



The Triglyceride-Glucose Index is Independently Associated with Chronic Kidney Disease in the Geriatric Population, Regardless of Obesity and Sex

Bokun Kim^{1,2}, Gwon-Min Kim^{2,3}, Kihoon Han⁴, Naoki Maki^{2,5}, Keisuke Taniguchi^{2,5}, Sechang Oh^{2,5}

¹Future Convergence Research Institute, Changwon National University, Changwon, Korea

²Human Community Renovation Research Center, R Professional University of Rehabilitation, Tsuchiura, Japan

³Medical Research Institute, Pusan National University, Busan, Korea

⁴Department of Physical Education, Pusan National University, Busan, Korea

⁵Faculty of Rehabilitation, R Professional University of Rehabilitation, Tsuchiura, Japan

Corresponding Author:

Sechang Oh, PhD

Division of Physical Therapy, Faculty of Rehabilitation, R Professional University of Rehabilitation, Tsuchiura, Ibaraki 300-0032, Japan

E-mail: ohsechang@u.a-ru.ac.jp

ORCID:

<https://orcid.org/0000-0002-7457-9305>

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Background: Insulin resistance (IR) negatively affects several risk factors of chronic kidney disease (CKD). This cross-sectional study investigated whether the triglyceride-glucose (TyG) index, which reflects IR, was independently associated with CKD in a geriatric population, regardless of obesity and sex. **Methods:** The analysis included 7,326 individuals (2,864 males and 4,462 females) aged ≥ 60 years. Non-obesity or obesity was evaluated using a body mass index cutoff of 25 kg/m^2 . The TyG index was calculated as $\ln [\text{triglyceride concentration (mg/dL)} \times \text{fasting plasma glucose concentration (mg/dL)}] / 2$. All participants were categorized into three groups according to TyG tertiles. Moderate-to-severe CKD ($_{\text{MS}}\text{CKD}$) was defined as an estimated glomerular filtration rate (eGFR) of $< 45.0 \text{ mL/min/1.73 m}^2$. **Results:** In males and females with or without obesity, a trend test showed a decreasing tendency in the eGFR from the lowest to highest TyG tertiles. Males without obesity and females with obesity in the middle and highest tertiles of the TyG index were 2.342 and 2.393, and were 2.313 and 3.516 times more likely to have $_{\text{MS}}\text{CKD}$, respectively. Those with or without obesity in the highest tertile of the TyG index were 1.736 and 2.374 times more likely to have $_{\text{MS}}\text{CKD}$, respectively. **Conclusion:** Geriatric populations with an increased TyG index have a high risk of $_{\text{MS}}\text{CKD}$ regardless of obesity and sex. Our findings suggest that increased IR is associated with CKD in the geriatric population independent of obesity and sex.

Key Words: Body mass index, Insulin resistance, Metabolic syndrome, Aged, Renal insufficiency

INTRODUCTION

Chronic kidney disease (CKD) is a common condition in older adults. CKD increases the mortality rate and risk of conditions including myocardial infarction, hypertension, and type 2 diabetes in the geriatric population. From a pathophysiological perspective, these health concerns share a common pathway mediated by insulin resistance (IR).¹⁻³ In 2017, CKD reportedly led to 1.2 million deaths globally. Owing to the aging of the global population, the prevalence and related mortality rate of CKD are expected to rise, with estimated CKD-related deaths increasing to 2.2 million or 4.0 million by 2040 in the best-case or worst-case scenarios, respec-

tively.⁴ Therefore, developing effective strategies for CKD screening, detection, and management is essential to prevent or suppress the development of severe CKD, particularly in the geriatric population.

The association between obesity and CKD has been globally recognized for decades, and studies have evaluated the risk of CKD by broadly using body mass index (BMI) as the obesity index.⁵⁻⁷ However, Kim et al.^{8,9} reported that high fat and low muscle mass are more closely related to CKD than BMI-based obesity evaluation. Additionally, the limitation of BMI is apparent in the early screening and detection of high-risk older adults with CKD. The reason for these findings is that BMI does not precisely re-

flect overall adiposity and does not distinguish visceral fat, which induces the onset of IR.^{10,11} IR, rather than BMI-based obesity evaluation, is strongly associated with CKD because IR induces CKD risk factors, including glomerular hyperfiltration, sodium retention, defective tubular reabsorption, tissue inflammation, and fibrosis.¹²⁻¹⁴ Therefore, IR is more likely related to CKD than obesity.

The homeostasis model assessment of insulin resistance (HOMA-IR) has been widely used to examine insulin sensitivity for many years.¹⁵ The triglyceride-glucose (TyG) index was strongly related to hyperinsulinemic-euglycemic clamp data collected in Brazil, Mexico, and South Korea.¹⁶⁻¹⁸ Additionally, the TyG index is better than the HOMA-IR index for identifying various IR-related health concerns such as arterial stiffness, hypertension, and non-alcoholic steatohepatitis.¹⁹⁻²¹ Therefore, the TyG index is a reliable and valid indicator of IR that is superior to the HOMA-IR.

We hypothesized that IR is associated with CKD independent of obesity and sex and that an increased TyG index can be used for the early screening and detection of high-risk geriatric populations with CKD. Based on this hypothesis, we conducted a population-based cross-sectional study to examine the association of the TyG index with CKD in the geriatric population, regardless of obesity and sex.

MATERIALS AND METHODS

Study Design and Subjects

We analyzed data from a database of South Koreans' general health, nutritional status, and lifestyle data from the Korea National Health and Nutritional Examination Survey (KNHANES) 2014–2018. The analysis included 7,326 participants (2,864 men and 4,462 women) among all participants aged ≥ 60 years from the 2014–2018 KNHANES. Fig. 1 shows a flowchart of participant recruitment. All participants provided written informed consent, and the study was conducted in accordance with the principles of the Declaration of Helsinki and approved by the Institutional Review Board of Silla University (No. 1041449-202203-HR-001).

This study complied the ethical guidelines for authorship and publishing in the *Annals of Geriatric Medicine and Research*.²²

TyG Index and eGFR

Blood samples were collected in the morning after a fast of at least 8 hours. Circulating glucose and triglyceride concentrations were measured by enzymatic methods using a Hitachi automatic analyzer 7600 (Hitachi, Tokyo, Japan). The TyG index was calculated as follows.¹⁷

$\ln [\text{triglyceride concentration (mg/dL)} \times \text{fasting plasma glucose concentration (mg/dL)}] / 2$.

Creatinine concentrations were analyzed using the Jaffe rate-blanked creatinine assay and compensated at a certified laboratory (Seegene Medical Foundation, Seoul, Korea). The estimated glomerular filtration rate (eGFR) was calculated using the new Japanese-coefficient modified MDRD (Modification of Diet in Renal Disease) study equation as follows.^{8,9,23-25}

$\text{eGFR (mL/min/1.73 m}^2) = 194 \times (\text{serum creatinine})^{-1.094} \times (\text{age})^{-0.287} (\times 0.739 \text{ for females})$.

All participants were assigned into groups according to TyG tertiles. Moderate-to-severe CKD ($_{MS}CKD$) was defined as an eGFR $< 45.0 \text{ mL/min/1.73 m}^2$.²³⁻²⁵

Statistical Analysis

All data are shown as mean \pm standard deviation. Independent t-test or Mann–Whitney U tests were used to compare male and female variables. A one-way analysis of variance (ANOVA) was used to compare the anthropometric and biochemical characteristics of the three TyG index groups. The Bonferroni post-hoc test was applied when ANOVA showed significant differences ($p < 0.05$). The Mann–Whitney U test was used to analyze differences between groups with non-normal data distributions ($p < 0.05$). The Jonckheere–Terpstra test was used to compare the values between the three groups (two-tailed, $p < 0.05$). The Jonckheere–Terpstra test generates standardized statistics (SS) that point to the strength of tendencies in variables that increase or decline across groups.²⁶⁻²⁸ We applied logistic regression to evaluate the obesity- and sex-specific associations between the TyG index

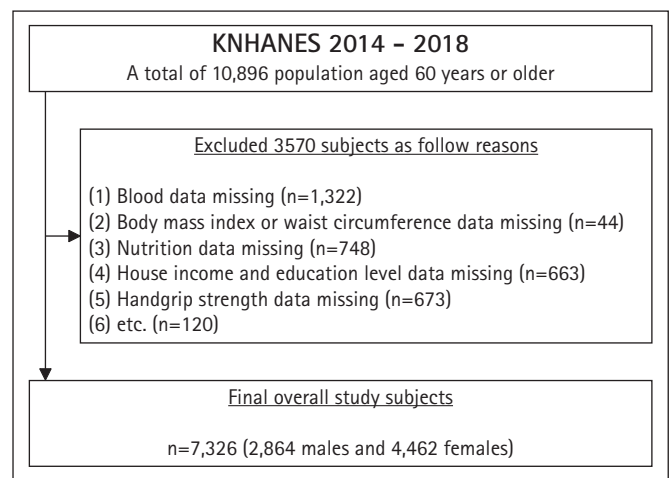


Fig. 1. Flowchart of the subjects. KNHANES, Korea National Health and Nutritional Examination Survey.

and $_{MS}CKD$. The fully adjusted model was adjusted for potential confounders such as education level, household income, smoking, drinking, handgrip strength, moderate-to-vigorous physical activity, total energy intake, and BMI, which are recognized or suspected factors associated with CKD. IBM SPSS Statistics for Windows, version 26.0 (IBM Corp., Armonk, NY, USA) was used to perform the statistical analyses. The optimal cutoff values for the TyG index in male and female participants with or without obesity to predict $_{MS}CKD$ were derived from receiver operating characteristic (ROC) curve analysis (area under the ROC curve [AUC] values). The sensitivity and specificity were also calculated. We used MedCalc for Windows version 9.1.0.1 (MedCalc, Ostend, Belgium).

RESULTS

Table 1 presents the participants' characteristics. The mean age of the participants was 66.1 ± 10.5 years and was significantly higher in male participants than in females ($p < 0.001$). The mean TyG index and eGFR were 8.6 ± 0.6 and 62.8 ± 13.4 mL/min/1.73m², respectively. The eGFR in male participants was significantly lower than that in females ($p < 0.001$). The TyG index did not differ significantly between the sexes. **Supplementary Table S1** provides more information on the subjects.

Table 2 displays the obesity- and sex-specific differences and tendencies based on the eGFR tertiles in male subjects. In subjects with and without obesity, the tendency test indicated a significant decrease in eGFR from the lowest to highest TyG tertiles (SS -5.61 and -3.59, respectively; both $p < 0.001$). Post hoc testing revealed that the mean eGFR in the lowest tertile was significantly higher

than that in the middle and highest tertile. Similar to the male participants, the eGFR values in female participants with and without obesity are shown in **Table 3**. **Supplementary Tables S2** and **S3** provide the results of the analyses of additional variables and several covariates for male and female participants, respectively.

Table 4 compares the obesity- and sex-specific odds ratios for an association between the TyG index and $_{MS}CKD$. Male and female participants with and without obesity were divided into tertiles based on TyG index values. For male participants without obesity, in the unadjusted model, the middle and highest tertiles displayed odds ratios of 2.392 (95% confidence interval [CI], 1.522–3.760) and 2.439 (95% CI, 1.552–3.835), respectively, compared to the lowest tertile in $_{MS}CKD$. In the fully adjusted model, the middle and highest tertiles showed odds ratios of 2.342 (95% CI, 1.464–3.747) and 2.393 (95% CI, 1.498–3.823), respectively, relative to the lowest tertile in $_{MS}CKD$. Regarding female participants without obesity, in the unadjusted model, the highest tertile displayed an odds ratio of 2.123, relative to the lowest tertile (95% CI, 1.411–3.194) in $_{MS}CKD$. In the fully adjusted model, the highest tertile showed an odds ratio of 2.374 relative to the lowest tertile (95% CI, 1.539–3.662) of $_{MS}CKD$. Among male subjects with obesity, in the unadjusted model, the highest tertile displayed an odds ratio of 1.620 relative to the lowest tertile (95% CI, 1.016–2.584) for $_{MS}CKD$. In the fully adjusted model, the highest tertile showed an odds ratio of 1.736 relative to the lowest tertile (95% CI, 1.053–2.863) in $_{MS}CKD$. Regarding obese female subjects, in the unadjusted model, the middle and highest tertiles had odds ratios of 2.216 (95% CI, 1.361–3.606) and 3.141 (95% CI, 1.975–4.974), respectively, relative to the lowest tertile of $_{MS}CKD$. In the fully ad-

Table 1. Characteristics of the subjects

	Overall (n = 7,326)	Male (n = 2,864)	Female (n = 4,462)	p-value
Age (y)	66.1 ± 10.5	69.4 ± 6.1	64.0 ± 12.1	< 0.001 ^{a)}
TyG index	8.6 ± 0.6	8.6 ± 0.5	8.6 ± 0.6	0.148
eGFR (mg/dL)	62.8 ± 13.4	60.6 ± 12.6	64.2 ± 13.7	< 0.001 ^{a)}
Height (cm)	158.9 ± 8.4	166.1 ± 5.8	154.3 ± 6.2	< 0.001 ^{a)}
Body mass (kg)	61.5 ± 9.2	67.4 ± 8.5	57.8 ± 7.4	< 0.001 ^{a)}
Body mass index (kg/m ²)	24.3 ± 2.8	24.4 ± 2.5	24.3 ± 2.9	< 0.001 ^{a)}
Waist circumference (cm)	84.7 ± 8.7	88.1 ± 7.6	82.6 ± 8.7	< 0.001 ^{a)}
FPG (mg/dL)	100.3 ± 20.2	100.3 ± 19.9	100.4 ± 20.3	0.868
HbA1c (%)	5.7 ± 0.7	5.7 ± 0.7	5.7 ± 0.7	0.385
Triglyceride (mg/dL)	124.5 ± 66.8	125.2 ± 65.0	124.0 ± 67.9	0.465
Creatinine (mg/dL)	0.85 ± 0.28	0.82 ± 0.29	0.85 ± 0.28	< 0.001 ^{a)}
Obesity status				< 0.01
Non-obese subjects	4,670 (63.7)	1,766 (61.7)	2,904 (65.1)	
Obese subjects	2,656 (36.2)	1,098 (38.3)	1,558 (34.9)	

Values are presented as mean ± standard deviation or number (%).

eGFR, estimated glomerular filtration rate; FPG, fasting plasma glucose; TyG index, triglyceride-glucose index.

^{a)}The Mann–Whitney U test was applied to assess differences between groups.

Table 2. Anthropometric data, the TyG index, and eGFR-related biochemical characteristics and trends according to TyG index tertiles in male subjects

	Male non-obese subjects				Male obese subjects							
	A (n = 604)	B (n = 590)	C (n = 593)	Post-hoc	ss ^{b)}	p-value ^{b)}	A (n = 359)	B (n = 362)	C (n = 356)	Post-hoc	ss ^{b)}	p-value ^{b)}
TyG index	8.00 ± 0.24	8.59 ± 0.15	9.22 ± 0.27	A < B < C	44.82	< 0.001	8.01 ± 0.25	8.61 ± 0.15	9.23 ± 0.25	A < B < C	34.79	< 0.001
eGFR (mg/dL)	64.0 ± 12.4	60.6 ± 12.6	60.0 ± 12.2	A > B, C	-5.61	< 0.001	61.1 ± 13.1	58.4 ± 12.1	57.7 ± 12.4	A > B, C	-3.59	< 0.001
Body mass index (kg/m ²)	22.9 ± 1.3	22.8 ± 1.3	22.8 ± 1.4	ns	-0.59	0.553	26.9 ± 1.7	27.0 ± 1.9	27.0 ± 1.7	ns	0.31	0.756
Age (yr)	69.8 ± 6.2	69.8 ± 6.3	70.0 ± 6.3	ns	0.58	0.564	68.0 ± 5.6	69.0 ± 5.9	69.2 ± 6.0	A < B, C	2.69	< 0.01 ^{a)}
Height (cm)	166.0 ± 5.8	165.9 ± 5.8	165.9 ± 5.7	ns	-0.44	0.66	166.3 ± 5.9	166.6 ± 5.7	166.0 ± 5.8	ns	-0.49	0.625
Body mass (kg)	63.1 ± 5.7	62.8 ± 5.9	62.9 ± 5.9	ns	-0.31	0.756	74.5 ± 7.2	75.0 ± 7.1	74.5 ± 7.2	ns	0.05	0.960
Waist circumference (cm)	84.5 ± 5.6	84.2 ± 5.3	84.3 ± 5.8	ns	-0.49	0.625	94.0 ± 6.2	94.5 ± 6.4	94.4 ± 6.0	ns	1.18	0.238
FPG (mg/dL)	92.0 ± 9.9	99.3 ± 15.8	108.6 ± 24.5	A < B < C	17.23	< 0.001 ^{a)}	92.8 ± 10.6	99.2 ± 13.7	110.9 ± 30.0	A < B < C	13.84	< 0.001 ^{a)}
Triglyceride (mg/dL)	67.0 ± 15.1	111.8 ± 20.9	198.0 ± 59.7	A < B < C	42.49	< 0.001 ^{a)}	67.5 ± 15.4	113.3 ± 21.0	195.2 ± 54.8	A < B < C	32.94	< 0.001 ^{a)}
Creatinine (mg/dL)	0.82 ± 0.16	0.86 ± 0.18	0.88 ± 0.18	A < B, C	6.16	< 0.001 ^{a)}	0.83 ± 0.15	0.88 ± 0.48	0.90 ± 0.17	A < B, C	5.15	< 0.001

Values are presented as mean ± standard deviation.

A group is lowest tertile; B, middle tertile; and C, highest tertile.

TyG index, triglyceride-glucose index; eGFR, estimated glomerular filtration rate; FPG, fasting plasma glucose; ss, standardized statistics; ns, not significant.

^{a)}The Mann-Whitney U test was applied to assess differences between the three groups.

^{b)}The Jonckheere-Terpstra test was used to assess the trend among the three groups.

Table 3. Anthropometric data, the TyG index, and eGFR-related biochemical characteristics and trends according to TyG index tertiles in female subjects

	Female non-obese subjects				Female obese subjects							
	A (n = 973)	B (n = 988)	C (n = 973)	Post-hoc	ss ^{b)}	p-value ^{b)}	A (n = 505)	B (n = 510)	C (n = 513)	Post-hoc	ss ^{b)}	p-value ^{b)}
TyG index	7.95 ± 0.25	8.57 ± 0.15	9.22 ± 0.28	A < B < C	57.44	< 0.001	7.97 ± 0.25	8.60 ± 0.15	9.24 ± 0.28	A < B < C	41.45	< 0.001
eGFR (mg/dL)	68.2 ± 13.9	65.6 ± 12.9	64.0 ± 13.5	A > B > C	-6.18	< 0.001 ^{a)}	63.6 ± 13.4	60.4 ± 13.3	59.1 ± 13.3	A > B, C	-4.54	< 0.001
Body mass index (kg/m ²)	22.6 ± 1.4	22.6 ± 1.3	22.6 ± 1.4	ns	0.41	0.686 ^{a)}	27.5 ± 2.3	27.6 ± 2.3	27.7 ± 2.3	ns	1.37	0.170
Age (yr)	60.4 ± 13.5	61.4 ± 12.9	61.5 ± 13.6	ns	1.79	0.074 ^{a)}	69.4 ± 6.3	69.4 ± 6.4	70.3 ± 6.2	ns	2.22	< 0.05
Height (cm)	155.3 ± 6.5	155.4 ± 6.5	155.0 ± 6.3	ns	-1.07	0.286	152.7 ± 5.4	152.5 ± 5.7	152.1 ± 5.1	ns	-1.51	0.131
Body mass (kg)	54.6 ± 5.2	54.6 ± 5.3	54.4 ± 5.4	ns	-0.71	0.478	64.2 ± 6.7	64.1 ± 7.0	64.1 ± 7.0	ns	-0.95	0.34
Waist circumference (cm)	78.2 ± 6.0	78.5 ± 6.0	78.1 ± 6.0	ns	-0.31	0.758	90.7 ± 6.8	90.5 ± 6.9	91.1 ± 7.0	ns	1.32	0.187
FPG (mg/dL)	91.8 ± 8.9	98.5 ± 14.2	109.4 ± 26.1	A < B < C	22.93	< 0.001 ^{a)}	92.4 ± 10.1	99.9 ± 15.2	111.5 ± 30.8	A < B < C	16.91	< 0.001 ^{a)}
Triglyceride (mg/dL)	63.9 ± 15.0	110.2 ± 19.4	197.5 ± 64.7	A < B < C	54.58	< 0.001 ^{a)}	64.9 ± 14.9	109.4 ± 19.4	198.4 ± 61.0	A < B < C	39.42	< 0.001 ^{a)}
Creatinine (mg/dL)	0.80 ± 0.45	0.83 ± 0.28	0.88 ± 0.27	A, B < C	10.99	< 0.001	0.82 ± 0.14	0.84 ± 0.17	0.88 ± 0.22	A < B < C	5.47	< 0.001 ^{a)}

Values are presented as mean ± standard deviation.

A group is lowest tertile; B, middle tertile; and C, highest tertile.

TyG index, triglyceride-glucose index; eGFR, estimated glomerular filtration rate; FPG, fasting plasma glucose; ss, standardized statistics; ns, not significant.

^{a)}The Mann-Whitney U test was applied to assess differences between the three groups.

^{b)}The Jonckheere-Terpstra test was used to assess the trend among the three groups.

Table 4. Obesity- and sex-specific odds ratios for the relationship between the triglyceride-glucose index and moderate-to-severe chronic kidney disease

	Unadjusted model	Fully adjusted model
Male non-obese subjects		
A (n = 593)	2.439 (1.552–3.835)***	2.393 (1.498–3.823)***
B (n = 590)	2.392 (1.522–3.760)***	2.342 (1.464–3.747)***
C (n = 604)	Reference	Reference
Male obese subjects		
A (n = 356)	1.620 (1.016–2.584)*	1.736 (1.053–2.863)*
B (n = 362)	1.355 (0.841–2.185)	1.485 (0.895–2.463)
C (n = 359)	Reference	Reference
Female non-obese subjects		
A (n = 973)	2.123 (1.411–3.194)***	2.374 (1.539–3.662)***
B (n = 988)	1.388 (0.900–2.141)	1.499 (0.952–2.360)
C (n = 973)	Reference	Reference
Female obese subjects		
A (n = 513)	3.141 (1.975–4.974)***	3.516 (2.164–5.713)***
B (n = 510)	2.216 (1.361–3.606)**	2.313 (1.397–3.828)**
C (n = 505)	Reference	Reference

Values are presented as odds ratio (95% confidence interval).

A group is highest tertile; B, middle tertile; and C, lowest tertile. The fully adjusted model was adjusted for education level, house income, medication, smoking, drinking, hand-grip strength, moderate to vigorous physical activity, total energy intake and body mass index.

* $p < 0.05$, ** $p < 0.01$, *** $p < 0.001$.

justed model, the middle and highest tertiles showed odds ratios of 2.313 (95% CI, 1.397–3.828) and 3.516 (95% CI, 2.164–5.713), respectively, relative to the lowest tertile in $_{MS}CKD$.

DISCUSSION

This study investigated whether the TyG index, which reflects IR, is associated with CKD in a geriatric population, independent of obesity and sex. The results showed that the TyG index was associated with CKD in the geriatric population, regardless of obesity. Additionally, the relationship between the TyG index and CKD was significant in both male and female participants. These findings suggested that IR is associated with CKD in the geriatric population independent of obesity and sex.

The long-standing consensus is that aging and obesity trigger a decline in kidney function. The effect of aging on CKD is undeniable due to the high global prevalence of CKD in the geriatric population. However, the association between obesity and CKD is increasing, mainly owing to the limitations of BMI. BMI is not an accurate indicator of overall adiposity and visceral fat, which induces the onset of IR.^{10,11} Recent studies published in 2019, 2020, and 2021 reported that IR was strongly related to a decline in kidney function rather than obesity per se.²⁹⁻³¹ Our findings on IR also support these recent reports. In the present study, both male and female participants (Tables 2, 3) showed a stronger decreasing trend in eGFR in subjects without obesity than in those with obe-

sity as the TyG index increased. If obesity causes a decline in kidney function, the decreasing trend in eGFR should be more pronounced in subjects with obesity than in those without. This suggests that the effect of IR on CKD is more significant than that on obesity.

We also found that male participants without obesity in the middle and highest tertiles of the TyG index were 2.342 and 2.393 times more likely to have $_{MS}CKD$, respectively. Similarly, female participants without obesity in the highest tertile were 2.374 times more likely to develop $_{MS}CKD$ (Table 4). Male participants with obesity in the highest tertile of the TyG index were 1.736 times more likely to develop $_{MS}CKD$. Additionally, female subjects with obesity in the middle and highest tertiles of the TyG index were 2.313 and 3.516 times more likely to have $_{MS}CKD$, respectively (Table 4). These findings suggest that increased IR is an independent risk factor for CKD in both men and women, regardless of obesity status. Therefore, IR is a primary pathophysiology that may be independently associated with CKD in geriatric populations regardless of obesity.

Several studies have reported a relationship between IR and CKD.^{32,33} Animal and human studies have reported that hyperinsulinemia leads to kidney vasodilatation, enhances sodium reabsorption, stimulates the renin-angiotensin system, and causes glomerular hyperfiltration, which increases the GFR.³⁴⁻³⁶ Increased filtration per nephron causes nephron loss and leads to glomerular hypertension, which causes glomerular sclerosis and a subsequent

decline in kidney function.³⁷⁾ In addition, clinical studies have shown pre-existing IR in individuals with a mild decline in kidney function.^{38,39)} The relationship between IR and CKD can be explained via biological mechanisms such as inflammation, oxidative stress, and metabolic acidosis.

Studies have demonstrated the good performance of the TyG index in predicting or discriminating IR-related health concerns. However, specific cutoff values have not been confirmed, and few studies have suggested the potential value of the TyG index. Shin⁴⁰⁾ studied 4,415 Korean adults aged 20–80 years and showed that a TyG index cutoff value of ≥ 8.81 discriminated individuals with IR with AUC of 0.894, sensitivity of 86.7%, and specificity of 80.1%. Endukuru et al.⁴¹⁾ studied 150 Indian adults aged 18–65 years and found that a TyG index cutoff value for IR of ≥ 9.88 showed AUC of 0.836, sensitivity of 76.0%, and specificity of 88.0%. The value suggested by Endukuru et al.⁴¹⁾ was derived from only 150 adults, and the AUC was relatively low compared with that suggested by Shin.⁴⁰⁾ In the present study, the potential cutoff value of the TyG index to distinguish individuals with _{MS}CKD, was > 8.62 (AUC 0.584; sensitivity 59.34%; specificity 53.26%) (Supplementary Fig. S1). Considering the differences in population age, race, and number, the potential cutoff value in the present study cannot be directly compared with those obtained in the two previous studies. Additionally, the relatively low AUC, sensitivity, and specificity of the potential TyG index cutoff value in the present study suggest the need for re-examination using another sample of the geriatric population. However, as a geriatric population-specific cutoff value to distinguish CKD in high-risk individuals in the early stages, the potential TyG index cutoff value identified in the present study may be appropriate for clinical practice.

Previous studies in the Korean population have provided TyG index cutoff values to discriminate high-risk individuals with several health concerns. Kim et al.²⁷⁾ found that TyG index values of ≥ 8.72 and 8.67 , respectively, were risk factors for sarcopenic obesity in men and women aged ≥ 60 years with health issues such as hypertension and hyperlipidemia. Kang et al.⁴²⁾ reported a TyG index value of ≥ 8.83 as a cutoff value for obstructive sleep apnea in men and women with health issues aged ≥ 55 years. Park et al.⁴³⁾ showed that a TyG index value of ≥ 8.44 was a cutoff value of coronary artery disease in men and women aged ≥ 65 years without health issues. Differences in age distribution, inconsistency in sex-specific populations, and differences in basic health status make comparing and determining a precise TyG index cutoff value for all health concerns in the Korean population difficult. However, these reports suggest that late middle-aged and older adults with health issues and a TyG index value of ≥ 8.6 require careful monitoring to suppress the progression of health concerns. In

healthy geriatric populations, a TyG index value of ≥ 8.44 may be applied as a cutoff for early-stage prevention of health concerns.

The present study had several strengths and limitations. This study's strength was the adjustment for potential covariates, such as demographic parameters and lifestyle factors that might affect the relationship between the TyG index and CKD. However, the study subjects were older Korean adults; thus, whether the findings of the present study can be applied to other ethnicities or nations is unclear. Further investigations in different races are needed to confirm the association between the TyG index and CKD.

In conclusion, the geriatric population with an increased TyG index has a high risk of CKD regardless of obesity and sex. This finding suggests that increased IR is associated with CKD in the geriatric population independent of obesity and sex.

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CONFLICT OF INTEREST

The researchers claim no conflicts of interest.

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AUTHOR CONTRIBUTIONS

Conceptualization, BK, SO; Data curation, BK, GK; Funding acquisition, BK, NM; Investigation, BK, SO, KH; Methodology, NM, KT; Project administration, BK, KH; Supervision, BK, GK; Writing-original draft, BK; Writing-review & editing, BK, SO.

SUPPLEMENTARY MATERIALS

Supplementary materials can be found via <https://doi.org/10.4235/agmr.23.0096>.

REFERENCES

1. Schefold JC, Filippatos G, Hasenfuss G, Anker SD, von Haehling S. Heart failure and kidney dysfunction: epidemiology, mechanisms and management. *Nat Rev Nephrol* 2016;12:610-23.
2. Mukai H, Ming P, Lindholm B, Heimburger O, Barany P, Stenvinkel P, et al. Lung dysfunction and mortality in patients with

- chronic kidney disease. *Kidney Blood Press Res* 2018;43:522-35.
3. Kalantar-Zadeh K, Jafar TH, Nitsch D, Neuen BL, Perkovic V. Chronic kidney disease. *Lancet* 2021;398:786-802.
 4. GBD Chronic Kidney Disease Collaboration. Global, regional, and national burden of chronic kidney disease, 1990-2017: a systematic analysis for the Global Burden of Disease Study 2017. *Lancet* 2020;395:709-33.
 5. Tsujimoto T, Sairenchi T, Iso H, Irie F, Yamagishi K, Watanabe H, et al. The dose-response relationship between body mass index and the risk of incident stage ≥ 3 chronic kidney disease in a general Japanese population: the Ibaraki prefectural health study (IPHS). *J Epidemiol* 2014;24:444-51.
 6. Lu JL, Molnar MZ, Naseer A, Mikkelsen MK, Kalantar-Zadeh K, Kovesdy CP. Association of age and BMI with kidney function and mortality: a cohort study. *Lancet Diabetes Endocrinol* 2015;3:704-14.
 7. Chang TJ, Zheng CM, Wu MY, Chen TT, Wu YC, Wu YL, et al. Relationship between body mass index and renal function deterioration among the Taiwanese chronic kidney disease population. *Sci Rep* 2018;8:6908.
 8. Kim B, Kim G, Kim E, Park J, Isobe T, Sakae T, et al. The A Body Shape Index Might Be a Stronger Predictor of Chronic Kidney Disease Than BMI in a Senior Population. *Int J Environ Res Public Health* 2021;18:12874.
 9. Kim B, Park H, Kim G, Isobe T, Sakae T, Oh S. Relationships of fat and muscle mass with chronic kidney disease in older adults: a cross-sectional pilot study. *Int J Environ Res Public Health* 2020;17:9124.
 10. de Mutsert R, Gast K, Widya R, de Koning E, Jazet I, Lamb H, et al. Associations of abdominal subcutaneous and visceral fat with insulin resistance and secretion differ between men and women: the Netherlands epidemiology of obesity study. *Metab Syndr Relat Disord* 2018;16:54-63.
 11. Rothman KJ. BMI-related errors in the measurement of obesity. *Int J Obes (Lond)* 2008;32 Suppl 3:S56-9.
 12. Moon JH, Kim JS. The cutoff value in body fat percentage for increased risk of metabolic syndrome in elderly people with normal body weight. *J Korean Geriatr Soc* 2015;19:16-24.
 13. Whaley-Connell A, Sowers JR. Insulin resistance in kidney disease: is there a distinct role separate from that of diabetes or obesity? *Cardiorenal Med* 2017;8:41-9.
 14. Artunc F, Schleicher E, Weigert C, Fritsche A, Stefan N, Haring HU. The impact of insulin resistance on the kidney and vasculature. *Nat Rev Nephrol* 2016;12:721-37.
 15. Matthews DR, Hosker JP, Rudenski AS, Naylor BA, Treacher DF, Turner RC. Homeostasis model assessment: insulin resistance and beta-cell function from fasting plasma glucose and insulin concentrations in man. *Diabetologia* 1985;28:412-9.
 16. Lee S, Choi S, Kim HJ, Chung YS, Lee KW, Lee HC, et al. Cutoff values of surrogate measures of insulin resistance for metabolic syndrome in Korean non-diabetic adults. *J Korean Med Sci* 2006;21:695-700.
 17. Simental-Mendia LE, Rodriguez-Moran M, Guerrero-Romero F. The product of fasting glucose and triglycerides as surrogate for identifying insulin resistance in apparently healthy subjects. *Metab Syndr Relat Disord* 2008;6:299-304.
 18. Vasques AC, Novaes FS, de Oliveira Mda S, Souza JR, Yamanaka A, Pareja JC, et al. TyG index performs better than HOMA in a Brazilian population: a hyperglycemic clamp validated study. *Diabetes Res Clin Pract* 2011;93:e98-100.
 19. Wang Y, Yang W, Jiang X. Association between triglyceride-glucose index and hypertension: a meta-analysis. *Front Cardiovasc Med* 2021;8:644035.
 20. Riviere B, Jaussent A, Macioce V, Faure S, Builles N, Lefebvre P, et al. The triglycerides and glucose (TyG) index: a new marker associated with nonalcoholic steatohepatitis (NASH) in obese patients. *Diabetes Metab* 2022;48:101345.
 21. Wu S, Xu L, Wu M, Chen S, Wang Y, Tian Y. Association between triglyceride-glucose index and risk of arterial stiffness: a cohort study. *Cardiovasc Diabetol* 2021;20:146.
 22. Noh JH, Jung HW, Ga H, Lim JY. Ethical guidelines for publishing in the *Annals of Geriatric Medicine and Research*. *Ann Geriatr Med Res* 2022;26:1-3.
 23. Matsuo S, Imai E, Horio M, Yasuda Y, Tomita K, Nitta K, et al. Revised equations for estimated GFR from serum creatinine in Japan. *Am J Kidney Dis* 2009;53:982-92.
 24. Kim B, Kim GM, Oh S. Use of the visceral adiposity index as an indicator of chronic kidney disease in older adults: comparison with body mass index. *J Clin Med* 2022;11:6297.
 25. Lee S, Shimada H, Park H, Makizako H, Lee S, Doi T, et al. The association between kidney function and cognitive decline in community-dwelling, elderly Japanese people. *J Am Med Dir Assoc* 2015;16:349.
 26. Kim B, Ku M, Kiyoji T, Isobe T, Sakae T, Oh S. Cardiorespiratory fitness is strongly linked to metabolic syndrome among physical fitness components: a retrospective cross-sectional study. *J Physiol Anthropol* 2020;39:30.
 27. Kim B, Kim G, Lee Y, Taniguchi K, Isobe T, Oh S. Triglyceride-glucose index as a potential indicator of sarcopenic obesity in older people. *Nutrients* 2023;15:555.
 28. Nakanishi Y, Tsugihashi Y, Akahane M, Noda T, Nishioka Y, Myojin T, et al. Comparison of Japanese centenarians' and non-centenarians' medical expenditures in the last year of life. *JAMA*

- Netw Open 2021;4:e2131884.
29. Chen HY, Lu FH, Chang CJ, Wang RS, Yang YC, Chang YF, et al. Metabolic abnormalities, but not obesity per se, associated with chronic kidney disease in a Taiwanese population. *Nutr Metab Cardiovasc Dis* 2020;30:418-25.
 30. Fritz J, Brozek W, Concin H, Nagel G, Kerschbaum J, Lhotta K, et al. The triglyceride-glucose index and obesity-related risk of end-stage kidney disease in Austrian adults. *JAMA Netw Open* 2021;4:e212612.
 31. Kawamoto R, Akase T, Ninomiya D, Kumagi T, Kikuchi A. Metabolic syndrome is a predictor of decreased renal function among community-dwelling middle-aged and elderly Japanese. *Int Urol Nephrol* 2019;51:2285-94.
 32. Kobayashi H, Tokudome G, Hara Y, Sugano N, Endo S, Suetsugu Y, et al. Insulin resistance is a risk factor for the progression of chronic kidney disease. *Clin Nephrol* 2009;71:643-51.
 33. Kobayashi S, Maesato K, Moriya H, Ohtake T, Ikeda T. Insulin resistance in patients with chronic kidney disease. *Am J Kidney Dis* 2005;45:275-80.
 34. Esteghamati A, Ashraf H, Nakhjavani M, Najafian B, Hamidi S, Abbasi M. Insulin resistance is an independent correlate of increased urine albumin excretion: a cross-sectional study in Iranian Type 2 diabetic patients. *Diabet Med* 2009;26:177-81.
 35. Tucker BJ, Anderson CM, Thies RS, Collins RC, Blantz RC. Glomerular hemodynamic alterations during acute hyperinsulinemia in normal and diabetic rats. *Kidney Int* 1992;42:1160-8.
 36. Dengel DR, Goldberg AP, Mayuga RS, Kairis GM, Weir MR. Insulin resistance, elevated glomerular filtration fraction, and renal injury. *Hypertension* 1996;28:127-32.
 37. Helal I, Fick-Brosnahan GM, Reed-Gitomer B, Schrier RW. Glomerular hyperfiltration: definitions, mechanisms and clinical implications. *Nat Rev Nephrol* 2012;8:293-300.
 38. Kato Y, Hayashi M, Ohno Y, Suzawa T, Sasaki T, Saruta T. Mild renal dysfunction is associated with insulin resistance in chronic glomerulonephritis. *Clin Nephrol* 2000;54:366-73.
 39. Chen J, Muntner P, Hamm LL, Fonseca V, Batuman V, Whelton PK, et al. Insulin resistance and risk of chronic kidney disease in nondiabetic US adults. *J Am Soc Nephrol* 2003;14:469-77.
 40. Shin KA. Triglyceride and glucose (TyG) index is a clinical surrogate marker for the diagnosis of metabolic syndrome. *Biomed Sci Lett* 2017;23:348-54.
 41. Endukuru CK, Gaur GS, Yerrabelli D, Sahoo J, Vairappan B. Cut-off values and clinical utility of surrogate markers for insulin resistance and beta-cell function to identify metabolic syndrome and its components among Southern Indian adults. *J Obes Metab Syndr* 2020;29:281-91.
 42. Kang HH, Kim SW, Lee SH. Association between triglyceride glucose index and obstructive sleep apnea risk in Korean adults: a cross-sectional cohort study. *Lipids Health Dis* 2020;19:182.
 43. Park GM, Cho YR, Won KB, Yang YJ, Park S, Ann SH, et al. Triglyceride glucose index is a useful marker for predicting subclinical coronary artery disease in the absence of traditional risk factors. *Lipids Health Dis* 2020;19:7.