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Objectives: The relationships between the triglyceride-glucose (TyG) index, TyG-related parameters of different obesity phenotypes and the risk of type 2 diabetes (T2D) remain unclear. We aimed to determine associations between TyG index, obesity-related TyG parameters and T2D risk in Chinese adults.

Methods: This cross-sectional study included 9489 participants aged \geq 40 years from a large scale, community-based cohort study. Multivariable logistic regression was performed to estimate odds ratios (ORs) and 95% confidence intervals (CIs). The receiver operating characteristic (ROC) curve was employed to test and compare the predictive power of obesity-related TyG parameters across different phenotypes for the risk of T2D.

Results: A total of 2081 (21.9%) participants with T2D were identified. When comparing with participants in the bottom quartile of TyG index, a heightened risk of T2D was observed among the highest quartile group, with an adjusted OR of 5.89 (95% CI: 4.98–6.98). Comparable relationships were found between obesity-related TyG indices and T2D, including TyG-waist circumference (TyG-WC), TyG-waist-to-height ratio (TyG-WHtR), TyG-body mass index (TyG-BMI) and TyG-body fat percentage (TyG-BFP). Abdominal obesity-related TyG indices had the highest predictive capability for T2D, with the area under the curve (AUC) was 0.711 (0.697–0.724) for TyG-WHtR and 0.705 (0.691–0.719) for TyG-WC, which was superior to the general obesity-related TyG indices, with the AUC were 0.683 (0.669–0.698) and 0.631 (0.616–0.646) for TyG-BMI and TyG-BFP, respectively.

Conclusions: Our findings demonstrate a positive associations between TyG index, obesity-related TyG indices and risk of T2D. Abdominal obesity-related TyG indices had a better predictive value to diabetes than general obesity-related TyG indices.

KEYWORDS

triglyceride-glucose index, obesity-related TyG indices, type 2 diabetes, abdominal obesity, middle-aged and older Chinese adults, cross-sectional study

1 Introduction

Diabetes stands as a prominent contributor to blindness, kidney failure, lower limb amputation and cerebrovascular accidents (1). It is estimated that 537 million adults suffer from diabetes in 2021, which is predicted to reach 783 million by 2045 (2). Type 2 diabetes (T2D) represents more than 95% of all diabetes cases. Interfering factors such as lifestyle, obesity, metabolic disorders are known to be recognized as key determinants of risks of T2D (3). Despite this, exploration into complex metabolism-related biomarkers on the risk of T2D has been limited.

Insulin resistance is the critical pathophysiological basic in development of T2D. The triglyceride-glucose (TyG) index, derived from the product of triglyceride (TG) and fasting plasma glucose (FPG) concentrations, has emerged as a non-insulin-based marker of insulin resistance in several studies (4-6). Given the widespread availability of TG testing compared to insulin, the TyG index may be a more convenient and practical method to assess insulin resistance in clinical practice (7), in comparison with the traditional indicator of homeostatic model assessment of insulin resistance (HOMA-IR) calculated using insulin and FPG levels. Furthermore, TyG index exhibit superior sensitivity in detecting insulin resistance in Chinese T2D patients compared to HOMA-IR (8). Associations between TyG index and prediabetes and T2D risk have been identified in previous studies (5, 9, 10). However, the robustness of TyG index and its-related indices remains controversial in the current studies (11-14).

Obesity is another modifiable and major cause for T2D, which usually interacts with other risk factors to elevate the risk of chronic conditions. Obesity could be categorized into general and abdominal adiposity, commonly characterized by body mass index (BMI), body fat percentage (BFP), waist circumference (WC), or waist-to-height ratio (WHtR). The latest research has demonstrated that general and abdominal obesity could discriminate people with or without hypertension (15). Abdominal obesity may pose a more important risk for pancreatic cancer compared to general adiposity (16). All of these suggests the differential impacts of distinct phenotypes of obesity on diseases outcomes. The existing studies have explored the relationships between TyG-BMI, TyG-WC, TyG-WHtR and T2D (17–19), but few of them were engaged in the interaction of TyG index and BFP on T2D. Therefore, the present study aimed to examine the relationship between TyG index, as well as obesity-related TyG indices, and the risk of T2D within a substantial sample size of middle-aged and elderly population in Chinese community.

2 Methods

2.1 Study population

A cross-sectional design was undertaken in a community in Guangzhou, China, utilizing a subgroup of participants from the Risk Evaluation of cAncers in Chinese diabeTic Individuals: A lONgitudinal (REACTION) study, which was established as a multicenter prospective cohort to assess diabetes and cancer in the Chinese population. Detailed protocols of this study have been reported previously (20, 21). In short, a total of 10,104 residents aged \geq 40 years were recruited via examination notices or home visits from June to November 2011, among which 9,916 signed written informed consent to participate. Study protocol was approved by the Ethical Committee of Sun Yat-sen Memorial Hospital of Sun Yat-sen University and Ruijin Hospital Ethics Committee of Shanghai Jiao Tong University School of Medicine with an approval number of 14 on March 10, 2011, and adhered to the declaration of Helsinki and its later amendments.

For the current analysis, participants were further excluded if they had no measurement data of FPG (n = 13), TG (n = 29), oral glucose tolerance test 2 h plasma glucose (OGTT 2h-PG, n = 68), glycated hemoglobin (HbA1c, n = 39), height or weight, (n = 200), and WC (n = 78). Finally, 9,489 eligible individuals were included in the present analysis, with 8,396 having undergone body fat assessment and obtained BFP measurements (Figure 1).

2.2 Sociodemographic data collection and anthropometric assessments

A standardized questionnaire was conducted to gather sociodemographic information, lifestyle habits, and medication use. Education levels were categorized as less than high school, high school, and college or above, while occupation was stratified



into light, medium and heavy physical labor. Smokers and alcohol drinkers were participants who smoked or drank regularly in the past six months.

Anthropometric measurements were conducted by experienced research personnel using standard procedure. BMI was calculated as weight/height squared (kg/m²), while WHtR was derived using the formula WC/height. Content of body fat was determined using a body fat meter (OMRON HBF-306, Omron Company, China) and BFP was calculated automatically. Blood pressure was recorded using the automated electronic device (OMRON, Omron Company, China). Individuals with elevated blood pressure (systolic \geq 140 mmHg or diastolic \geq 90 mmHg), self-reported diagnosis of hypertension by a clinician, or prescribed antihypertensive drugs were considered to existence of hypertension.

2.3 Blood sample collection and laboratory measurements

With a minimum of 10 hours of overnight fasting, peripheral venous blood was collected for laboratory analysis. FPG, fasting serum insulin (FINS), TG, total cholesterol (TC), high-density lipoprotein cholesterol (HDL-C), low-density lipoprotein cholesterol (LDL-C) were determined using the autoanalyzer (Beckman CX-7 Biochemical Autoanalyzer, Brea, CA, USA). HbA1c was measured by high-performance liquid chromatography (Bio-Rad, Hercules, CA). OGTT was carried out according to standard procedure and OGTT 2h-PG was determined. TyG index and obesity-related TyG indices were defined as described previously (22, 23): TyG = ln (TG [mg/dL] × FPG [mg/dL]/2), TyG-BMI = TyG × BMI, TyG-WC = TyG × WC, TyG-WHtR = TyG × WHtR, TyG-BFP = TyG × BFP.

2.4 Outcome definition

T2D was defined (24) if any of the following conditions was met: (1) FPG \geq 7.0 mmol/L, (2) OGTT 2h-PG \geq 11.1 mmol/L, (3) HbA1c \geq 6.5%, (4) a self-reported diagnosis of diabetes by a clinician, (5)the use of diabetic medications, including insulin. Prediabetes was defined as the existence of one of the following: (1) concentrations of FPG were between 6.1 mmol/L and 7.0 mmol/ L, (2) concentrations of OGTT 2h-PG were between 7.8 mmol/L and 11.1 mmol/L, (3) HbA1c was between 5.7% and 6.5%. Impaired glucose metabolism encompassed the presence of either T2D or prediabetes.

2.5 Statistical analysis

Participants were clustered into fours groups based on quartiles of TyG index. Continuous variables were described as mean and standard deviation (SD), whereas categorical variables were summarized by counts and frequencies, respectively. Comparisons of basic characteristics, including demographic, anthropometric, lifestyles, diseases information and laboratory measurements across TyG index quartiles, were conducted via one-way ANOVA, Kruskal-Wallis tests or Pearson's Chi-Squared tests as appropriate.

Logistic regression was employed to investigate the relationship between both TyG index and obesity-related TyG indices (including TyG-BMI, TyG-WC, TyG-WHtR, and TyG-BFP), and T2D risk without adjustment in model 1. Model 2 adjusted for non-modifiable factors including sex (men, women) and age (continuous). Model 3 further adjusted for modifiable risk factors of T2D, including education levels (categorized as mentioned above), occupation (categorized as mentioned above), smoking (smoker, non-smoker), alcohol drinking (alcohol drinker, non-alcohol drinker) and BMI (continuous). Odds ratios (ORs) and 95% confidence intervals (CIs) were calculated, with the lowest quartile serving as the reference. Linear trends were assessed by replacing quartiles of TyG index with median values within subgroups and assigning as continuous variables in the models. Linear regression was also conducted to assess relationships between TyG index, obesity-related TyG-parameters, and indicators of glucose metabolism without and with adjustment for potential covariates mentioned above. Stratified analysis was performed to explore whether the correction between TyG index quartiles and T2D risk varied among subjects with different characteristics, with interactions estimated by including multiplicative interaction terms. The receiver operating characteristic (ROC) curve was performed to assess the predictive capability of TyG index and obesity-related TyG indices for T2D risk. Pearson's Correlation analysis was conducted to explore relationships among TyG index and obesity-related TyG indices. All of the correlation coefficients calculated were greater than 0.4, indicating moderate to strong positive correlations among them (Supplementary Table S1). Missing covariates data were handled through the multiple imputation method.

All statistical analyses were performed using R 4.4.2 and were two-sided. P < 0.05 was considered statistically significant.

3 Results

3.1 Basic characteristics

Of the 9,489 study participants, there were 2,692 (28.4%) men and 6,797 (71.6%) women. Mean age was 56.0 \pm 8.0 years. Table 1 listed the basic characteristics of participants stratified by quartiles of TyG index. Participants in the highest quartile were older, had higher BMI, BFP, WC and blood pressure, high levels of FPG, OGTT 2h-PG, HbA1c, FINS, TC, TG and LDL-C, had lower education levels and lower HDL-C levels (all *P* < 0.001, all *P*-trend < 0.001) compared to those in the lowest quartile. Participants in the higher quartiles of TyG index tended to be men, smokers, and had high prevalence of T2D, impaired glucose metabolism and hypertension (all *P* < 0.001, all *P*-trend < 0.001). No significant differences were noted in alcohol drinking status among participants with various quartiles of TyG index (*P* = 0.937, *P*-trend = 0.924).

3.2 TyG index, obesity-related TyG indices and risk of T2D

Table 2 presented the relationships between TyG index quartiles, obesity-related TyG indices and T2D. As for TyG index, the number of cases of T2D were lowest in the bottom quartile and highest in the fourth quartile of TyG index. A positive correction between TyG index and T2D risk was identified with an unadjusted OR of 7.88 (95% CI: 6.69-9.27, P-trend < 0.001) in the highest quartile in comparison with the first quartile. Multivariable analyses, adjusted for sex and age, revealed an adjusted OR of 7.08 (95% CI: 6.00-8.35, *P*-trend < 0.001) in the highest TyG index quartile compared to the lowest. This association remained statistically significant after further adjusting for modifiable T2D risk factors, with a fully adjusted OR of 5.89 (95% CI: 4.98-6.98, P-trend < 0.001) in the fourth quartile. When considering TyG index as a continuous variable, the risk of T2D also sharply increased for each SD increment in TyG index, irrespective of potential covariate adjustments (fully adjusted OR = 2.07, 95% CI: 1.96-2.19, P < 0.001). With regard to obesity-related TyG indices, including TyG-BMI, TyG-WC, TyG-WHtR and TyG-BFP, similar findings were noted between obesity-related TyG indices and T2D without or with adjustments for potential covariates (all Ptrend < 0.001). Consistent results were also found between TyG index, obesity-related TyG indices and risk of impaired glucose metabolism (Supplementary Table S2).

3.3 TyG index, obesity-related TyG indices and indicators of glucose metabolism

Linear corrections of TyG index, obesity-related TyG indices with glucose metabolism were shown in Table 3. Positive associations between TyG index and indicators of glucose metabolism, including FPG (β = 0.41, standard error (SE) = 0.01), OGTT 2h-PG (β = 0.85, SE = 0.03) and HbA1c (β = 0.23, SE = 0.01), were found (all *P* < 0.001), following adjustment for potential confounding factors. As for obesity-related TyG indices, identical positive associations were observed with FPG (β : 0.28–0.41, *P* < 0.001), OGTT 2h-PG (β : 0.50–0.91, *P* < 0.001) and HbA1c (β : 0.14–0.24, *P* < 0.001) after additionally adjusting for potential covariates.

3.4 Interactions and stratified analyses

Table 4 delineated the influences of TyG index on T2D risk across strata of selected potential risk factors. A statistically significant multiplicative interaction was demonstrated between quartiles of TyG index and sex on associations with T2D risk (*P*-interaction = 0.024). Stratified by sex, stronger associations of TyG index with T2D risk were found in women (OR = 6.51, 95% CI: 5.32–7.98, *P*-trend < 0.001) than in men (OR = 4.41, 95% CI: 3.23–6.01, *P*-trend < 0.001). No notable multiplicative interactions were identified between smoking status, alcohol drinking, different phenotypes of obesity and quartiles of TyG index (all *P*-interaction > 0.05). The influences of TyG index on risk of impaired glucose metabolism stratified by confounders mentioned above were shown in Supplementary Table S3 and consistent results were found.

3.5 The predictive value of TyG index, obesity-related TyG indices for T2D

ROC curve for TyG and obesity-related TyG indices were presented in Figure 2. TyG had the highest area under the curve (AUC = 0.722, 95% CI: 0.709-0.736), followed by TyG-WHtR (0.711, 95% CI: 0.697-0.724), TyG-WC (0.705, 95% CI: 0.691-0.719), TyG-BMI (0.683, 95% CI: 0.669-0.698) and TyG-BFP (0.631, 95% CI: 0.616-0.646). As for impaired glucose metabolism, similar results were observed in Supplementary Figure S1, with the AUC of 0.682 (95% CI: 0.669-0.696), 0.682(95% CI: 0.668-0.695), 0.673 (95% CI: 0.659-0.686), 0.665 (95% CI: 0.651-0.678) and 0.614 (95% CI: 0.600-0.628) for TyG, TyG-WHtR, TyG-WC, TyG-BMI and TyG-BFP, respectively.

4 Discussion

In this representative cross-sectional study with 9,489 middleaged and older Chinese adults, we delved into the corrections between T2D risk and both TyG index and obesity-related TyG indices. We observed that TyG index, as well as TyG-BMI, TyG-WC, TyG-WHtR and TyG-BFP, were positively associated with risk of T2D. These associations persisted even after accounting for potential confounders. Positive associations between TyG index, obesity-related TyG indices and FPG, OGTT 2h-PG and HbA1c were also found. In addition, TyG index had the higher predictive ability for T2D, followed by abdominal obesity-related TyG indices (TyG-WHtR and TyG-WC), and general obesity-related TyG indices (TyG-BMI and TyG-BFP).

A majority of previous researches have examined the associations between the TyG index and disorder glucose

TABLE 1 Basic characteristics of participants by quartiles of TyG index¹.

Characteristics	Total	Quartiles of TyG index					D_trond
Characteristics	TOLAL	Q1 (n = 2505)	Q2 (n = 2501)	Q3 (n = 2498)	Q4 (n = 2501)	P	P-trend
Range of TyG index	6.16-12.07	≤8.70	8.27-8.63	8.64-9.04	≥9.05	0.001	0.001
Mean of TyG index	8.69 ± 0.61	7.99 ± 0.24	8.45 ± 0.10	8.82 ± 0.12	9.50 ± 0.43	<0.001	\U.UI
Age, years	56.0 ± 8.0	54.0 ± 7.7	55.7 ± 7.7	56.7 ± 7.9	57.5 ± 8.0	< 0.001	< 0.001
Sex, n (%)						< 0.001	< 0.001
Men	2692 (28.4)	563 (23.5)	615 (25.9)	681 (28.9)	833 (35.4)		
Women	6797 (71.6)	1835 (76.5)	1763 (74.1)	1677 (71.1)	1522 (64.6)		
Occupation, n (%)						< 0.001	< 0.001
Light labor	6999 (73.8)	1672 (69.7)	1751 (73.6)	1784 (75.7)	1792 (76.1)		
Medium labor	1426 (15.0)	444 (18.5)	354 (14.9)	321 (13.6)	307 (13.0)		
Heavy labor	1064 (11.2)	282 (11.8)	273 (11.5)	253 (10.7)	256 (10.9)		
Education levels, n (%)						< 0.001	< 0.001
Less than high school	3715 (39.2)	845 (35.2)	897 (37.7)	944 (40.0)	1029 (43.7)		
High school	4893 (51.6)	1319 (55.0)	1266 (53.2)	1192 (50.6)	1116 (47.4)		
College or above	881 (9.3)	234 (9.8)	215 (9.0)	222 (9.4)	210 (8.9)		
Smoking status, n (%)						< 0.001	< 0.001
Non-smoker	7750 (81.7)	2038 (85.0)	1976 (83.1)	1925 (81.6)	1811 (76.9)		
Smoker	1739 (18.3)	360 (15.0)	402 (16.9)	433 (18.4)	544 (23.1)		
Alcohol drinking status, n (%)						0.937	0.924
Non-alcohol drinker	6832 (72.0)	1732 (72.2)	1700 (71.5)	1701 (72.1)	1699 (72.1)		
Alcohol drinker	2657 (28.0)	666 (27.8)	678 (28.5)	657 (27.9)	656 (27.9)		
Height, cm	158.3 ± 7.5	158.0 ± 7.3	158.1 ± 7.3	158.2 ± 7.6	159.0 ± 7.9	< 0.001	< 0.001
Weight, kg	59.3 ± 9.6	56.2 ± 8.9	57.9 ± 8.9	60.2 ± 9.6	62.9 ± 9.6	< 0.001	< 0.001
BMI, kg/m ²	23.6 ± 3.1	22.5 ± 3.1	23.1 ± 3.0	24.0 ± 3.1	24.8 ± 2.9	< 0.001	< 0.001
WC, cm	81.6 ± 9.2	77.7 ± 8.7	80.1 ± 8.8	82.9 ± 8.8	85.8 ± 8.3	< 0.001	< 0.001
Hip circumference, cm	94.0 ± 7.0	92.4 ± 6.7	93.3 ± 6.9	94.5 ± 7.0	95.7 ± 7.0	< 0.001	< 0.001
WHtR	0.52 ± 0.06	0.49 ± 0.05	0.51 ± 0.06	0.52 ± 0.05	0.54 ± 0.05	< 0.001	< 0.001
Body fat percentage, %	29.4 ± 6.0	27.9 ± 6.1	29.0 ± 6.1	30.0 ± 5.8	30.5 ± 5.8	< 0.001	< 0.001
SBP, mmHg	126.1 ± 16.5	120.4 ± 15.4	124.2 ± 15.6	127.4 ± 16.1	132.5 ± 16.5	< 0.001	< 0.001
DBP, mmHg	75.3 ± 9.8	72.4 ± 9.6	74.3 ± 9.6	76.0 ± 9.4	78.6 ± 9.6	< 0.001	< 0.001
FPG, mmol/L	5.7 ± 1.3	5.1 ± 0.6	5.4 ± 0.7	5.7 ± 1.0	6.6 ± 2.1	< 0.001	< 0.001
OGTT 2h-PG, mmol/L	8.1 ± 3.1	6.9 ± 2.1	7.5 ± 2.3	8.3 ± 2.9	9.8 ± 3.9	< 0.001	< 0.001
HbA1c, %	6.0 ± 0.9	5.7 ± 0.5	5.9 ± 0.5	6.0 ± 0.7	6.5 ± 1.4	< 0.001	< 0.001
FINS, µU/ml	8.3 ± 5.8	6.2 ± 3.6	7.3 ± 5.9	8.8 ± 5.1	10.9 ± 7.1	<0.001	<0.001
TC, mmol/L	5.2 ± 1.3	4.5 ± 1.3	5.2 ± 1.1	5.5 ± 1.1	5.7 ± 1.2	< 0.001	< 0.001
TG, mmol/L	1.6 ± 1.2	0.7 ± 0.2	1.1 ± 0.2	1.5 ± 0.3	3.0 ± 1.7	<0.001	< 0.001
HDL-C, mmol/L	1.3 ± 0.4	1.4 ± 0.4	1.4 ± 0.4	1.3 ± 0.3	1.2 ± 0.3	<0.001	<0.001
LDL-C, mmol/L	3.1 ± 1.0	2.7 ± 0.9	3.2 ± 0.9	3.4 ± 0.9	3.3 ± 1.0	<0.001	< 0.001

(Continued)

	Total	Quartiles of TyG index					Detwored
Characteristics		Q1 (n = 2505)	Q2 (n = 2501)	Q3 (n = 2498)	Q4 (n = 2501)	Ρ	- trenu
TyG-WC	710.8 ± 107.6	621.2 ± 73.7	676.8 ± 75.5	731.4 ± 79.8	815.9 ± 90.1	< 0.001	<0.001
TyG-WHtR	4.49 ± 0.67	3.94 ± 0.46	4.29 ± 0.48	4.63 ± 0.49	5.14 ± 0.56	< 0.001	< 0.001
TyG-BMI	205.6 ± 34.3	179.7 ± 25.4	195.4 ± 25.8	211.9 ± 27.5	235.9 ± 30.4	< 0.001	< 0.001
TyG-BFP	255.5 ± 57.8	223.2 ± 50.0	245.4 ± 51.4	264.7 ± 51.6	289.7 ± 56.0	< 0.001	< 0.001
Presence of hypertension, n (%)						< 0.001	< 0.001
No	7525 (79.8)	2096 (87.9)	1983 (83.7)	1842 (78.8)	1604 (68.5)		
Yes	1906 (20.2)	288 (12.1)	385 (16.3)	496 (21.2)	737 (31.5)		
Presence of type 2 diabetes, n (%)						< 0.001	< 0.001
No	7408 (78.1)	2187 (91.2)	2049 (86.2)	1834 (77.8)	1338 (56.8)		
Yes	2081 (21.9)	211 (8.8)	329 (13.8)	524 (22.2)	1017 (43.2)		
Presence of impaired glucose metabolism, n (%)						< 0.001	< 0.001
No	1877 (19.8)	813 (33.9)	528 (22.2)	356 (15.1)	180 (7.6)		
Yes	7612 (80.2)	1585 (66.1)	1850 (77.8)	2002 (84.9)	2175 (92.4)		
Status of glucose metabolism, n (%)						< 0.001	< 0.001
Normal	1877 (19.8)	813 (33.9)	528 (22.2)	356 (15.1)	180 (7.6)		
Prediabetes	5531 (58.3)	1374 (57.3)	1521 (64.0)	1478 (62.7)	1158 (49.2)		
Diabetes	2081 (21.9)	211 (8.8)	329 (13.8)	524 (22.2)	1017 (43.2)		

TABLE 1 Continued

¹ Data are mean \pm SD or n (%).

TyG index, triglyceride-glucose index; Q1, first quartile; Q2, second quartile; Q3, third quartile; Q4, fourth quartile; BMI, body mass index; WC, Waist circumference; WHtR, waist-to-height ratio; SBP, systolic blood pressure; DBP, diastolic blood pressure; FPG, fasting plasma glucose; OGTT 2h-PG, oral glucose tolerance test 2 h plasma glucose; HbA1c, glycated hemoglobin; FINS, fasting serum insulin; TC, total cholesterol; TG, triglycerides; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; WC, waist circumference; BFP, body fat percentage.

metabolism, including prediabetes and diabetes. In a cross-sectional study conducted in representative American adults, in comparison to the bottom group, participants with higher TyG index were associated with the higher risk of both prediabetes and diabetes (25). Another prospective cohort study undertaken in Japan also demonstrated a positive correction between TyG index and the risk of T2D (26). In the Chinese population, several studies conducted in geriatric (5, 27), rural area (9), or general individuals (28) found consistently that increasing TyG index was related with the higher incidence risk of prediabetes and diabetes. The above results revealed that TyG index served as a potential marker or predictor of risk of both prediabetes and T2D. Corresponding with these studies, the current investigation also observed a positive relationship between TyG index and T2D, as well as impaired glucose metabolism. Disorder of glucose and lipid metabolism was an crucial cause for diabetes. The increase of glucose and TG levels disrupted the function of pancreatic islet β cells through elevating reactive oxygen, inflammation, endoplasmic reticulum stress and ectopic fat deposition, therefore resulting to pancreatic dysfunction and insulin resistance (29-32). Therefore, the TyG index could be a complex and effective indicator to reflect diabetes or disorder of glucose metabolism, which aligned with the description of the present and previous researches mentioned above.

Obesity was another important and modifiable risk factor for diabetes. It was reported that obesity accounted for 52.2% of the disability-adjusted life-years (DALYs) linked to T2D in 2021, with a more than 20% increase since 1990 (33), which emphasized its significant impact on T2D. The existing studies have examined the relationships between T2D and obesity-related TyG indices, such as TyG-BMI, TyG-WC and TyG-WHtR (17-19). BMI was the most common indicator for measuring general obesity, and many studies combined it with the TyG index to investigate the correction between obesity-related TyG indices and impaired glucose metabolism. TyG-BMI was discovered to be associated with the risk of T2D, whatever prediabetes or general individuals from people of different ethnic groups (19, 34-37). Moreover, reduction of TyG-BMI via active management was possible to help to convert prediabetes into normoglycemia (38, 39). Body fat was additionally another indicator on behalf of general obesity, however, there was a paucity of studies that addressed the interaction of TyG index and body fat on T2D. In the present research, we found that the TyG-BFP index, the product of TyG and BFP, was as important as other effective obesity-indicator to predict T2D and impaired glucose metabolism. In regard to abdominal obesity, the latest research have found that general and abdominal adiposity could discriminate people with or without hypertension (15), which suggested that the abdominal adiposity, usually described by

Quartiles of TyG and obesity-related TyG indices TyG index P-trend Per SD increment Q1 02 Q3 Q4 TyG Case/total. n 211/2398 329/2378 524/2358 1017/2355 Model 11 1.00 (ref.) 1.66 (1.39, 2.00) 2.96 (2.49, 3.52) 7 88 (6 69, 9 27) < 0.001 2.24 (2.13, 2.37) < 0.001 Model 2² 1.00 (ref.) < 0.001 < 0.001 1.55 (1.29, 1.87) 2.66 (2.23, 3.16) 7.08 (6.00, 8.35) 2.21 (2.09, 2.33) Model 3³ 1.00 (ref.) 1.47 (1.22, 1.77) 2.35 (1.97, 2.80) 5.89 (4.98, 6.98) < 0.001 2.07 (1.96, 2.19) < 0.001 Abdominal obesity-related TyG indices TyG-WC 210/2379 563/2378 951/2363 Case/total, n 357/2369 Model 11 1.00 (ref.) 1.83 (1.53, 2.20) 3.20 (2.70, 3.80) 6.96 (5.91, 8.19) < 0.001 2.15 (2.04, 2.27) < 0.001 Model 2² 1.00 (ref.) 1.78 (1.48, 2.13) 2.98 (2.51, 3.54) 6.51 (5.51, 7.71) < 0.001 2.13 (2.01, 2.25) < 0.001 Model 3³ 1.00 (ref.) 1.76 (1.45, 2.12) 2.93 (2.42, 3.54) 6.30 (5.11, 7.76) < 0.001 2.34 (2.17, 2.52) < 0.001 TyG-WHtR Case/total, n 214/2379 350/2367 529/2372 988/2371 Model 11 1.00 (ref.) 1.76 (1.47, 2.10) 2.90 (2.45, 3.44) 7.23 (6.14, 8.50) < 0.001 2.23 (2.11, 2.35) < 0.001 Model 2² 1.00 (ref.) 1.64 (1.37, 1.97) 2.65 (2.23, 3.15) 6.22 (5.28, 7.34) < 0.001 2.11 (2.00, 2.24) < 0.001 Model 3³ 1.00 (ref.) 1.65 (1.37, 1.99) 2.67 (2.21, 3.23) 6.26 (5.09, 7.71) < 0.001 2.38 (2.20, 2.57) < 0.001 General obesity-related TyG indices TyG-BMI Case/total, n 237/2381 386/2373 538/2365 920/2370 Model 11 1.00 (ref.) 1.76 (1.48, 2.09) 2.66 (2.26, 3.14) 5.74 (4.90, 6.72) < 0.001 1.98 (1.88, 2.08) < 0.001 Model 2² 1.00 (ref.) 1.71 (1.44, 2.04) 2.58 (2.18, 3.05) 5.63 (4.80, 6.61) < 0.001 1.98 (1.88, 2.09) < 0.001 Model 3⁴ 1.00 (ref.) 1.71 (1.44, 2.04) 2.58 (2.18, 3.05) 5.60 (4.77, 6.57) < 0.001 1.98 (1.87, 2.08) < 0.001 TyG-BFP Case/total, n 297/2100 364/2104 424/2103 721/2089 Model 11 1.27 (1.07, 1.50) 1.00 (ref.) 1.53 (1.30, 1.80) 3.20 (2.75, 3.73) < 0.001 1.65 (1.56, 1.75) < 0.001 Model 2² 1.00 (ref.) 1.48 (1.24, 1.76) 1.98 (1.66, 2.37) 4.28 (3.57, 5.13) < 0.0011.86 (1.74, 1.99) < 0.001 Model 3³ 1.00 (ref.) 1.30 (1.08, 1.55) 1.58 (1.30, 1.91) 2.86 (2.30, 3.56) < 0.001 1.67 (1.53, 1.83) < 0.001

TABLE 2 Odds ratios and 95% CIs of T2D by quartiles of TyG and obesity-related TyG indices.

¹ Unadjusted. ² Adjusted for age (continuous) and sex (men, women). ³ Adjusted additionally for education levels (less than high school, high school, college or above), occupation (light, medium, heavy physical labor), smoking status (smoker, non-smoker), alcohol drinking (alcohol drinker, non-alcohol drinker) and BMI (continuous). ⁴ Adjusted for covariates mentioned above except for BMI.

CI, confidence interval; T2D, type 2 diabetes; TyG index, triglyceride-glucose index; Q1, first quartile; Q2, second quartile; Q3, third quartile; Q4, fourth quartile; SD, standard deviation; WC, waist circumference; WHtR, waist-to-height ratio; BMI, body mass index; BFP, body fat percentage.

WC and WHtR, may be as a important risk factor as the general adiposity to diabetes. As the findings from previous studies shown, TyG-WC and TyG-WHtR were also effective markers to identify the diabetes risk (17, 18). Furthermore, they seem to be better indicators for predicting prediabetes in Indian (40) and Qatari (41) population in Asia. In this study, significant positive corrections were found between TyG-WC, TyG-WHtR, and risk of T2D and impaired glucose metabolism. TyG-WC and TyG-WHtR demonstrated superior predictive efficacy for diabetes than TyG-BMI and TyG-BFP, which were aligned with the previous findings.

Our study boasts notable strengths. It was undertaken in a representative, community-dwelling population with a wide range of age (equal or more than 40 years) in China. It was also a frontier research to combine different phenotypes of obesity, including general and abdominal obesity, with TyG index to discuss the relationship between obesity-related TyG indices and T2D, and compared their predictive value ulteriorly.

Several limitations need to be considered in our study. First, despite our efforts to adjust to potential risk factors, residual confounding cannot be eradicated fully. Moreover, the cross-

TyG index	Model 1 ¹			Model 2 ²			Model 3 ³		
	eta (95% CI)	SE	Р	eta (95% CI)	SE	Р	eta (95% CI)	SE	Р
FPG			1						
TyG	0.45 (0.42, 0.47)	0.01	<0.001	0.43 (0.41, 0.45)	0.01	< 0.001	0.41 (0.39, 0.44)	0.01	<0.001
TyG-WC	0.36 (0.34, 0.39)	0.01	< 0.001	0.35 (0.32, 0.37)	0.01	< 0.001	0.41 (0.38, 0.44)	0.02	< 0.001
TyG-WHtR	0.36 (0.33, 0.38)	0.01	< 0.001	0.34 (0.32, 0.36)	0.01	< 0.001	0.41 (0.38, 0.44)	0.02	< 0.001
TyG-BMI	0.32 (0.29, 0.34)	0.01	< 0.001	0.31 (0.28, 0.33)	0.01	< 0.001	$0.30 \ (0.28, \ 0.33)^4$	0.01	< 0.001
TyG-BFP	0.21 (0.18, 0.23)	0.01	< 0.001	0.29 (0.26, 0.32)	0.02	<0.001	0.28 (0.24, 0.32)	0.02	< 0.001
OGTT 2h-PG									
TyG	0.96 (0.91, 1.01)	0.03	<0.001	0.91 (0.86, 0.96)	0.03	<0.001	0.85 (0.79, 0.90)	0.03	<0.001
TyG-WC	0.83 (0.77, 0.88)	0.03	< 0.001	0.79 (0.74, 0.85)	0.03	< 0.001	0.86 (0.79, 0.94)	0.04	< 0.001
TyG-WHtR	0.87 (0.82, 0.93)	0.03	< 0.001	0.81 (0.75, 0.86)	0.03	< 0.001	0.91 (0.84, 0.98)	0.04	< 0.001
TyG-BMI	0.76 (0.70, 0.81)	0.03	< 0.001	0.73 (0.67, 0.78)	0.03	< 0.001	$0.72 \ (0.67, \ 0.77)^4$	0.03	< 0.001
TyG-BFP	0.54 (0.48, 0.60)	0.03	< 0.001	0.62 (0.55, 0.68)	0.03	< 0.001	0.50 (0.42, 0.58)	0.04	< 0.001
HbA1c									
TyG	0.26 (0.24, 0.27)	0.01	<0.001	0.24 (0.23, 0.26)	0.01	<0.001	0.23 (0.21, 0.24)	0.01	<0.001
TyG-WC	0.22 (0.21, 0.24)	0.01	< 0.001	0.21 (0.20, 0.23)	0.01	< 0.001	0.24 (0.21, 0.26)	0.01	< 0.001
TyG-WHtR	0.23 (0.21, 0.24)	0.01	< 0.001	0.21 (0.19, 0.23)	0.01	< 0.001	0.24 (0.22, 0.26)	0.01	< 0.001
TyG-BMI	0.20 (0.18, 0.21)	0.01	< 0.001	0.19 (0.17, 0.21)	0.01	< 0.001	0.19 (0.17, 0.20) ⁴	0.01	< 0.001
TyG-BFP	0.14 (0.12, 0.16)	0.01	<0.001	0.17 (0.15, 0.19)	0.01	< 0.001	0.14 (0.11, 0.16)	0.01	< 0.001

TABLE 3 Associations between TyG index, obesity-related TyG indices and indicators of glucose metabolism.

¹ Unadjusted. ² Adjusted for age (continuous) and sex (men, women). ³ Adjusted additionally for education levels (less than high school, high school, college or above), occupation (light, medium, heavy physical labor), smoking status (smoker, non-smoker), alcohol drinking (alcohol drinker, non-alcohol drinker) and BMI (continuous). ⁴ Adjusted for covariates mentioned above except for BMI.

TyG index, triglyceride-glucose index; Q1, first quartile; Q2, second quartile; Q3, third quartile; Q4, fourth quartile; CI, confidence interval; SE, standard error; FPG, fasting plasma glucose; WC, waist circumference; WHtR, waist-to-height ratio; BMI, body mass index; BFP, body fat percentage; OGTT 2h-PG, oral glucose tolerance test 2 h plasma glucose; HbA1c, glycated hemoglobin.

TABLE 4 Odds ratios and 95% CIs of T2D by quartiles of TyG index stratified by covariates¹.

			Quarti	Dtrand			
ryG index	n	Q1	Q2	Q3	Q4	<i>P</i> -trend	P-Interaction
Sex							0.024
Men	2692	1.00	1.46 (1.04, 2.07)	2.14 (1.55, 2.97)	4.41 (3.23, 6.01)	< 0.001	
Women	6797	1.00	1.45 (1.16, 1.81)	2.38 (1.93, 2.93)	6.51 (5.32, 7.98)	< 0.001	
Smoking status							0.058
Non-smoker	7750	1.00	1.42 (1.15, 1.74)	2.40 (1.97, 2.91)	6.27 (5.20, 7.57)	< 0.001	
Smoker	1739	1.00	1.65 (1.08, 2.52)	1.99 (1.32, 3.01)	4.22 (2.86, 6.23)	< 0.001	
Alcohol drinking status							0.683
Non-alcohol drinker	6832	1.00	1.39 (1.11, 1.73)	2.32 (1.89, 2.85)	5.86 (4.80, 7.14)	< 0.001	
Alcohol drinker	2657	1.00	1.71 (1.21, 2.41)	2.38 (1.70, 3.33)	5.83 (4.21, 8.05)	< 0.001	
WC ² , cm							0.165
< 90 or 80	5044	1.00	1.44 (1.13, 1.84)	2.59 (2.04, 3.28)	5.66 (4.45, 7.19)	< 0.001	

(Continued)

	2	Quartiles of TyG index				Dtrand	Distoraction
TyG index		Q1	Q2	Q3	Q4	P-trend	P-Interaction
≥ 90 or 80	4445	1.00	1.57 (1.18, 2.09)	2.24 (1.72, 2.93)	6.27 (4.87, 8.09)	<0.001	
WHtR							0.170
≤ 0.5	3781	1.00	1.43 (1.07, 1.92)	2.93 (2.21, 3.89)	7.02 (5.24, 9.42)	<0.001	
> 0.5	5708	1.00	1.48 (1.16, 1.89)	2.07 (1.65, 2.60)	5.40 (4.35, 6.69)	<0.001	
BMI, kg/m ²							0.599
< 24	5458	1.00	1.54 (1.22, 1.96)	2.69 (2.14, 3.38)	6.31 (5.05, 7.89)	< 0.001	
≥ 24	4031	1.00	1.40 (1.04, 1.89)	2.19 (1.66, 2.89)	6.04 (4.65, 7.84)	<0.001	
BFP ³ , %							0.709
≤ 25 or 30	3469	1.00	1.54 (1.16, 2.07)	2.42 (1.81, 3.22)	6.10 (4.62, 8.07)	< 0.001	
> 25 or 30	4927	1.00	1.46 (1.11, 1.92)	2.35 (1.82, 3.03)	6.26 (4.91, 7.99)	<0.001	

TABLE 4 Continued

¹ Adjusted for age (continuous) and sex (men, women), education levels (less than high school, high school, college or above), occupation (light, medium, heavy physical labor), smoking status (smoker, non-smoker), alcohol drinking (alcohol drinker, non-alcohol drinker) and BMI (continuous). Stratified factors were not included in the corresponding models. ² Cut-off points of WC were 90 cm for men and 80 cm for women. ³ Cut-off points of BFP were 25% for men and 30% for women.

CI, confidence interval; T2D, type 2 diabetes; TyG index, triglyceride-glucose index; Q1, first quartile; Q2, second quartile; Q3, third quartile; Q4, fourth quartile; WC, waist circumference; WHtR, waist-to-height ratio; BMI, body mass index; BFP, body fat percentage.



sectional design constrained our ability to definitively rule out the possibility that reverse causality and residual confounding may have skewed our findings. Further prospective or interventional researches were required to solidify our understanding of associations between obesity-related TyG parameters and diabetes.

5 Conclusions

In conclusion, positive associations were observed between TyG index, as well as obesity-related TyG indices including TyG-BMI, TyG-WC, TyG-WHtR and TyG-BFP, and risk of T2D. Abdominal obesity-related TyG index had enhanced predictive value to

diabetes than general obesity-related TyG index. Further large prospective researches were required to validate our findings.

Data availability statement

The data analyzed in this study is subject to the following licenses/restrictions: Data described in the article are available from the corresponding author upon reasonable request. Requests to access these datasets should be directed to Li Yan, yanli@mail.sysu.edu.cn.

Ethics statement

The studies involving humans were approved by the Ethical Committee of Sun Yat-sen Memorial Hospital of Sun Yat-sen University and Ruijin Hospital Ethics Committee of Shanghai Jiao Tong University School of Medicine. The studies were conducted in accordance with the local legislation and institutional requirements. The participants provided their written informed consent to participate in this study.

Author contributions

XTL: Conceptualization, Data curation, Formal analysis, Investigation, Methodology, Software, Writing – original draft. SC: Data curation, Investigation, Methodology, Validation, Writing – review & editing. CW: Funding acquisition, Investigation, Validation, Writing – review & editing. DL: Funding acquisition, Investigation, Project administration, Writing – review & editing. FH: Investigation, Writing – review & editing. XHL: Investigation, Writing – review & editing. HW: Investigation, Writing – review & editing. PL: Investigation, Writing – review & editing. LY: Conceptualization, Funding acquisition, Project administration, Supervision, Visualization, Writing – review & editing. MR: Conceptualization, Funding acquisition, Project administration, Supervision, Visualization, Writing – review & editing. CC: Conceptualization, Funding acquisition, Project administration, Supervision, Visualization, Writing – review & editing.

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Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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Supplementary material

The Supplementary Material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fendo.2025. 1526849/full#supplementary-material

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Glossary

T2D	type 2 diabetes
TyG index	triglyceride-glucose index
TG	triglyceride
FPG	fasting plasma glucose
HOMA-IR	homeostatic model assessment of insulin resistance
BMI	body mass index
BFP	body fat percentage
WC	waist circumference
WHtR	waist-to-height ratio
REACTION study	the Risk Evaluation of cAncers in Chinese diabeTic Individuals, A lONgitudinal study
OGTT 2h-PG	oral glucose tolerance test 2 h plasma glucose
HbA1c	glycated hemoglobin
FINS	fasting serum insulin
TC	total cholesterol
HDL-C	high-density lipoprotein cholesterol
LDL-C	low-density lipoprotein cholesterol
Q1	first quartile
Q2	second quartile
Q3	third quartile
Q4	fourth quartile
SD	standard deviation
OR	odds ratio
CI	confidence interval
ROC	receiver operating characteristic
SE	standard error
AUC	area under the curve
NHANES	National Health and Nutrition Examination Survey
NAGALA	NAfld in the Gifu Area, Longitudinal Analysis
DALYs	disability-adjusted life-years