

OBESITY MAY INFLUENCE ON MEASUREMENTS OF CHOROID THICKNESS IN OPTICAL COHERENCE TOMOGRAPHY

Erhan YUMUSAK*, Kemal ORNEK*, Senay Arıkan DURMAZ, Aydin CIFCI, *, Hatice Ayhan GULER*, Zehra Bacanlı*****

*Kırıkkale University, School of Medicine, Department of Ophthalmology, Kırıkkale, Turkey., **Kırıkkale University, School of Medicine, Department of Endocrinology, Kırıkkale, Turkey. ***Kırıkkale University School of Medicine Department of Internal Medicine, Kırıkkale, Turkey

Corresponding author: Dr. Erhan Yumusak

Address: Kırıkkale University Hospital Department of Ophthalmology Yahşihan, Kırıkkale, Turkey.

Abstract

Background: Excessive weight is a well-known risk factor for microvascular diseases. Changes in thickness in a vascular tissue, such as the choroid, can be useful to evaluate the effect of obesity on the microvascular system. The aim of this study was to evaluate the choroidal thickness (CT) changes in obese women, using optical coherence tomography (OCT).

Methods: The prospective clinical study included examination of the right eyes of 72 patients. The right eyes of 68 patients were examined and served as the controls. A complete ophthalmological examination and OCT imaging were performed for each group studied. The CT in each eye was measured using OCT.

Results: The obese group consisted of 72 female patients with a mean age of 37.27±1.18 years. The control group included 68 female subjects with a mean age of 37.85±7.98 years (p>0.05). There was no statistical significant difference for the foveal retinal thickness measurements between the two groups (p>0.5). Our study revealed significant choroidal tissue thickening subfoveally and at areas 500 µm temporal, 500 µm nasal, and 1500 µm nasal to the fovea in the obese group (all p<0.05). There was a positive correlation between body mass index (BMI) and CT changes.

Conclusions: CT may increase in obese women and a positive correlation was found between BMI and CT.

Keywords: choroidal thickness, obese women, optical coherence tomography

Introduction

Obesity is a common health problem and its prevalence is increasing worldwide[1-3]. The association of obesity with cataract formation, glaucoma, and age-related macular degeneration has been shown in varying degrees. Researchers have hypothesized that retinal microvascular changes are precursors to developing obesity based on experimental and clinical observations [4-5]. In the Blue Mountains Eye Study, retinal vessel diameter was associated with the prevalence of higher body mass index (BMI) and the increased risk of incident obesity [6].

In the eye, the choroid, the posterior portion of the uveal tract, nourishes the outer portion of the retina. It contributes to the blood supplied to the prelaminar portion of the optic nerve [1], is an integral constituent in the functioning of the eye, and is involved in important diseases affecting the optic nerve, retinal pigment epithelium, and the retina. Previous studies have suggested that a higher BMI can trigger structural changes in the retinal vascular system that could provoke retinal dysfunction, as shown in aged-related macular degeneration or diabetic retinopathy. Therefore, knowledge of the thickness changes in a vascular tissue, such as the choroid, may help to evaluate the effect of obesity on the microvascular system. By using enhanced depth imaging optical coherence tomography (EDI-OCT), choroid images can be obtained and the choroidal thickness (CT) can be measured [3].

Therefore, in the present study, we hypothesized that obesity is correlated with CT changes, particularly in women. To the best of our knowledge, this is the first study evaluating CT in obese female patients.

Materials and Methods

This prospective clinical study included the examination of the right eyes of 72 patients. In total, 68 right eyes of 68 patients were examined and served as controls. The study was conducted between 2015 and 2016 in accordance with the tenets of the Declaration of Helsinki. The trial protocol was approved by the Local Ethical Committee of the University of Kırıkkale. Registration of the trial was requested on April 27th, 2015 (decision no:10/11). All patients and control subjects voluntarily participated in the study and signed an informed consent form. The obese group was classified according to the World Health Organisation criteria; (BMI 18.5–24.9 kg/m²= normal; 25.0–29.9 kg/m²= pre-obese/overweight, and ≥30.0 kg/m²= obese).

In the study, the obese group included patients who had a BMI> 30 kg/m², without any other disease, whereas healthy adults with BMI <25 kg/m² constituted the control group. Obese patients were randomly selected from those monitored by the Department of Endocrinology. The exclusion criteria were as follows: a previous systemic or chronic disease such as hypertension, smoking, ocular surgery in one or both eyes; axial length >24 ± 1.0 mm; and a refractive measurement > 2.0 diopters.

All participants underwent a complete ocular examination, including a best-corrected visual acuity measurement, slit-lamp examination, intraocular pressure measurement, and dilated funduscopy. Only the right eyes of each of the patients were selected to avoid any intra-individual bias.

The CT was measured as close to noon as possible to avoid diurnal variations. The measurements were performed using an EDI-OCT scanning system (OCT Advance Nidek RS-3000; Nidek Co. Ltd., Gamagori, Japan). Prior to evaluation using EDI-OCT scanning, the central macular thickness was measured in the right eye of each patient. Choroidal and scleral boundaries were drawn with the assistance of software programs. The boundaries limited the Bruch membrane, between the subfoveal points (FCT), to 500 and 1500 µm in the nasal regions (N500, N1500) and 500 and 1500 µm in the temporal regions (T500, T1500), for CT measurements. All measurements including the demarcation of the choroid and sclera were made by two independent (masked) observers. There were no significant differences between the results of the two observers (p = 0.317: Paired t-test, r = 0.716 and p<0.001:Pearson's correlation), and the average of the two results was used in our analyses.

Statistical analyses were performed using the SPSS statistical software (SPSS for Windows 23.0, Inc., Chicago, USA). The results of the descriptive analysis were provided in numbers, percentages, mean, median, and standard deviations. A paired t-test was used to assess the difference in the means of the observers' measurements to test the repeatability and accuracy of the two independent measurements. The independent t-test was used to compare the variables between the obese group and the control group, and correlations were performed using Pearson's correlation coefficient. A multiple linear regression analysis (forward) was used to determine confounding factors among the variables. p<0.05 was considered statistically significant.

Results

The study group consisted of 140 female (100%) subjects, with a mean age of 37.55±1.01 years (median:38; range:21–59 years). There were 72 patients in the obese group, with a mean age of 37.27±1.18 years (median: 38.5; range 21–59 years). The control group included 68 subjects, with a mean age of 37.85±7.98 years (median:38; range 24–54years). There was no significant difference, in terms of age, between the two groups (p>0.5). Demographics of the study groups are shown in Table-1.

There was no significant difference found for foveal retinal thickness (FT) when the two groups were compared (p>0.5). In contrast, the CT revealed significant differences at FCT, T500, N500, and N1500 between the two groups (all p<0.05). Changes in both FT and CT are demonstrated in Table-2.

There was a positive correlation found between BMI and CT at FCT, T500, and N500 (Table-3). Multiple linear regression analysis revealed that CT had been affected by BMI independently from the aspect of age of the patient (Table-4).

Table 1. Demographics of the groups

Group	Age	BMI (Kg/m ²)	Weight (kg)	Height(cm)
1(N=72)	37.27±1.18	39.16±6.88	100.63±16.49	160.42±6.20
2(N=68)	37.85±7.98	21.95±5.9	58.97±16.49	168.36±4.94
	P=0.738	P<0.001	P=0.004	P<0.001

Table 2. Changes in foveal thickness and choroidal thickness

Group	FT	FCT	T500	T1500	N500	N1500
1(N=72)	250.1±19.8	349.2±58.7	346.5±55.1	335.8±55.1	345.5±59.7	327.0±55.3
2(N=68)	256.0±18.9	322.7±37.8	317.3±39.7	313.6±39.7	323.3±39.5	322.2±37.7
	P=0.185	P=0.002	P=0.014	P=0.230	P<0.001	P<0.001

FCT: choroidal thickness at fovea; N500, choroidal thickness at 500 µm nasal to the fovea; N1500, choroidal thickness at 1500 µm nasal to the fovea; T500, choroidal thickness at 500 µm temporal to the fovea; T1500, choroidal thickness at 1500 µm temporal to the fovea; FT: central macular thickness

Table 3. The Pearson Correlation analysis between body mass index - Choroidal thickness and foveal thickness

AGE	FCT	N500	N1500	T500	T1500	FT
r = 0.125	r = 0.198	r = 0.193	r = 0.082	r = 0.206	r = 0.151	r = -0.162
P = 0.140	P = 0.019	P = 0.022	P = 0.334	P = 0.015	P = 0.076	P = 0.056

FCT: choroidal thickness at fovea; N500, choroidal thickness at 500 µm nasal to the fovea; N1500, choroidal thickness at 1500 µm nasal to the fovea; T500, choroidal thickness at 500 µm temporal to the fovea; T1500, choroidal thickness at 1500 µm temporal to the fovea; FT: central macular thickness

Table 4. Multiple linear regression analysis between choroidal thickness, age and Body mass index.

		Beta(β)	P
N500	BMI	0.233	0.004
	Age	-0.317	<0.001
N1500	BMI	0.249	0.011
	Age	-0.325	0.012
T500	BMI	-0.402	0.001
	Age	-0.289	<0.001
T1500	BMI	0.204	0.009
	age	-0.423	<0.001
FCT	BMI	0.243	0.002
	Age	-0.364	<0.001

FCT: choroidal thickness at fovea; N500, choroidal thickness at 500 µm nasal to the fovea; N1500, choroidal thickness at 1500 µm nasal to the fovea; T500, choroidal thickness at 500 µm temporal to the fovea; T1500, choroidal thickness at 1500 µm temporal to the fovea; FT: central macular thickness

Discussion

In the eye, CT may be affected by several factors, such as age, axial length, and refractive errors [7-8]. Diurnal changes in CT have also been reported [9]. It is believed that systemic blood pressure and intraocular pressure induce choroidal tissue changes through an autoregulatory mechanism [10]. Therefore, because the choroid possesses a rich vascular structure, all of the aforementioned factors have the potential to alter the CT [11]. A number of studies have found that CT plays a prognostic or predictive role in various local (for example, diabetic retinopathy, and AMD), and systemic diseases (for example, hypertension, anemia, and rheumatoid arthritis) [12-14]. Jongh et al., reported the effects of obesity on the microvascular system; hyperinsulinemia and elevated blood pressure were found to be the major causes of the vascular alterations in obese women [15]. In another study by Kawasaki et al., both retinal venous and arterial dilatation were found in hypertensive patients [16]. Research by Saito et al., studied the retinal venous system in 900 subjects and reported an incidence of 5 years of obesity in some patients [17]. The authors found a positive correlation between vessel caliber and BMI; however, no correlation was shown between these changes and the development of obesity.

In this study, CT was found to be significantly reduced in the non-obese controls, except for the temporal measurement of 1500 µm. In the light of previous reports, we hypothesized that there is a relationship between obesity and the choroidal layer of the eye. In the present study, the obesity group consisted of patients with a BMI > 30, and subjects with a BMI < 25 constituted the control group. To avoid any diurnal effect, we performed all the measurements at noon for each patient. We also excluded patients with a history of local and systemic diseases. Although no significant differences were found for FT between the groups, there was a significant increase in CT at certain points (CFT, nasal 500, and 1500µm, and temporal 500µm) in the obese group. The results indicated that there was a positive correlation between BMI and CT, and multiple linear regression analysis revealed that CT was independently affected by the age of the patients. There were some limitations in the study, such as the pathogenesis of obesity, which included several unknown hormonal and genetic factors; moreover, because choroid is a vascular tissue, it may be affected by local and systemic factors

Conclusion

In summary, our data provides evidence for a relationship between CT and obesity in female patients. Vascular abnormalities may occur at early stages in obesity and ocular circulation may be a preferred target for the disease process. The assessment of CT is a quick and non-invasive technique, which can be utilized to determine such abnormalities. Meanwhile, it is unclear how this data may be applied to individual patients and how it can benefit obesity management. The data suggests that CT measurement has a predictive role and BMI should be included among the parameters that may affect CT results in obese women.

REFERENCES

- Berghöfer A, Pischon T, Reinhold T, Apovian CM, Sharma AM, Willich SN. Obesity prevalence from a European perspective: a systematic review. BMC Public Health 2008;8: 200.
- Flegal KM, Carroll MD, Kit BK, Ogden CL. Prevalence and trends in obesity among US adults, 1999–2000. JAMA 2002; 288: 1723–1727.
- Must A, Spadano J, Cookley EH, Field AE, Colditz G, Dietz WH. The disease burden associated with overweight and obesity. JAMA 1999; 282:1523–1529.
- Hayashi K, Kanda T, Homma K, Tokuyama H, Okubo K, Takamatsu J, Tatamatsu S, Kumagai H, Saruta T. Altered renal microvascular response in Zucker obese rats. Metabolism 2002; 51: 1553–1561.
- Ikram MK, de Jong FJ, Vinglerling JR, Witteman JC, Hofman A, Breteler MM, de Jong PT. Are retinal arteriolar or venular diameters associated with markers for cardiovascular disorders? The Rotterdam Study Invest Ophthalmol Vis Sci 2004;45:2129–2134.
- Wang JJ, Taylor B, Wong TY, Chua B, Rochtchina E, Klein B, Mitchell P. Retinal vessel diameters and obesity: a population-based study in older persons. Obesity 2006;14: 206–214.
- Ikuno Y, Kawaguchi K, Nouchi T, Yasuno Y. Choroidal thickness in healthy Japanese subjects. Invest Ophthalmol Vis Sci 2010;51:2173–2176.
- Agawa T, Miura M, Ikuno Y, Makita S, Fabritius T, Iwasaki T, Goto H, Nishida K, Yasuno Y. Choroidal thickness measurement in healthy Japanese subjects by three-dimensional high-penetration optical coherence tomography. Graefes Arch Clin Exp Ophthalmol 2011;249(10):1485–92.
- Brown JS, Flitcroft DI, Ying GS, Francis EL, Schmid GF, Quinn GE, Stone RA. In vivo human choroidal thickness measurements: evidence for diurnal fluctuations. Invest Ophthalmol Vis Sci 2009;50:5-1236.
- Reiner A, Li C, Del Mar N, Fitzgerald ME. Choroidal blood flow compensation in rats for arterial blood pressure decreases is neuronal nitric oxide-dependent but compensation for arterial blood pressure increases is not. Exp Eye Res 2010;90:734–74137.
- Kiel JW. Choroidal myogenic autoregulation and intraocular pressure. Exp Eye Res 1994;58:529–543.
- Yiu G, Chiu SJ, Petrou PA, Stinnett S, Sarin N, Farsi S, Chew EY, Wong WT, Toth CA. Relationship of central choroidal thickness with age-related macular degeneration status. Am J Ophthalmol. 2015;159(4):617-26.
- Yumusak E, Ciftci A, Yalcin S, Sayan CD, Dikel NH, Ornek K. Changes in the choroidal thickness in reproductive-aged women with iron-deficiency anemia. BMC Ophthalmol. 2015 Dec 29;15(1):186.
- Duru N, Altinkaynak H, Erten S, Can ME, Duru Z, Uğurlu FG, Çağrı N. Thinning of Choroidal Thickness in Patients with Rheumatoid Arthritis Unrelated to Disease Activity. Ocul Immunol Inflamm. 2015 Jul 31;1-8. [Epub ahead of print].
- de Jongh RT, Semé EH, IJzerman RG, de Vries G, Stehouwer CD. Impaired microvascular function in obesity: implications for obesity-associated microangiopathy, hypertension, and insulin resistance. Circulation 2004; 109: 2529–2535.
- Kawasaki R, Tielsch JM, Wang JJ, Wong TY, Mitchell P, Tano Y, Tominaga M, Ozumi T, Daimon M, Kato T, Kawata S, Kayama T, Yamashita H. The metabolic syndrome and retinal microvascular signs in a Japanese population: the Funagata study. Br J Ophthalmol 2008; 92: 161–166.
- Saito K, Tanabe Y, Kawasaki R, Daimon M, Ozumi T, Kato T, Kawata S, Kayama T, Yamashita H. Is retinal vasculature change associated with risk of obesity? Longitudinal cohort study in Japanese adults: The Funagata study. J Diabetes Investig. 2011 Jun 5;2(3):225-32.