Study the Comparison of Gonadotropin Levels in Diabetes Mellitus Females.

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Abstract
A cross sectional study was designed to assess the correlation of LH, FSH and HbA1c levels in patients with and without DM in premenopausal women, and the correlation of age with LH and FSH hormones in diabetic women, using the data that collected. It was conducted in the specialized centre for Endocrinology and Diabetes (SCED) in Baghdad city from December 2013 to September 2014. Hundred women with type II diabetes (30-49) and hundred healthy women in the same age and BMI as control during the luteal phase of menstrual cycle were enrolled. Patients assessed for HbA1C, FSH and LH levels. Results showed that HbA1c, LH and FSH were higher in diabetic premenopausal women compared with control individuals.

Keywords: Diabetes mellitus, Gonadotropin hormones, Glycated hemoglobin.

Introduction
Diabetes mellitus is an endocrine disorder in which human body suffers a lack of insulin or resistance to insulin. Currently, 150 million people in the world suffer from diabetes. Diabetes causes various complications and usually its chronic complications develop 5 to 10 years after the diagnosis of both types of the diabetes [1]. Type II is characterized by decreased sensitivity to the effect of insulin, in other word relative lack of insulin. Worldwide 246 million people are affected by DM [2]. Among them One hundred million persons have a history of diabetes (DM) worldwide [3]. Complications of diabetes are uniquely and often more severely, affect women [4].

Prevention of diabetes include Lifestyle modifications, dietary modification, regular physical activity and weight reduction are indicated [5,6]. Long-term glycemic control routinely uses Glycated hemoglobin (HbA1c) as a marker. In accordance with its function as an indicator for the mean blood glucose level, in diabetes patients HbA1c predicts the risk for the development of diabetic complications [7]. According to American Diabetes Association (ADA) criteria HbA1C value of (≥ 6.5%) or FPG≥126 mg/dl (7.0 mmol/l) or 2-hour plasma glucose≥200mg/dl(11.1mmol/l) or in a patient with classic symptoms of hyperglycemia or hyperglycemic crisis a random plasma glucose ≥200 mg/dl (11.1mmol/l), is strongly predicted as pharmacotherapy for diabetes[8].

The gonadotrophin hormones LH and FSH from the interior pituitary gland controlled the growth and the reproductive activities of the gonadal tissue. LH and FSH are called gonadotrophins because they regulate the function of the gonads (ovaries and testes), in both (male, female, FSH stimulates gametes (sperm or egg) production, while LH promotes production of gonadal hormones [9]. Gonadotrophin releasing hormone from the hypothalamus stimulates the secretion of both LH and FSH, which are subjected to feed back loops regulation by the ovarian hormones [10]. Part of the workup of infertility in women, LH along with FSH are ordered, also its useful in the investigation of menstrual irregularities, and to aid in the diagnosis delayed and precocious puberty[11].

Patients and Methods
A total of 100 type II diabetic premenopausal women (30-49) year and 100 healthy women in the same age and BMI visiting the specialized center for Endocrinology and Diabetes (SCED) from December 2013 to September 2014 during luteal phase of menstrual cycle were included in this study. Venous blood samples were collected from all the subjects. The serum was later used for analyzing HbA1C, LH and FSH. HbA1c was measured using immunoturbidimetric test according DCCT/NSGP protocol by Biolzyer (Turbitex/Germany Kit), LH and FSH using enzyme linked fluorescent assay technique by vidas (Biomerieux /FranceKit). Diabetes was defined according to American Diabetes Association (ADA) criteria. We excluded from
this study pregnant women, menopausal women, lactating, Patients on hormonal therapy, single women, Infertile women, poly cystic ovarian syndrome, smokers, Patients with endocrine disease e.g. Cushing's syndrome and family history of diabetes.

**Statistical Analysis**
Minitab (version 16) was used to analyse the collected data. These data analysed using the following measure:
1. Descriptive statistics: frequency, percentages, mean and standard deviation.
2. Inferential statistics: student t-test was used to define the difference between the means of two groups in quantitative variables. P value less than 0.05 considered statistically significant.

**Results and Discussion**
From hundred diabetic premenopausal women and hundred healthy premenopausal women, the result in Table (1) showed that the Mean± SD value of age was 38.31±4.08 in diabetic women and 37.45±4.97 in healthy women with p value of ≥ 0.183 (not significant), the level of BMI in diabetic and control was 27.42±2.20, 27.18±1.38 respectively and p ≥ 0.362 (not significant), while the Mean± SD of HbA1c was 9.42±2.33 in diabetic women and 4.657±0.447 in control with p≤0.05 (significant), the mean levels of LH and FSH in diabetic women was 5.37±0.87 and 5.66±1.25 respectively compared the levels in the control 0.576±0.092 and 1.855±0.353 respectively and difference is significant due to the p value of ≤0.05, this comparison shows that there is a significant increase in the level of these hormones in diabetic patients.

**Table (1)**
**Biochemical parameters for the diabetic and control.**

<table>
<thead>
<tr>
<th>Parameters</th>
<th>DIABETIC No=100</th>
<th>CONTROL No=100</th>
<th>P value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (years)</td>
<td>38.31±4.08</td>
<td>37.45±4.97</td>
<td>0.183</td>
</tr>
<tr>
<td>BMI(Kg/m²)</td>
<td>27.42±2.20</td>
<td>27.18±1.38</td>
<td>0.362</td>
</tr>
<tr>
<td>LH(mIU/ml)</td>
<td>5.37±0.87</td>
<td>0.576±0.092</td>
<td>0.000</td>
</tr>
<tr>
<td>FSH(mIU/ml)</td>
<td>5.66±1.25</td>
<td>1.855±0.353</td>
<td>0.000</td>
</tr>
<tr>
<td>HbA1C (%)</td>
<td>9.42±2.33</td>
<td>4.657±0.447</td>
<td>0.000</td>
</tr>
</tbody>
</table>

*P value is significant ≤ 0.05 level.*

The biochemical findings of this study showed a significant elevation of HbA1c in diabetic female compared to control Fig.(1) and this is similar to the findings in a group of patients studied by other researchers [12-16]. When the groups were investigated, the mean HbA1C were higher in diabetic premenopausal women.

Also it was found Fig.(1) a significant increase of LH and FSH in diabetic premenopausal women compared to control. When corpus leutum starts lysing during the last period of leuteal phase (in cases no pregnancy) the progesterone and estrogen levels will decrease while LH, FSH levels increase regularly, also the decrease of progesterone and estrogen may cause a positive feedback mechanism at the pituitary gland level [17].

And result of this, an increment of gonadotropin (LH, FSH) will be occurring. On the other hand, LH hormone is considered as a principle factor in stimulating theca interna, causing an elevating production of pregnenolone compound which is converted by granulose cells to progesterone during leuteal phase [18].

The results of the present study are consistent with other results [19, 20].

Which showed that higher (LH, FSH) levels in diabetic premenopausal women compared with the control.
Fig.(1) Comparison of mean value of gonadotropins and HbA1C in diabetic and control group.

Table (2)
Variation of LH, FSH and HbA1C with Age.

<table>
<thead>
<tr>
<th>Parameters</th>
<th>Group I Diabetic women Age(30-38) years No=(48)</th>
<th>Group II Diabetic women Age(38-48) years No=(52)</th>
<th>P Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>HbA1C%</td>
<td>Mean± SD 7.490 ± 0.968</td>
<td>Mean± SD 11.20 ± 1.72</td>
<td>0.00</td>
</tr>
<tr>
<td>LH(mIU/ml)</td>
<td>3.88 ±0.36</td>
<td>6.73 ± 0.69</td>
<td>0.077</td>
</tr>
<tr>
<td>FSH(mIU/ml)</td>
<td>4.85±1.14</td>
<td>6.414 ± 0.799</td>
<td>0.00</td>
</tr>
</tbody>
</table>

P value is significant ≤ 0.05 level.

Table (2) shows that the mean HbA1C was 7.490± 0.968 in group I and 11.206± 1.72 in group II, while the Mean±SD of both LH, FSH were 3.883±0.36, 4.8517±1.14 respectively in group I and 6.73± 0.694,6.4140± 0.799 respectively in group II with p value of ≥ 0.081(not significant) for LH and ≤0.05 (significant) for FSH.

It was found in the present study that there were changes in sex hormones in patients according to their age group. As shown in Table (2) and Fig.(2) that the changes in FSH hormone between age groups were significant except for LH where the changes between age groups were not significant.

The age of women is one of factors which restricted the ovarian steroid secretion capacity, these hormones was decline around the age of 30 years old, so that gonadotropin (LH, FSH) will be increase [21, 22].
Conclusion
A significant elevation of HbA1c in diabetic female compared to control. Also a significant increase of LH and FSH in diabetic premenopausal women compared to control. According to the age of women a significant elevated in FSH level and not significant elevated level of LH in diabetic women. In conclusion could be drawn that LH and FSH increase in diabetes mellitus type II premenopausal women.

References


