

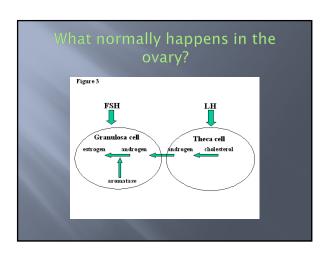


PCOS is the most common endocrinopathy in women

7% of all women 18-45

Obesity 1/3 of all US women

Incidence of PCOS is increasing with increase obesity



Signs of PCOS Obesity Irregular menses Normal gonadotropins Not pregnant Insulin resistance Hirsutism Acne/ hair loss



HAIR-AN Hyper-Androgenism Insulin Resistance Acanthosis Nigricans

3 PCOS Criteria Schemes NIHCD 1990 Rotterdam 2003 AES 2006

NIHCD 1990 Hyper-androgenism Irregular menses Excludes other diagnoses Weaknesses: not all PCOS have either of these, but a broad spectrum of disease from mild to severe



Androgen Excess Society

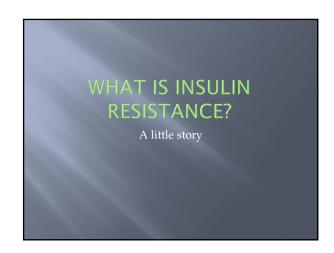
- Hyperandrogenism with hirsutism
- Oligoovulation
- Excludes others diseases
- Weakness, may make treatment pool too small
- Doesn't account for obesity, insulin resistance

Obesity in PCOS

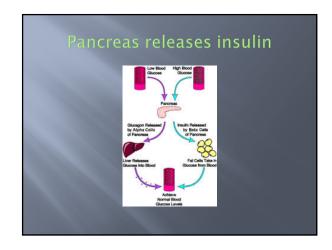
- Present in 35-60% of PCOS patients
- Difficulty losing weight major presenting symptom
- PCOS worsens with increase weight
- Seems to be a set point over which things go downhill

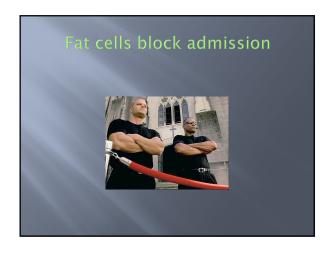
Insulin Resistance

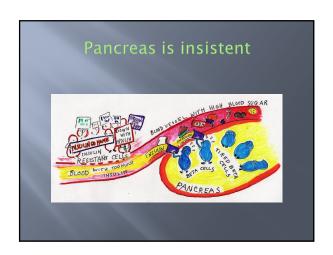
- Prevalence is 50-75% in PCOS
- Type II DM in 10 % of PCOS
- A HUGE PROBLEM: Diabetes now costs \$276
 Billion per year direct and indirect costs
- \$174 Billion just for hospitalization and meds

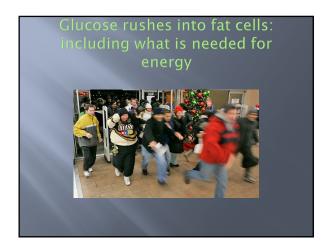
















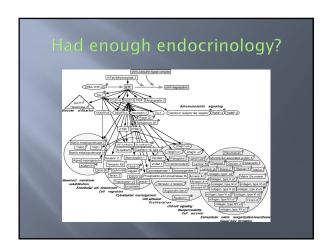


What does this have to do with ovulation?

- Increase pulses of LH in PCOS
- Increased estrone made in fat cells
- E1 inhibits FSH
- Lower FSH fails to stimulate follicles appropriately
- Elevated LH tone fails to signal LH surge
- No ovulation of mid-sized follicles

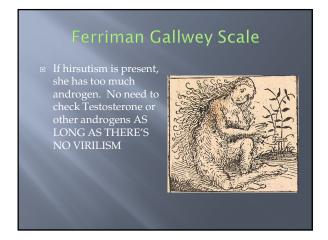
There's more

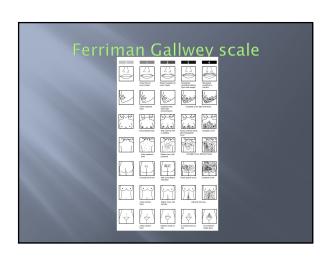
- IGF 1 and 2 are important hormones in ovarian control of ovulation
- Excess circulating insulin binds to their receptors and blocks their action
- Excess fat results in increase places to make estrone from androstenedione
- Higher LH results in more androstenedione and less estradiol in the ovary, inhibiting follicle growth

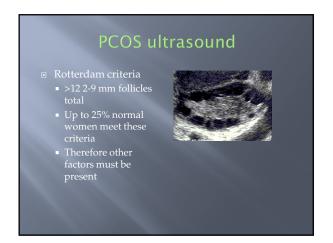


The Work Up

- H&P
- Weight BMI BP abdominal circumference
- Breast staging if patient is young, look for discharge
- Thyroid
- Acanthosis nigricans, acne, male hair loss, hirsutism, skin tags, striae purple or white
- Buffalo hump, thin limbs, skin and muscle breakdown







Testing for Insulin Resistance

- No standardized criteria
- 2 hour GTT is often recommended but may not be sensitive enough to detect patient who may benefit from metformin
- 75g, 2 hour glucose less than 140, 2hour insulin > 80-100
- Inconvenient, uncomfortable, expensive

Adrenal testing not needed unless

- Hirsutism occurs young
- Positive family history
- High risk ethnic groups
- 17OHP (am)
 - <200 is normal</p>
 - >800 is diagnostic
 - 200-800 more testing is required (ACTH stim)

Work up in review

- Pregnancy test
- Prolactin, Day 3 FSH (if cycling), TSH
- Ultrasound of ovaries
- If all bloods are normal and ultrasound meets criteria with symptoms, you've got a patient with PCOS

Rare diseases

- Leprechaunism is severe congenital insulin resistance caused by a mutation in the insulin receptor
- Death in infancy is common
- Elfin features, LBW, failure to thrive



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Rare diseases

- Cushing disease
- Truncal obesity, thin limbs
- Diabetes, hypertension
- Excessive cortisol
- Dexamethasone suppression test is best screen
- 1:1 million



Other tests that may be worthwhile

- Endometrial biopsy if menses less than 3 per year for several years
- Fasting lipids
- Complete metabolic panel if metformin planned
- Testosterone total and/or free if virilism or no response to therapy

Treatment

- Weight loss: even 2-5% will help
- Exercise reduces insulin resistance and increases sensitivity, even without weight loss
- Hormone therapy
- Metformir
 - Increase dose gradually
 - Dietary counseling: no sugar, syrup, HCFS
 - If not tolerating, try XR, food diary

PCO=OCP

- Contraception
- Increases SHBG
- Decreases androgens

- Protection from endometrial hyperplasia

Ovulation induction

- Clomid start day 3-5, continue for 5 days
- Do pregnancy test before taking
- Do ultrasound day 10-12 if no ovulation
- LH kit for timing intercourse
- Baseline ultrasound to rule out cysts

- Eflornithine (Vaniqa) faster than laser but not permanent

- Antiandrogens need a good contraception!!

Summary

- Rotterdam Criteria need oligoovulation, hyperandrogenism, PCOS ovaries 2/3
 Insulin resistance and obesity are most important comorbidities
 Diet and exercise are mainstays of treatment
 OCP for cycle control, hyperandrogenism

- Antiandrogens for hyperandrogenism, but need birth control
- Metformin for insulin resistance and help with weight loss