Evaluation of the relationship between vitamin D3 level and sex hormones in women with PCOS

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Abstract---The current study included collection of 90 serum samples to assess the relationship between the level of vitamin D3 and some biochemical variables in women with polycystic ovarian syndrome. The samples were collected from external laboratories in Salah al-Din Governorate for the period from 10/29/2021 to 12/25/2021. 60 samples were taken for women with PCOS, and 30 samples were taken from the control group, their ages ranged between (18-40) years for healthy women and patients. Vitamin D3 level and some hormones concentrations (luteinizing hormone, follicle-stimulating hormone, and testosterone) were estimated in the blood serum of the study groups. The results of the current study showed that the level of vitamin D3 decreased significantly (P≤0.05) in women with PCOS compared to healthy women as a control group, the results showed a significant increase (P≤0.05) in the level of studding hormones (luteinizing hormone, follicle-stimulating hormone, and testosterone) in women with PCOS compared to healthy women as a control group.

Keywords---PCOS, vitamin D3, luteinizing hormone, follicle-stimulating hormone, testosterone.

Introduction

PCOS is one of the most common endocrine and metabolic disorders in premenopausal women. PCOS is defined as heterogeneous disorder, with a combination of signs and symptoms of excess androgen and ovarian dysfunction in the absence of other specific diagnoses. The etiology of this syndrome is still largely unknown, but mounting evidence indicates that suggest that PCOS may be a complex polygenic disorder with strong genetic and environmental
influences, including diet and lifestyle factors. PCOS is often associated with abdominal fat, insulin resistance, obesity, metabolic disorders, and cardiovascular risk factors \(^{(1)}\). More than 1 in 10 women worldwide have PCOS, which is the leading cause of female reproductive and metabolic dysfunction \(^{(2)}\). PCOS is believed to be the most common endocrine disorder in women. Common symptoms include irregular menstruation, PCOS, and hirsutism, as well as an increased risk of several conditions, including insulin resistance, dyslipidemia, and infertility \(^{(3)}\).

Vitamin D3 is a fat-soluble vitamin \(^{(4)}\). There are two forms of it, the first form D2 and it is called ergocalciferol. It can be obtained from the raw material ergosterol, a substance found in plants, and it does not have a great nutritional value because it cannot be absorbed through the intestine \(^{(5)}\). The second form is D3 cholecalciferol, which can be obtained from 7-dehydrocholesterol, which is found in the skin when exposed to sunlight, and can also be obtained from animal foods \(^{(6)}\). Vitamin D3 is also called the sunshine vitamin because it can be synthesized inside the body from cholesterol when the body is exposed to sunlight \(^{(7)}\). There are many sources of vitamin D, some natural and some synthetic that can be obtained naturally from food \(^{(8)}\). It was found that exposure to sunlight is one of the main natural sources for obtaining the vitamin \(^{(9)}\). While foods rich in vitamin D2 and D3 such as milk, yogurt and vegetable ghee are important industrial sources for obtaining the vitamin \(^{(10)}\).

The ovaries in women with PCOS do not respond to the pituitary hormones appropriately as in the natural ovaries. Both LH and FSH hormones affect the ovaries and their hormones directly, so FSH is the main hormone in the follicular phase of the menstrual cycle if it works to stimulate the formation of the maturation of the eggs and thus the secretion of estrogen from the mature egg \(^{(11)}\).

While LH is the main hormone in the luteal phase that stimulates ovulation and then the formation of estrogen and progesterone in the luteal phase after ovulation, and any imbalance in the levels of FSH, LH is reflected in the level of the hormones progesterone and estrogen as well as on the production of the male hormone and thus causes disturbances in the reproduction function of the ovaries, such as the difference in the menstrual cycle, delayed childbearing, or infertility \(^{(12)}\).

**Materials and Methods**

**The samples**
The blood samples were (90) which distributed to tow groups, 60 samples were from women with PCOS and 30 blood samples from healthy women as a control group, their ages ranged between (18-40) these samples were collected from external laboratories in Salah El-Din Governorate for the period between 10/29/2021 to 12/2021.
Estimation the level of Vitamin D3

Vitamin D3 was calculated by following the steps attached to the vitamin analysis kit, and according to the instructions of the kit, to estimating the concentration of vitamin D3.

Estimation the concentration of sex hormones

The study included estimation of the levels of sex hormones (luteinizing hormone, follicle-stimulating hormone, and testosterone) by means of an examination kit supplied by the American company Mybiosource by means of the Sandwich enzyme-linked immunosorbent assay (ELISA) technique.

Results and Discussion

The Level of Vitamin D3

The results showed that the mean ± standard deviation of vitamin D level was (17.063 ± 3.912) ng/ml for women with PCOS, compared to (35.399 ± 9.828) ng/ml for healthy women as a control group, as shown in Table (1).

Table (1) the level of vitamin D3 in the blood serum of POC female patients and control group.

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Mean ± SD</th>
</tr>
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<tbody>
<tr>
<td>Vitamin D ng/ml</td>
<td></td>
</tr>
<tr>
<td>Control</td>
<td>35.399±9.828</td>
</tr>
<tr>
<td>Patients</td>
<td>17.063±3.912</td>
</tr>
<tr>
<td>P≤0.05</td>
<td>*</td>
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</tbody>
</table>

Table (1) shows that the level of vitamin D showed a significant decrease (P≤0.05) in the blood serum of women with PCOS compared to the control, as shown in Figure (1).
Polycystic ovary syndrome (PCOS) is the most common endocrine disorder in women of childbearing age with an incidence of 10%, many studies have shown that vitamin D3 has a positive effect on female reproductive diseases, as well as PCOS, androgen excess and neuroendocrine causes and insulin resistance is all part of the pathophysiology, Some female reproductive organs have been found to contain vitamin D3 receptors, so vitamin D3 is essential in the human body (13). The results of the current study agree with the results of the study of Ibrahim et al., (14), which showed a significant decrease in the level of vitamin D3 in women with PCOS compared to healthy women as a control group. The results of the current study are also in agreement with the results of the study of Li and his group (15), which showed a decrease in the level of vitamin D3 among women with PCOS compared with healthy women as a control group.

The reason for the decrease in the level of vitamin D3 in the current study may be attributed to malnutrition and dependence on fast foods, as well as the poor table for seafood and the lack of interest in health institutions to educate people about the importance of eating foods fortified with vitamin, and the importance of dietary diversity and interest in eating foods rich in vitamin such as egg yolk, beef meat and liver, Milk and cheese, and since the vitamin is one of the fat-soluble vitamins, any factor that affects the absorption of fats in the intestines can affect the absorption of the vitamin, in addition to the way of life in Iraqi society, as the lack of exposure to sunlight within the required peak periods.

**The levels of sex hormones**

The level of sex hormones, which includes follicle-stimulating hormone, luteinizing hormone and testosterone, was measured in the blood serum of women with polycystic ovary syndrome (PCOS) and healthy women as a control group, and the mean standard deviation of hormones was shown in Table (2).
Table (2) shows the level of FSH, LH, and testosterone in the blood serum of the study groups

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Mean ± SD</th>
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<tbody>
<tr>
<td></td>
<td>FSH (mlU/ml)</td>
</tr>
<tr>
<td>Control</td>
<td>17.284±7.972</td>
</tr>
<tr>
<td>Patients</td>
<td>22.564±6.926</td>
</tr>
<tr>
<td>P≤0.05</td>
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</tbody>
</table>

*High significant

The level of follicle-stimulating hormone

The results of the current study showed that the level of FSH hormone was (22.564 ± 6.926) mIU/cm³ in the blood serum of women with PCOS, while its level was (17.284 ± 7.972) mIU/cm³ in healthy subjects (control group), as shown in the table (2). It was obtained from the above table that the level of FSH was significantly increased (P≤0.05) in women with PCOS compared to healthy subjects as a control group, as shown in Figure (2).

**Figure (2) shows the level of FSH in PCOS groups compared to the control group**

The results of the current study do not agree with ElSirgany et al., (18) who indicated that the FSH level in the group of women with PCOS was lower than in the control group. Also, the results of the current study did not agree with the results of the study of Al-Tikriti et al., (19), which they showed no significant differences in the level of FSH in women with PCOS compared with healthy women. The results of the current study agree with the results of the study by Sak et al., (20), which they found high levels of FSH in women with PCOS compared to healthy women as a control group.
The growth of ovarian follicles is under the control of both hormones FSH and LH, and the response of the ovaries is when the level of FSH reaches the required level, and the level of FSH is variable during the follicular phase, due to the growth of the follicle and sensitivity to the stimulating of the gonadotropins (21). Thus, high levels of sex hormones lead to hormonal disruption and thus the lack of ovulation. The pituitary gland and the hypothalamus are among the main organs that regulate the mechanism of action of the endocrine system (22). In the anterior lobe of the pituitary gland, the portal system works to produce peptides that bind with specific receptors located on the surfaces of cells, thus either inhibiting hormones or releasing hormones (23).

The hypothalamus also stimulates the production of gonadotropins from the pituitary gland through the pulsating production of gonadotropin-releasing hormones, and this leads to the stimulation of genetic transcription of gonadotropins LH and FSH (24).

**Level of serum LH**

The results of the current study showed that the level of LH hormone was (24.594 ± 3.820) mIU/cm³ in the blood serum of women with PCOS, while its level was (20.944 ± 5.899) mIU/cm³ in healthy subjects as a control group as shown in the table (2). It is clear from the table (2) that the level of LH was significantly increased, Ps0.05, in women with PCOS compared to healthy women as a control group, as shown in Figure (3).

![Figure (2) shows the level of LH in PCOS groups compared to the control group](image)

The results of the current study agree with many studies (20, 26, 25) that showed a significant increase in the level of LH in patients with PCOS. The reason for the high level of the hormone may be due to the increased sensitivity of the pituitary gland to LH-GnR, and this leads to an increase in the concentration of -Gonadotropin-releasing hormones GnRH or to changes in its secretion patterns.
So, LH stimulates ovulation and then stimulates the corpus luteum to form steroid hormones \(^{(28)}\), but high levels of LH suppress the activity of aromatase and inhibit oocyte growth \(^{(29)}\). Thus, the high levels of sex hormones lead to hormonal disruption and thus the lack of ovulation.

**Level of serum Testosterone**

The results of the current study showed that the level of Testosterone hormone was \((0.579 \pm 0.221)\) mIU/cm\(^3\) in the blood serum of women with PCOS, while its level was \((0.368 \pm 0.106)\) mIU/cm\(^3\) of healthy women (control group) as shown in Table (2). The results which obtained from the study indicated that the level of Testosterone was significantly increased \((P \leq 0.05)\) among women with PCOS compared to the healthy control group as shown in Figure (4) and table (2).

![Testosterone levels in PCOS and control groups](image)

Figure (4) shows the level of testosterone in PCOS groups compared to the control group.

The results of the current study agree with the results of several last studies \(^{(20, 25, 30)}\) that showed a significant increase in the level of testosterone in women with PCOS compared to healthy women as a control group. Androgens are usually derived from cholesterol in the ovaries and adrenal glands, and can be produced in muscle, liver, skin, and adipose tissue through the conversion of certain types of steroids \(^{(31)}\).

In muscle and adipose tissue, a chemical group is added to androgens to turn into estrogens, thus converting testosterone and androstenedion into estrogens, and this agrees with the study of Frhan et al., \(^{(32)}\) which showed that the concentration of testosterone hormone of women with PCOS is higher than the control group and that the increase in the concentration of LH hormone results in polycystic ovary syndrome, which causes high testosterone and other hormone disorder.
References


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