CONTEMPORARY AND AYURVEDIC PERSPECTIVE OF POLYCYSTIC OVARIAN SYNDROME (PCOS): A CRITICAL REVIEW.

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ABSTRACT:
Polycystic ovary syndrome (PCOS) is a highly prevalent heterogeneous syndrome of clinical and biochemical androgen excess, ovulatory dysfunction and polycystic ovaries. It affects about 6–10% of women worldwide and is thought to be one of the leading causes of female sub-fertility. Androgen excess, insulin resistance play important role in pathogenesis of PCOS along with genetic factors. It is now increasingly perceived as disorder of changed life styles. In Ayurveda, PCOS is described under the headings of Yonivyapad and Artavadushti. This paper reviews the contemporary and ayurvedic perspectives of PCOS to develop holistic approach for the prevention and treatment of PCOS.

KEY WORDS: polycystic ovarian syndrome, Ayurved, Yonivyapad, insulin resistance.

INTRODUCTION:
Polycystic ovary syndrome (PCOS, is a highly prevalent heterogeneous syndrome of clinical and/or biochemical androgen excess, ovulatory dysfunction and polycystic ovaries (PCO).¹ Women with PCOS are at increased risk of reproductive abnormalities. They also have an increased risk of developing type 2 diabetes mellitus (T2DM) and cardiovascular disease (CVD). Stein and Leventhal were the first to describe the triad of menstrual dysfunction, PCO and androgenic features.² Worldwide, PCOS affects 6–10% of women ³⁻⁶. In India, PCOS affects 9⁻ 22% of young girls in their reproductive age ⁷⁻⁸ which makes it most common endocrinopathy in women of reproductive age group. The National Institute for Health (NIH) Criteria 1990 was revised in 2003 and Rotterdam criteria⁹ has been adopted world over to diagnose PCOS. However, recently in 2006, Androgen Excess Society (AES) has come up with a consensus statement, defining PCOS as a hyperandrogenic state and emphasizes the presence of either clinical and/or biochemical features of hyperandrogenism along with other features of PCOS for diagnosis.¹⁰

Pathogenesis:
Androgen excess: Androgen excess is considered by some investigators to be the sine qua non of PCOS; however, only 80–85% of women with clinical hyperandrogenism have PCOS.¹¹

Ovulatory dysfunction and polycystic ovaries: In PCOS, ovarian hyperandrogenism, hyperinsulinemia from insulin resistance and altered intraovarian paracrine signaling can disrupt follicle growth. The consequent follicular arrest in PCOS is accompanied by menstrual irregularity, anovulatory subfertility and the accumulation of small antral follicles within
the periphery of the ovary, giving it a polycystic morphology.\textsuperscript{9, 12} Follicular arrest in PCOS develops when granulosa cells in antral follicles normally begin to express aromatase (at a size of 7 mm).\textsuperscript{13} An excess intraovarian 5α-reduced androgens has shown to inhibit granulosa cell aromatase activity in vitro and impair follicle growth.\textsuperscript{14}

**Insulin resistance and PCOS:** Many women with PCOS have insulin resistance beyond that predicted by their BMI, with 50–70% of these women demonstrating insulin resistance by various measures.\textsuperscript{15} Most women with PCOS are young and develop compensatory hyperinsulinemia, with impaired glucose tolerance detectable more readily by oral or intravenous glucose testing than by basal glucose measures.\textsuperscript{16} Chen YH et al reported that reduced glucose transporter 4 (GLUT 4) expression is one of the reason of insulin resistance in women with PCOS.\textsuperscript{17}

**Gonadotropin abnormalities:** LH hypersecretion increases serum immunoactive and bioactive LH levels in about 70% of women with PCOS,\textsuperscript{18} and elevated LH pulse amplitude and frequency induces a twofold to threefold elevation in circulating LH versus FSH levels. Increased LH pulse frequency in PCOS, from enhanced hypothalamic gonadotropin releasing-hormone (GnRH) pulsatile release, occurs owing to reduced steroid hormone negative feedback on LH secretion because of androgen excess.\textsuperscript{19}

**Genetics:** With 20–40% of first-degree female relatives of women with PCOS affected by the syndrome, PCOS is more prevalent among family members\textsuperscript{20} than in the general population.\textsuperscript{4} Genes for which association with PCOS or its component traits have been replicated include fibrillin 3 (\textit{FBN3}) and 17β-hydroxysteroid dehydrogenase type 6 (\textit{HSD17B6}).\textsuperscript{21, 22}

**Environmental factors:** Lifestyle profoundly affects the phenotypic expression of PCOS. Weight gain worsens metabolic and reproductive abnormalities of PCOS, as evidenced by increased total and abdominal obesity as well as insulin resistance, menstrual irregularity and hyperandrogenism in women with the most severe PCOS phenotype.\textsuperscript{23} A sedentary lifestyle alone also contributes to metabolic dysfunction in PCOS because moderate-intensity exercise without weight loss improves insulin resistance and decreases body adipose tissue.\textsuperscript{24} Bisphenol A (BPA), a widely used estrogenic industrial plasticizer accumulation in susceptible women might exaggerate the severity of the PCOS phenotype.\textsuperscript{25}

**Clinical features:** Women with PCOS often seek care for menstrual disturbances, clinical manifestations of hyperandrogenism, and infertility. Menstrual disturbances commonly observed in PCOS include oligomenorrhea, amenorrhea, and prolonged erratic menstrual bleeding. Hirsutism is a common clinical presentation of hyperandrogenism occurring in up to 70% of women with PCOS.\textsuperscript{26}

**Diagnosis:** Women with possible PCOS and other androgen excess disorders should undergo a relatively straightforward evaluation. Women to be evaluated would present with unwanted or excess body and/or facial hair growth, irregular menstruation, polycystic-appearing ovaries detected incidentally on ultrasonography, alopecia and/or acne. The evaluation should confirm the presence of features of PCOS and also exclude related disorders.

The \textit{Rotterdam criteria}\textsuperscript{27} of assessment require the presence of two out of the following three criteria: Oligomenorrhea and/or anovulation. Hyperandrogenism (clinical and/or biochemical). Polycystic ovaries on ultrasound, with the exclusion of other etiologies such as congenital adrenal hyperplasia, androgen secreting tumours, Cushing syndrome, thyroid dysfunction and hyperprolactinaemia.

**Management:** Current treatment of PCOS can be summarized as follows:

**Treatment of infertility:** Weight loss is recommended as first-line therapy for the management of infertility in overweight and obese women with PCOS. This is followed by induction of ovulation (OI) with Clomiphene citrate. Subsequently, administration of insulin sensitizer with Clomiphene is advisable. Gonadotropin therapy and FSH hormone are the next option followed by Gonadotropins with insulin sensitizer. A study reported that treatment with metformin, a
antihyperglycemic drug increased live birth rates in obese women.28

**Treatment of androgen-related symptoms:** Combined hormonal contraceptives (CHC) are a good treatment option for those patients that do not wish to become pregnant. Anti-androgenic therapy is used to reduce the masculine effects of testosterone like alopecia, hirsutism.29

**Ayurvedic Perspective of PCOS**

Ayurveda classifies PCOS as a kapha disorder. Ayurveda describes Polycystic Ovarian Syndrome to have an equal involvement of the Dosha, Dhatu and Upadhathu. In Ayurveda, PCOS is not defined as single disease, rather its symptoms bear a resemblance to the terminologies defined as ‘Anartava’ (Amenorrhoea), ‘Yoniyapad’ (anatomical and physiological disorders of the reproductive system) like – Arajaska (Oligomenorrhoea due to vitiation of Vata Doshha), Lohitakshaya (Oligomenorrhoea due to vitiation of Vata-Pitta Doshas), Vandhya (infertile), Pushpaghni Revati (Idiosyncratic anovulatory menstruation), Abeejata (anovulation), Rajodushti and Ashtartava Dushti (menstrual flow disorder due to vitiation of Doshas) etc. The terms Raja and Artava have been used synonymously or otherwise in the classics. Usually Raja is considered as the Upadhathu of Rakta dhatu whereas Artava as the Saptam Dhatu itself. Similarly, their Srotas (channels) are also two entirely different entities. In the present paper, Raja has been considered as the menstrual flow while Artava is indicative of the ovum.

**CORRELATION OF PCOS WITH CERTAIN AYURVEDIC TERMINOLOGIES:**

**ArajaskaYoniyapad:** When Pitta situated in Yoni and uterus vitiates Raka, the women becomes extremely emaciated and discolored, this condition is known as Arajaska.30 Acharya Chakrapani has described amenorrhea as a symptom.

**LohitakshayaYoniyapad:** The Nidan Sevan of Vata-Pitta Pradhana Aahar-Vihar causes a vitiation of these Dosa resulting in Rajakshenata (scanty menstruation), the lady suffers from burning sensation, emaciation and discoloration. This may be presented in either of the previously discussed ways. Again, a similarity to the contemporary symptom of menstrual irregularity is noted but it fails to clarify oligo/anovulation.

**VandhyaYoniyapad:** Sushrutacharya quotes this type of Yoniyapad presenting as Nashtartava (loss of menstruation).31 Charakacharya states this condition to arise due to loss of ovulation. Harita elaborates on six types of Vandhyayoni, each having specific features, management and prognosis. One of them is Anapatya Vandhya (infertility) wherein Dhatukshaya is etiological factor of Nashtartava. Here, Artava is considered as the Saptadhatu or ovum and its loss results in infertility. However this type is incurable. The above mentioned Anapatya Vandhyayoni can be fairly compared with PCOS due to the similar features of anovulation and absence/irregularity of menstruation thereby resulting in sterility. However, other clinical features tend to vary.

**Abeejata (Anovulation):** Sushrutacharya states the aetiological factors of Shukradushti (vitiation of sperm) in males to be similar to those of Rajodushti in females leading to Abeejata. The same factors are also responsible for the vitiation of Doshas in females causing the vitiation of Raja/ Artava. Hence, just as ‘Shukrambeejata’ (azoospermia) is seen as a result of vitiation of Shukra, a condition of ‘Artavaabeejata’ (anovulation) is noted in females due to vitiation of Artava. Charakacharya too quotes frequent or untimely coitus, over-exercise, unbalanced diet that includes Ruksha (dry), Tikta (bitter), Kashaya (astringent), Atilavana (excessively salty), Amla (sour) and Ushna (hot) Aahar, as also Chinta / Shoka (stress-related tension), Bhaya (fear), Krodha (anger) and Aghata i.e. injuries due to Shastra (weapon) or Kshara (alkali) as the causative factors of Shukradoshti.30 These can be correlated with the current lifestyle changes.

**Ashtartava Dushti:** Vagbhatacharya states that like Shukra, Artava can too be vitiiated by the Doshas resulting in eight types of Artavadoshti. Such vitiation leads to Abeejata.32

**Rajodushti:** This terminology, put forth by Sushrut acharya is a result of the vitiation of Raja by the Dosa, primarily Vata and Pitta resulting in its Ksheenata (Oligomenorrhoea). The other clinical features of PCOS are however not observed.
AYURVEDIC MANAGEMENT OF PCOS:
1) "Nidana Parivarjana" (avoid the causative factors) is said to be the very first step towards the management of PCOS. As Agnimandya, Medovriddhi, Apana Vayu and Kapha dushti plays the major role in the pathogenesis of the syndrome, so taking above fact into the consideration, Pathya Ahara-Vihara (dietary regimen & exercise) is to be used.
2) For Agnimandya and Aampachana, use of Trikatu Churna, Chitrakadi Gutika, Shadushana Churna, Haritaki Churna (Terminalia Chebula), Hingwashtaka Churna is to be done in order to palliate the Srotovarodha and to facilitate the Apana-Vatanulomana.
3) For Medovriddhi (Obesity), use of Takrarirshita, Madhu like lekhanadrayyas (scrapping agents) along with Yava, yavaka, kulattha etc. as aahara (diet) is mentioned by Acharya Charaka in chikitsa of Atistula (obese). Moreover, lifestyle modification as well as regular exercise is also emphasized.
4) To remove the Sanga (obstruction) of Aartavavah Srotas, Uttar-Basti (douche) is given with Dhanvantari Taila.
5) Vamana Karma (emesis) – as it alleviates the Srotovarodha by eliminating vitiated Kapha. As Kapha is soumya in prakriti, decrease in Kapha consequently increases Aartava of Aagneya nature.
6) Shatpushpa and Shatavari Churna (Asparagus racemosus Wild) are to be used in females with deficiency or loss of Aartava, women getting their menstruation but not conceiving.
7) Kanchanara Guggulu, Sukumara Ghrita for reducing the size of formed ovarian cysts.
8) Pathadi Kwath mentioned by Acharya Sushruta in Vatakaphaja Aartava dushti given orally along with the matrabasti of Shatpushpa taila after the cessation of menstrual cycle for seven days is found efficient due to its properties of Aampachana, agnideepana, Vatanulomana, Srotoshodhana and Vata-Kaphashamana.

The main objective of the chikitsa is-
- To flame the Jathragni and dhatvagni with the use of Deepana- pachana dravyas to correct the Agnimandya.
- To remove the Sanga of Aartava-vaha srotas and Srotoshodhana through various Samshodhana karmas to balance the imbalanced doshas i.e. reduction of Kapha and Anulomana of Apana Vata; to nourish the dushta dhatus and to regulate the irregular menstrual cycle.
- To reduce the weight through Lekhana dravyas, Pathya aahara- vihara and regular exercise to regulate the hormonal imbalance
- To reduce the size of already formed ovarian cysts through Kanchanara Gugglu & Sukumara Ghrita as Kanchanara is very useful in treating extra growths or tumors.
- To enhance the chances of conception with the use of Shatpushpa and Shatavari Churna. As Shatavari (Asparagus racemosus Wild) is known for its phytoestrogenic properties.

CONCLUSION:
Integration of Ayurvedic treatments into modern medical approaches for PCOS has the potential to improve patient outcomes. Encouraging results can be obtained with Shastrokta formulations, Panchakarma, Pathya aahara and regular exercise due to its holistic approach towards-Samprapti-Vighatana.

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