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# Study of the Association between Thyroid Hormonal Abnormalities and Polycystic Ovary Syndrome among Egyptian Females

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## Abstract

**Background:** Polycystic Ovary Syndrome (PCOS) is marked by hyperandrogenism, anovulation with menstrual irregularities, and morphologic polycystic ovarian alterations. The implicated hormonal abnormalities may extend to other endocrine glands.

**Aim of the Work:** To investigate whether thyroid hormone abnormalities and PCOS are related in Egyptian women.

**Patients and Methods:** There were 500 women in the cohort diagnosed with PCOS based on the 2003 Rotterdam Criteria. Clinical, menstrual, and reproductive factors and ovarian morphology were analyzed in relation to the underlying thyroid gland's clinical and hormonal status.

**Results:** The prevalence of amenorrhea, oligomenorrhea, and hirsutism/acne was found to be 16.2%, 54.8%, and 70.6%, respectively, with a mean Ferriman-Gallwey hirsutism score of  $8.04 \pm 4.101$ . Elevated Body Mass Index ( $30.40 \pm 11.98$  kg/m<sup>2</sup>) and waist circumference ( $91.77 \pm 9.93$  cm) suggested underlying insulin resistance. Gravidity, parity, and miscarriage analysis revealed 69.2% nulligravida, 78.2% nullipara, and 7.8% miscarriage among the studied patients. The thyroid gland examination indicated that 70.4% had a normal appearance, 10.4% had a diffuse goiter, 16.4% had a multi-nodular goiter, and 2.8% had a solitary thyroid nodule. Functional assessment revealed 69.6% euthyroid, 14.8% with subclinical hypothyroidism, 9.6% hypothyroid, 4.8% hyperthyroid, and 1.2% subclinical hyperthyroidism. Thyroid-stimulating hormone (TSH) levels correlated significantly with body weight, BMI, and miscarriage in PCOS patients ( $P$ -value  $<0.001$ ,  $<0.001$ , and  $0.012$ , respectively).

**Conclusion:** This study underscores the connection between thyroid abnormalities and reproductive parameters in PCOS.

**Keywords:** Thyroid Hormonal Abnormalities; Polycystic Ovary Syndrome; Egyptian Females

## 1. Introduction

Polycystic ovarian syndrome (PCOS) is a prevalent hormonal condition affecting females. Approximately 6-10% of women get Polycystic Ovary Syndrome (PCOS) during their reproductive years. One in ten women suffers signs of the condition during her reproductive years.<sup>1</sup>

Several studies have attempted to determine the prevalence of PCOS in Egypt. It is estimated that around 13% of females have given birth, and 37.5 percent of patients have primary or secondary infertility.<sup>2</sup>

Furthermore, there is mounting evidence indicating a correlation between PCOS and the heightened occurrence of thyroid disorders, including nodular goiter and autoimmune

thyroiditis.<sup>3</sup>

Thyroid problems and polycystic ovary syndrome (PCOS) are prevalent endocrine illnesses in the general population. Despite having distinct etiopathogenesis, hypothyroidism, and PCOS share numerous similar traits.<sup>4</sup> Primary hypothyroidism has been associated with an elevation in ovarian volume and the occurrence of cystic abnormalities in the ovaries. Conversely, there is a growing recognition that thyroid issues are more prevalent among women with PCOS than in the general population.<sup>5,6</sup>

The objective of the present study is to assess the correlation between thyroid hormonal irregularities and PCOS in Egyptian individuals with PCOS and to investigate its influence on the clinical results.

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## 2. Patients and methods

This study was conducted on 500 females diagnosed with PCOS who attended obstetrics, gynecology, and Endocrine outpatient clinics of Al-Azhar University Hospital from July 2022 to July 2023.

2.1. Inclusion Criteria: All females who were married  $\geq 2$  years with the diagnosis of PCOS based on Rotterdam Criteria 2003<sup>7</sup>, which defined the presence of two of the following three characteristics at least: oligo-ovulation, increased androgen levels (clinical and or laboratory), morphological features of polycystic ovary detected by ultrasound were valid for inclusion.

2.2. Exclusion criteria: Females with past known thyroidal illness before enrollment, females with past thyroid surgery, females on L-thyroxin replacement, Cushing's syndrome, and active malignancy

2.3. Methods: All enrolled patients were subjected to the following:

2.4. Full history taking: Full menstrual history, fertility history including gravidity, parity, and miscarriage, history suggestive for hyperandrogenism including hirsutism and acne, and history suggestive for thyroidal abnormalities including Symptoms of hypo/hyperthyroidism and goiter.

2.5. Complete clinical examination with special emphasis on Thyroid gland examination, clinical evaluation of thyroid status, clinical stigmata of insulin resistance, and clinical features of hyperandrogenism, particularly hirsutism and acne. The degree of hirsutism was evaluated according to the Ferriman and Gallwey score.

2.6. Thyroid hormone measurements: TSH, free T3, and free T4

A venipuncture-drawn 2 ml of venous blood was centrifuged for 20 minutes at 3000 rpm, yielding serum samples frozen at -20 degrees Celsius until needed. Following the manufacturer's instructions, TSH, FT3, and FT4 were quantitatively measured on a Cobas e411 using electrochemiluminescence immunoassay (ECLIA). According to the National Academy of Clinical Biochemistry's recommendations for laboratory diagnosis and thyroid disease monitoring, thyroid abnormalities were diagnosed and classified.<sup>8</sup>

2.7. Radiological evaluation of the ovaries by transvaginal ultrasonography. A single transvaginal sonographer used an UltraSonix RP ultrasound scanner with a 9-MHz transvaginal transducer to do an ultrasound scan at random times during the menstrual periods.

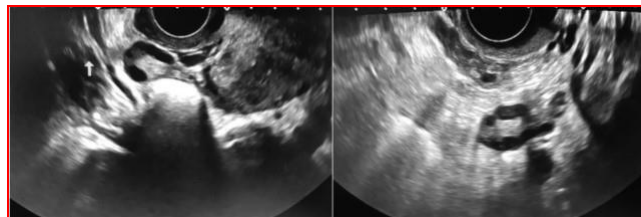


Figure 1. Trans vaginal Ultrasound showing PCO.

The sonographic criteria of PCOS were included according to Rotterdam Criteria.<sup>7</sup> The evaluation involved considering all the follicles in each ovary, independent of their arrangement, from the inner edge to the outside edge. Additionally, sections on different planes were examined to ensure comprehensive research. The follicular diameter refers to the average diameter measured across three sections or the follicle's diameter in the scan when it appears round. We employed the ellipsoid formula  $0.52 \times (D1 \times D2 \times D3)$  to compute the volume. A criterion of ovarian volume above 10mm<sup>3</sup> was established.

2.8. Statistical analysis: The analysis was conducted using the software package STATA (version 12; STATA Inc., College Station, TX, USA). The significance threshold was  $p < 0.05$ , and the confidence interval was 95%.

## 3. Results

Table 1. Demographic and anthropometric characteristics of the patients.

| VARIABLES                | MEAN  | ± SD  |
|--------------------------|-------|-------|
| AGE (YEARS)              | 27.50 | 1.86  |
| WAIST CIRCUMFERENCE (CM) | 91.77 | 9.93  |
| HEIGHT (CM)              | 164.2 | 7.00  |
| WEIGHT (KG)              | 81.04 | 11.91 |
| BMI (KG/M <sup>2</sup> ) | 30.40 | 11.98 |

The age distribution of the selected patients spanned from 22 to 35 years, with a mean  $\pm$  SD of  $27.5 \pm 1.86$ . Their waist circumference ranged from 76 to 118 cm, with a mean  $\pm$  SD of  $91.77 \pm 9.93$ . Heights ranged from 150 to 179 cm, with a mean  $\pm$  SD of  $164.2 \pm 7.00$ . Their weight varied from 55.1 to 118.9 kg, with a mean  $\pm$  SD of  $81.04 \pm 11.91$ . Consequently, their body mass index (BMI) covered a range from 22.8 to 39.5 kg/m<sup>2</sup>, with a mean  $\pm$  SD of  $30.40 \pm 11.98$ , Table 1.

Table 2. Gravidity, Parity, and Miscarriage frequencies among the studied patients.

|             |              | NUMBER | PERCENTAGE |
|-------------|--------------|--------|------------|
| GRAVIDITY   | Nulligravida | 346    | 69.2%      |
|             | Gravida-1    | 117    | 23.4%      |
|             | Multigravida | 37     | 7.4%       |
| PARITY      | Nullipara    | 391    | 78.2%      |
|             | Para-1       | 98     | 19.6%      |
|             | Multipara    | 11     | 2.2%       |
| MISCARRIAGE | Absent       | 461    | 92.2       |
|             | present      | 39     | 7.8        |

Among the studied patients, 346 (69.2%) were nulligravida, while 117 (23.4%) were gravida-1, and the remaining 37 (7.4%) were multigravida. 391 (78.2%) of them were nullipara, 98 (19.6%) were para-1, and the remaining 11 (2.2%) were multipara. A 7.8% experienced miscarriage, [Table 2](#)

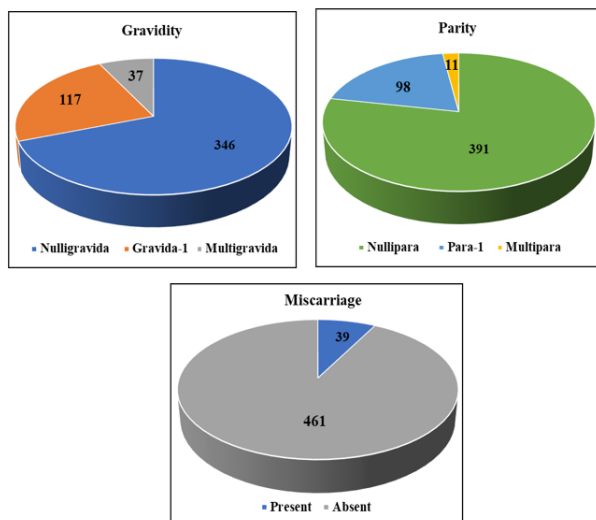


Figure 2. The frequencies of the Gravidity, Parity, and miscarriage among the studied patients.

Table 3. Clinical characteristics of the patients in the study group.

|                                   | NUMBER  | PERCENTAGE |      |       |
|-----------------------------------|---------|------------|------|-------|
| <b>MENSTRUAL ABNORMALITIES</b>    |         |            |      |       |
| NORMAL                            | 145     | 29.0%      |      |       |
| AMENORRHEA                        | 81      | 16.2%      |      |       |
| OLIGOMENORRHEA                    | 274     | 54.8%      |      |       |
| <b>HYPERANDROGENIC SYMPTOM</b>    |         |            |      |       |
| ASYMPTOMATIC                      | 147     | 29.4%      |      |       |
| HIRSUTISM & ACNE                  | 353     | 70.6%      |      |       |
| <b>FERRIMAN AND GALLWEY SCORE</b> |         |            |      |       |
|                                   | Minimum | Maximum    | Mean | SD    |
|                                   | 1       | 14         | 8.04 | 4.101 |

In terms of Menstrual cycle abnormalities, 145 (29%) had normal menstrual cycles, 81 (16.2%) experienced amenorrhea, and 274 (54.8%) had oligomenorrhea. Regarding Hyperandrogenic symptoms, 147 (29.4%) were asymptomatic, while 353 (70.6%) exhibited hirsutism and acne. The results of Ferriman and Gallwey scores ranged from 1 to 14, with a mean  $\pm$  SD of  $8.04 \pm 4.101$ , [Table 3](#).

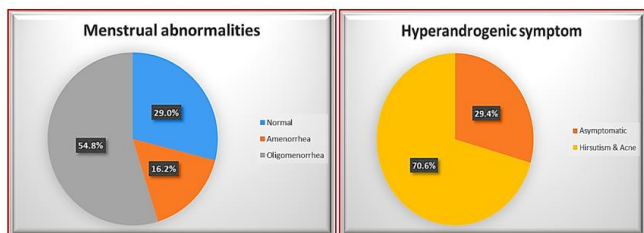


Figure 3. distribution of menstrual status and HA symptoms.

Table 4. Evaluation of number of follicles in both ovaries and ovarian volumes.

| VARIABLES                               | MINIMUM | MAXIMUM | MEDIAN | IQR         |
|---|---------|---------|--------|-------------|
| LEFT OVARY FOLLICLE NUMBERS             | 12      | 21      | 16.64  | 15 - 18     |
| LEFT OVARIAN VOLUME (ML <sup>3</sup> )  | 9.7     | 16.2    | 13.5   | 12.2 - 14.8 |
| RIGHT OVARY FOLLICLE NUMBERS            | 14      | 25      | 20.56  | 19 - 22     |
| RIGHT OVARIAN VOLUME (ML <sup>3</sup> ) | 10.3    | 16.7    | 14.1   | 12.9 - 15.5 |
| BOTH OVARIES FOLLICLE NUMBERS           | 13      | 23      | 18.58  | 17 - 20     |
| BOTH OVARIAN VOLUME (ML <sup>3</sup> )  | 10.1    | 17.4    | 13.8   | 12.5 - 15.1 |

The average number of follicles of the left ovaries was 16.64 with IQR of 15-18. Similarly, the right ovaries harbored an average of 20.56 follicles with IQR of 19-22 moreover, the average number of follicles in both ovaries was 18.58 with IQR of 17-20. Regarding the ovarian volume, the average left ovarian volume was 13.5 ml<sup>3</sup> with IQR of 12.2-14.8 Similarly, the average right ovarian volume 14.1 ml<sup>3</sup> follicles with IQR of 12.9-15.5 moreover, the average volume of both ovaries was 13.8 ml<sup>3</sup> with IQR of 12.5-15.1, [Table 4](#).

Table 5. Prevalence of Thyroid Disorders: Clinical Examination and Functional Status.

| THYROID GLAND CLINICAL EXAMINATION |        |            |
|------------------------------------|--------|------------|
|                                    | Number | Percentage |
| NORMAL THYROID GLAND               | 352    | 70.4%      |
| DIFFUSE GOITER                     | 52     | 10.4%      |
| MULTI NODULAR GOITER               | 82     | 16.4%      |
| SOLITARY THYROID NODULE            | 14     | 2.8%       |
| THYROID GLAND FUNCTIONAL STATUS    |        |            |
| EUTHYROID                          | 348    | 69.6%      |
| HYPERTHYROIDISM                    | 24     | 4.8%       |
| HYPOTHYROIDISM                     | 48     | 9.6%       |
| SUBCLINICAL HYPERTHYROIDISM        | 6      | 1.2%       |
| SUBCLINICAL HYPOTHYROIDISM         | 74     | 14.8%      |

Regarding thyroid gland clinical examination, 70.4% (n=352) had a normal thyroid gland examination, 10.4% (n=52) had a diffuse goiter, 16.4% (n=82) had a multi-nodular goiter, and 2.8% (n=14) had a solitary thyroid nodule. Regarding Thyroid gland functional status, 69.6% (n = 348) were euthyroid, 4.8% (n=24) were hyperthyroid, 9.6% (n=48) were hypothyroid, 1.2% (n=6) had subclinical hyperthyroidism, and 14.8% (n=74)

had subclinical hypothyroidism, [Table 5](#).

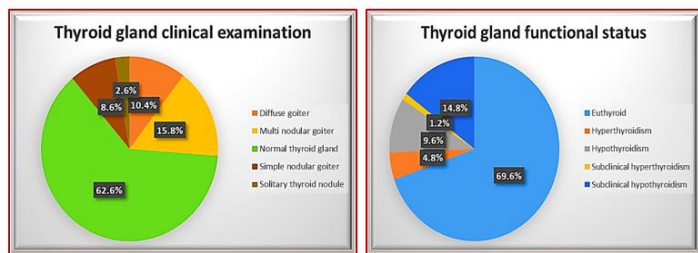


Figure 4. Prevalence of Thyroid Disorders: Clinical Examination and Functional Status.

Table 6. Thyroid hormonal profile levels of the studied patients.

| HORMONES        | MINIMUM | MAXIMUM | MEDIAN | IQR         |
|-----------------|---------|---------|--------|-------------|
| TSH (UIU/ML)    | 0.01    | 16.40   | 3.40   | 2.52 - 4.90 |
| FREE T3 (PG/ML) | 1.2     | 6.7     | 2.80   | 2.50 - 3.0  |
| FREE T4 (NG/DL) | 0.2     | 2.9     | 1.30   | 1.0 - 1.47  |

The TSH levels ranged from 0.01 to 16.4 ( $\mu$ IU/ml) with Median value 3.4 and IQR (2.52-4.9). The Free T4 levels ranged from 0.2 to 2.9 (ng/dl) with a Median value of 1.30 and IQR (1.0-1.47). The Free T3 levels ranged from 1.2 to 6.7 (pg/dl) with Median value 2.8 and IQR (2.5-3), [Table 6](#).

Table 7. Correlation between TSH level and variables among studied patients.

| VARIABLES                                   | CORRELATION COEFFICIENTS | P-VALUE |
|---|--------------------------|---------|
| AGE   | 0.055                    | 0.216   |
| WEIGHT (KG)                                 | 0.254                    | <0.001  |
| BMI (KG/M2)                                 | 0.222                    | <0.001  |
| FERRIMAN AND GALLWEY SCORES                 | 0.087                    | 0.052   |
| AVERAGE NUMBER OF FOLLICLES IN BOTH OVARIES | -0.034                   | 0.442   |
| AVERAGE OVARIAN VOLUME (ML <sup>3</sup> )   | -0.007                   | 0.875   |
| NULLIGRAVIDITY                              | 0.066                    | 0.142   |
| NULLIPARITY                                 | 0.073                    | 0.102   |
| MISCARRIAGE                                 | 0.112                    | 0.012   |
| MENSTRUAL ABNORMALITIES                     | 0.073                    | 0.105   |
| HYPERANDROGENISM SYMPTOM                    | 0.070                    | 0.119   |

There is a strong positive relationship between TSH levels and Body The weight, BMI, and the likelihood of Miscarriage, with statistically significant P-values of <0.001, <0.001, and 0.012, respectively. However, there was no statistically significant link found between TSH levels and the other covariates, [Table 7](#).

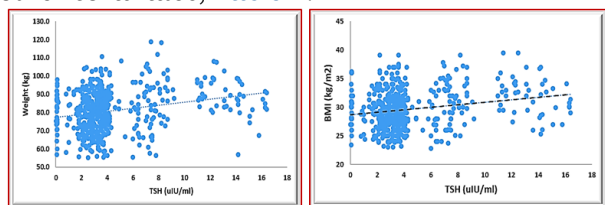


Figure 5. Correlation between TSH levels and the patient's weight & BMI.

#### 4. Discussion

The relationship between PCOS and other endocrine disorders, particularly thyroid abnormalities, has been a longstanding area of interest. Our study represents the first Egyptian trial to illuminate this intricate relationship by including a substantial cohort of 500 PCOS patients.

Among the 500 studied patients, oligomenorrhea was prevalent in more than half of the cases (54.8%). This finding was aligned with previous studies suggesting that a high percentage of females with PCOS experience oligomenorrhea. For example, Salih et al.<sup>9</sup> found that 72% of PCOS patients had oligomenorrhea.

Similarly, Jesmin et al.<sup>10</sup> Additionally, these results were corroborated by a study that revealed that oligomenorrhea was present in 64.4% to 71.8% of patients diagnosed with PCOS. Furthermore, it was discovered that oligomenorrhea manifested in 75-85% of women diagnosed with Polycystic Ovary Syndrome (PCOS).

Concerning fertility negative results among the patients under investigation, 69.2% had never been pregnant, and 78.2% had never given birth. In addition, a total of 7.8% of the patients suffered from miscarriage.

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The previous research lacked definitive data on the proportion of females diagnosed with PCOS who experienced pregnancy or childbirth. However, studies have demonstrated a correlation between PCOS and an increase in subfertility, ectopic pregnancy, and early pregnancy loss.<sup>11</sup>

Furthermore, an additional study revealed that there was no notable disparity in the number of live births (parity) between women diagnosed with PCOS (1.9  $\pm$  1.3 children) and the control group (1.7  $\pm$  1.0 children).<sup>12</sup> A recent study found that PCOS is linked to 75% of cases of ovulatory infertility, and the infertility rate in PCOS ranges from 2% to 5%.<sup>13</sup>

Additionally, Acharya et al.<sup>14</sup> highlighted that PCOS is a common cause of anovulatory fertility.

Furthermore, the present findings indicate a significant incidence of Hirsutism and acne, with a prevalence rate of 70.6%. The incidence of hirsutism in PCOS varies between 65% and 75%, whereas the occurrence of acne in adult females with PCOS is approximately 76% according to the NIH definition and 36% according to the Rotterdam criteria.<sup>15</sup>

The Ferriman and Gallwey mean score in our series was 8.04. Previously, Gurbüz & Alanya Tosun,<sup>16</sup> reported a mean Score of 7.4 in patients with PCOS, which is aligned with the current study. Alternatively, Espinós et al.<sup>17</sup> found that the mean score using the modified Ferriman-Gallwey method was 15.1 in women with PCOS.

Furthermore, the current study examined the morphological characteristics of ovaries, including the average number of follicles (ranged from 13 to 23 (Median= 18.58, IQR= (17-20)) and volume of both ovaries (ranged from 10.1 to 17.4 ml<sup>3</sup> (Median= 13.8, IQR= (12.5-15.1))).

Previous investigations have corroborated the findings of our research regarding the ovarian volume in patients with polycystic ovary syndrome (PCOS). These investigations have found that the average volume in persons with PCOS is between 10.6 and 16.7 ml, while healthy women usually have volumes ranging from 5.2 to 8.7 ml.<sup>18</sup> Top of Form

Regarding thyroidal hormonal disturbances and clinical examination of the thyroid gland among the selected patients, although most of our cases demonstrated euthyroid status and normal thyroid gland clinical examination, a subset of participants exhibited variations. Specifically, 14.8% had subclinical hypothyroidism, emerging as the most common thyroid dysfunction.

This finding emphasizes the importance of comprehensive evaluation for potential endocrine disparities, especially thyroidal abnormalities, and highlights the diversity in thyroidal health within the studied cohort.

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Recent research has revealed the complex connection between PCOS and thyroid function, specifically focusing on the involvement of TSH. For example, Lee et al.<sup>19</sup> revealed elevated TSH levels, confirming SCH diagnosis in the PCOS group compared to the control group.

A Chinese metanalysis by Ding et al.<sup>20</sup> A study comparing 692 people diagnosed with Polycystic Ovary Syndrome (PCOS) to 540 patients without PCOS revealed that women with PCOS have a greater risk of developing Subclinical Hypothyroidism (SCH). Likewise, research conducted in Pakistan compared cases and controls by Raj et al.<sup>21</sup> was Conducted on a sample of two hundred patients diagnosed with PCOS compared to another sample of two hundred patients without PCOS. Their data indicates a robust correlation between subclinical hypothyroidism (SCH) in women with polycystic ovary syndrome (PCOS) compared to women without any health issues.

On the contrary, Abdollahi et al.<sup>22</sup> Discovered that all PCOS patients had normal levels of TSH, T3, and T4.

In our study, TSH demonstrated a significant positive correlation with case weights, BMI, and occurrence of miscarriage. In contrast, no statistically significant differences were found between TSH and other variables, including age, the Ferriman and Gallwey scores, follicle numbers, ovarian volume, nulla-gravidity, and

nulliparity.

Rojhani et al.<sup>23</sup> A recent study proposed a potential link between TSH levels and weight gain as well as excessive body mass in people with PCOS. However, Fatima et al.<sup>24</sup> o correlation was observed between obesity and subclinical hypothyroidism in patients with polycystic ovary syndrome (PCOS).

The diverse findings underscore the complexity of the relationship between thyroid dysfunctions and various phenotypes in PCOS.

#### 4. Conclusion

Monitoring women with PCOS for thyroid problems is pertinent and indispensable, considering the substantial evidence indicating an elevated susceptibility to thyroid dysfunction in this demographic.

#### Disclosure

The authors have no financial interest to declare in relation to the content of this article.

#### Authorship

All authors have a substantial contribution to the article

#### Funding

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#### Conflicts of interest

There are no conflicts of interest.

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