

# PCOS

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# INTRODUCTION

- Heterogeneous, complex genetic trait
- Aetiology is unclear
- Common cause of menstrual irregularity
- Common cause of subfertility
- Also a cause of hyperandrogenism
  - Acne
  - Hirsutism
- First described by Stein and Leventhal in 1935

# INTRODUCTION

- Polycystic ovaries- does not mean Polycystic Ovarian Syndrome
- 4-12% of population
  
- Diagnosis: 2003 Rotterdam criteria
  - 2 of the following 3 required
    - Oligo or anovulation (menstrual irregularity)
    - Hyperandrogenism
    - USS appearance of Polycystic ovaries

# DIAGNOSIS

## ⊙ Menstrual dysfunction

- Typically begins in prepubertal period
- Menarche may be delayed
- Normally oligomenorrhoea
  - Less than 9 periods a year
- Amenorrhoea
  - No periods for 3 or more consecutive months
  
- Anovulation
- Infertility

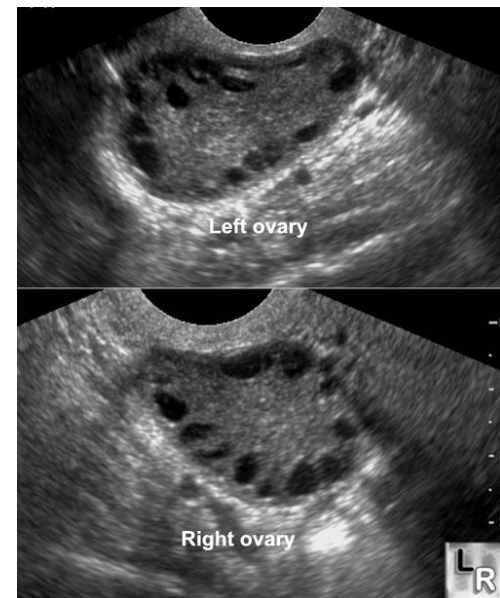
# DIAGNOSIS

## ○ Androgen excess

- Clinical signs
- Acne
- Hirsutism
  - Excess thick pigmented body hair
  - Upper lip, chin, midsternum, lower abdomen
- Male pattern hair loss
- Less common virilisation
  - Clitoromegaly
  - Deepening of voice
  - More likely above due to androgen secreting tumour

# DIAGNOSIS

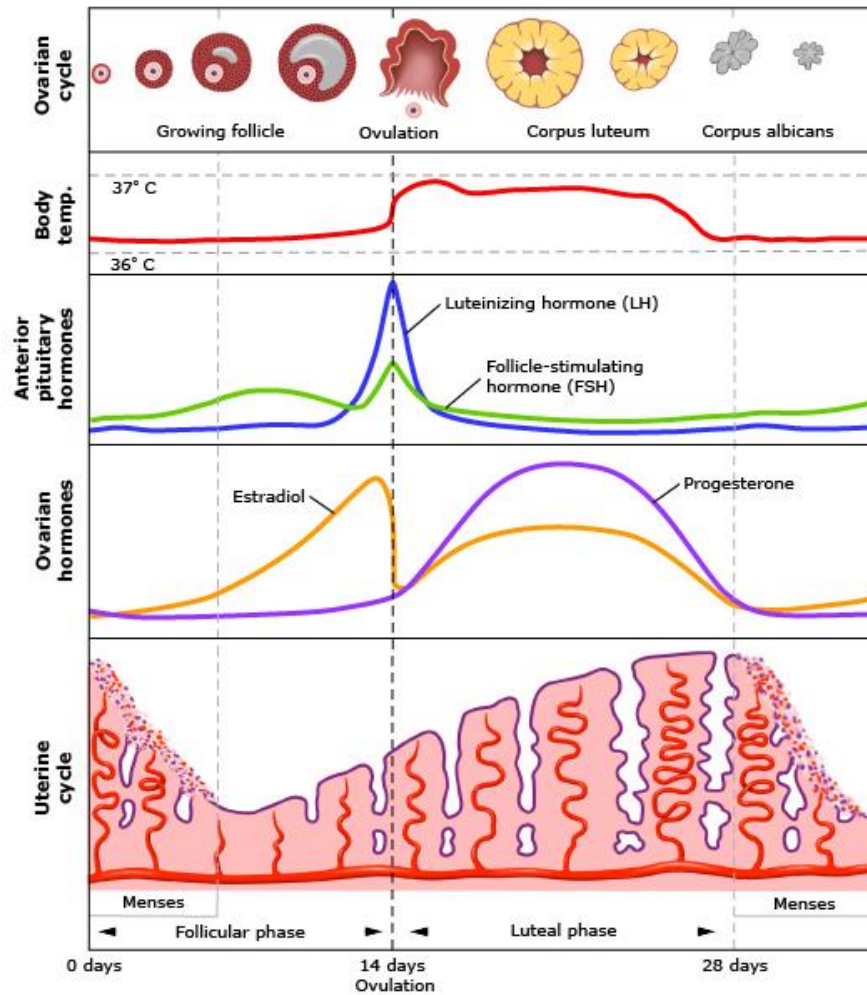
- Polycystic appearance of Ovaries
  - Multiple follicles
  - Thick stroma
- However this may also be seen  
In normal cycling women
- Isolated finding is of no  
clinical significance



# CLINICAL MANIFESTATIONS

- ◉ 40-85% of women with PCOS are overweight
- ◉ Not a diagnostic criteria
- ◉ Insulin resistance is present
  - 30% of lean
  - 70% of overweight
- ◉ Increased risk of type 2 Diabetes
- ◉ Increased risk of CHD
- ◉ Sleep apnea
- ◉ Metabolic syndrome
- ◉ Mood disorder- anxiety, depression, binge eating

## Menstrual cycle





# CLINICAL MANIFESTATION

- Ovulation leads to progesterone secretion and menstruation
- Chronic anovulation results in lack of progesterone and increased estrogen
- Obesity contributes to hyperestrogenic state
- Chronic stimulation of endometrium with lack of menstruation can lead to endometrial hyperplasia
- If untreated, this can lead to cancer

# EVALUATION

- Clinical picture
  - BMI measurement
  - Glucose intolerance
  - Dyslipidaemia
  - Fatty liver
  - Obstructive sleep apnea
  
- Based on Rotterdam criteria, history and examination should be enough in most women

# EVALUATION

## ⦿ Hirsutism

- Beware ethnic variation
- East asian women least hairy
- White european and black women intermediate
- Middle eastern, south asian and mediterranean most hairy

## ⦿ Diagnosis of exclusion

- Exclude other cases of amenorrhoea
- Menopause
- Pregnancy
- Thyroid disorder, hyperprolactinoma

# OTHER CONDITIONS

- **Androgen excess**
  - Tumours
  - Ovarian hyperthecosis
  - Congenital adrenal hyperplasia

# INVESTIGATIONS

## ○ Biochemistry

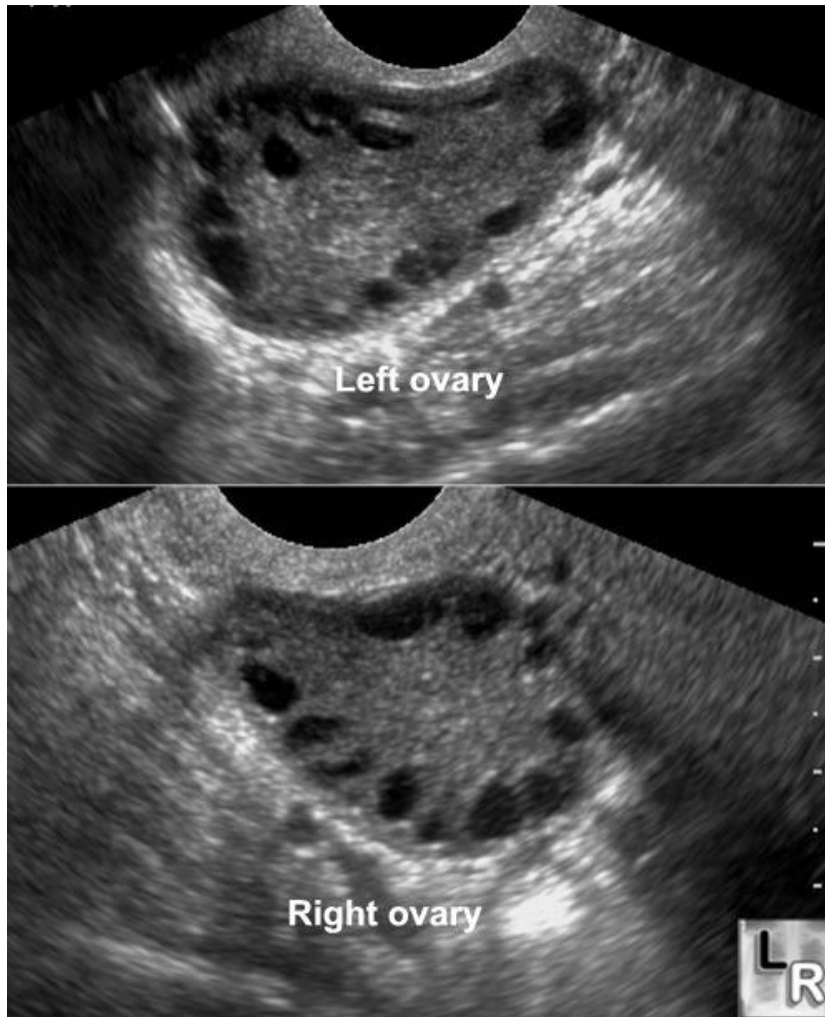
- TFT
- Prolactin
- FSH- to rule out menopause
- LH/FSH ratio- can be helpful but not diagnostic criteria
- Must be  $>2$  to 3:1
- Must be done in follicular phase but most women are oligomenorrhoeic
- Serum total Testosterone levels if very raised may be due to other causes
- Possibly SHBG- if low then confirms diagnosis

# INVESTIGATIONS

## ○ Ultrasound

- Performed as part of diagnosis
- However not always needed
- Absence of Poly cystic morphology does not exclude
- Useful if only one either criteria present
- If scan is to be done then must be Transvaginal
- Remember these are follicles and not cysts
- 12 or more follicles with large ovaries
- Unilateral PCO is sufficient
- Can also check for endometrial thickness

# IGNORE INCIDENTAL FINDING



# MANAGEMENT

- Management depends on goals
- Goals are
  - Amelioration of hyperandrogenic symptoms
  - Metabolic disorders
    - Obesity
    - Type 2 diabetes
    - Cardiovascular disease
  - Prevention of Endometrial hyperplasia
    - Due to chronic anovulation and estrogen excess
  - Contraception
  - Ovulation induction



# MANAGEMENT

## ○ Lifestyle changes

- Diet and exercise for weight reduction in Obese women
  - Improves insulin resistance and hyperandrogenism
  - Fertility improves
- Multidisciplinary management
  - Gynaecology
  - Dietician
  - Endocrinologist
  - Clinical psychologist

# MANAGEMENT

- Oral Contraceptive Pill
  - Mainstay of treatment
  - Treats hyperandrogenism
    - Increases SHBG which mops up excess testosterone
    - Yasmin- contains drospirenone or Dianette which has cyproterone acetate- anti androgenic
  - Regulates Menses
  - Endometrial protection
  - Improve insulin sensitivity
- However, not useful if goal is to peruse a pregnancy

# MANAGEMENT

- Oral Contraceptive Pill- risks
  - Increased risk of VTE particularly if obese
  - Risk assessment important
    - BMI <30
    - Age
    - Smoking status
    - Family history of VTE
- Can use Progesterone only pill
  - Mini pill ie cerazette
- Or cyclical progestogen therapy to induce menstruation

# MANAGEMENT

## ○ Goals- endometrial protection

- Progestagen therapy ie Norethisterone of Provera cyclically
- Mirena IUCD

## ○ Goals- Hirsutism

- OCP
- Antiandrogens
  - Spironolactone
  - Cyproterone acetate
  - Finasterite
  - Flutamide- hepatotoxicity
  - Laser hair removal/ waxing
  - Vaniqa - Topical antiandrogen that prevents hair growth but must be used indefinitely

# MANAGEMENT

## ○ Goals- Weight loss

- Lifestyle changes
  - Even 5-10% weight loss can result in ovulation
- No diet superior to other
- Low carb diets however very popular
- Bariatric surgery

## ○ Goals - Metabolic disorders

- Weight loss
- Metformin
- Statins

# MANAGEMENT

- Goals- Anovulatory infertility
  - Weight loss
  - Ovulation induction
    - Clomiphene citrate from D2-D6 of cycle
    - Ensure semenalysis normal
    - Letrozole
    - Metformin
    - Exogenous gonadotrophins (IVF/IUI)
  - Laparoscopic Ovarian drilling
    - Too invasive but effective if all else fails

# OVARIAN DRILLING



# METFORMIN

- ⊙ Inhibits production of hepatic glucose
- ⊙ Increases fatty acid oxidation
- ⊙ Enhances Insulin sensitivity
- ⊙ Observational studies
  - Reducing serum androgens and menstrual cycle
  - Effective in achieving ovulation
- ⊙ Randomised studies had conflicting results
  - No consensus on dose and duration in the latest Cochrane review



# METFORMIN

## ○ Body Weight

- May achieve weight loss
- Randomised control trials failed to confirm this
- Combined with lifestyle changes may work

## ○ Ovulation Induction

- One study found higher live birth rates with Metformin- but only just reached statistical significance

# RCOG SCIENTIFIC PAPER

- ◉ August 2017

- ◉ Conclusion

‘ Metformin appears to have limited role in improving reproductive outcomes in PCOS but in patient specific groups- ie obesity or those with higher impaired glucose tolerance or type 2 diabetes’

# CONCLUSION

- ◉ Common problem
- ◉ Long term sequelae
- ◉ Treatment depends on goal
- ◉ Mainstay is weight loss and lifestyle changes
- ◉ Remember psychological impact
- ◉ Avoid labelling