



Association between Serum Level of Uric Acid and Polycystic Ovarian Syndrome in Females with and without Insulin Resistance -Baghdad Medical City / 2024

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ABSTRACT:

BACKGROUND:

Polycystic ovarian syndrome (PCOS) is a hormonal disorder of females, the elevated of serum uric acid is linked to metabolic disease as PCOS

OBJECTIVE:

To estimate the prevalence of hyperuricemia in females of PCOS with or without insulin resistance and compare serum level of uric acid in patients of PCOS whether the insulin resistance is present or not.

PATINETS AND METHODS:

Cross-sectional; comparative stud; non-probability convenience sampling was conducted at Out-patient's department of Gynecology of Baghdad Medical City, from March 15th till September 15th 2024. The study tools used measuring serum level of uric acid and HOMA-IR (homeostatic model assessment-insulin resistant) which is the gold standard in assessment of insulin resistance. Also used questionnaire that cover socio-demographic of study participants and used Rotterdam criteria for diagnosis of polycystic ovarian syndrome (PCOS). **Inclusion criteria** all females, that are 20-40 years old diagnosed with PCOS, are included . **Exclusion criteria** include females of PCOS with and without insulin resistance and had diabetes mellitus; hypertension; Cushing's syndrome; coronary heart disease; liver cirrhosis; renal diseases women who are pregnant, lactated, smoker, taken vitamin supplementation and medications such as (diuretic, hormonal, hypoglycemic).

RESULTS:

The study showed that serum level of uric acid was highest among PCOS with insulin resistance 78.1% with P. value 0.008 and there was statistical significant association.

CONCLUSION:

More than two third of PCOS with insulin resistance had increase in serum level of uric acid and approximately less than one third of PCOS without insulin resistance had increase serum level of uric and there was statistical significant association.

KEYWORDS: polycystic ovarian syndrome, insulin resistance, uric acid .

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INTRODUCTION:

Polycystic Ovarian Syndrome is a hormonal disorder of females, the anovulation; infertility; hyperandrogenism and metabolic dysfunctions which are the characteristics of the disease ⁽¹⁾. Main cause of this disease is insulin resistance that cause hyperinsulinemia; altered response of gonadotropins; irregular metabolism of ovarian androgen ^(2,3,4,5).

There are some unique features of PCOS in adolescents as decline in insulin sensitivity during puberty that may developed of insulin

resistance (IR) and hyperinsulinemia ⁽⁶⁾. The clinical features of insulin resistance (IR) and hyperandrogenism usually begin during adolescent in females with PCOS and these may be increased with the presence of obesity. ⁽⁷⁾ There are several criteria used in diagnosis of PCOS; The most common is the National Institute of Health (NIH) and Rotterdam criteria ⁽⁸⁾. These criteria were expanded in 2003 with the introduction of the Rotterdam criteria; considered gold standard which require the two out of the following features of PCOS in absence of

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congenital adrenal hyperplasia; androgen-secreting neoplasms; hyperprolactinemia, (9):

1. Oligo – and /or anovulation.
2. Signs of hyperandrogenism, whether clinical or biochemical (serum testosterone and dehydroepiandrosterone sulfate (DHEAS)).
3. Polycystic ovaries on ultrasonography (the presence of eight or more subcapsular follicular cysts measuring less than 10 mm and an increase in ovarian stroma. (9)

PCOS is estimated to occur about 6–7% of global population (10). The World Health Organization (WHO) estimates that PCOS affects 8–13% of women of reproductive age (11). It is suggested that this increased prevalence is likely due to changes in lifestyle, diet, and stress (12,13).

An organic acid called uric acid (UA) is end product of purine nucleotides metabolism, elevated serum uric acid (SUA) levels may have a pro-oxidative and pro-inflammatory effect that lead to metabolic syndrome including; polycystic ovarian syndrome, diabetic patients, obese people and adiposity (14,15). The pro-oxidant effect of uric acid has been observed to be of notable involve itself in the pathogenesis of the metabolic syndrome. Existing literature has reported a positive correlation between polycystic ovarian syndrome (PCOS) in women and elevated levels of oxidative stress (16,17). Furthermore, investigations revealed an association between total testosterone levels and serum uric acid levels among women with polycystic ovarian syndrome (PCOS), thus showed the association between hyperandrogenism and hyperuricemia (18).

PATIENTS AND METHODS:

Cross-sectional comparative study, non-probability convenience sampling conducted at Out-patient's department of Gynecology in Baghdad Medical City, from March 15th till September 15th 2024.

The target population of this study was patients of PCOS with or without insulin resistance. This study was take two groups 200 patients of PCOS participated depend on their impaired insulin response:

Group (A): In this group females were obtained due to their PCOS and insulin resistance.

Group (B): In this group females were obtained due to their PCOS and without insulin resistance.

Inclusion Criteria:

all females that are 20-40 years old diagnosed with PCOS using Rotterdam criteria (Oligo- and/or anovulation, Signs of hyperandrogenism, polycystic ovaries on ultrasonography (9)).

Exclusion Criteria: females of PCOS with or without insulin resistance and had diabetes

mellitus; hypertension; Cushing's syndrome; coronary heart disease; liver cirrhosis; renal diseases women who are pregnant, lactated, smoker, taken vitamin supplementation and medications such as (diuretic, hormonal, hypoglycemic).

The questionnaire encompassed two parts:

Part one: Cover socio-demographic of study participants by researcher-reported which included: Age, height, weight, Body Mass Index (BMI) is a medical screening tool that measures the ratio of height to weight to estimate the amount of body fat that classified as (underweight <18.8, normal weight 18.5-24.9, overweight 25-29.9, obesity ≥30) (19), marital state (single, married, divorced, widow), number of children, history of infertility.

Part two: Females of polycystic ovarian syndrome diagnosed with Rotterdam criteria (9):

1. Oligo- and/or anovulation.
2. Signs of hyperandrogenism; whether clinical or biochemical [serum testosterone and dehydroepiandrosterone sulfate (DHEAS)].
3. Polycystic ovaries on ultrasonography (the presence of eight or more subcapsular follicular cysts measuring less than 10 mm and an increase in ovarian stroma). (9)

Serum uric acid measurement by a health care professional will take a blood sample from a vein in arm, using a small needle (cut point defined for hyperuricemia >360 µmol/L [6.0 mg/dL]) in women (20).

Determine insulin resistance; by measures plasma insulin and glucose level that drawn after an overnight fast to calculate HOMA-IR (homeostatic model assessment-insulin resistant) which is the gold standard in assessment of insulin resistance: $HOMA1-IR = [Fasting Insulin (\mu U/ml) \times fasting glucose (mg/dl)] \div 405$

(HOMA1-IR level = 0.7- 2 normal reference range; and level >2 insulin resistance) (21).

RESULTS:

Figure 1: Distribution of study groups PCOS with insulin resistance was 65% (130 out of 200 sample size) and PCOS without insulin resistance was 35% (70 out of 200 sample size).

Table 1: Shows that Polycystic Ovarian Syndrome with insulin resistance was 61% highest among aged 20-30 years and there was significant association between Group A and age. Mean age of females was 29.15 years ±4.75 years, the mean age with Group A was 29.95 ± 4.57 years and mean age in Group B was 27.66 ± 4.75 years.

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Regarding BMI level, Polycystic Ovarian Syndrome with insulin resistance was highest among participants who were obese 71.5% and there was significant association found between BMI level and Group A. Moreover, Polycystic Ovarian Syndrome with insulin resistance was highest among patients who had positive family history 71.2% and there was statistical significant association between Group A and positive family history. Also, Polycystic Ovarian Syndrome with insulin resistance was found higher 67.1% among married and there was no significant association between Group A and marital state. Polycystic ovarian syndrome with

insulin resistance was found in 90% of patient who had no children and there was significant association between Group A and number of children. Finally, the Polycystic Ovarian Syndrome with insulin resistance was highest among patients who had history of infertility 73.8% and there was statistical significant association between Group A and history of infertility.

Table 2: shows that serum level of uric acid was highest among PCOS with insulin resistance 78.1% and there was significant association between serum level of uric acid and PCOS with insulin resistance.

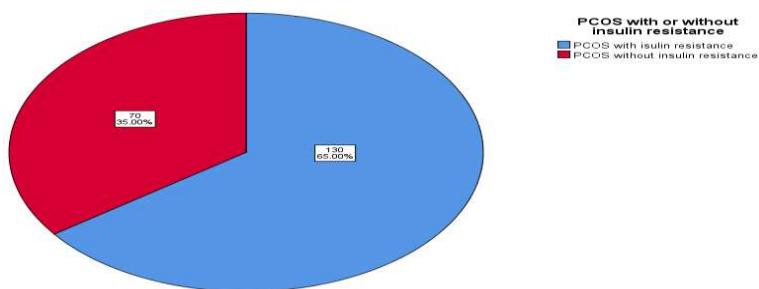


Figure 1: Distribution of the studied Groups

Table 1: Distribution of the studied Groups with Socio-demographic and health characteristics..

| Variables | Group PCOS with insulin resistance (n=130) | | Group without insulin resistance (n=70) | | Total (n=200) | | P-value |
|-------------------------------|--|------|---|------|---------------|------|---------|
| | No. | % | No. | % | No. | % | |
| Age(years) | | | | | | | |
| 20-30 | 69 | 56.5 | 53 | 43.4 | 122 | 61 | 0.002 |
| 31-40 | 61 | 78.2 | 17 | 21.7 | 78 | 39 | |
| BMI (kg/m²) | | | | | | | |
| normal weight (18.5-24.9) | 21 | 70 | 9 | 30 | 30 | 15 | 0.02 |
| overweight (25-29.9) | 31 | 50.8 | 30 | 49.2 | 61 | 30.5 | |
| obesity (>30) | 78 | 71.5 | 31 | 28.4 | 109 | 54.5 | |
| Family history | | | | | | | |
| Positive | 104 | 71.2 | 42 | 28.7 | 146 | 73 | 0.002 |
| negative | 26 | 48.1 | 28 | 51.8 | 54 | 27 | |
| Marital state | | | | | | | |
| single | 15 | 65.2 | 8 | 34.7 | 23 | 11.5 | 0.674 |
| married | 94 | 67.1 | 46 | 32.8 | 140 | 70 | |
| divorced | 16 | 55.1 | 13 | 44.8 | 29 | 14.5 | |
| widow | 5 | 62.5 | 3 | 37.5 | 8 | 4 | |
| Number of children* | | | | | | | |
| none | 36 | 90 | 4 | 10 | 40 | 22.5 | 0.01 |
| 1-3 | 60 | 60 | 40 | 40 | 100 | 56.4 | |
| >3 | 19 | 51.3 | 18 | 48.6 | 37 | 20.9 | |
| History of infertility | | | | | | | |
| Yes | 82 | 73.8 | 29 | 26.1 | 111 | 62.7 | 0.01 |
| No | 33 | 50 | 33 | 50 | 66 | 37.2 | |

*(Group PCOS with insulin resistance n=115, Group PCOS without insulin resistance n=62, Total n=177)

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Table 2: Distribution of the studied groups with serum level of uric acid.

| Serum level of uric acid | Group PCOS with insulin resistance (n=130) | | Group PCOS without insulin resistance (n=70) | | Total (n=200) | | P-value |
|--------------------------|--|------|--|------|---------------|----|---------|
| | No. | % | No. | % | No. | % | |
| increase | 50 | 78.1 | 14 | 21.8 | 64 | 32 | 0.008 |
| normal | 80 | 58.8 | 56 | 41.1 | 136 | 68 | |

Mean of serum level of uric acid is $5.67 \pm SD 1.12$

DISCUSSION:

The current study had shown that more than half 61% of polycystic ovarian syndrome aged 20-30 years, The mean age of cases was 29.15 years ± 4.75 years, there was significant association between polycystic ovarian syndrome and age and this was agreed with a study in Pakistan⁽²²⁾.

The prevalence of polycystic ovarian syndrome in this study was higher among obese participants which was similar to the results obtained by a study conducted in Iraq shown that body weight above normal BMI has positive relation with polycystic ovarian syndrome⁽²³⁾.

A positive Family history of polycystic ovarian syndrome and the occurrence of this disease was found higher among participants and this was correlated with the study done in Oman and United Arab Emirates⁽²⁴⁾.

The study was shown that the polycystic ovarian syndrome was found more among married than divorced, widow and less among single which was agreed with study done in Tehran, Iran that the marital state did not differ significantly by group with polycystic ovarian syndrome (PCOS)⁽²⁵⁾.

The number of children differ between polycystic ovarian syndrome with or without reduced insulin response and this is agreed with Western Australia that some women with polycystic ovarian syndrome are at a greater risk of obstetric problems may arise in pregnancy⁽²⁶⁾.

Infertility was found higher among PCOS and there was a significant association between prevalence of infertility and PCOS this concordance with a study done in Yemen that has shown Polycystic ovary syndrome (PCOS) is a significant cause of infertility due to ovulation dysfunction in women of childbearing age⁽²⁷⁾.

This study showed increase serum level of uric acid among patients of polycystic ovarian syndrome with insulin resistance and there is significant association which was agreed with study done in China showed that Polycystic ovarian syndrome is associated with a significant elevation of serum uric acid level and insulin resistance (IR) may be the mediators in the pathogenesis of hyperuricemia in women with

Polycystic ovarian syndrome⁽²⁸⁾, also concordance with study done by Sun R. and Shen J. that found the Serum uric acid is one of the characteristic factors of metabolic syndrome, the occurrence and development of hyperuricemia is closely associated with insulin resistance and hyperinsulinemia in patients with polycystic ovarian syndrome⁽²⁹⁾.

The serum uric acid may play a crucial role in polycystic ovarian syndrome and its related metabolic disorders, as they share common risk factors, elucidating the relationship between hyperuricemia and polycystic ovarian syndrome may offer valuable insights into pathogenesis mechanisms of polycystic ovarian syndrome⁽³⁰⁾. The study In India showed the incidence of hyperuricemia in polycystic ovarian syndrome (PCOS), The HOMA-IR was positively correlated with both hyper and non-hyper uricemic groups⁽³¹⁾, this was agreed with the current study.

CONCLUSION:

The study showed that more than two third of Group A had increase in serum level of uric acid and approximately less than third of Group B had increase serum level of uric acid and there was significant association found between polycystic ovarian syndrome with insulin resistance and serum level of uric acid. Dietary recommendations and lifestyle changes with exercise and weight reduction can help PCOS that reduce risk of hyperuricemia.

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