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NECK CIRCUMFERENCE MEASUREMENT: A SIMPLE METHOD TO ASSESS THE RISK OF CARDIOVASCULAR DISEASE IN TYPE 2 DIABETES MELLITUS

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ABSTRACT

Adipose tissue distribution and biochemical parameters of the type 2 diabetes mellitus (T2DM) patients have been investigated in association to cardiovascular disease (CVD) risk factors. Neck circumference (NC) shown to have a strong association with CVD. The aim of this study is to assess the relation between NC measurement and clinical characteristics involved in risk of CVD in T2DM. A case-control and cross sectional study were conducted with 100 T2DM patients and 100 healthy individuals of both male and female with 30-60 years of age. NC was measured using a flexible tape at the upper margin of the laryngeal prominence, HbA1C was estimated by immunoturbidimetry method, serum cholesterol was estimated by cholesterol oxidase, Triglycerides by Glycerol phosphate oxidase/ peroxidase method, HDL-C by precipitation method, high sensitive C reactive protein (hsCRP) was estimated by latex immunoturbidimetry method and Homocysteine was estimated by enzymatic colorimetric method. Significant difference was observed in mean levels of body mass index (BMI), NC, waist circumference (WC), diastolic blood pressure (DBP), systolic blood pressure (SBP), fasting blood glucose (FBS), total cholesterol (TC), Triglycerides (TGL), high density lipoprotein (HDL-C), Glycated hemoglobin HbA1C, C reactive hsCRP and homocysteine (Hcy) of T2DM patients when compared with healthy controls (p<0.05). NC was positively correlated with BMI, DBP, WC, TGL and Hcy (p<0.05). NC as a novel indicator of CVD showed good predictive ability for adverse CVD events in T2DM individuals.

KEY WORDS: obesity, neck circumference, inflammatory marker.

INTRODUCTION

Body obesity is related with an elevated risk of developing metabolic syndrome (MS), T2DM, and hypertension and is consider as a major risk factor for CVD events and mortality. Excessive adiposity is involved in the pathogenesis of coronary heart disease (CHD), as it is related with the occurrence of conditions like dyslipidiemia, hypertension and T2DM¹. The distribution of body weight, in addition to obesity consider as a vital aspect of MS, and is a great predictor of cardiovascular risk. The accumulation of body fat in particular regions is associated to disease risk². Generally, subcutaneous and visceral fat are related with IR³. Visceral adipose tissue is a distinctive pathogenic fat deposit that is highly implicated in the pathogenesis of T2DM, and ischemic heart disease (IHD)⁴. Upper-body subcutaneous adipose tissue, evaluated by neck circumference (NC), is a fat depot that might be additional risk for metabolic risk factors over central and general adiposity⁵ and has shown a positive correlation with biochemical parameters of the MS⁶. NC is a strong marker to identify obese persons and correlates with various anthropometric measurements. NC has been correlated with cardio-metabolic risk factors apart from the relationship with other adiposity measurements $^{7, 8}$. NC is a simple and convenient anthropometric measurement, which is correlates with WC

and BMI and has been related with components of MS in and cohort studies^{9, 10} and cross-sectional ^{11, 12} in different populations. The relation between neck fat and MS and its components might be pertaining to an excess release of fatty acids from upper body subcutaneous fat into plasma⁸. High concentration of plasma free fatty acids, in turn related with oxidative stress markers and IR², which may impact glycemia. Fat in the neck region may be highly similar to visceral fat, which is strongly associated with cardiometabolic risks when compared with subcutaneous fat⁸. An increase in NC found to be an independent predictor of nonalcoholic fatty liver disease also¹³. The aim of this study is to observe the relation between NC and clinical characteristics involved in risk of CVD in T2DM.

MATERIALS AND METHODS

A case-control and cross sectional study was conducted with 100 T2DM patients and 100 healthy individuals of both male and female with 30-60 years of age. NC (cm) was measured using a flexible tape positioned horizontally at the upper margin of the laryngeal prominence. Height and body weight were measured with participants standing without footwear. Body weight was measured to the nearest 0.1 kg and height to the nearest 0.5 cm. BMI was calculated as weight (in kilograms) divided by height (in

meters square). WC was the minimum abdominal girth measured to the nearest 0.1 cm. BP was measured by using a mercury sphygmomanometer after the subject had rested for at least 10 min. SBP 140 mmHg, DBP 90 mmHg and participants who were on antihypertensive drugs was considered as hypertensive patients. Overnight fasting venous blood samples were collected from all subjects. The blood was transferred into glass tubes without anticoagulant and allowed to clot at room temperature. But for HbA1c estimation 0.5 ml of blood was transferred separately into glass tube containing EDTA. The serum was obtained by centrifugation of blood at 3500 rpm for 10 min. Blood glucose as estimated by glucose oxidase and peroxidase¹⁴, HbA1C was estimated by immunoturbidimetry method¹⁵, Cholesterol by cholesterol oxidase¹⁶, Triglycerides by Glycerol phosphate oxidase/peroxidase method¹⁷, HDL-C by precipitation method¹⁸, hsCRP was estimated by latex immunoturbidimetry method¹⁹ and Homocysteine was measured by enzymatic colorimetric method²⁰. Data was

expressed as mean \pm standard deviation (SD). Comparisons between two groups were performed by unpaired Student t-test. Correlation was performed by Pearson's calculation. A statistical analysis was performed by SPSS (Statistical Package for Social Sciences) software 17 trail version. *p* value <0.05 was considered as statistically significant.

RESULTS

The mean levels of BMI, NC, WC, diastolic blood pressure (DBP), systolic blood pressure (SBP), fasting blood glucose (FBS), Cholesterol, Triglycerides (TGL), Glycated hemoglobin HbA1C, high sensitive C reactive protein (hsCRP) and homocysteine (Hcy) were higher but high density lipoprotein (HDL-C) levels were lower in T2DM patients than healthy controls (p<0.05). Table 2 shows the correlation between NC and various clinical parameters. There was significant positive correlation was found between NC and BMI, DBP, WC, TGL and Hcy (p<0.05).

IABLE1. Clinical characteristics of all the study population				
Clinical characteristics	T2DM	Healthy control	p value	
BMI (kg/m^2)	26.4 ± 4.8	24.6 ± 3.7	< 0.05	
SBP (mmHg)	128.4 ± 14.1	115.6 ± 7.2	< 0.05	
DBP (mmHg)	82.8 9.0	76.2 ± 5.6	< 0.05	
NC (cm)	38.1 ± 3.7	36.7 ±2.9	< 0.05	
WC (cm)	98.2 ± 13.3	89.5 ±9.3	< 0.05	
FBS (mg/dl)	$188.0 \pm \! 55.0$	85.7 ± 8.2	< 0.05	
TC (mg/dl)	196.9 ± 41.8	179.8 ± 27.1	< 0.05	
TGL (mg/dl)	210.5 ± 112.7	140.1 ± 37.1	< 0.05	
HDL-C (mg/dl)	45.4 ± 5.2	46.8 ± 6.8	< 0.05	
HbA1C (%)	8.71 ± 1.4	5.16 ±0.34	< 0.05	
hsCRP (mg/L)	1.1 ± 0.7	0.48 ± 0.3	< 0.05	
Hcy (µmol/L)	14.12 ± 7.1	9.71 ±2.5	< 0.05	

TABLE1. Clinical characteristics of all the study population

Data are shown as mean \pm SD, maximum and minimum levels. *p* value was calculated by student, t-test

p <0.05 was consider as statistically significant

TABLE 2. Simple correlation of NC with various parameters

	r	р
WC (cm)	0.608	< 0.05
BMI (kg/m ²)	0.553	$<\!0.05$
DBP (mmHg)	0.201	$<\!0.05$
TGL (mg/dl)	0.239	< 0.05
Hcy (µmol/L)	0.405	< 0.05

p < 0.05 was consider as statistically significant

DISCUSSION

In the current study an attempt was made to investigate the relation between NC and various parameters for cardiovascular disease risk. This study had shown significant positive relation between NC and WC. This result were consistent with a study conducted with a total of 3307 adults aged 20-65 years from two communities of Tongzhou, Beijing²¹ and a cross-sectional study was conducted with 350 T2DM and 350 non-diabetics of >30 years of age²². This study had shown a significant positive relation between NC and BMI, This result were consistent with longitudinal cohort study with 364 subjects who were

not on major medical conditions and not receiving any medication¹⁰ and a study with total of 3,182 diabetic subjects with 20–80 years of age were from 15 different community health centers in Beijing⁹. There was a significant positive relation between NC and DBP in the present study and the similar results were also reported in a community-based cross-sectional study of descriptive design where 500 students from urban and rural areas of northern Tamilnadu²³ and in meta-analysis, that examined the association of NC with risk of MS, or at minimum, one of its components as outcomes²⁴. Significant positive relation was found between NC and TGL and this result was consistent with cross-sectional study in 1053 Brazilian

adults (18-60 years) with BMI 18.5-40.0kg/m², with normal glucose tolerance or T2DM¹¹ and a study from Calcutta from India with total of 451 individuals, 30–80 years age, without any co-morbid state who gave informed written consent underwent clinical, anthropometric, and biochemical assessment²⁵. To the best of my knowledge this is the first study investigated the relationship between NC and Hcy in T2DM patients of Andhra Pradesh, India and shown positive significant correlation. As Hcy is one of the marker for inflammation status and there is role development of CVD.

Brazilian population-based study showed that NC was correlated with fasting glucose levels, high TGL, and low high-density lipoprotein (HDL)-cholesterol levels, and insulin resistance index¹¹. NC was associated with metabolic syndrome more strongly than WC in Turkish population⁶. Framingham Heart Study offspring participants shown NC was related with increased carotid intima-media thickness (IMT) but neither BMI / WC was associated²⁶. Several mechanisms reported that underlying the relation between large NC and impaired glucose metabolism. Higher NC levels alter peripheral blood flow and results to endothelial function²⁷ which may decrease insulin delivery and promote insulin resistance in the whole body²⁸.

In the Heinz Nixdorf Recall Study, epicardial fat associated with nonfatal and fatal coronary events, as well as CVD risk factors and occurrence of myocardial infarction in the general population²⁹. Not only accumulation of fat in visceral region but also in other regions like pericardial, liver, muscle, even perirenal and perivascular areas was associated with heart function and CVD ^{30,31,32,33,34}. Free fatty acid secreted from upper-body subcutaneous fat was reported to be more than that from subcutaneous fat form lowerbody³⁵. Framingham Heart Study reported that NC is associated with CVD risk factors even after adjustment for BMI and visceral adipose tissue, and considered NC to be a measure of cardio metabolic risk^{8.} In the Beijing Community Diabetes Study4, NC was positively correlated with the MS in T2DM Chinese population⁹. In other study NC was significantly correlated with all outcomes of cardiometabolic risk in males and females¹². Cardio metabolic Risk study with healthy adults, significant associations were observed between high NC and elevated risk of IR and several CVD risk factors in Chinese individuals³⁶. Dia et al.³⁷ reported that a higher NC indicated a higher incidence of future fatal and non-fatal CVD events and allcause mortality in both male and female high-risk participants. CVD risk factors increased more in the higher NC group. NC as a novel indicator of CVD showed good predictive ability for CVD events and mortality in a highrisk population.

CONCLUSION

Raised Neck circumference shown relationship with CVD risk factors such as high levels of BMI, TGL, and also with elevated levels of inflammatory marker such as homocysteine. So, neck circumference measurement may be considered as a marker to asses CVD risk in T2DM.

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REFERENCES

- [1]. Logue, J., Murray, H.M., Welsh, P., Shepherd, J., Packard, C., Macfarlane, P. (2011) Obesity is associated with fatal coronary heart disease independently of traditional risk factors and deprivation. *Heart* 2011; 97:564–568.
- [2]. Santosa, S., Jensen, M.D. Why are we shaped differently, and why does it matter? American journal of physiology. *Endocrinology and metabolism* 2008; 295: E531–535
- [3]. Ramachandran, A., Wan, Ma R.C., Snehalatha, C. Diabetes in Asia. *The Lancet* 2010; 375:408–418.
- [4]. Kuk, J.L., Katzmarzyk, P.T., Nichaman, M.Z., Church, T.S., Blair, S.N., Ross, R.. Visceral fat is an independent predictor of all-cause mortality in men. *Obesity* 2006; 14:336–341.
- [5]. Grunfeld, C., Rimland, D., Gibert, C.L., Powderly, W.G., Sidney, S., Shlipak, M.G. Association of upper trunk and visceral adipose tissue volume with insulin resistance in control and HIV-infected subjects in the FRAM study. *J Acquir Immune Defic Syndr* 2007;46: 283–290.
- [6]. Onat, A., Hergenç, G., Yüksel, H., Can, G., Ayhan, E., Kaya, Z. Neck circumference as a measure of central obesity: associations with metabolic syndrome and obstructive sleep apnea syndrome beyond waist circumference. *Clinical nutrition* 2009; 28: 46–51.
- [7]. Fitch, K.V., Stanley, T.L., Looby, S.E., Rope, A.M., Grinspoon, S.K. Relationship between neck circumference and cardiometabolic parameters in HIV-infected and non–HIV-infected adults. *Diabetes care* 2011; 34: 1026–1031.
- [8]. Preis, S.R., Massaro, J.M., Hoffmann, U. Neck circumference as a novel measure of cardiometabolic risk: the Framingham Heart study. J Clin Endocrinol Metab 2010; 95:3701–3710
- [9]. Yang, G.R., Yuan, S.Y., Fu, H.J. Neck circumference positively related with central obesity, overweight, and metabolic syndrome in Chinese subjects with type 2 diabetes: Beijing Community Diabetes Study 4. *Diabetes Care* 2010;33(11):2465–2467
- [10]. Ben-Noun, L., Laor A. Relationship between changes in neck circumference and cardiovascular risk factors. *Experimental and Clinical Cardiology* 2006; 11(1):14–20.
- [11]. Stabe, C., Vasques, A.C., Lima, M.M., Tambascia, M.A., Parej,a J.C., Yamanaka, A. Neck circumference as a simple tool for identifying the metabolic syndrome and insulin resistance: results from the Brazilian Metabolic Syndrome Study. *Clinical Endocrinology* 2013;78(6):874–881.
- [12]. Zhou, J. Neck circumference as an independent predictive contributor to cardio-metabolic syndrome. *Cardiovasc Diabetol* 2013;12:76

- [13]. Huang, B.X., Zhu, M.F., Wu, T. Neck circumference, along with other anthropometric indices, has an independent and additional contribution in predicting fatty liver disease. *PLoS ONE*. 2015; 10(2):e0118071
- [14]. Trinder, P. Determination of glucose in blood using glucose oxidase with an alternative oxygen receptor. *Ann Clin Biochem* 1969; 6(1):24-27
- [15]. Sachs, D.B. Carbohydrates. In: Burtis CA, Ashwood ER, editors. Tietz Textbook of Clinical Chemistry. 3rd ed. Philadelphia: WB Saunders Company; 1999:790-796.
- [16]. Allain, C.C., Poon, L.S., Chan, C.S., Richmond, W., Fu.P.C. Enzymatic determination of total serum cholesterol. *Clin Chem* 1974; 20(4):470-475.
- [17]. McGowan, M.W., Artiss, J.D., Strandbergh, D.R., Zak, B. A peroxidase-coupled method for the colorimetric determination of serum triglycerides. *Clin Chem* 1983; 29(3):538-542.
- [18]. Dominiczak, M., McNamara, J. The system of cardiovascular prevention. 103–125; Nauk M, Wiebe D, Warnick G. Measurement of High-Density-Lipoprotein Cholesterol. 221–244. In: Handbook of Lipoprotein Testing (eds. Rifai, Warnick and Dominiczak), 2nd edition
- [19]. Eda, S., Kaufmann, J., Roos, W., Pohl, S. Development of a new micro particle enhanced turbidimetric assay for C-reactive protein with superior features in analytical sensitivity and dynamic range. J Clin Lab Anal 1998; 12:137-44.
- [20]. Refsum, H., Macko, R.F., Kittner, S.J., Ivey, F.M., Epstein, A., Sparks, M.J. Effects of vitamin therapy on plasma total homocysteine, endothelial injury markers, and fibrinolysis in stroke patients. *J Stroke Cerebrovasc Dis* 2002; 11(1):01-08.
- [21]. Wang, X., Zhang, N., Yu, C., Ji, Z. Evaluation of neck circumference as a predictor of central obesity and insulin resistance in Chinese adults. *Int J Clin Exp Med* 2015; 8(10): 19107–19113.
- [22]. Aswathappa, J., Garg, S., Kutty, K., Shankar, V. Neck Circumference as an Anthropometric Measure of Obesity in Diabetics. *N Am J Med Sci* 2013; 5(1): 28–31.
- [23]. Rajagopalan, A., Balaji, N. Association of neck circumference and obesity with blood pressure among adolescents in urban and rural population in North Tamilnadu. J Nat Sc Biol Med 2017; 8:144-149.
- [24]. Namazi, N., Larijani, B., Surkan, P.J., Azadbakht, L. The association of neck circumference with risk of metabolic syndrome and its components in adults: A systematic review and meta-analysis. *Nutrition, Metabolism and Cardiovascular Diseases*; 21 March 2018. (IN PRESS]
- [25]. Selvan, C., Dutta, D., Thukral, A., Nargis, T., Kumar M., Mukhopadhyay, S., Chowdhury, S. Neck height ratio is an important predictor of metabolic syndrome among Asian Indians. *Indian J Endocrinol Metab* 2016; 20(6):831-837.

- [26]. Rosenquist KJ, Massaro JM, Pencina KM, D'Agostino RB, Beiser A, O'Connor GT, O'Donnell CJ, *et al.* Neck circumference, carotid wall intimamedia thickness, and incident stroke. *Diabetes Care* 2013; 36: e153–e154.
- [27]. Chung S. Yoon IY, Shin YK, Lee CH, Kim JW, Lee T, Choi DJ, Ahn HJ. Endothelial dysfunction and Creactive protein in relation with the severity of obstructive sleep apnea syndrome. *Sleep* 2007; 30: 997–1001.
- [28]. Hsueh WA, Quinones MJ. Role of endothelial dysfunction in insulin resistance. Am J Cardiol2003; 921:0J–17J.
- [29]. Mahabadi AA, Berg MH, Lehmann N, Kälsch H, Bauer M, Kara K, et al. Association of epicardial fat with cardiovascular risk factors and incident myocardial infarction in the general population: the Heinz Nixdorf Recall Study. *Journal of the American College of Cardiology* 2013;61: 1388–1395.
- [30]. Lim S. Ectopic Fat Assessment Focusing on Cardiometabolic and Renal Risk. *Endocrinology and Metabolism* 2014; 29:1–4.
- [31]. Wang CY, Ou HY, Chen MF, Chang TC, Chang CJ. Enigmatic Ectopic Fat: Prevalence of Nonalcoholic Fatty Pancreas Disease and Its Associated Factors in a Chinese Population. *Journal of the American Heart* Association 2014; 3:e000297.
- [32]. Okauchi Y, Nishizawa H, Funahashi T, Ogawa T, Noguchi M, Miwa Ryo M, Kihara S, et al. Reduction of visceral fat is associated with decrease in the number of metabolic risk factors in Japanese men. *Diabetes Care* 2007; 30:2392–2394.
- [33]. Hiuge-Shimizu A, Kishida K, Funahashi T, Ishizaka Y, Oka R, Okada M, Suzuki S, et al. Reduction of visceral fat correlates with the decrease in the number of obesity-related cardiovascular risk factors in Japanese with Abdominal Obesity (VACATION-J Study) Journal of atherosclerosis and thrombosis 2011;19:1006–1018.
- [34]. Neeland IJ, Gupta S, Ayers CR, Turer AT, Rame JE, Das SR, et al . Relation of regional fat distribution to left ventricular structure and function. Circulation: *Cardiovascular Imaging* 2013; 6: 800–807.
- [35]. Guo, Z., Hensrud, D.D., Johnson, C.M., Jensen, M.D. Regional postprandial fatty acid metabolism in differentobesity phenotypes. *Diabetes* 1999; 48:1586– 1592
- [36]. Liang, J., Teng, F., Li, Y., Liu, X., Zou, C., Wang, Y., Li, H, Lu, Q. Neck Circumference and Insulin Resistance in Chinese Adults: The Cardiometabolic Risk in Chinese (CRC) Study. *Diabetes Care* 2013; 36: e145–e146.
- [37]. Dai, Y., Wan, X., Li, X., Jin, E., Li, X. Neck circumference and future cardiovascular events in a high-risk population—A prospective cohort study. *Lipids Health Dis* 2016; 15: 46.