

Nutritional Management Of Pcos: A Review Article

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Abstract

Polycystic ovarian syndrome (PCOS) is an endocrine disorder characterized by a marked pattern of ovarian cysts, rupture of anovulation, and endocrine differences affecting women. According to figures from the World Health Organization (WHO), more than 116 million women (3.4%) are affected by PCOS worldwide. Risk factors considered include genetic, neuroendocrine, lifestyle / environmental, obesity that contribute to the development of PCOS. The pathophysiological feature of PCOS is mainly focused on hormonal dysfunction, insulin resistance and hyperandrogenism leading to damage to folliculogenesis that triggers the risk of related factors such as endometrial cancer, type II diabetes. This review highlights a brief overview of risk and pathophysiological treatment with drugs that work to relieve anovulation, infertility and clinical symptoms of PCOS.

Introduction:

Polycystic ovarian syndrome (PCOS) is an endocrine disorder associated with metabolic syndrome. (1) .It is characterized by manifestations of anovulation, ovarian cyst and endocrine variables that affect a woman's health and life (2). This condition can be morphological i.e. by the presence of polycystic ovaries or may be biochemical (hyperandrogenemia). Hyperandrogenism is considered a clinical manifestation of PCOS that interferes with the growth of follicles, micro-cysts in the ovaries and menopause (3, 4).

This complex disease usually affects older women by about 6-10% (5). According to the WHO, the prevalence of PCOS is higher than the 116 million (3.4%) affected women worldwide (6). In India an estimated 8.2% to 22.5% of women are affected depending on the method of diagnosis (7).

Aetiology:

PCOS is mainly represented by the signs and symptoms of many androgens and ovulatory dysfunction disrupts the function of the HPO axis (hypothalamic pituitary ovarian). The female HPO axis is closely aligned and controlled by a network responsible for reproduction and survival of animal species. The HPO axis responds to internal signals such as hormonal and external signals as natural influences. These traits affect future generations through epigenetic factors that affect the brain and

cells of growing viruses (8, 9). The most common clinical symptoms are hirsutism, chronic anovulation, irregular menstruation and infertility. Chronic hyperandrogenism is associated with hypothalamic-pituitary dysfunction, LH hypersecretion, premature granulosa cell luteinization, abnormal oocyte maturation, and premature binding of activated primary follicles (10).

The genetic etiology of PCOS is unknown, family history is common, but family links to PCOS remain unclear. The present study suggests that the collection of PCOS in families resembles a prominent autosomal pattern (11). Natural factors such as obesity can be exacerbated by poor nutrition and exercise, infectious diseases and toxins may also play a role. These factors can be reversed by lifestyle changes such as weight loss and exercise (12).

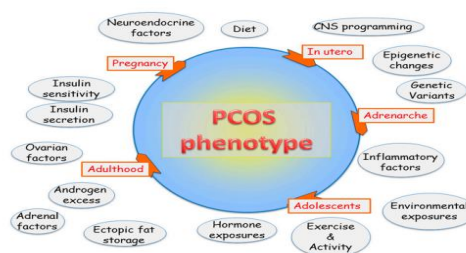


Fig.1 Factors contribute to PCOS phenotype. PCOS encompasses a woman's life cycle (8).

Pathophysiology:

The pathophysiology of PCOS treats it as a multifactorial disease and includes uncontrolled ovarian steroidogenesis, unexplained insulin expression, excessive oxidative stress and genetic and environmental factors (14). An internal factor in theca cells explains hyperandrogenemia in patients with PCOS. This theca cell promotes high levels of androgens due to the internal functioning of steroidogenesis even when there are no trophic factors. This intrinsic dysfunction affects granulosa cells producing high levels of antimullerian hormone via PCOS compared to healthy controls. Studies have shown that the higher the number of follicles, the more pre-antral and small antral follicles, in women with PCOS. Also impairment in apoptotic processes in the mature follicles increases the number in PCOS patients (15, 16, 17).

Decreased insulin sensitivity caused by the binding factor of the post receptor in insulin signaling pathways leading to the internal component of PCOS (18). The study noted that alterations in the genetic expression of other athletes in insulin signaling pathways by microarray genetic analysis. In addition, PCOS is also associated with an increase in secondary glycooxidative stress for mitochondrial dysfunction (19).

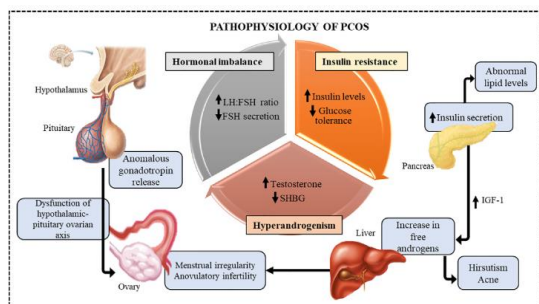
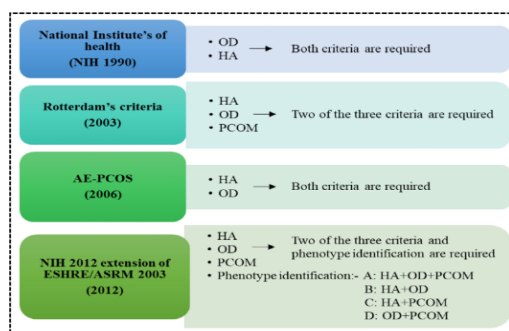


Fig 2. schematic depiction of PCOS linked mechanism (1,13)

PCOS TEST:

According to the old features of PCOS, the current consensus is that the use of the Rotterdam procedure is appropriate for older women. To diagnose PCOS, women should complete two-thirds: oligo-ovulation, anovulation, clinical hyperandrogenism and biochemical or polycystic ovary morphology on ultrasound without any other interference (20).



Summary of PCOS diagnostic criteria. (2,21)

Abbreviations: AE-PCOS: Androgen Excess and PCOS community; ASRM: American Reproductive Organization; ESHRE: European Society for Human Reproduction and Embryology; HA: Hyperandrogenism; OD: Ovulatory dysfunction; PCOM: Polycystic ovarian morphology (12 follicles with 2-9 mm per uterus).

PCOS treatment:

The most commonly used treatments for women with PCOS include lifestyle changes (i.e., diet, exercise and behavioral therapy) and medications such as clomiphene citrate, aromatase inhibitors, low-dose human menopausal gonadotropin or FSH, insulin sensors, laparoscopic ovarian drilling and in vitro . pregnancy (IVF) (22).

NUTRITION INTERVENTION:

Studies have reported that there is no proper diet or diet pattern such as calorie restriction and modified macronutrient dietary composition of PCOS (23, 24). There is a difference in weight loss with different diets, and this variation depends on how the body reacts to different macro- or micronutrients. Negative energy balance appears to be an important factor leading to active weight loss and fat loss, improvement of the menstrual cycle and insulin sensitivity, regardless of the accepted dietary pattern (25, 26).

Low Glycemic Foods:

Studies have shown that low-glycemic diets play an important role in reducing risk and complications and at the same time enhance the clinical and biochemical features of PCOS (27). Insulin resistance is a major cause in obese and non-obese patients with PCOS. Since insulin plays an important role in PCOS and its metabolic factors, controlling PCOS diet is the best step (28,29). About 50% of patients with PCOS develop diabetes or Non-diabetes before the age of 40 (30),

so maintaining weight with a special diet and lifestyle significantly reduces insulin levels and associated symptoms (31).

Low calorie foods:

Researchers believe that a high-calorie diet leads to reversible hormonal disorders and menopause (32). High fiber diets reduce weight and some androgens like testosterone in obese and overweight women suffering from hirsutism. Taking a high fiber diet for 12 weeks results in a significant decrease in free testosterone and LH levels and an increase in SHBG, a decrease in BMI and weight loss in people with PCOS (33).

Role of calcium and vitamin D:

Vitamin D deficiency has been reported in about 67-85% of people with PCOS and there are good associations for PCOS and other well-known joint diseases such as type 2 diabetes, insulin resistance, metabolic syndrome and heart disease. (34). There is a direct effect of calcium and vitamin D on the ovarian tract and the adrenal steroidogenesis may be linked to the observed decrease in circulating androgens (35). Studies have shown a positive link between serum calcium content and insulin levels and insulin resistance and fasting glucose levels in many healthy people. 1,25 (OH) 2D contributes to insulin production with a significant increase in intracellular ionic calcium level following 1,25 (OH) 2D which stimulates insulin secretion by islet cells. Along with a significant decrease in Vitamin D, glucose and phosphorus levels in obese and overweight women with PCOS are negatively associated with insulin and insulin resistance (36). Calcium and vitamin D metabolism affect oocyte maturation and production of androgens (37). Based on research (Razavi et al 2016) describing vitamin D-K-calcium co-supplementation for 8 weeks among vitamin D-deficient women with PCOS has beneficial effects on serum dehydroepiandrosterone sulfate (DHEAS), free testosterone, plasma antioxidant capacity, malondialdehyde (MDA) concentration (38).

Prebiotics and PCOS:

Studies in humans and mice have shown that there is a link between mutations in the gut microbiome and metabolic parameters and in the PCOS clinic. Additionally it has been suggested that dysbiosis of the gut microbiota is a possible pathogenic factor in the development of PCOS. Therefore prebiotics, probiotics and symbiotic products may serve as a new treatment for PCOS (39).

In the gut microbiota a decrease in α diversity and β variability and the formation of micro intestinal biota may lead to changes in intestinal function which may increase PCOS (40). One of the most important mechanisms involved in gut microbiota dysbiosis is activating the immune system. Activating the immune system also disrupts the function of the insulin receptor which causes hyperinsulinemia which increases the production of ovarian androgens and inhibits the growth of normal follicle. This leads to features of the PCOS feature. This can be due to normal obesity and high-fat diets - low fiber in PCOS (41).

Cinnamon:

It is a promising agent in the treatment of PCOS by increasing the activity of phosphatidylinositol 3-kinase in insulin signaling pathways, thereby stimulating insulin action (42). Studies have provided evidence that the presence of inositol is necessary for insulin sensitivity and function, which play an important role in patients with PCOS. They regulate the use of insulin-dependent glucose in the cell by various means such as cellular glucose uptake or glycogen synthesis. In addition inositol plays a role in fertility recovery in women with PCOS (43). Myo-inositol is the second messenger of LH and FSH signaling pathways in oocytes and follicular cells. Follicular maturation and oocyte quality are determined by myo-inositol concentration in human follicular cells (44).

Flax seeds:

Flaxseeds contain high levels of dietary lignan and lignan secoisolariciresinol diglucoside (SDG). A typical dose of 15 gm of flax powder will improve menstrual frequency by significantly reducing ovarian volume and the number of follicles in the ovary (46).

Conclusion:

PCOS is a complex problem due to insulin resistance and obesity. Therapies should therefore include dietary and exercise changes as well as medication interventions. Management of certain conditions such as menstrual irregularities, infertility and hirsutism play an important role in PCOS. Elements of nutrition such as LGD, high fiber diets, balanced diets, low-calorie diets, probiotics have significant effects on PCOS control.

Reference:

- [1]. Behboudi-Gandevani S, Amiri M, Bidhendi Yarandi R, Noroozadeh M, Farahmand M, Rostami Dovom M, Ramezani Tehrani F. The risk of metabolic syndrome in polycystic ovary syndrome: A systematic review and meta-analysis. *Clin Endocrinol (Oxf)*. 2018 Feb;88(2):169-184. doi: 10.1111/cen.13477. Epub 2017 Oct 16. PMID: 28930378.
- [2]. Bulsara, Jeshica et al. "A Review: Brief Insight Into Polycystic Ovarian Syndrome". *Endocrine And Metabolic Science*, vol 3, 2021, p. 100085. Elsevier BV, <https://doi.org/10.1016/j.endmts.2021.100085>.
- [3]. Lin LH, Baracat MC, Gustavo AR, et al. Androgen receptor gene polymorphism and polycystic ovary syndrome. *Int J Gynaecol Obstet*. 2013;120:115–118. [PubMed] [Google Scholar]
- [4]. Patten, Rhiannon K. et al. "Exercise Interventions In Polycystic Ovary Syndrome: A Systematic Review And Meta-Analysis". *Frontiers In Physiology*, vol 11, 2020. Frontiers Media SA, <https://doi.org/10.3389/fphys.2020.00606>.
- [5]. Bozdogan G, Mumusoglu S, Zengin D, Karabulut E, Yildiz BO. The prevalence and phenotypic features of polycystic ovary syndrome: a systematic review and meta-analysis. *Hum Reprod Oxf Engl*. 2016;31:2841–2855. pmid:27664216
- [6]. Bharathi, R.V., Swetha, S., Neerajaa, J., Madhavica, J.V., Janani, D.M., Rekha, S.N., Ramya, S., Usha, B., 2017 Dec 1. An epidemiological survey: Effect of predisposing factors for PCOS in Indian urban and rural population. *Middle East Fertility Society Journal* 22 (4), 313–316.
- [7]. Gupta M SD, Toppo M, Priya A, Sethia S, Gupta P. A cross sectional study of polycystic ovarian syndrome among young women in Bhopal, Central India. *Int J Community Med Public Health*2018;5:95-100.
- [8]. Selma Feldman Witchel, Sharon E Oberfield, Alexia S Peña, Polycystic Ovary Syndrome: Pathophysiology, Presentation, and Treatment With Emphasis on Adolescent Girls, *Journal of the Endocrine Society*, Volume 3, Issue 8, August 2019, Pages 1545–1573, <https://doi.org/10.1210/js.2019-00078>
- [9]. Hochberg Z, Feil R, Constanica M, Fraga M, Junien C, Carel JC, Boileau P, Le Bouc Y, Deal CL, Lillycrop K, Scharfmann R, Sheppard A, Skinner M, Szyf M, Waterland RA, Waxman DJ, Whitelaw E, Ong K, Albertsson-Wikland K. Child health, developmental plasticity, and epigenetic programming. *Endocr Rev*. 2011;32(2):159–224.
- [10]. Palomba S, Daolio J, La Sala GB. Oocyte competence in women with polycystic ovary syndrome. *Trends Endocrinol Metab*. 2017;28(3):186–198.
- [11]. Diamanti-Kandarakis E, Kandarakis H, Legro RS. The role of genes and environment in the etiology of PCOS. *Endocrine*. 2006; 30:19–26.
- [12]. Shannon M, Wang Y. Polycystic ovary syndrome: A common but often unrecognized condition. *J Midwifery Womens Health*. 2012; 57:221–230.
- [13]. Walters, K.A., Gilchrist, R.B., Ledger, W.L., Teede, H.J., Handelsman, D.J., Campbell, R.E., 2018 Dec 1. New perspectives on the pathogenesis of PCOS: neuroendocrine origins. *Trends in Endocrinol . Metabol*. 29 (12), 841–852. doi: 10.1016/j.tem.2018.08.005.
- [14]. Patel, Royal et al. "POLY CYSTIC OVARIAN SYNDROME: AN UPDATED REVIEW". *Journal Of Applied Pharmaceutical Sciences And Research*, 2020, pp. 7-10. *Journal Of Applied Pharmaceutical Sciences And Research*, <https://doi.org/10.31069/japsr.v3i1.2>.
- [15]. Nelson, V. L., Legro, R. S., Strauss, J. F. III., and McAllister, J. M. (1999). Augmented androgen production is a stable steroidogenic phenotype of propagated theca cells from polycystic ovaries. *Mol. Endocrinol*. 13, 946–957. doi: 10.1210/mend.13.6.0311.
- [16]. Azziz, R., Marin, C., Hoq, L., Badamgarav, E., and Song, P. (2005). Health care-related economic burden of the polycystic ovary syndrome during the reproductive life span. *J. Clin. Endocrinol. Metab*. 90, 4650–4658. doi: 10.1210/jc.2005-0628.
- [17]. Pellatt, L., Hanna, L., Brincat, M., Galea, R., Brain, H., Whitehead, S., et al. (2007). Granulosa cell production of anti-Mullerian hormone is increased in polycystic ovaries. *J. Clin. Endocrinol. Metab*. 92, 240–245. doi: 10.1210/jc.2006-1582.
- [18]. Cortón, M., Botella-Carretero, J. I., López, J. A., Camafeita, E., San Millán, J. L., Escobar-Morreale, H. F., et al. (2008). Proteomic analysis of human omental adipose tissue in the polycystic ovary syndrome using two-dimensional difference gel electrophoresis and mass spectrometry. *Hum. Reprod*. 23, 651–661. doi: 10.1093/humrep/dem380.

- [19].Victor, V. M., Rocha, M., Bañuls, C., Sanchez-Serrano, M., Sola, E., Gomez, M., et al. (2009). Mitochondrial complex I impairment in leukocytes from polycystic ovary syndrome patients with insulin resistance. *J. Clin. Endocrinol. Metab.* 94, 3505–3512. doi: 10.1210/jc.2009-0466.
- [20].Powers, S. E., Uliassi, N. W., Sullivan, S. D., Tuchman, L. K., Mehra, R., and Gomez-Lobo, V. (2015). Trends in standard workup performed by pediatric subspecialists for the diagnosis of adolescent polycystic ovary syndrome. *J. Pediatr. Adolesc. Gynecol.* 28, 43–46. doi: 10.1016/j.jpag.2014.03.002.
- [21].Lizneva, D., Suturina, L., Walker, W., Brakta, S., Gavrilova-Jordan, L., Azziz, R., 2016 Jul 1. Criteria, prevalence, and phenotypes of polycystic ovary syndrome. *Fertil. Steril.* 106 (1), 6–15.
- [22].Messinis IE, Messini CI, Anifandis G, Dafopoulos K. Polycystic ovaries and obesity. *Best Pract Res Clin Obstet Gynaecol.* 2015;29(4):479–488.
- [23].Moran LJ, Pasquali R, Teede HJ, Hoeger KM, Norman RJ. Treatment of obesity in polycystic ovary syndrome: a position statement of the Androgen Excess and Polycystic Ovary Syndrome Society. *Fertil Steril.* 2009;92(6):1966–1982.
- [24].Papavasiliou K, Papakonstantinou E. Nutritional support and dietary interventions for women with polycystic ovary syndrome. *Nutrition and Dietary Supplements.* 2017;9:63-85
<https://doi.org/10.2147/NDS.S119738>.
- [25].Stamets K, Taylor DS, Kunselman A, Demers LM, Pelkman CL, Legro RS. A randomized trial of the effects of two types of short-term hypocaloric diets on weight loss in women with polycystic ovary syndrome. *Fertil Steril.* 2004;81(3):630–637.
- [26].Soares NP, Santos AC, Costa EC, et al. Diet-induced weight loss reduces DNA damage and cardiometabolic risk factors in overweight/obese women with polycystic ovary syndrome. *Ann Nutr Metab.* 2016;68(3):220–227.
- [27].Najmieh Saadati, Fatemeh Haidari, Mojgan Barati, Roshan Nikbakht, Golshan Mirmomeni, Fakher Rahim, The effect of low glycemic index diet on the reproductive and clinical profile in women with polycystic ovarian syndrome: A systematic review and meta-analysis, *Heliyon.* 2021;7(11): 2405-8440.
- [28].A.J. Morales, et al., Insulin, somatotrophic, and luteinizing hormone axes in lean and obese women with polycystic ovary syndrome: common and distinct features, *J ClinEndocrinol Metab* 81 (8) (1996) 2854–2864.
- [29].E. Diamanti-Kandarakis, A. Dunaif, Insulin resistance and the polycystic ovarysyndrome revisited: an update on mechanisms and implications, *Endocrine reviews* 33 (6) (2012) 981–1030.
- [30].J. Rojas, et al., Polycystic ovary syndrome, insulin resistance, and obesity:navigating the pathophysiologic labyrinth, *International journal of reproductivemedicine* 2014 (2014) 719050.
- [31].E.T. Wang, et al., Polycystic ovary syndrome and risk for long-term diabetes and dyslipidemia, *Obstetrics and gynecology* 117 (1) (2011) 6–13.
- [32].Forouhari S, Hosseini R, Salehi M, Namavar Jahromi MS, Sayadi M. Effect of vitamin C supplementation on the levels of related hormones in infertile women with polycystic ovary syndrome (PCOS) in Shiraz City. *Int J Health Sci.* 2014;2(1):61-70.
- [33].Krouni A, Forouhari S, Akbarzadeh M, Dabbaghmanesh M, Jowkar F, Salehi M et al. Effect of High Fibre, Low Calorie Balanced Diet in Obese Women with Hirsutism: A Randomised Clinical Trail. *JOURNAL OF CLINICAL AND DIAGNOSTIC RESEARCH.* 2018; Vol-12(6): IC06-IC09.
- [34].Shojaeian, Zahra et al. “Calcium and vitamin D supplementation effects on metabolic factors, menstrual cycles and follicular responses in women with polycystic ovary syndrome: A systematic review and meta-analysis.” *Caspian journal of internal medicine* vol. 10,4 (2019): 359-369. doi:10.22088/cjim.10.4.359.
- [35].De Groot PC, Dekkers OM, Romijn JA, Dieben SW, Helmerhorst FM. PCOS, coronary heart disease, stroke and the influence of obesity: A systematic review and meta-analysis. *Hum Reprod.* 2011;17:495–500. [PubMed] [Google Scholar].
- [36].Mahmoudi T, Gourabi H, Ashrafi M, Salman Yazdi R. Calcitropic hormones, insulin resistance, and the polycystic ovary syndrome. *Fertil Steril.* 2010;93:1208–14.
- [37].Bonakdaran S, Mazloom Khorasani Z, Davachi B, Mazloom Khorasani J. The effects of calcitriol on improvement of insulin resistance, ovulation and comparison with metformin therapy in PCOS patients: a randomized placebo- controlled clinical trial. *Iran J Reprod Med.* 2012;10:465–72. [PMC free article] [PubMed] [Google Scholar]

- [38]. Razavi M, Jamilian M, Karamali M, et al. The effects of vitamin d-k-calcium co-supplementation on endocrine, inflammation, and oxidative stress biomarkers in vitamin d-deficient women with polycystic ovary syndrome: a randomized, double-blind, placebo-controlled trial. *Horm Metab Res.* 2016;48:446–51. [PubMed] [Google Scholar].
- [39]. Gamze Yurtdas, and Yasemin Akdevelioglu. A New Approach to Polycystic Ovary Syndrome: The Gut Microbiota :a review. *JOURNAL OF THE AMERICAN COLLEGE OF NUTRITION.* 2019. pages 1-13. DOI: 10.1080/07315724.2019.1657515.
- [40]. Thackray VG. Sex, microbes, and polycystic ovary syndrome. *Trends Endocrinol Metab.* 2019;30(1):54–65. doi:10.1016/j.tem.2018.11.001
- [41]. Tremellen K, Pearce K. Dysbiosis of Gut Microbiota (DOGMA)—a novel theory for the development of Polycystic *JOURNAL OF THE AMERICAN COLLEGE OF NUTRITION* 9 Ovarian Syndrome. *Med Hypotheses.* 2012;79(1):104–112. doi:10.1016/j.mehy.2012.04.016.
- [42]. Qin B, Nagasaki M, Ren M, Bajotto G, Oshida Y, Sato Y. Cinnamon extract (traditional herb) potentiates in vivo insulin-regulated glucose utilization via enhancing insulin signaling in rats. *Diabetes Res Clin Pract.* 2003;62(3):139–48.
- [43]. Laganà AS, Garzon S, Casarin J, Franchi M, Ghezzi F. Inositol in polycystic ovary syndrome: restoring fertility through a pathophysiology-based approach. *Trends Endocrinol Metab.* 2018;29(11):768–80.
- [44]. Garzon S, Laganà AS, Monastra G. Risk of reduced intestinal absorption of myo-inositol caused by D-chiro-inositol or by glucose transporter inhibitors. *Expert Opin Drug Metab Toxicol.* 2019;15(9):697–703.
- [45]. Fatima farzana K' Abubacker sulaiman F' Ruckmani A, Vijayalakshmi K, karunya Lakshmi . effect of flax seed supplementation in PCOS. *Int. J. Pharm. Sci. Rev. Res.*, 31(1), March – April 2015; Article No. 23, Pages: 113-119.
- [46]. Brooks JD, Ward WE, Lewis JE, Hilditch J, Nickell L. Supplementation with flaxseed alters estrogen metabolism in postmenopausal women to greater extent than does supplementations with equal amount soy. *Am J Clin Nutr*, 79, 2004, 318–325.