

Original Article / Orijinal Araştırma

Carotid blood-flow velocity and intima-media thickness changes in patients with obesity

Obez hastalarda karotis arter akım hızı ve intima-media kalınlık değişiklikleri

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Abstract

Aim. Carotid ultrasound scanning including carotid intima-media thickness (IMT) and Doppler velocity measurements is commonly used modality to assess carotid artery disease related to atherosclerosis as an important result of obesity. The aim of the present study was to assess the effect of obesity on the carotid ultrasonographic parameters including lumen diameters, IMTs, peak systolic velocities (PSVs), and end-diastolic velocities (EDVs) of common carotid arteries (CCAs) and internal carotid arteries (ICAs) in an asymptomatic population. **Methods.** The lumen diameters, IMTs, PSVs, and EDVs of CCAs and ICAs were performed in the study participants grouped according to their body mass index (BMI) in an age-adjusted manner: normal group with BMI 18.50-24.99; overweight group with BMI 25.00-24.99; obesity classes I&II group with BMI 30.00-39.99; obesity class III group with BMI \geq 40.00. **Results.** Overall, obesity class I-III increased the lumen diameters of CCAs and ICAs ($p < 0.05$); increased the IMTs of CCAs but not of the ICAs. ($p < 0.05$); and decreased the PSVs of CCAs but not of the ICAs ($P < 0.05$). Overall, in accordance with the severity of obesity, the EDVs were decreased in the CCAs but not in the ICAs ($p < 0.05$). **Conclusions.** Obesity may have a potential to increase the lumen diameter of CCAs and ICAs, to increase the IMTs of CCA but not ICA, and to decrease the PSVs and EDVs of CCA but not ICA. There is a need to determine the effect of obesity on the wall and function of carotid arteries in further studies in a broader population subclassified according to the severity and duration of obesity.

Keywords: Carotid artery, intima-media thickness, Doppler blood-flow velocity, obesity

Özet

Amaç. Doppler akım ölçümü ve karotis intima-media kalınlığını (İMK) içeren karotis ultrason taraması, obezitenin önemli sonucu olarak arterioskleroz ile ilişkili karotis arter

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hastalığının değerlendirilmesinde yaygın olarak kullanılmaktadır. Bu çalışmanın amacı, asemptomatik toplulukta internal karotis arterin (İKA) ve kommon karotis arterin (KKA); lümen çapını, İMK, pik-sistolik (PSH) ve end-diastolik hızını (EDH) içeren karotid ultrasonografik bulgularına obezitenin etkilerini değerlendirmektir. **Yöntem.** KKA ile İKA'nın damar lümen çapı, İMK, PSH ve EDH'ları yaşlara göre ve body mass indekslerine (BMİ) göre gruplara ayrıldı. Normal grubun BMİ'si: 18.50-24.99; kilolu grubun BMİ'si 25.00-24.99; obezite I&II grubun BMİ'si 30.00-39.99; obezite III grubun BMİ'si ≥ 40.00 olacak şekilde gruplara ayrıldı. **Bulgular.** Genel olarak, obezite I-III grubunda KKA ve İKA damar lümen çapı artmıştı ($p < 0.05$); KKA'nın İMK artmasına rağmen İKA'da artış saptanmadı ($p < 0.05$) ve KKA'nın PSH'ında azalmaya rağmen, İKA'de azalma olarak saptanmadı ($p < 0.05$). Genel olarak obezitenin şiddeti göz önüne alındığı zaman, EDH'da, KKA'da azalma saptanırken, İKA'da saptanmadı ($p < 0.05$). **Sonuçlar.** Obezite, İKA'nın değil ama KKA'nın İMK'indeki artış ile PSH ve EDH azalışında ve KKA ile İKA damar lümen çapının artışına sebep olabilir. Karotid arterin duvar ve fonksiyonlarının obeziteden etkilenmesinin saptanması için daha geniş çalışmalara ihtiyaç vardır.

Anahtar sözcükler: Karotis arter, intima-media kalınlığı, Doppler kan akım hızı, obezite

Introduction

Obesity has become one of the most important public health problems worldwide. Obesity, primarily abdominal obesity, is a significant risk factor for the development of atherosclerosis. The clinically defined metabolic syndrome is the most prominent atherosclerotic risk factor based on adipose tissue dysfunction producing low-grade inflammation and endothelial dysfunction [1, 2]. Atherosclerosis is a systemic pathologic process of the large and medium-sized arteries such as carotid and cerebral arteries. It causes luminal narrowing as a result of the accumulation of lipid and fibrous material between the intimal and medial layers of the vessel [3-5]. Atherosclerosis as a chronic disease can remain asymptomatic for decades. Approximately 25% of strokes are due to carotid artery disease because of slowly progressive and cumulative narrowing and decreasing blood flow [6, 7]. Stroke is one of the leading cause of death and the most common cause of long-term neurologic disability.

Carotid artery intima-media thickness (IMT) is a well-accepted marker of subclinical atherosclerosis associated with cardiovascular risk factors and predictive of incident stroke [8-10]. Carotid ultrasonography imaging including evaluations of lumen diameter and IMT and color Doppler measurements of peak systolic velocity (PSV) and end diastolic velocity (EDV) has gain important role in the prevention and management of cardiovascular and neurologic disorders. Carotid Doppler ultrasound via B-mode imaging detects focal increases in blood flow velocity indicative of high-grade carotid stenosis [11].

Carotid ultrasonography as a noninvasive, accurate, and cost-effective imaging modality has gained important place to obtain morphologic and functional information for the investigation of carotid abnormalities. It is increasingly becoming the first and often the sole imaging approach for prediction and management of cardiovascular disorders such as determining requirement of endarterectomy, although in special cases, costly and

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invasive procedures may be required [12]. The PSV is the most frequently used measurement to gauge the severity of the stenosis, but the EDV and other Doppler parameters provide additional information [13, 14].

According to our knowledge, there is not sufficient data to determine the effect of obesity with a body mass index from >25.00 to >40.00 kg/m² in a population asymptomatic for cardiovascular and neurologic disorders. The aim of the present study was to assess the effect of obesity on the carotid ultrasonographic parameters including the lumen diameters, IMTs, PSVs, and EDVs of common carotid arteries (CCAs) and internal carotid arteries (ICAs) in an asymptomatic population.

Materials and methods

Written informed consent was obtained from each subject, and the study protocol was approved by the Human Ethics Committee of our university. Study participants are a Turkish population of 160 asymptomatic adults aged from 30 to 61 (mean [SD]: 46.8 [5.8]) years. Participants were not included if they had diagnosed as having heart attack, stroke, transient ischemic attack, heart failure, angina, atrial fibrillation, or history of any cardiovascular procedure. Age, gender, BMI, fasting plasma glucose, total LDL and HDL cholesterol and triglyceride levels of the study population were recorded.

Study participants were included to study groups according to their body mass index (BMI) [15] in an age-adjusted manner: normal group with BMI 18.50-24.99; overweight group with BMI 25.00-24.99; obesity classes I&II group with BMI 30.00-39.99; obesity class III group with BMI ≥40.00.

The patients fasted overnight and not given any medication before blood drawn. Blood samples were obtained for determination of glucose and lipids and full blood count. Blood pressures were measured at the right brachial artery after 10-min supine rest. The participants studied underwent ultrasonography of the common, right, and left carotid arteries for Doppler flow and IMT measurements. Heart rate and blood pressure were recorded prior to US examination. One radiologist performed all ultrasound examinations by an ultrasonography device (General Electric Logic 9; Waukesha, WI, USA) equipped with a 10 MHz linear probe. Intraobserver reliability was determined as Kappa coefficient ranged from 0.67-0.87. In all patients, routine carotid US studies were performed, which included gray-scale, pulsed Doppler, and color Doppler flow US examinations of the left and right CCAs and ICAs. Lumen diameters of left and right CCAs and ICAs were recorded.

Assessment of carotid IMT

Participants were examined supine with the head rotated 45° towards the left or right side. Imaging was done in the plane parallel to the neck with the jugular vein lying immediately above the common carotid artery. Images of the right common carotid artery were centered 10 to 15 mm below (caudad to) the right common carotid artery bulb. End-diastolic images (smallest diameter of the artery) were captured. IMT of the common carotid artery (CCA) and presence of plaques were assessed. IMT of the CCA was

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determined in the far wall according 1 cm proximal to the bifurcation. Carotid artery plaque was excluded [10, 16].

Assessment of carotid Doppler ultrasonography

Patients were examined while they were in a supine position. Doppler spectral waveforms were obtained in the CCA and ICA. The measured angle of insonation was less than 60° for all measurements [17].

Statistical analysis

Data were expressed as mean ± SD. The age, blood test, ultrasonographic and Doppler ultrasonographic findings were analyzed with ANOVA with post hoc Tukey test. Significance was determined at the $p < 0.05$ level.

Results

Table 1 presents age, sex ratio, BMI, and fasting plasma glucose, total cholesterol, LDL cholesterol, HDL cholesterol, and triglyceride levels of the normal, overweight, obesity classes I&II, and obesity class III groups. The age and fasting plasma glucose level of obesity class III group were significantly higher than those of the normal, overweight, obesity classes I&II groups ($p < 0.05$). The ages of overweight and obesity classes I&II groups were significantly higher than that of the normal group ($p < 0.05$). Total, LDL, HDL cholesterol and triglyceride levels were found comparable among the study groups.

Table 1. Selected clinical data of study groups.

	Normal (n=50)	Overweight (n=40)	Obesity classes I&II (n=40)	Obesity class III (n=30)
Age (y)	46.6±4.5	45.8±6.1c	46.6±7.5b	48.9±4.3a
Sex ratio (F/M)	25/25	20/20	20/20	20/10
BMI (kg/m ²)	23.1±0.2	27.7±0.3	32.2±0.3	42.0±0.3
Fasting plasma glucose (mg/dL)	93.4±7.6	96.2±12.1	94.4±8.7	103.4±9.1d
Total cholesterol (mg/dL)	197.7±33.7	199.1±44.4	195.5±35.3	198.2±33.6
LDL cholesterol (mg/dL)	115.3±37.9	127.7±33.0	120.0±35.7	120.6±24.4
HDL cholesterol (mg/dL)	47.2±24.2	39.4±8.2	40.4±17.5	44.8±7.1
Triglyceride (mg/dL)	108.6±52.6	127.3±60.1	134.9±60.9	120.1±46.2

^{a,d}P < 0.05 vs. normal, overweight, obesity classes I&II groups.
^{b,c}P < 0.05 vs. normal group.

Figure 1 shows the lumen diameters of left common and internal carotid arteries of the normal, overweight, obesity classes I&II, and obesity class III groups. The lumen diameter of left common carotid artery of obesity class III group was significantly higher than those of the normal, overweight and obesity CI&II groups ($p < 0.05$).

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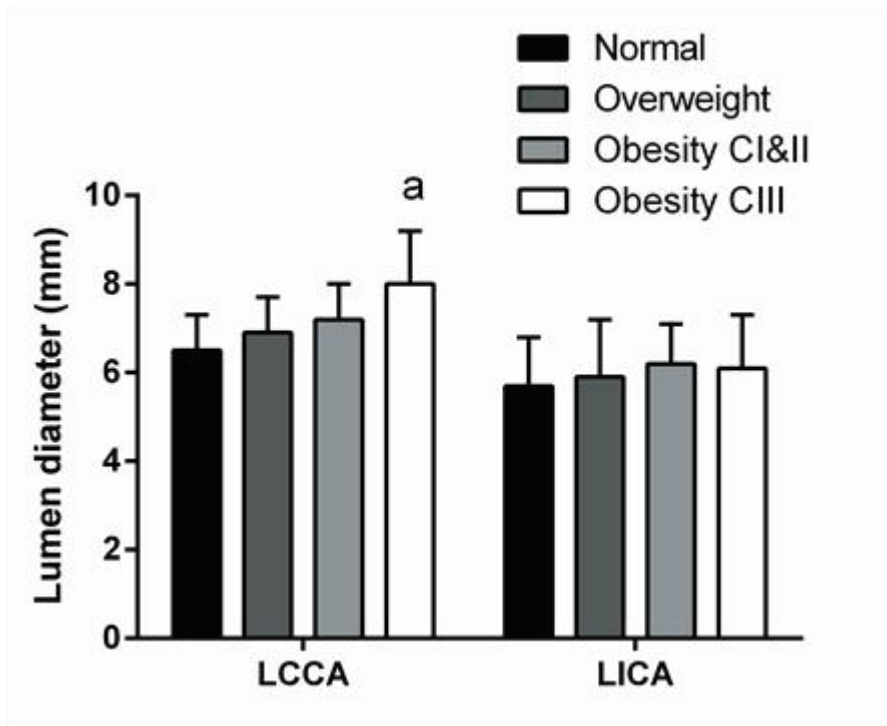


Figure 1. Lumen diameters of left common and internal carotid arteries of study groups. Data were expressed as mean±SD. LCCA, left common carotid artery; LICA, left internal carotid artery; obesity CI&II, obesity classes I&II; obesity CIII, obesity class III. aP<0.05 vs. normal, overweight, and obesity CI&II groups.

Figure 2 displays the lumen diameters of right common and internal carotid arteries of the normal, overweight, obesity classes I&II, and obesity class III groups. The lumen diameters of right common and internal carotid arteries of obesity class III group were significantly higher than those of the normal, overweight, and obesity CI&II groups ($p<0.05$). The lumen diameters of right common and internal carotid artery of obesity classes I&II group were significantly higher than those of the normal and overweight groups ($p<0.05$).

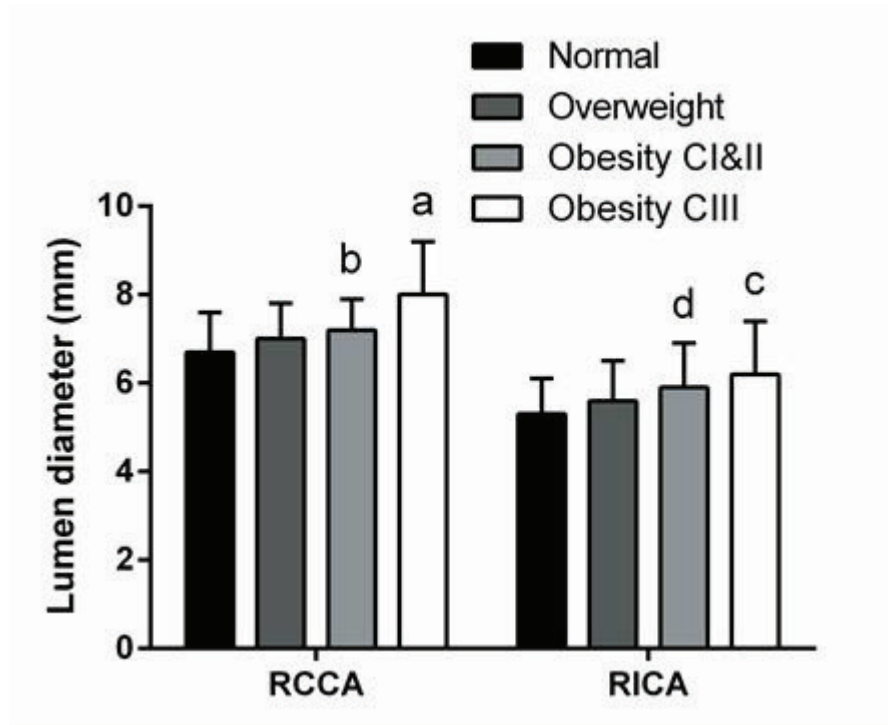


Figure 2. Lumen diameters of right common and internal carotid arteries of study groups. Data were expressed as mean±SD. RCCA, right common carotid artery; RICA, right internal carotid artery; obesity CI&II, obesity classes I&II; obesity CIII, obesity class III.

a,c $p<0.05$ vs. normal, overweight, and obesity CI&II groups.

b,d $p<0.05$ vs. normal and overweight groups.

Figure 3 displays the IMTs of left common and internal carotid arteries of the normal, overweight, obesity classes I&II, and obesity class III groups. The IMTs of left common and internal carotid arteries of obesity class III group were significantly higher than those of the normal, overweight, and obesity CI&II groups ($p < 0.05$). The IMTs of left common and internal carotid arteries of obesity classes I&II group were significantly higher than those of the normal and overweight groups ($p < 0.05$). The IMT of left common carotid artery of overweight group was significantly higher than that of the normal group ($p < 0.05$).

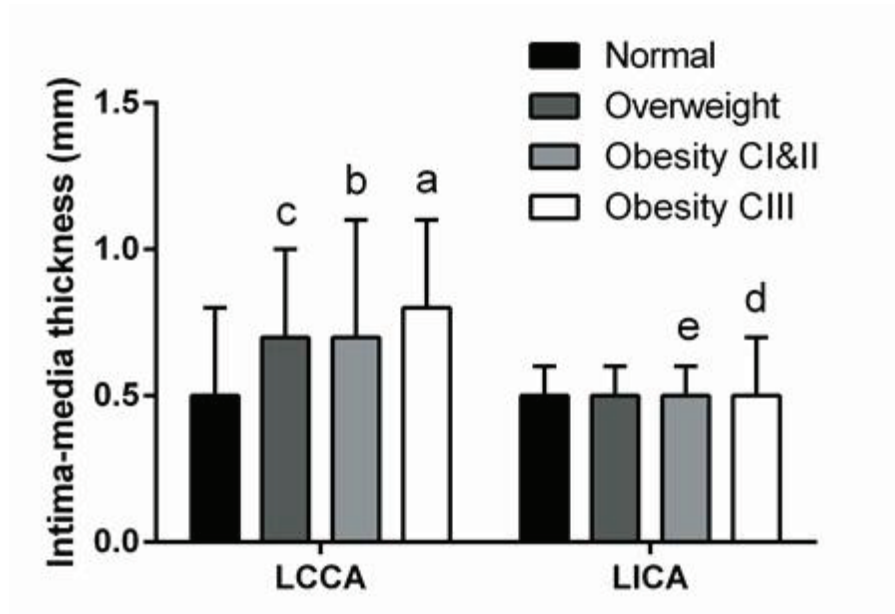


Figure 3. Intima-media thicknesses of left common and internal carotid arteries of study groups. Data were expressed as mean±SD. LCCA, left common carotid artery; LICA, left internal carotid artery; obesity CI&II, obesity classes I&II; obesity CIII, obesity class III.

a, d $P < 0.05$ vs. normal, overweight, and obesity CI&II groups.

b, e $P < 0.05$ vs. normal and overweight groups.

c $P < 0.05$ vs. normal group.

Figure 4 presents the IMTs of right common and internal carotid arteries of the normal, overweight, obesity classes I&II, and obesity class III groups. The IMT of right common carotid artery of obesity class III group was significantly higher than those of the normal, overweight, and obesity CI&II groups ($p < 0.05$). The IMT of right common carotid artery of obesity classes I&II group was significantly higher than those of the normal and overweight groups ($p < 0.05$).

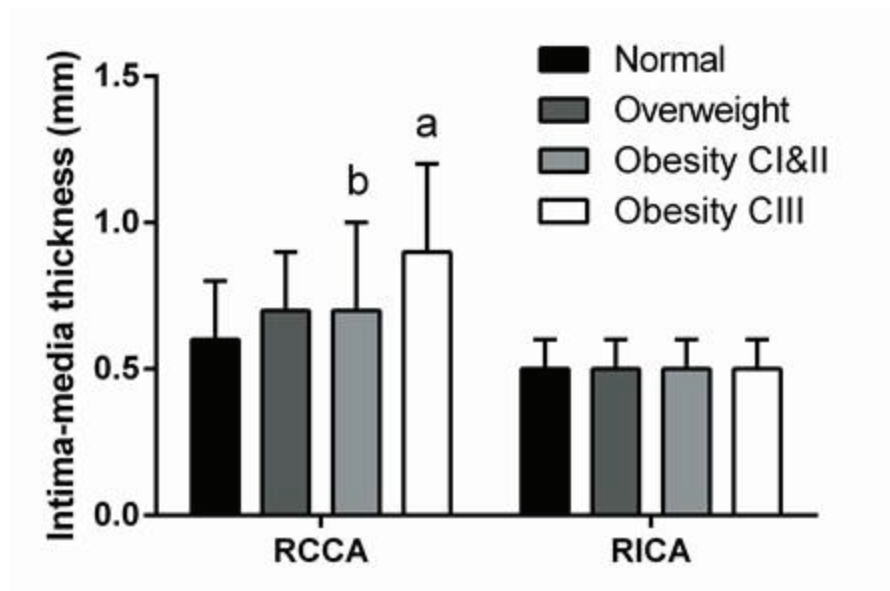


Figure 4. Intima-media thicknesses of right common and internal carotid arteries of study groups. Data were expressed as mean±SD. RCCA, right common carotid artery; RICA, right internal carotid artery; obesity CI&II, obesity classes I&II; obesity CIII, obesity class III. aP<0.05 vs. normal, overweight, and obesity CI&II groups. a,bP<0.05 vs. normal and overweight groups.

Figure 5 presents the peak systolic velocities of left common and internal carotid arteries of the normal, overweight, obesity classes I&II, and obesity class III groups. The PSV of left common carotid artery of normal group was significantly higher than those of the other groups ($p < 0.05$).

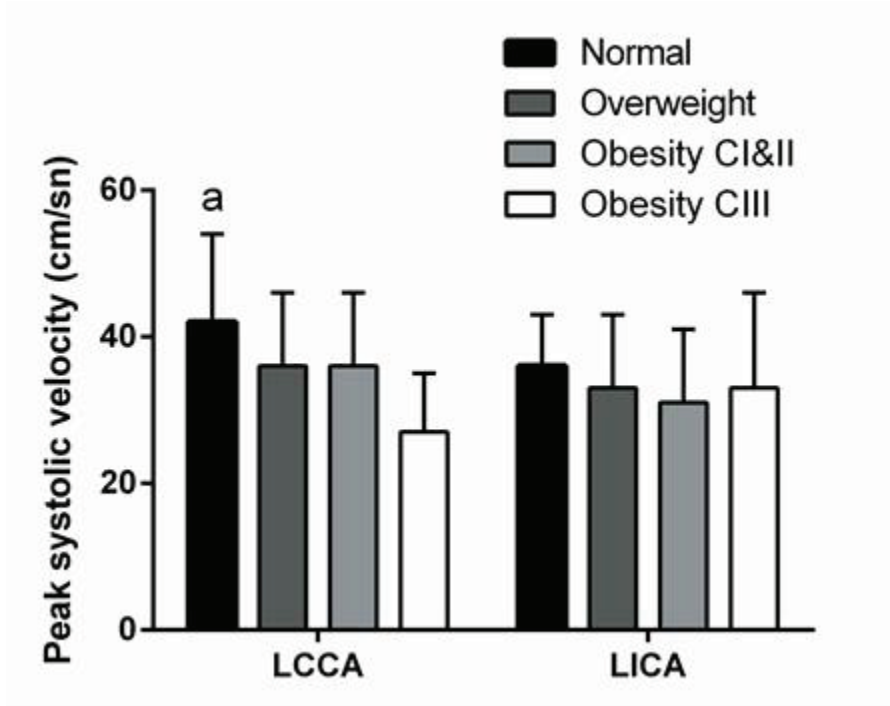


Figure 5. Peak systolic velocities of left common and internal carotid arteries of study groups. Data were expressed as mean \pm SD. LCCA, left common carotid artery; LICA, left internal carotid artery; obesity CI&II, obesity classes I&II; obesity CIII, obesity class III. $aP < 0.05$ vs. overweight, obesity CI&II, and obesity CIII groups.

Figure 6 shows the peak systolic velocities of right common and internal carotid arteries of the normal, overweight, obesity classes I&II, and obesity class III groups. The PSV of right CCA and ICA of normal group was significantly higher than those of the other groups ($p < 0.05$).

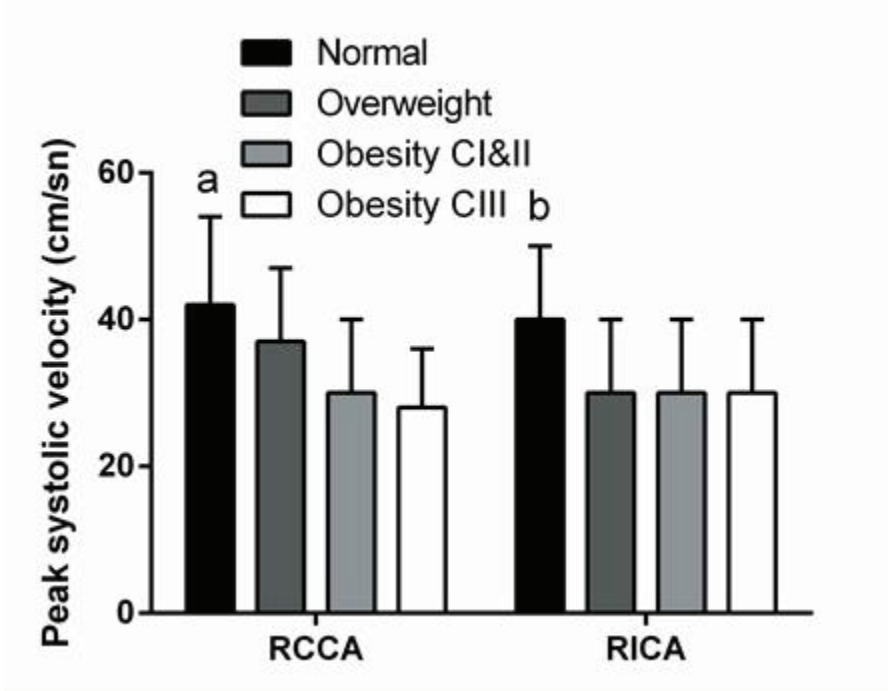


Figure 6. Peak systolic velocities of right common and internal carotid arteries of study groups. Data were expressed as mean±SD. RCCA, right common carotid artery; RICA, right internal carotid artery; obesity CI&II, obesity classes I&II; obesity CIII, obesity class III. a,bP<0.05 vs. overweight, obesity CI&II, and obesity CIII groups.

Figure 7 shows the end diastolic velocities of left common and internal carotid arteries of the normal, overweight, obesity classes I&II, and obesity class III groups. The EDV of left common carotid artery of obesity class III group was significantly lower than those of the other groups ($p < 0.05$).

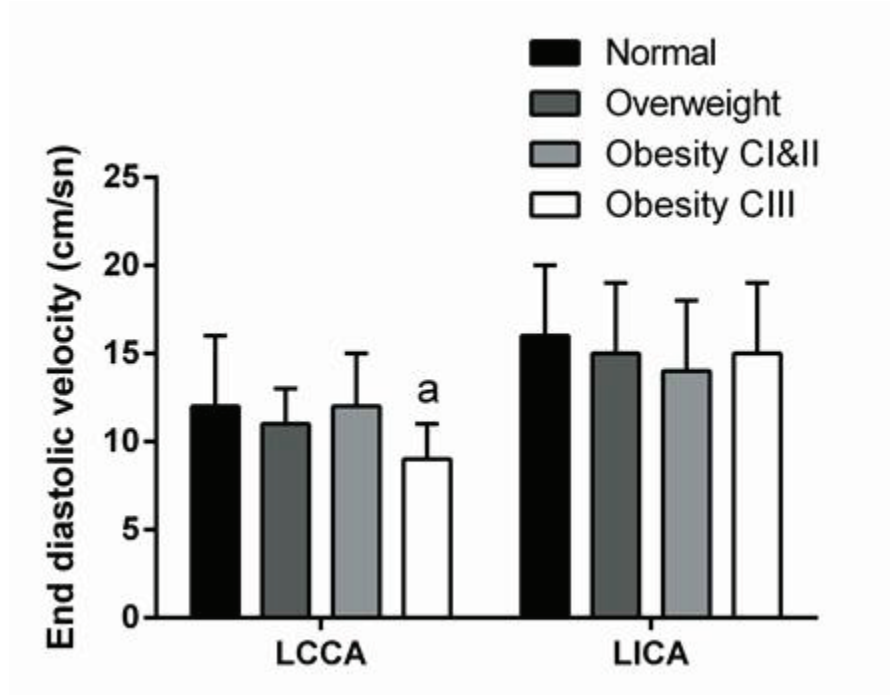


Figure 7. End diastolic velocities of left common and internal carotid arteries of study groups. Data were expressed as mean±SD. LCCA, left common carotid artery; LICA, left internal carotid artery; obesity CI&II, obesity classes I&II; obesity CIII, obesity class III. $aP < 0.05$ vs. normal, overweight, and obesity CI&II groups

Figure 8 shows the end diastolic velocities of right common and internal carotid arteries of the normal, overweight, obesity classes I&II, and obesity class III groups. The EDV of right common carotid artery of obesity class III group was significantly lower than those of the other groups ($p < 0.05$).

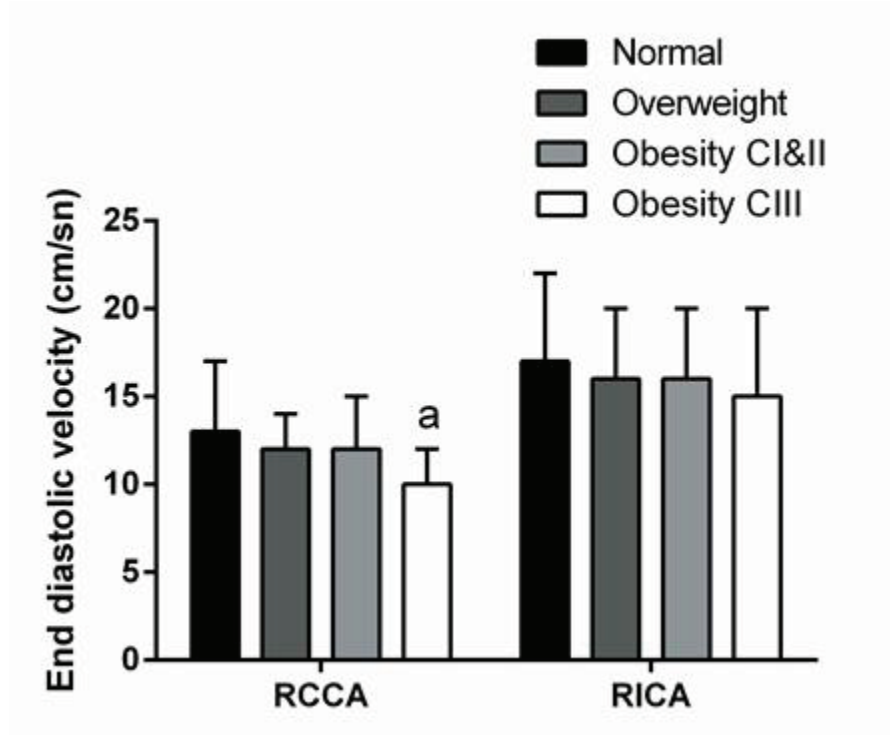


Figure 8. End diastolic velocities of right common and internal carotid arteries of study groups. Data were expressed as mean±SD. RCCA, right common carotid artery; RICA, right internal carotid artery; obesity CI&II, obesity classes I&II; obesity CIII, obesity class III. aP<0.05 vs. normal, overweight, and obesity CI&II groups

Discussion

In this study, the IMT, PSV, and EDV of left and right CCAs and ICAs were evaluated in adult subjects according to the degree of obesity. Normal, overweight, obesity classes I&II, and obesity class III subjects were found considerable comparable with regard to the fasting plasma glucose and lipid levels. In accordance with the findings of previous study of Wildman et al. [18], overall, we found that the lumen diameter of left and right CCAs and ICAs increased as parallel to the severity of obesity. Overall, according to our findings, obesity increased the IMTs of left and right CCAs but not of the left and right

ICAs. Overall, according to our findings, obesity decreased the PSVs and EDVs of left and right CCAs but not of the left and right ICAs.

The carotid artery provides itself an easy way to study by high-resolution ultrasound devices. It is located superficially and found as relatively stationary and parallel to the surface of the neck, at least to the level of the carotid bifurcation [19]. Generally, it is classified into three segments during ultrasound examination, each approximately 1 cm in length: common carotid as the most proximal segment, the 1-cm straight segment of the extracranial carotid artery immediately prior to the bifurcation. Its distal boundary is identified at where it begins to divide into its internal and external ICAs. Of these anatomic segments, the easiest to examine by ultrasound is the CCA, making it a favorable target to perform ultrasonographic measurements. Polak et al. [20] have concluded that carotid IMT measurements made in the CCA, carotid artery bulb and ICA were all associated with cardiovascular risk factors. According to their findings, they were also noted that while some cardiovascular risk factors showed qualitatively stronger associations with IMT measured in the bulb or internal carotid artery, the common carotid artery IMT best reflected overall exposure to traditional cardiovascular risk factors. O'Leary and Bots [19] indicated that the carotid artery has proven to be an extremely complex vessel, with differing associations for each segment to risk factors and outcomes, complicating the search for a standard imaging protocol that best meets every clinical and investigational requirement.

Carotid IMT is a well-known surrogate finding of atherosclerosis and can be used both to detect an accelerated disease process with clinical findings and subclinical conditions [21, 22]. Furthermore, carotid IMT can be used as a surrogate endpoint to monitor the efficacy of therapy against atherosclerosis in clinical practice [23, 24]. Cobble and Bale [25] noted that advantages of carotid IMT were that it was noninvasive, relatively inexpensive, and can be repeatedly performed with no adverse effects on the patient. They have concluded that carotid IMT was associated with cardiovascular disease and was an independent predictor of stroke and myocardial infarction. They have indicated that carotid IMT was currently limited by the lack of standardized protocols that may affect reproducibility from measurement to measurement.

Grau et al. [26] were determined their population reference ranges of carotid IMT for individuals aged 35-84 years in Spain and analyzed the association of carotid IMT with cardiovascular risk factors (age, smoking, diabetes, pulse pressure, lipid profile, and body mass index). They concluded that the main determinants of carotid IMT were age and pulse pressure.

Aldridge et al. [27] were noted that the training of nonsonographer physicians was required to practice carotid IMT and plaque screening by ultrasound for the assessment of subclinical atherosclerosis. They have conducted a study to assess the effect of formal training on carotid IMT evaluation and plaque detection by medical residents. They have found that medical residents performed carotid IMT measurement and plaque visualization to detect subclinical atherosclerosis in a successful manner.

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Ultrasonography with Doppler modality is the method of choice for the noninvasive evaluation of carotid arteries [28, 29]. The degree of carotid artery stenosis is mainly depends on the measurement of the PSV or EDV, or both, of the carotid arteries [30]. AbuRahma et al. [30] have validated the ultrasonographic imaging consensus criteria published in 2003 and have analyzed 376 carotid arteries. They have found that the consensus criteria used a PSV of 125 to 230 cm/s for detecting angiographic stenosis of 50% to 69%, which had a sensitivity of 93%, specificity of 68%, and overall accuracy of 85% and that a PSV of ≥ 230 cm/s for $\geq 70\%$ stenosis had a sensitivity of 99%, specificity of 86%, and overall accuracy of 95%. They have indicated that the consensus criteria for diagnosing 50% to 69% stenosis could be significantly improved by using an ICA PSV of 140 to 230 cm/s, with a sensitivity of 94%, specificity of 92%, and overall accuracy of 92%. They have concluded that the consensus criteria could be accurately used for diagnosing $\geq 70\%$ stenosis; however, the accuracy could be improved for detecting 50% to 69% stenosis if the ICA PSV was changed to 140 to < 230 cm/s.

During comparison of findings of color Doppler ultrasound examinations, differences in equipment, abilities, and experiences of ultrasonographers or radiologists, could cause considerable variability from laboratory to laboratory [28]. Therefore, each laboratory must adapt a method according to its equipment and validate their method when using Carotid Duplex Consensus criteria [29]. As mentioned in the review by Oates et al. [31] including Joint Recommendations for Reporting Carotid Ultrasound Investigations in the United Kingdom, there is need to create general agreement of acquisition, interpretation and reporting of the ultrasound and Doppler ultrasound data related to carotid arteries.

Several potential limitations of our study need to be considered. This study used a small sample size, especially in the obesity class III group. Our study groups were not similar with respect to sex in the obesity class III group and there was a significant male preponderance in the obesity class III group. Due to the small number of patients, sex adjusted analysis could not be conducted. Because of these limitations mentioned, we could not found meaningful differences among the obesity groups with regard to the IMTs, PSVs, and EDVs of ICAs, especially, in accordance with the severity of obesity as expected.

In conclusion, according to our findings, obesity may have a potential to increase the lumen diameter of CCAs and ICAs, to increase the IMTs of CCA but not ICA, and to decrease the PSVs and EDVs of CCA but not ICA. There is a need to determine the effect of obesity on the wall and function of carotid arteries in further studies in a broader population subclassified according to the severity and duration of obesity.

Conflicts of interest

The authors declare that no scientific and/or financial conflicts of interest exists with other people or institutions.

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