or recurrent depressive symptoms
- A thorough physical examination should be done with special attention to pelvic organs, breasts and hepatic function
- A Papanicolaou (Pap) test should be done prior to initiation of therapy and every 6-12 months while client is taking medicines
- Monitor vital signs including BP
- Monitor I/O

# **Patient Education**

- Instruct client about dosing and timing of the medication
- Warn client about possible side effects
- Inform postmenopausal women of possibility of resumption of cyclical vaginal bleeding
- Instruct client to monitor BP and I/O
- Caution client to exposure to UV lights
- Instruct client to monitor the glucose level closely
- Instruct client to take medication with food

# Antiprogestins

- Antiprogestin, first discovered in 1981, is mifepristone, used to terminate pregnancy
- In the presence of progesterone, mifepristone acts as a competitive receptor antagonist for both progesterone receptors
- When administered in the early stages of pregnancy, mifepristone causes decidual breakdown by blocking uterine progesterone receptors, which leads to detachment of the blastocyst, decreasing hCG production.

# **ORAL CONTRACEPTIVE AND HORMONE REPLACEMENT THERAPY**

# **Hormonal Contraceptives**

These are hormonal preparations used for suppression of fertility. The word 'contraception' means interception in the birth process at any stage ranging from ovulations to ovum implantation.

# Definition

These are the birth control methods that act on the endocrine system. Hormonal contraceptives when properly used are the most effective spacing method contraception. Hormonal contraceptive may be estrogen, progesterone or testosterone.

SECTION 1: Pharmacology-II

# Classification (Fig. 4.2)



Fig. 4.2: Types of hormonal contraceptives.

# **Oral Contraceptives**

#### Types of Oral Contraceptive

- Combined pills
- Mini pill (progesterone only pill)
- Post coital pill (emergency contraceptive)
- Centchroman (non-hormonal estrogen receptor antagonists)

## Combined Pills

It contains estrogens and progestin. It is the most effective and popular method for contraception with 99%–99.5% success rate.

Combined pills can be:

*Monophasic:* No phasic increase or decrease in the estrogen, progestin content during 21 days of pill administration.

Biphasic/Triphasic: Level of estrogen remains same, but progestin level are altered.

The goal of these therapies are to minimize the occurrence of irregular bleeding while maintaining the efficacy. One tablet is taken daily for 21 days, starting on 5th day of menstruation. The next course is started after a gap of 7 days in which bleeding occur. Thus a cycle of 28 days is maintained.

# Mini Pill (Progestin only Pill/Ezy Pill)

- It has been devised to eliminate the estrogen, because of many long-term risks associated with estrogen
- The efficacy is 96%–98%
- The menstrual cycle tends to become irregular and ovulation occurs in 20%–30% women, but other mechanisms contribute to contraceptive action
- Not associated with deep vein thrombosis (DVT) or heart disease.

# Candidate for mini pill

- Cigarette smokers over age 35
- Women with H/O blood clots
- Women with H/O high BP
- Women who experience extreme migraine.

#### Postcoital Contraceptive (Morning after Pill)

- They are recommended within 48 hours of an unprotected intercourse
- High dose estrogen/progestins are quite effective in emergency contraception, when given immediately after unprotected coitus

**CHAPTER 4:** Drugs Used for Hormonal Disorders and Supplementation...

- Acts by preventing ovulation and post-fertilization and implantation of blastocyst
- Delay in maturation of endometrium.

The following are the three regimens available:

- Levonorgestrel 0.5 mg + ethinyl estradiol 0.1 mg—taken as early as within 75 hours of unprotected intercourse and repeated after 12 hours. Women usually experience nausea and vomiting
- Levonorgestrel 0.75 mg taken twice with 12 hours:
  - Gap within 72 hours of intercourse. Effective mainly as post-ovulatory methods of fertility control
- Mifepristone 600 mg single dose taken within 72 hours of intercourse have high success rate and fewer side effects:

Have high failure rate and side effects than oral combined pills.

#### Centchroman (Ormeloxifene/ Chhaya)

- It is a non-steroidal estrogen antagonist or selective estrogen receptor modulator (SERM) introduced in national family welfare program to be distributed as an oral contraceptive under the brand name Saheli
- Available as a tablet containing 30 mg of centchroman taken twice a week for first 3 months and then once a week subsequently to be continued irrespective of the following menstrual cycle, as long as contraceptive is desired
- The missed tablet should be taken as soon as possible, if the dose is missed for more than 7 days then reinitiate the therapy
- Contraceptive effects usually reversible within 6 months
- Has long plasma t<sup>1</sup>/<sub>2</sub> about 1 week
- Acts as an anti-implantation agent by inducing embryo-uterine asynchrony, accelerated tubal transport and suppression of deciduation.

# **Preparation and Doses**

Preparations and doses of oral contraceptives is presented in Table 4.3.

# Advantages

- One of the most effective reversible method for birth control
- Simple and easy to use
- Regulate menstrual cycle
- Decrease acne
- Reduces the risk of ovarian and endometrial cancer
- May reduce perimenopausal symptoms

# Disadvantages

- May be taken everyday
- May cause irregular bleeding
- Effectiveness may be reduced by other medication
- Not be used by women over age 35
- Does not protect against sexually transmitted infections (STIs)
- May increase the number of headaches
- May not be suitable for breastfeeding women

#### SECTION 1: Pharmacology-II

122

#### Table 4.3: Oral contraceptives with their doses.

| Drug                              | Dose                   |  |  |
|-----------------------------------|------------------------|--|--|
| Combined pills                    |                        |  |  |
| Norgestrel, ethinyl estradiol     | 0.3 mg, 30 μg          | Mala-D (21 tabs+ 7 ferrous sulfate<br>60 mg tablet |  |
| Levonorgestrel, ethinyl estradiol | 0.25 mg, 50 μg         | Ovral  |  |
| Desogestrel, ethinyl estradiol    | 0.15 mg, 30 μg         | Novelon  |  |
| Desogestrel, ethinyl estradiol    | 0.15 mg, 20 μg         | Femilon  |  |
| Phased pills                      |                        |  |  |
| Levonorgestrel, ethinyl estradiol | 50–75–125 mg, 30 μg    | Triqular   |  |
| Norethindrone, ethinyl estradiol  | 0.5–0.75–1.0 mg, 35 μg | Orthonovum   |  |
| Postcoital pills                  |                        |  |  |
| Levonorgestrel, ethinyl estradiol | 0.25 mg, 50 μg         | Ovral (2+2)  |  |
| Levonorgestrel                    | 0.75 mg                | Norlevo  |  |
| Mini pills                        |                        |  |  |
| Norethindrone                     | 0.35 mg                | Nor-qd   |  |
| Norgestrel                        | <b>75</b> μg           | Ovrette  |  |

# **Hormone Replacement Therapy**

Hormone replacement therapy is a system of medical treatment for surgical menopausal, perimenopausal and to a lesser extends postmenopausal women. It includes administrations of estrogen and progesterone combinations. Due to cessation of ovarian function at menopause women suffer a number of physical, psychological and emotional consequences. Also aid in prolong life and may reduce incidence of dementia.

The benefits and risks of HRT are considered below:

- · Reduction of vasomotor menopausal symptoms and vaginal atrophic changes
- Reduction in risk of osteoporosis and fractures
- Cardiovascular events—improves HDL/LDL RATIO, retard atherogenesis, risk of stroke increase
- Neuroprotective and CNS effects—insomnia and fatigue are reduced, improvement in cognitive abilities
- Cancers—predisposes to endometrial cancer and cause breast cancer. Protective effect on colorectal carcinoma
- Increase the risk of developing gall stones
- Trigger the migraine
- Improvement in quality of life
- Newer benefits:
  - Decreased risk of colorectal cancer by 37%
  - Decreased age-related tooth loss

123

- Decreased age-related macular degeneration
- Delay of onset and progression of Alzheimer's disease when started early postmenopausal

#### Indications

- Relief of menopausal symptoms
- Prevention of osteoporosis
- To maintain the quality of life in menopausal years
- **Special group of women to whom HRT should be prescribed:** Premature ovarian failure, Gonadal dysgenesis, Surgical or radiation menopause

# Dosage, Route and Indication of HRT (Table 4.4)

Low dose oral conjugated estrogen 0.3 mg daily is effective and has got minimal side effects. Dose interval may be modified as daily for initial 2-3 months then it may be changed to every other day for another 2-3 months and then every third day for the next 2-3 months. It may be stopped thereafter if symptoms are controlled.

| Forms                                | Indication  | Dosage  | Route   |
|--------------------------------------|---|---|---|
| Oral estrogen<br>regime              | Hysterectomy  | Estrogen: Conjugated equine estrogen 0.3 mg or 0.625 mg                       | Orally  |
| Estrogen and<br>cyclic progestin     | Intact uterus   | Estrogen for 25 days and progesterone is added for last 12-14 days            | Orally  |
|                                      | Endometrial<br>hyperplasia<br>Hysterectomy  | 17 beta estradiol implants 25 mg, 50 mg or<br>100 mg (for 6 months)           | Subdermal<br>implants   |
| Continuous                           | Endometrial 1 g applicator of gel, delivering 1 mg of<br>hyperplasia estradiol daily                                  |   | Percutaneous<br>estrogen gel over<br>skin and anterior<br>wall of thigh |
| estrogen and<br>progestin<br>therapy | Endometrial<br>hyperplasia  | 3.2 mg of 17 betaestradiol, releasing about<br>50 ug of estradiol in 24 hours | Transdermal<br>patch applied<br>below the<br>waistline                  |
| 5                                    | AtrophicConjugated equine vaginal estrogenVaginitiscream 1.25 mg dailyUrogenitalatrophyContraindicatedto systemic HRT |   | Vaginal cream   |
|                                      | Breast carcinoma<br>Endometrial<br>carcinoma<br>Progestins  | Medroxyprogesterone acetate 2.5 – 5 mg/<br>day                                | Orally  |

#### Table 4.4: Dosage, route and indication of HRT.

SECTION 1: Pharmacology-II

# Contraindication

124

- · Known or suspected pregnancy or breast cancer
- Undiagnosed genital tract bleeding
- Presence estrogen dependent neoplasm in the body
- History of venous thromboembolism
- Active liver disease
- Gallbladder disease

## Side Effects

- Fluid retention
- Bloating
- Breast tenderness or swelling
- Headaches
- Indigestion
- Depression
- Acne
- Backache
- Endometrial cancer
- Breast cancer
- Venous thromboembolic (VTE) disease
- Coronary heart disease (CHD)
- Lipid metabolism—increase the cholesterol level
- Dementia, Alzheimer's disease

# Nursing Responsibilities (Table 4.5)

Table 4.5: Nursing responsibilities in process of hormone replacement therapy.

| Prior to<br>adminis-<br>tration | <ul> <li>Obtain a complete history including personal or familial history of breast cancer, gallbladder disease, diabetes mellitus, liver or kidney disease.</li> <li>Obtain a drug history to determine possible drug interactions and allergies.</li> <li>Assess cardiovascular status including hypertension, history of MI, cerebrovascular accident, or thromboembolic disease.</li> </ul>   |
|---------------------------------|---|
| During<br>HRT                   | <ul> <li>Monitor for thromboembolic disease. (Estrogen increase risk for thromboembolism).</li> <li>Monitor for abnormal uterine bleeding. (If undiagnosed tumor is present, these drugs can increase its size and cause uterine bleeding).</li> <li>Monitor breast health. (Estrogen promote the growth of certain breast cancer).</li> <li>Monitor for vision changes. (These drugs may worsen myopia or astigmatism and cause intolerance of contact lenses).</li> <li>Encourage client not to smoke. (Smoking increases risk of cardiovascular disease)</li> <li>Encourage client to avoid caffeine. (Estrogens and caffeine may lead to increased CNS stimulation).</li> <li>Monitor for seizure activity. (Estrogen- induced fluid retention may increases risk of seizures).</li> <li>Monitor client's understanding and proper self- administration. (Improper administration may incidence of adverse effects).</li> </ul> |
| Evaluation phase                | <ul> <li>The client verbalizes relief of unpleasant symptoms of menopause.</li> <li>The client demonstrates an understanding of the drug's actions by accurately describing drug side effects and precautions.</li> <li>The client accurately states signs and symptoms to be reported to the healthcare provider.</li> </ul>   |

# **VAGINAL CONTRACEPTIVES (FIGS. 4.3A TO D)**

# **Vaginal Rings**

Vaginal rings containing levonorgestrel have been found to be effective. The hormone is slowly absorbed through vaginal mucosa. The ring is worn in the vagina for 3 weeks and removed for the fourth weeks (Fig. 4.3A).

# Female Condom

Female condom (FEMIDOM) is also a sheath made up of thin, transparent, soft plastic, closed at smaller end and opened at the wider end. It is of single time usage (Fig. 4.3B).

# Diaphragm (Vaginal Diaphragm and Dough Diaphragm)

Dough cap named after a German physician Dutch Neo Mathusians, 1882. It is a shallow, soft rubber cup, with a stiff but flexible rim, made up of coiled spring, which helps in retention. Size varies from 5 to 10 cm in diameter (**Fig. 4.3C**).

# **Cervical Cap**

It is thimble-shaped like diaphragm but smaller. It covers the vaginal portion of the cervix, thus acting as a barrier. The woman inserts the cervical cap with spermicidal, in the proper position in the vagina before having sexual intercourse **(Fig. 4.3D)**.



Figs. 4.3A to D: Types of vaginal contraceptives: A: Vaginal ring; B: Female condom; C: Diaphragm; D: Cap contraceptive.

1<u>25</u>

# Nursing Responsibilities

- β-adrenergics: If client continues on to deliver after receiving uterine relaxant medications, be prepared with oxytocic for treatment of PPH
- Monitor vital signs and I/O
- Nifedipine: Avoid grapefruit juice during administration
- Use indomethacin for short period of time
- If mother is using terbutaline during pregnancy, monitor the neonate for hypoglycemia.

# **MULTIPLE CHOICE QUESTIONS**

| 1. | Which of the following is a non-steroidal | for  | m of estrogen?   |
|----|---|------|------------------|
|    | a. Diethylstilbestrol                     | b.   | Ethinylestradiol |
|    | c. Mestranol                              | d.   | Tibolone         |
| 2. | What is the effect of estrogen on cervix? |      |                  |
|    | a. Proliferation                          | b.   | Thickening       |
|    | c. Watery secretion                       | d.   | Both a and c     |
| 3. | Preferred route of administration of estr | adio | ol.              |
|    | a. Topical                                | b.   | Oral             |
|    | c. IM                                     | d.   | IV               |
| 4. | Clomiphene is used in treatment of:       |      |                  |
|    | a. Dysmenorrhea                           | b.   | Infertility      |
|    | c. Gynecomastia                           | d.   | Delayed puberty  |
| 5. | Route of administration of progesterone   | est  | ers.             |
|    | a. Oral                                   | b.   | Topical          |
|    | c. IM                                     | d.   | IV               |
|    |   |      |                  |
| An | swer Key                                  |      |                  |
| 1. | a 2. c 3.                                 | с    | 4. b 5. c        |
|    |   |      |                  |

# **FURTHER READING**

- 1. Acconcia F, et al. Palmitoylation-dependent estrogen receptor alpha membrane localization: regulation by 17 beta-estradiol. Mol Biol Cell. 2005;16:231.
- 2. Action to control cardiovascular risks in Diabetes study group: effects of intensive glucose lowering in type 2 diabetes. N Engl J Med. 2008;358:2545.
- 3. Adler AL, et al. Association of systolic blood pressure with macrovascular and microvascular complications of type 2 diabetes (UKPDS 36): prospective observational study. By Med J. 2000;321:412.
- 4. Advance collaborative group: Intensive blood glucose control and vascular outcomes in patients with type 2 diabetes. N Engl J Med. 2008:358:2560.
- 5. Alesci S, et al. Glucocorticoid-induced ospeoporosis: from basic mechanisms to clinical aspects. Neuroimmunomodulation. 2005;12:1.
- 6. American thyroid association (http://www.thyroid.org).
- 7. Anderson GL. et al. Women's Health Initiative Steering Committee: effects of conjugated equine estrogen in postmenopausal women with hysterectomy. JAMA. 2004;291:1701.
- 8. Bacopoulou F, Greydanus DE, Chrousos GP. Reproductive and contraceptive issues in chronically ill adolescents. Eur J Contracept Reprod Health Care. 2010;15:389.
- 9. Baggio LA, et al. Biology of incretins: GLP-1 and GIP. Gastroenterology. 2007;132:2131.
- 10. Bamberger CM, Schulte HM, Chrousos GP. Molecular determinants of glucocorticoid receptor function and tissue sensitivity. Endocr Rev. 1996;17:221.

# Textbook of Pharmacology, Pathology and Genetics for Nurses-II

The Textbook of Pharmacology, Pathology & Genetics for Nurses-II has been precisely written as per the revised syllabus of Indian Nursing Council for BSc Nursing Program. This textbook is an excellent attempt towards presentation of comprehensive, lucid, illustrative, and example-oriented content of Pharmacology, Pathology and Genetics.

#### **Salient Features**

- Simple and lucid content: This textbook provides clear concise information about current concepts and principles
  of pharmacology, pathology and genetics in simple and lucid manner incorporating their applications to healthcare
  and nursing practices.
- Easy-to-follow: This is an applied, user-friendly, and self-explanatory textbook with simple language and
  presentation for the students.
- Comprehensive presentations: The textbook provides in-depth coverage of all aspects of pharmacology, pathology, and genetics in a comprehensive and concise manner.
- An applied textbook: This book will equip students to prepare for exams as well as independently apply the knowledge of pharmacology, pathology and genetics into their clinical practices and patient care.
- An example-oriented book: This is an example-oriented book where numerous nursing examples are cited throughout the textbook for enriching the understanding about the applicability of pharmacology, pathology and genetics knowledge into nursing practices.
- Illustrative presentation: Presentation of content has been supplemented with illustrations to facilitate quick
  review and recall important concepts.
- Systemic and logical organization: Content organized in systemic and logical manner to gain better understanding of the presented content.
- Revision MCQs: This textbook covers the MCQs for quick revision of topics.
- Authentic content: Content has been contributed and reviewed by a panel of experts in the field of pharmacology, pathology and genetics.

Suresh Sharma MSC (N) PhD FNRS RN (USA) is currently Professor and Principal, College of Nursing, All India Institute of Medical Sciences, Jodhpur, Rajasthan, India. He is also ICN-Global Nursing Leadership Fellow-2020 at Global Nursing Leadership Institute, Geneva, Switzerland. A gold medalist in nursing education, he did his postgraduate studies at PGIMER, Chandigarh, and received his doctorate in nursing administration from Punjab University, Chandigarh. He is registered as an RN with Board of Registered Nursing, California, USA, and received the coveted Florence Nightingale Award in 2001. He is a prolific writer and has published seven textbooks in nursing, beside a number of chapters contributed and more than 100 research papers published in international and national journals of repute. Dr Sharma is also Founder President, Society of Perioperative Nurses, India and Society for Critical Care Nursing, India. He is also a section editor of Journal of Medical Evidence.

Printed in India

Available at all medical bookstores or buy online at www.jaypeebrothers.com



JAYPEE BROTHERS Medical Publishers (P) Ltd. EMCA House, 23/23-B, Ansari Road, Daryaganj, New Delhi - 110 002, INDIA www.jaypeebrothers.com

Join us on f facebook.com/JaypeeMedicalPublishers

