Purpose: To evaluate whether pregnancy have an effect on the foveal thickness (FT) and volume (FV) in Saudi women.

Methods: This study involved 120 pregnant women (age range: 19-37 years) in the first, second and third trimester as the study groups and compared to 40 healthy non-pregnant women with similar age range. The assessment of FT and FV in each group was done using the optical coherence tomography (OCT). Statistics such as the mean, percentiles and range were used. A $P$-value of 0.05 or less was set to test significance.

Results: Average foveal thickness and foveal volume were significantly larger ($P \leq 0.05$) in the first, second and third trimesters compared to controls. The mean FT was 226.57±23.3 μm, 231.37±19.21 μm, 240.3±21.76 μm in group 2, 3, and 4 respectively compared to control mean of 214.2±19.73 μm. Mean FV was 0.259±0.07 mm$^3$, 0.241±0.05 mm$^3$, 0.300±0.11 mm$^3$ in group 2, 3, and 4 respectively compared to control mean of 0.169±0.015 mm$^3$. The 3rd trimester was associated with significantly ($P \leq 0.05$) higher FV compared to the 2nd trimester.

Conclusion: The changes associated with pregnancy such as the increase of body fluids, can cause significant alteration in FT and FV. Although such alteration may not affect the quality of vision in healthy subjects, pregnant women with conditions that may cause fluid retention, or those with pre-existing systemic or eye disease or who are predisposed to risk factors should be monitored during pregnancy to avoid unexpected complications.

Keywords: Foveal thickness; Foveal volume; Pregnancy; Optical coherence tomography

Introduction

Pregnancy is associated with many physiological and physical changes that are attributed to the shift in hormonal levels, mainly the estrogen and progesterone during the reproductive phases of the first, second and third trimesters. These include gastrointestinal complaints, breast tenderness, fatigue, dizziness, gaining weight, frequent urination and pregnancy mask [1]. Moreover, the estrogen and progesterone were related to somatopsychologic disturbances in pregnancy as early as the first trimester [2,3].

The pregnancy imposes an effect on each organ system of the body, and the eye is no exception. Pregnancy-induced ocular changes include reduction in the intraocular pressure (IOP) [4], unilateral ptosis [5], dry eye [4] and alteration in lens and corneal curvature have been observed and attributed to fluid and hormonal changes [6-8].

The ophthalmic changes could start as early as the first trimester and recovery to the pre-pregnancy levels could take weeks to months post-partum [6].

In pregnancy, plasma volume expands to support the uteroplacental blood flow to the fetus. The increase in total body water can be detected early in the pregnancy and continue till the delivery [9]. Pregnancy-induced corneal thickness due to edema has been reported [7]. An alteration in the refractive error and transient loss of accommodation may follow [7]. Fluid retention during pregnancy may affect other ocular tissue such as the choroid, increasing its thickness in the second or third trimesters [10,11]. Subfoveal choroidal areas were also found to be of higher thickness in healthy pregnant subjects [12,13]. Theses changes were detected using optical coherence tomography (OCT) [10-12].

On the other hand, pregnancy may exacerbate retinal diseases such as diabetic retinopathy (DR). Pregnant women with diabetes have a risk of developing either non-proliferative or proliferative DR [14]. One suggestion to the increased risk of DR during pregnancy is the increase in retinal blood flow and hyper-perfusion induced-stress on the compromised retina [15]. It has also been found that progesterone may influence the release of vascular endothelial growth factor (VEGF), a well-known angiogenic factor that exacerbates DR [16]. As the macular edema may develop or worsen during pregnancy, early detection is essential for the management and prevention of further complications.
Clinical examination involves the use of OCT, a non-invasive procedure, which gives high resolution cross-sectional, imaging that, enables the clinician to scan accurately, the subclinical changes in the retina. The measurement of central foveal thickness correlates with the visual acuity and provides a mean of monitoring the macula in different conditions [17]. In this study, we used OCT to quantify foveal thickness and volume in healthy pregnant women during the first, second and third trimesters and to assess whether pregnancy-induced foveal changes contribute to any visual disturbances in healthy subjects.

Methods

Study population and examination

All pregnant women were recruited from the gynecology and obstetrics department and ophthalmology unit in King Khalid University between March and May 2014. This study included 120 healthy pregnant women (120 eyes) and 40 healthy non-pregnant women (40 eyes). Group 1 (control group) included 40 healthy non-pregnant women (40 eyes), Group 2 consisted of 40 healthy pregnant women (40 eyes) in first trimester; group 3 included 40 healthy pregnant women (40 eyes) in the second trimester; and group 4 included 40 healthy pregnant women (40 eyes) in third trimester.

Exclusion criteria for all participants included any history of diabetes mellitus, glaucoma, any corneal and lens pathologies preventing ocular fundus examination, laser therapy, and refractive error greater than 0.50 or less than –0.50 dioptres.

All participants underwent full ophthalmic examination in the form of: visual acuity test using E chart, auto refraction KR.8800, and complete slit-lamp examination of the anterior and posterior segment of the eye.

The values of FT and FV were measured by OCT RTVue model – RT100 (optovueInc, Fremont, CA) after mydriasis with tropicamide drop 1%. Three measurements were taken for the right eye and used for statistical analysis. The 6 macular map was used to evaluate the values of FT and FV. All measurements were carried out by same operator on all subjects.

Data from this study showed that FT and FV values were increased significantly (P≤0.05) during the pregnancy as early as the first trimester (Figure 1B and 2B). The mean FT was 226.57±23.3 μm, 231.37±19.21 μm, 240.3±21.76 μm in group 2,3, and 4 respectively compared to control mean of 214.2±19.73 μm. Mean FV was 0.259±0.07 mm³, 0.241±0.05 mm³, 0.300±0.11 mm³ in group 2,3, and 4 respectively compared to control mean of 0.169±0.015 mm³. Moreover, the mean FT was altered significantly (P≤0.05) in the third trimester as compared to the control and the first trimester (Figure 1B). Similarly, the mean of the FV was significantly higher (P≤0.05) compared to the second trimester (Figure 2B). No correlation was found between the FT and FV, the age and trimesters in all groups.

Discussion

The assessment of foveal thickness and volume are important for the diagnosis, treatment, and follow-up of retinal diseases such as diabetic retinopathy, age-related macular degeneration and glaucoma. In this study, OCT was utilized to detect changes to the fovea during pregnancy.

Statistical analysis

The results were analyzed using Graphpad prism software version 7 (GraphPad Software, Inc., La Jolla, USA) and variables are presented as mean ± SD. D’Agostino & Pearson test was used to assess the normality of the data. One- way analysis of variance (ANOVA) was used to compare datasets. The level of significance was set at P value of 0.05 or less.

Ethical consideration

The protocol of the study was explained to each participant at the time of recruitment and informed consent was obtained according to the Declaration of Helsinki. The research was approved by the research ethical committee at King Khalid University Hospital (KKUH).

Results

The study included 120 eyes of 120 healthy pregnant women. The control group consisted of 40 eyes of 40 healthy non-pregnant women. There was no variation in age between the four groups. The mean age was: 30.4±4.1, 29.1±4.7, 28.03±5.4 and 27.4±4.6 years, in group 1,2,3 and 4 respectively. Table 1 shows the results of mean age, FT, and FV values in all participants.

Table 1: Mean age, FT and FV values of the groups.

<table>
<thead>
<tr>
<th>Group 1</th>
<th>Group 2</th>
<th>Group 3</th>
<th>Group 4</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean± SD</td>
<td>30.4±4.1</td>
<td>29.1±4.7</td>
<td>28.03±5.4</td>
</tr>
<tr>
<td>FT (μm)</td>
<td>214.2±19.7</td>
<td>226.6±23.3</td>
<td>231.4±19.2</td>
</tr>
<tr>
<td>Mean± SD</td>
<td>0.169±0.015</td>
<td>0.259±0.070</td>
<td>0.241±0.051</td>
</tr>
<tr>
<td>FV (mm³)</td>
<td>0.19</td>
<td>0.26</td>
<td>0.24</td>
</tr>
</tbody>
</table>

FT: Foveal Thickness; FV: Foveal Volume; SD: Standard Deviation.
pregnancy trimesters compared to healthy non-pregnant women. Several studies have shown alteration in the choroidal thickness during pregnancy especially in the third trimester [10-13,18,19]. The affected regions include the central subfield, inferior inner macular choroid, temporal, nasal and sub-foveal areas [12,19]. On the other hand, others reported that pregnancy has no effect on the choroidal thickness and it is comparable to healthy non-pregnant women [20,21].

The current results showed that pregnancy could alter significantly the FT and FV as compared to healthy non-pregnant values. This is in accordance with Demir et al., (2011) [22] findings in which women in their third trimester had higher mean values for the foveal and parafoveal thickness. One may expect that the third trimester is associated with more changes compared to earlier gestational periods; our study shows that earlier changes can also be detected in the first and second trimester using OCT. These were significantly higher in our sample. Similarly, there was a significant 53% and 77% increase in the FV in the first and third trimester respectively as compared to control. This is in parallel with Cankaya et al., (2013) [23] who found that total macular volume and FT were increased in the first, second and third trimesters in healthy women. As no pathological findings were found, this could be explained by the expansion of plasma volume in pregnancy, which is reported to increase by 45% [9,24]. One of the main limitations of the current study was the small sample size. In addition, it is worth monitoring one group from the first trimester till post-partum and monitor the time needed for the FT and FV values to return to the baseline.

In conclusion, pregnancy can result in an alteration to different ocular structures without visual complaints. However, routine examination of the retina using OCT is recommended for detecting early signs of macular edema especially if the patients are predisposed to any risk factors for developing hypertension, preeclampsia, diabetes or any other conditions that affect the retina.

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