## RESEARCH



# The correlation of triglyceride/high-density lipoprotein cholesterol ratio with muscle mass in type 2 diabetes patients



Qingsong Fu<sup>1+</sup>, Zhenwen Zhang<sup>2+</sup>, Wenchao Hu<sup>3</sup> and Yinrong Yang<sup>4\*</sup>

## Abstract

**Objective** Triglyceride/high-density lipoprotein cholesterol (TG/HDL-C) ratio is correlated with metabolic diseases. The prevalence of sarcopenia is significantly higher in type 2 diabetes mellitus (T2DM) patients compared with healthy controls. The purpose of our study is to evaluate the correlation of TG/HDL-C ratio with muscle mass in T2DM patients.

**Method** Our study consists of 1048 T2DM inpatients recruited from the department of endocrinology. Skeletal muscle index (SMI) was detected with a dual energy X-ray absorptiometry method. Low muscle mass was diagnosed using the criteria of SMI less than 7.0 kg/m<sup>2</sup> (in male subjects) or 5.4 kg/m<sup>2</sup> (in female subjects).

**Result** The prevalence of low muscle mass was 20.9% and 14.5% in male and female groups respectively. SMI was correlated with TG/HDL ratio after adjustment for age, duration of diabetes, diastolic blood pressure (DBP), and HbA1c in male subgroup. In female subgroup, SMI was associated with TG/HDL ratio after adjustment for age and DBP.

**Conclusion** Higher TG/HDL-C ratio is correlated with muscle mass in T2DM patients.

Keywords Triglyceride/high-density lipoprotein cholesterol ratio, Skeletal muscle index, Type 2 diabetes mellitus

<sup>†</sup>Qingsong Fu and Zhenwen Zhang contribute equally to this article.

## \*Correspondence:

Yinrong Yang

yangyinrong1207@126.com

<sup>1</sup>Department of Central Laboratory and Mitochondrial Medicine Laboratory, Qilu Hospital (Qingdao), Cheeloo College of Medicine, Shandong University, Qingdao, China

<sup>2</sup>Department of Cadre Health Care, Qilu Hospital (Qingdao), Cheeloo

College of Medicine, Shandong University, Qingdao, China <sup>3</sup>Department of Endocrinology, Qilu Hospital (Qingdao), Cheeloo College

of Medicine, Shandong University, Qingdao, China <sup>4</sup>Department of Laboratory, Qilu Hospital (Qingdao), Cheeloo College of

Medicine, Shandong University, 758 Hefei Road, Shibei District, Qingdao 266035, China

## Introduction

Sarcopenia is defined as a disease with the characteristics of progressive loss of skeletal muscle mass and strength [1]. Sarcopenia is demonstrated to increase the risk of disabilities, infection, metabolic disorders, falls and fractures, and mortality [2]. Traditional factors including aging, physical inactivity, and malnutrition are correlated with the development of sarcopenia [3]. Type 2 diabetes mellitus (T2DM), one of the most common metabolic diseases, is reported to significantly increase the risk of developing sarcopenia compared with control subjects in Korean population [4]. The prevalence of sarcopenia was dramatically higher in the T2DM patients compared with healthy controls [5].

Dyslipidemia, represented by elevated blood lowdensity lipoprotein cholesterol (LDL), decreased



© The Author(s) 2023. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/. The Creative Commons Public Domain Dedication waiver (http://creativecommons.org/publicdomain/zero/1.0/) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

high-density lipoprotein cholesterol (HDL), and elevated blood triglyceride (TG) levels, is a well-known marker for metabolic syndrome [6], T2DM [7], and cardiovascular disease [8]. Among lipid profiles, the TG/HDL ratio has emerged as a good predictive index for insulin resistance [9], diabetes [10], and cardiovascular diseases [11]. We hypothesized that higher TG/HDL-C ratio may be correlated with muscle mass.

Therefore, the purpose of our study is to evaluate the correlation of higher TG/HDL-C ratio with muscle mass in T2DM patients.

## **Materials and methods**

## Patients

Our study was cross-sectional designed and enrolled 1048 T2DM inpatients recruited from the Department of Endocrinology of our hospital from September of 2017 to September of 2019. T2DM was diagnosed according to the American Diabetic Association criteria with a fasting glucose level $\geq$ 7.0 mmol/L or 2-hour postprandial plasma glucose level $\geq$ 11.1 mmol/L. Inclusion criterion was age $\geq$ 20 year. Patients who were pregnant, had infectious diseases, cancer, severe hip or knee osteoarthritis, and a history of stroke were excluded. This study was approved by the Hospital ethics board and all patients provided written informed consent.

## Low muscle mass definition

A dual energy X-ray absorptiometry (Hologic Discovery A, Waltham, MA, USA) was utilized to detect the skeletal muscle index (SMI). SMI was calculated with the formula of appendicular skeletal muscle mass in kg divided by the square of the body height. Low muscle mass was diagnosed using the criteria of SMI less than 7.0 kg/m<sup>2</sup> (in male subjects) or 5.4 kg/m<sup>2</sup> (in female subjects) [12].

## Measurements

Information of height, weight and blood pressures, the duration of diabetes, comorbidity disease history, and medications were recorded. Body mass index (BMI) was computed as weight in kilograms divided by height squared in meters  $(kg/m^2)$ . Blood was obtained from all the subjects after an overnight fasting.

## Statistical analysis

Data are displayed as means  $\pm$  SD. Chi-square tests and unpaired t test were utilized to compare the statistical significance of the differences between T2DM patients with and without low muscle mass. Data were analyzed by univariate simple and multiple linear regression models looking for significant associations between SMI or TG/HDL ratio and other variables. A *P* value of less than 0.05 was considered as statistically meaningful.

## Results

## The differences between subjects with and without low muscle mass

As shown in Table 1, the prevalence of low muscle mass was 20.9% and 14.5% in male and female groups respectively. In male subjects, low muscle mass group showed higher age, HDL, HbA1c, and percentage of sulfonylureas treatment, as well as lower BMI, SMI, diastolic blood pressure (DBP), TG, and TG/HDL ratio compared with normal muscle mass group. In female subjects, age was increased, whereas BMI, SMI, TG, TG/HDL ratio, and percentage of metformin treatment were decreased in low muscle mass group compared with normal muscle mass group.

## The association between SMI and other characteristics

As shown in Table 2, SMI was correlated with age, duration of diabetes, DBP, TG, HDL, HbA1c, and TG/HDL ratio in male subjects after simple linear regression analysis. Age, DBP, HbA1c, and TG/HDL ratio were still correlated with SMI after a multiple linear regression analysis.

In female subjects, SMI was correlated with age, DBP, and TG/HDL ratio after simple linear regression analysis (Table 3). Multiple linear regression analysis showed that age, DBP, and TG/HDL ratio were still correlated with SMI (Table 3).

## The association between TG/HDL ratio and other characteristics

Simple linear regression analysis showed that TG/HDL ratio was correlated with age, duration of diabetes, BMI, DBP, total cholesterol, LDL, and fasting plasma glucose (FPG) in male subjects (Table 4). Age, duration of diabetes, BMI, and FPG were still correlated with TG/HDL ratio after a multiple linear regression analysis (Table 4).

As shown in Table 5, TG/HDL ratio was correlated with FPG and HbA1c in female subjects after simple linear regression analysis. Multiple linear regression analysis showed that FPG was still correlated with TG/HDL ratio.

## Discussion

Our investigation indicated higher TG/HDL-C ratio is negatively correlated with muscle mass in T2DM patients. Wang also demonstrated that TG/HDL-C ratio was negatively associated with sarcopenia occurrence rate in community-dwelling Chinese adults [13]. However, other studies performed in Korea and Japan reported inconsistent results. The prevalence of low muscle mass significantly increased in accordance with TG/ HDL ratio quartiles in elderly Korean males [14]. The atherogenic dyslipidemia ratio [log(TG)/HDL-C] was significantly related to skeletal sarcopenia in T2DM females of Japan [15]. The reason for these conflicting data is

Table 1         The characteristic differences between T2DN	patients with and without low muscle mass
---	---

Characteristics	Male (n = 558)			Female (n = 490)		
	Normal muscle	Low muscle mass	P value	Normal muscle	Low muscle mass	Р
	mass (n=441)	(n = 117)		mass (n=419)	(n=71)	value
Age (years)	54.73±12.07	60.78±13.53	< 0.001	$61.13 \pm 10.97$	65.52±11.35	0.002
Duration (years)	$8.04 \pm 6.82$	$8.93 \pm 5.79$	0.192	$8.65 \pm 5.97$	$9.24 \pm 6.75$	0.446
BMI (kg/m <sup>2</sup> )	$27.4 \pm 4.74$	$23.58 \pm 5.43$	< 0.001	$27.14 \pm 5.03$	$22.17 \pm 4.1$	< 0.001
SMI	$8.19 \pm 1.54$	$6.45 \pm 0.48$	< 0.001	$6.5 \pm 0.89$	$5.07 \pm 0.33$	< 0.001
SBP (mmHg)	$140.83 \pm 19.96$	$138.13 \pm 20.75$	0.197	143.23±20.87	143.07±22.11	0.953
DBP (mmHg)	82.61±13	78.97±12.85	0.007	76.54±12.87	$76.45 \pm 11.16$	0.958
TG (mmol/L)	$1.83 \pm 1.08$	1.39±0.77	< 0.001	$1.72 \pm 0.94$	$1.39 \pm 0.69$	0.006
TC (mmol/L)	$4.44 \pm 1.04$	$4.45 \pm 1.08$	0.912	$4.65 \pm 1.16$	4.6±1.17	0.753
LDL (mmol/L)	$2.97 \pm 0.9$	$2.97 \pm 0.95$	0.962	$3.01 \pm 0.92$	$2.89 \pm 0.99$	0.319
HDL (mmol/L)	$1.14 \pm 0.26$	$1.22 \pm 0.3$	0.005	1.28±0.32	$1.36 \pm 0.35$	0.051
FPG (mmol/L)	$8.02 \pm 2.65$	$7.94 \pm 3.17$	0.766	$7.74 \pm 2.9$	$7.04 \pm 2.76$	0.075
HbA1c (%)	$8.41 \pm 2.1$	$8.86 \pm 2.09$	0.043	$8.29 \pm 2.02$	8.19±2.13	0.697
TG/HDL ratio	$1.74 \pm 1.15$	$1.26 \pm 0.87$	< 0.001	1.49±0.16	1.12±0.7	0.011
Cardiovascular disease (n, %)	105 (23.8%)	34 (29.1%)	0.243	156 (37.2%)	30 (42.3%)	0.42
Renal disease (n, %)	9 (2%)	2 (1.7%)	0.819	15 (3.6%)	3 (4.2%)	0.789
Pulmonary disease (n, %)	32 (7.3%)	9 (7.7%)	0.872	13 (3.1%)	4 (5.6%)	0.281
Treatment						
Metformin (n, %)	302 (68.5%)	75 (64.1%)	0.335	302 (72.1%)	42 (59.2%)	0.022
Acarbose (n, %)	150 (34%)	41 (35%)	0.86	166 (39.6%)	24 (33.8%)	0.337
Sulfonylureas (n, %)	105 (23.8%)	44 (37.6%)	0.003	127 (30.3%)	21 (29.6%)	0.872
DPP-IV inhibitor (n, %)	87 (19.7%)	32 (27.4%)	0.078	75 (17.9%)	12 (16.9%)	0.825
Insulin (n, %)	167 (37.9%)	44 (37.6%)	0.959	162 (38.7%)	20 (33.8%)	0.435
Statin (n, %)	123 (27.9%)	37 (31.6%)	0.603	138 (32.9%)	28 (39.4%)	0.597

**Table 2** The association between clinical characteristics and SMI in male subjects

Table 3	The association	between	clinical	characteristics and SMI
in female	e subiects			

	simple regression analysis		multiple regression analysis	
	β (95% Cl)	P value	β (95% CI)	Р
				value
Age (years)	-0.27 (-0.037, -0.017)	< 0.001	-0.023 (-0.035, -0.012)	< 0.001
Duration (years)	-0.02 (-0.04, -0.001)	0.04	-0.007 (-0.027, 0.014)	0.529
SBP (mmHg)	0.005 (-0.002, 0.011)	0.137		
DBP (mmHg)	0.017 (0.007, 0.027)	0.001	0.01 (0.000, 0.021)	0.049
TG (mmol/L)	0.29 (0.167, 0.413)	< 0.001	-	-
TC (mmol/L)	0.028 (-0.096, 0.151)	0.659		
LDL (mmol/L)	0.033 (-0.109, 0.176)	0.645		
HDL (mmol/L)	-0.561 (-1.035, -0.088)	0.02	-	-
FPG (mmol/L)	0.013 (-0.034, 0.059)	0.596		
HbA1c (%)	-0.071 (-0.136, -0.006)	0.032	-0.091 (-0.155, -0.028)	0.005
TG/HDL ratio	0.262 (0.148, 0.377)	< 0.001	0.177 (0.054, 0.3)	0.005

unclear but may be due to differences in disease advancement, ethnic populations or assays applied.

Our results indicated that HDL was significantly higher in low muscle mass group of T2DM patients than in

	simple regression analysis		multiple regression analysis	
	β (95% Cl)	P value	β (95% Cl)	P value
Age (years)	-0.019 (-0.027, -0.012)	< 0.001	-0.017 (-0.025, -0.01)	< 0.001
Duration (years)	0.001 (-0.014, 0.014)	0.974		
SBP (mmHg)	0.003 (-0.001, 0.007)	0.191		
DBP (mmHg)	0.011 (0.004, 0.017)	0.002	0.007 (0.000, 0.014)	0.037
TG (mmol/L)	-0.005 (-0.079, 0.07)	0.903	-	-
TC (mmol/L)	-0.006 (-0.07, 0.058)	0.857		
LDL (mmol/L)	0.011 (-0.082, 0.103)	0.822		
HDL (mmol/L)	-0.386 (-0.651, -0.121)	0.004	-	-
FPG (mmol/L)	0.006 (-0.025, 0.037)	0.691		
HbA1c (%)	-0.009 (-0.053, 0.035)	0.691		
TG/HDL ratio	0.103 (0.026, 0.018)	0.009	0.084 (0.009, 0.159)	0.028

normal muscle mass group. HDL was negatively correlated with SMI in T2DM patients. Tuzun reported that muscle-related index was inversely associated with HDL [16]. A cross-sectional study performed in Brazil demonstrated that HDL was higher in T2DM patients with 
 Table 4
 The association between TG/HDL ratio and other clinical characteristics in male subjects

	simple regression analysis		multiple regression analysis	
	β (95% Cl)	P value	β (95% Cl)	P value
Age (years)	-0.022 (-0.029, -0.015)	<0.001	-0.014 (-0.022, -0.007)	< 0.001
Duration (years)	-0.026 (-0.04, -0.012)	<0.001	-0.018 (-0.032, -0.004)	0.009
BMI (kg/m²)	0.063 (0.046, 0.081)	<0.001	0.062 (0.044, 0.08)	< 0.001
SBP (mmHg)	0.004 (-0.001, 0.009)	0.084		
DBP (mmHg)	0.011 (0.004, 0.018)	0.002	0.000 (-0.007, 0.007)	0.91
TC (mmol/L)	0.134 (0.046, 0.222)	0.003	0.166 (-0.035, 0.368)	0.106
LDL (mmol/L)	0.137 (0.036, 0.239)	0.008	-0.167 (-0.4, 0.066)	0.16
FPG (mmol/L)	0.065 (0.032, 0.098)	<0.001	0.049 (0.016, 0.081)	0.003
HbA1c (%)	0.031 (-0.015, 0.076)	0.186		

**Table 5** The association between TG/HDL and other clinical characteristics in female subjects

	simple regression analysis		multiple regression analysis	
	β (95% Cl)	P value	β (95% Cl)	P value
Age (years)	-0.007 (-0.016, 0.002)	0.123		
Duration (years)	0.008 (-0.008, 0.024)	0.343		
BMI (kg/m <sup>2</sup> )	0.014 (-0.005, 0.034)	0.151		
SBP (mmHg)	0.001 (-0.003, 0.006)	0.581		
DBP (mmHg)	0.007 (0.000, 0.015)	0.066		
TC (mmol/L)	-0.013 (-0.099, 0.073)	0.766		
LDL (mmol/L)	0.027 (-0.079, 0.134)	0.617		
FPG (mmol/L)	0.088 (0.053, 0.122)	<0.001	0.072 (0.029, 0.115)	0.001
HbA1c (%)	0.087 (0.037, 0.137)	0.001	0.032 (-0.027, 0.092)	0.281

sarcopenia compared with non-sarcopenia group [17]. In addition, other investigators also reported higher HDL in that sarcopenia group showed than in non-sarcopenia group among T2DM patients; however, the difference was not statistically meaningful [18–20]. Previous investigations showed that T2DM patients with sarcopenia had lower TG levels than those without sarcopenia [17– 20]. Our investigation arrived at similar conclusions. This may be contradicted with the traditional beliefs. Obesity and hyperlipidemia are considered to be risk factors of developing metabolic disease such as diabetic complication and cardiovascular disorders. But when it comes to sarcopenia or low muscle mass, obesity and hyperlipidemia are associated with a lower risk of sarcopenia or low muscle mass development in T2DM patients. In addition, TG/HDL-C ratio may be utilized to be a biomarker to diagnose or predict low muscle mass in T2DM patients.

The present study had several limitations. First of all, the sample size of this study was relatively small. Secondly, this study was a cross-sectional study, which limited its causal conclusions. Causality must be assessed by further longitudinal researches.

## Conclusions

In conclusion, higher TG/HDL-C ratio is negatively correlated with low muscle mass in type 2 diabetes patients.

## Acknowledgements

No acknowledgments

### Authors' contributions

Yinrong Yang performed the design. Qingsong Fu collected the clinical data. Zhenwen Zhang and Wenchao Hu did the statistical work.

#### Funding

Funded by Qingdao Outstanding Health Professional Development Fund, and Qingdao Key Health Discipline Development Fund.

## Data Availability

The datasets used during the current study available from the corresponding author on reasonable request.

## Declarations

#### Ethics approval and consent to participate

This study was approved by the Hospital ethics board of Qilu Hospital (Qingdao) and all patients provided written informed consent. All methods were performed in accordance with the relevant guidelines and regulations or in accordance with the Declaration of Helsinki.

## **Consent for publication**

Not Applicable.

#### **Competing interests**

The authors declare no competing interests.

Received: 26 July 2022 / Accepted: 19 April 2023 Published online: 26 April 2023

#### References

- Umegaki H. Sarcopenia and frailty in older patients with diabetes mellitus. Geriatr Gerontol Int. 2016;16:293–9.
- Hanna JS. Sarcopenia and critical illness: a deadly combination in the elderly. JPEN J Parenter Enteral Nutr. 2015;39:273–81.
- Simsek H, Meseri R, Sahin S, Kilavuz A, Bicakli DH, Uyar M, et al. Prevalence of sarcopenia and related factors in community-dwelling elderly individuals. Saudi Med J. 2019;40:568–74.
- Park SW, Goodpaster BH, Lee JS, Kuller LH, Boudreau R, de Rekeneire N, et al. Excessive loss of skeletal muscle mass in older adults with type 2 diabetes. Diabetes Care. 2009;32:1993–7.
- Veronese N, Stubbs B, Punzi L, Soysal P, Incalzi RA, Saller A, et al. Effect of nutritional supplementations on physical performance and muscle strength parameters in older people: a systematic review and meta-analysis. Ageing Res Rev. 2019;51:48–54.
- Cordero A, Laclaustra M, León M, Casasnovas JA, Grima A, Luengo E, et al. Comparison of serum lipid values in subjects with and without the metabolic syndrome. Am J Cardiol. 2008;102:424–8.

- Prasad M, Sara J, Widmer RJ, Lennon R, Lerman LO, Lerman A. Triglyceride and Triglyceride/ HDL (high density lipoprotein) ratio Predict Major adverse Cardiovascular Outcomes in Women with Non-Obstructive Coronary Artery Disease. J Am Heart Assoc. 2019;8:e009442.
- González-Chávez A, Simental-Mendía LE, Elizondo-Argueta S. Elevated triglycerides/HDL-cholesterol ratio associated with insulin resistance. Cir Cir. 2011;79:126–31.
- Qin H, Chen Z, Zhang Y, Wang L, Ouyang P, Cheng L, et al. Triglyceride to high-density lipoprotein cholesterol ratio is associated with incident diabetes in men: a retrospective study of chinese individuals. J Diabetes Investig. 2020;11:192–8.
- Hadaegh F, Khalili D, Ghasemi A, Tohidi M, Sheikholeslami F, Azizi F. Triglyceride/HDL-cholesterol ratio is an independent predictor for coronary heart disease in a population of iranian men. Nutr Metab Cardiovasc Dis. 2009;19:401–8.
- Chen LK, Liu LK, Woo J, Assantachai P, Auyeung TW, Bahyah KS, et al. Sarcopenia in Asia: consensus report of the asian Working Group for Sarcopenia. J Am Med Dir Assoc. 2014;15:95–101.
- 13. Wang N, Chen M, Fang D. Relationship between serum triglyceride to high-density lipoprotein cholesterol ratio and sarcopenia occurrence rate in community-dwelling chinese adults. Lipids Health Dis. 2020;19:248.
- Chung TH, Kwon YJ, Shim JY, Lee YJ. Association between serum triglyceride to high-density lipoprotein cholesterol ratio and sarcopenia in elderly korean males: the Korean National Health and Nutrition Examination Survey. Clin Chim Acta. 2016;463:165–8.

- Hermans MP, Ahn SA, Rousseau MF. The atherogenic dyslipidemia ratio [log(TG)/HDL-C] is associated with residual vascular risk, beta-cell function loss and microangiopathy in type 2 diabetes females. Lipids Health Dis. 2012;11:132.
- Tuzun S, Oner C, Dabak MR, Kasikci HO, Sargin M. Relation of muscle indices with metabolic parameters and C-Peptide in type 2 diabetes Mellitus. J Coll Physicians Surg Pak. 2017;27:673–7.
- de Freitas MM, de Oliveira VLP, Grassi T, Valduga K, Miller MEP, Schuchmann RA, et al. Difference in sarcopenia prevalence and associated factors according to 2010 and 2018 european consensus (EWGSOP) in elderly patients with type 2 diabetes mellitus. Exp Gerontol. 2020;132:110835.
- Chen F, Xu S, Wang Y, Chen F, Cao L, Liu T, et al. Risk factors for Sarcopenia in the Elderly with type 2 diabetes Mellitus and the Effect of Metformin. J Diabetes Res. 2020;2020:3950404.
- Bouchi R, Fukuda T, Takeuchi T, Minami I, Yoshimoto T, Ogawa Y. Sarcopenia is associated with incident albuminuria in patients with type 2 diabetes: a retrospective observational study. J Diabetes Investig. 2017;8:783–7.
- Sung MJ, Lim TS, Jeon MY, Lee HW, Kim BK, Kim DY, et al. Sarcopenia is independently Associated with the degree of liver fibrosis in patients with type 2 diabetes Mellitus. Gut Liver. 2020;14:626–35.

## **Publisher's Note**

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.