Polycystic ovarian syndrome

PCOS is a common complex endocrine disorder , affecting woman at reproductive age group and most common cause of chronic anovulation . It is also called stein – Leventhal syndrome .

Definition

It is chronic anovulation syndrome, its diagnosis made when two out of three of the following criteria are found (Roherdom's criteria 2003).

- 1. Clinical or biochemical evidence of androgen excess after exclusion of others related disorders .
- 2. Oligo and / or anovulation.
- 3. Ultrasound appearance of polycystic ovaries :- presence of ≥ 12 follicles in each or one ovary measuring 2 9 mm surrounding hyperandrogenic stroma and / or increased ovarian volume (> 10 ml) which is called Pearl's of sting .

This ultrasonic features are not applicable to women take Cocp. Obesity, with insulin resistance, elevated LH level, reversed LH to FSH ratio and subfertility are common features.

Prevalence

It is thought that PCOS occurs in about $6-10\,\%$ of women in reproductive age group worldwide . The prevalence is much more higher in south Asian women .

Pathophysiology and etiology

The etiology of PCOS is largely unknown and still understood but seems to involve certain complex interactions between environmental (diet & exercise) & multiple genetic factors. The mode of inheritance is autosomal dominant pattern.

Several factors are implicated in pathophysiology of PCOS involving ovarian dysfunction characterized by increased production of ovarian androgen

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hypothalamic dysfunction LH hypersecretion stimulate androgen production by theca cells and peripheral insulin resistance in both obese & non obese PCOS patients leads to insulin hyper secretion ______ more hyperandrogenism by inhibition of hepatic production of SHBG & IGF – BP1 causing anovulation .

Other PCOS manifestations

- Obesity: 50 % or more patients of PCOS are obese.
- Metabolic syndrome :- Type II diabetes , hypertension , dyslipidaemia , therosclerosis and is hemic heart disease .
- Dermatological manifestations :- hirustism , alopecia , acne , oily skin , seborhoce & acanthosis negricans .

Acamthosis negricans is area of increased skin pigamentation that are velvetry in texture & occurs in axilla and other flextures . It occurs in 2 % of PCOS women & it is a marker of insulin resistance .

- Oligamenonhoea am enorhoea :- occurs in 65 75 % of patients with PCOS.
- Hirustism: 30 % 70 % of patients.
- Subfertility in 75 % of patients .
- Recurrent miscarriage in 50 60 % (why?)

Laboratory tests:-

- 1- Elevated testaserone level.
- 2- Decreased SHBG level.
- 3- Elevated LH level.
- 4- Elevated LH / FSH ratio .
- 5- Increased fasting insulin level.

Management:

PCOS treatment depends whether or not fertility is desired , and the patients have cyclical disturbance or not , so the treatment should be directed at patients symptoms .

- 1- Changes of certain life styles such as diet and exercise are considered first line treatment for adolescent girl and women with PCOS.
- 2- Treatment of obesity :- weight lose is the cornerstone of treatment of PCOS , wherever a small amount of weight loss ($\stackrel{\sim}{-}$ 5 kg) can lead to spontaneous resumption of ovulation which minimize the risk of PCOS during pregnancy regarding fetal & maternal morbidities .

Weight loss is best achieved by diet & excersise or by weight losing agents such as centrally acting serotonin (sibutramine) & peripherally acting lipase inhibitor (orlistat) .

3- Treatment of oligo and / or amenohoca :- without treatment , there is theortical risk of unopposed estrogen stimulation of endometrium causing endometrial cancer .

Cyclical progesterone is useful in treatment of PCOS. oral progesterone should be given at least 10 days from day $16^{th} - 25^{th}$ day of cycle or for 21 days from $5^{th} - 25^{th}$ day of cycle for 3 - 6 months oral progesteronis are either .

No rithe sterong acetate (primolut n 5 mg bd) .

Or Medroxyprogesterone acetate (10 mg oncedaily).

Or cyclical COCP . which is the first line treatment for cyclical regulation in adolesunt females .

- 4- Treatment of Hirustim.
- 5- Ovulation induction.

Antiestrogen (clomiphene citrate) is used for ovulation induction on hypothalamus , clomid binds to and blocks estrogen receptors leading to increased GnRH. Pulse amplitude & FSH level leads to better follicular growth . 50-150 mg of elamid frem 2^{nd} day or 5^{th} day of cycle for 5 days is given .

It can be offered for 6-12 months clomiphene citrate ferilure or resistance or recumont obotions are beit indications for use of aromatase inhibitor (Letrazol) or gonadotrophines use or laproswpical ovarion drilling.

Tamoxifen :- 20 – 40 mg daily for 5 days.

Gonadotrophines: FSH use in patients with clomid resistance or failure.

Gonadotrophines agonists: - used by subcutancous pump.

Aromatase inhibitors: letrazol and anstrazole decrease ovarian estrogen production causing endogeneous FSH stimulation & spontaneous ovulation & pregnancy with less aboution rate.

6- Insulin sensitizing agents

Metformin , a biguanide , is the most common sensitizing agent used for ovulation induction in PCOS patients either as a first line theropy or second line second line or in combination with ovulatory drugs . It has been shown that non- obese patients respond better than obese one . Metformin if continued in the first trimester of gestation may decrease miscarriage risk in obese women . Metoformin (Glucophage) :- 500mg b.d or T.d.s. or 850 mg once or twice daily

7- Laproswpical ovarian drilling (LOD)

LOD is mainly indicated in clomiphene – resstant patient which have similar efficacy to gonadotrophines with advantage of low risk of multiple pregnancy rate & OHSS. LOD can cause ovulation in 80 & pregnancy rate of 60x. It is also correct the biochemical abnormalities such as high LH & androgen.

In LOD only minimal ovarion thermal damage is required to stmulate ovulation (four points diathermy set at 40 watt for 40 seconds at each point).

Long term sequale of PCOS

- 1- Pregnancy induced Hypertension & CHT.
- 2- Type II D..M. & gestational D.M.
- 3- Dyslipidaemia.
- 4- Endometrial cancer with 5 folds increase.
- 5- Ovarian cancer in 2 5 % of PCOS patients.
- 6- Breast cancer.

- 7- Recurrent miscoviage .
- 8- Stroke & myocardial infarction .