# Polycystic Ovarian Disease and Endometriosis

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## Objectives

At the end of this lecture, the student should be able to:

- Know the clinicopathologic features of endometriosis with special emphasis on: definition, typical sites and theories behind its pathogenesis.
- Understand the clinical manifestations and pathologic features of polycystic ovarian disease.

• It is characterized by bilateral enlargement of ovaries by multiple small cysts, chronic anovulation and clinical manifestations secondary to excessive production of estrogens and androgens, mainly **androgens**.

• Other names for this disease include **polycystic ovarian syndrome** and **Stein-Leventhal syndrome**.



- The initial abnormality resulting in the syndrome is not known but is believed to be related to hypothalamus-pituitary dysfunction leading to over-secretion of luteinizing hormone (LH).
- LH in turn stimulates the ovary to produce excess androgens. The secretion of follicle stimulating hormone (FSH) is inhibited → leading to suppression of ovulation and formation of cystic follicles in the ovary.
- Patients have:
  - high levels of LH
  - low FSH
  - high testosterone
  - high estrogen



- It usually affects young women (between 15 and 30 years) and present with:
  - secondary amenorrhea with anovulation
  - oligomenorrhea or irregular menses
  - infertility
  - hirsutism
  - virilism due to increased androgenic (masculinizing) hormones
  - obesity
  - acne



- A: The ovarian surface reveals numerous nodular elevations of clear cysts.
- B: Cut section shows several subcortical cystic follicles in the ovary.



• C: Cystic follicles seen in a low-power microphotograph.



- Ovaries:
  - Ovaries are 2 times the normal size with many subcortical cysts measuring 0.5 to 1.5 cm in diameter.
  - The outer portion of the cortex is thickened and fibrotic (cortical stromal fibrosis) with multiple cysts underneath. The follicular cysts usually have a prominent theca interna layer.
  - Corpora lutea are frequently absent (because no ovulation, women with PCOD have anovulatory cycles).



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- Endometrium:
  - Chronic anovulation → unopposed estrogen → hyper estrogenic state → endometrium may develop estrogen associated hyperplasia and show any of the following:
    - simple with or without atypia
    - complex hyperplasia, with or without atypia
    - or even endometrial adenocarcinoma

#### • Women with PCOD are at risk for the following:

- Endometrial hyperplasia and endometrial cancer
- Insulin resistance/Type II diabetes
- High blood pressure
- Depression/Anxiety
- Dyslipidemia
- Cardiovascular disease
- Strokes
- Weight gain
- Miscarriage
- Acanthosis nigricans (patches of darkened skin under the arms, in the groin area, on the back of the neck)
- Autoimmune thyroiditis

- Treatment with drugs that either induce ovulation (clomiphene or hCG) or regulate the menstrual cycle restores fertility.
- Reduction of ovarian volume by wedge resection of the ovaries is also successful in initiating ovulation and restoring fertility.
- The endometrial changes usually regress once ovulation is achieved.



• Normally endometrial glands and endometrial stroma are found in the endometrium of the uterus.

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- Endometriosis is the presence of ectopic endometrial glands and stroma outside the uterus.
- It is non-neoplastic condition.

- Endometriosis is usually found on the peritoneal surfaces of the reproductive organs and adjacent pelvic organs. The most frequent locations are:
  - ovary (the most common site, approx. 50%)
  - Pouch of Douglas and uterine ligaments (second most common)
  - Occasionally: cervix, vagina, perineum, bladder, large bowel and umbilicus
  - Rarely: small bowel, kidneys, lungs, nose and brain
- It has been reported in men. The sites involved have been the bladder, scrotum and prostate.



- Like the uterine endometrium it is responsive to the hormonal variations of the menstrual cycle, and bleeds during menstruation.
- Therefore in endometriosis there is menstrual type bleeding at the site of the ectopic endometrium, resulting in blood filled areas (e.g. chocolate cysts).

- The clinical presentation depends on the site of endometriosis:
  - Dysmenorrhea
  - Dyspareunia
  - Cyclic abdominal pain (there is usually a severe menstrual-related pain)
  - Infertility
  - Fibrous adhesions

• The behavior of endometriosis is benign with no malignant potential. It may recur after surgical excision but the risk is low.

Endometriosis usually appears as multiple red or brown (due to hemosiderin)
1 mm to 5 mm nodules (some may form larger masses or cysts). Dense fibrous adhesions may surround the foci.

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- Repeated hemorrhage into the ovary with each menstrual cycle produces cysts, filled with chocolate-brown material. The cyst is called "chocolate cyst".
- With time the ovaries become totally cystic and turn into large cystic masses filled with chocolate brown fluid.

#### Chocolate cyst of ovary (endometriotic cyst)



#### Fibrous adhesions



- Ectopic endometrial glands and endometrial stroma are present.
- Denatured blood from previous bleeding is present.
- Macrophages containing hemosiderin (siderophages) are present.
- When endometriosis develops in a muscular organ, the smooth muscle around it is often hyperplastic.



## Adenomyosis

- This is defined as the presence of endometrial glands and endometrial stroma in the myometrium of the uterus.
- It is more common in the posterior wall than the anterior wall (but it may affect both walls in the same uterus).
- The disease is primarily a disorder of parous women and is uncommon in the nullipara.

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## Adenomyosis



- It is associated with menorrhagia and severe dysmenorrhea, however it is asymptomatic in 1/3 of the patients.
- When extensive, the lesions cause myometrial thickening with small yellow or brown cystic spaces containing fluid or blood.

• The behavior of this conditions is benign with no known malignant potential and it usually regresses after the menopause.

## Adenomyosis

 Cross section through the wall of a hysterectomy specimen of a 30year-old woman who reported chronic pelvic pain and abnormal uterine bleeding. The endometrial surface is at the top of the image, and the serosa is at the bottom.



#### Reference

• Kumar V, Abbas AK, Aster JC. Robbins Basic Pathology. 10<sup>th</sup> ed. Elsevier; 2018. Philadelphia, PA.

## Thank You

End of Lecture