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Association between estimated glucose disposal rate and metabolic syndrome: a cross-sectional analysis of the National Health and Nutrition Examination Survey

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Metabolic syndrome (MS) is a complex metabolic disorder that is often closely associated with the development of chronic diseases such as cardiovascular disease and diabetes. This study aimed to explore the relationship between estimated glucose metabolic rate (eGDR) and MS. The correlation between eGDR levels and the prevalence of metabolic syndrome was analyzed here based on data from the National Health and Nutrition Examination Survey from 2005 to 2020. The study sample consisted of 63,131 adult participants, and the results showed that lower eGDR levels were significantly associated with a higher prevalence of metabolic syndrome. Further regression analyses showed that eGDR acted as a protective factor and that the risk of MS significantly decreased as its level increased. Subgroup analyses showed that this trend held across gender, age, and BMI categories, and that the protective effect of eGDR was weaker in the higher BMI group. Based on the nonlinear relationship between subjects' eGDR levels and MS prevalence, RCS analyses further confirmed a significant correlation between lower eGDR levels and increased risk of MS. In conclusion, the present study suggests that eGDR levels could serve as a potential biomarker for predicting metabolic syndrome, providing new perspectives for early screening and intervention of MS.

KEYWORDS

metabolic syndrome, NHANES, eGDR, logistic regression, RCS

1 Introduction

Metabolic syndrome (MS) is a complex metabolic disorder characterized by central obesity, atherosclerotic dyslipidemia, hypertension, insulin resistance, prothrombotic and proinflammatory states, among other features (1, 2). Approximately 20 to 25% of adults worldwide are affected by MS (3, 4). According to data from the National Health and Nutrition Examination Survey (NHANES), about one-third of adults were diagnosed with MS between 1988 and 2010 (5). In China, the prevalence of MS among adults was 24.2% in 2018, with a higher rate of 24.6% in males compared to 23.8% in females (6). Early prediction and diagnosis of MS are crucial for timely intervention, which can not only effectively reduce health risks

associated with MS but also alleviate the burden on the healthcare system (7, 8).

Currently, the diagnostic criteria for MS include waist circumference, blood glucose, blood pressure, triglycerides, and highdensity lipoprotein cholesterol (HDL-C) (2). In addition, composite indices based on combinations of different metrics, such as the glucose–glucose index (TyG) and TyG-waist circumference, have demonstrated better results in the diagnosis and prognosis of MS (9, 10). Therefore, establishing and evaluating the impact of various physiological and biochemical indicators on the prevalence of MS is of significant theoretical and practical importance.

Williams et al. developed an index for assessing insulin resistance, known as the estimated glucose disposal rate (eGDR), based on the euglycaemic hyperinsulinaemic clamp (11). The adjusted eGDR formula proposed by Epstein and colleagues has since been more widely applied (12). The index integrates clinical data such as blood pressure, hemoglobin A1c levels, and waist circumference, and is simple to calculate. Although eGDR was originally developed to assess insulin resistance in individuals with type 1 diabetes, it has since been recognized as a general biomarker for metabolic syndrome and is closely associated with various diseases, including cardiovascular diseases and diabetes (13-15). However, research on the relationship between eGDR levels and the prevalence of MS is still insufficient. Therefore, this study will conduct a retrospective analysis using data from the National Health and Nutrition Examination Survey conducted between 2005 and 2020, focusing on individuals with MS. The primary objective is to systematically explore the potential of eGDR levels as a predictive marker for MS prevalence, providing a scientific foundation for its potential role in the prevention and treatment of MS.

2 Materials and methods

2.1 Sample sources

The data used in this study were derived from the National Health and Nutrition Examination Survey, a major program of the National Center for Health Statistics that provides national health and nutrition statistics. The NHANES aims to assess the health and nutritional status of adults and children in the United States, utilizing a combination of interviews and physical examinations (16). This study collected data on eGDR levels and metabolic syndrome across three survey cycles: 2005–2006, 2007–2008, and 2009–2010. The NHANES protocol has been approved by the NCHS Research Ethics Review Board and by all participants, eliminating the need for additional institutional review board approval. The primary objective of this study was to investigate the relationship between eGDR levels and the prevalence of metabolic syndrome.

2.2 About eGDR levels

The eGDR is a clinical indicator used to assess an individual's insulin sensitivity and the degree of insulin resistance, reflecting the effectiveness of insulin metabolism. The calculation of eGDR levels incorporates easily accessible clinical parameters, such as glycated hemoglobin (HbA1c), blood pressure status (hypertension or normal blood pressure), and waist circumference. The formula for calculating eGDR is as follows: eGDR (mg/kg/min) = $21.158 - (0.09 \times \text{Waist}$ circumference in cm) - ($3.407 \times \text{Hypertension}$, 1 = Yes, 0 = No) - ($0.551 \times \text{HbA1c\%}$) (12). A decrease in eGDR levels typically indicates an increase in insulin resistance, while higher eGDR values are associated with better insulin sensitivity.

2.3 Definition of metabolic syndrome

MS is a pathological condition characterized by a cluster of metabolic abnormalities that are closely associated with an increased risk of chronic diseases such as cardiovascular disease and type 2 diabetes. It consists of a series of interrelated metabolic disturbances, including abdominal obesity, hyperglycemia, dyslipidemia, and hypertension. In this study, the diagnosis of MS was based on the modified criteria of the National Cholesterol Education Program Adult Treatment Panel III (NCEP: ATP III). MS is diagnosed if three or more of the following five conditions are met: (1) Abdominal Obesity: Waist circumference ≥ 40 inches (102 cm) in men; ≥ 35 inches (88 cm) in women. (2) Triglycerides: \geq 150 mg/dL (1.7 mmol/L). (3) Low HDL Cholesterol (HDL-C): < 40 mg/dL (1.0 mmol/L) in men; < 50 mg/dL (1.3 mmol/L) in women. (4) Elevated Blood Pressure: Systolic blood pressure \geq 130 mmHg or diastolic blood pressure \geq 85 mmHg, or currently receiving treatment for hypertension. (5) Elevated Fasting Glucose: ≥ 100 mg/dL (5.6 mmol/L), or currently receiving treatment for diabetes. The final outcome of this study was the presence or absence of MS.

2.4 Selection and processing of covariates

In addition to eGDR levels, this study incorporated several demographic variables, including age, sex, race, marital status, education level, poverty status, physical activity, body mass index (BMI), and alcohol and tobacco use. Race was categorized into four groups: White, Mexican American, Black individuals, and Other. Marital status was classified as married, divorced, widowed, separated, never married, or living with a partner. Education level was categorized as high school or below, high school or equivalent, and above high school. Past medical history data included cancer, comorbidities, hypertension, and diabetes, all of which were treated as dichotomous variables. Additionally, laboratory variables such as triglycerides (TG, mmol/L), total cholesterol (TC, mmol/L), high-density lipoprotein cholesterol (HDL-C, mmol/L), and low-density lipoprotein cholesterol (LDL-C, mmol/L) were also collected. These covariates will be adjusted for in the statistical analysis.

2.5 Statistical methods

Data analysis was conducted using complex sampling weights recommended by the Centers for Disease Control and Prevention. Following the methodology outlined on the NHANES website, we combined the sample weights from the 2005–2006, 2007–2008, and 2009–2010 cycles. eGDR levels were categorized into tertiles based on prior literature. Continuous variables with a normal distribution were expressed as means (SD), while skewed continuous variables were reported as medians (IQR) and analyzed using the T-test or rank-sum test. Categorical variables were presented as proportions and compared between groups using the χ^2 test.

To examine the relationship between eGDR levels and the prevalence of MS, we performed both univariate and multivariate logistic regression analyses. In Model 1, only the relationship between eGDR tertiles and MS prevalence was examined. Model 2 adjusted for age, sex, and race/ethnicity, and Model 3 further adjusted for all covariates. To explore the nonlinear relationship between eGDR levels and MS, we applied the restricted cubic spline (RCS) method. All statistical analyses and visualizations were conducted using R software (https://www.r-project.org/, Version 4.3.1), with a significance level set at p < 0.05.

3 Results

3.1 Baseline demographic and clinical characterization

Our study initially extracted data from the NHANES database, comprising 63,131 individuals with metabolic syndrome information

and 81,664 individuals with eGDR data. Following data integration, a combined cohort of 84,104 participants was established. After excluding 32,769 individuals with missing values, the remaining sample consisted of 51,335 participants. The covariates exhibited varying sample sizes, ranging from 81,227 to 95,872 across different parameters. Subsequent integration of covariate information yielded a comprehensive dataset of 95,872 individuals. Following the exclusion of 70,748 participants with any missing values, our final analytical cohort comprised 25,124 individuals. Quality control measures confirmed the absence of outliers in the final dataset. Weighted to represent 138,013,430 participants. The screening flowchart is shown in Figure 1.

Based on eGDR levels, all samples were categorized into three groups by tertiles: Q1 (eGDR range: -4.232 to 6.42), Q2 (6.42 to 9.558), and Q3 (9.558 to 13.106). Table 1 demonstrates the between-group differences in baseline demographic characteristics and eGDR level subgroups. The study population was 52% male and 48% female, with a mean age of 45.7 years. Higher proportions were White individuals, married, had a high school education or higher, had never

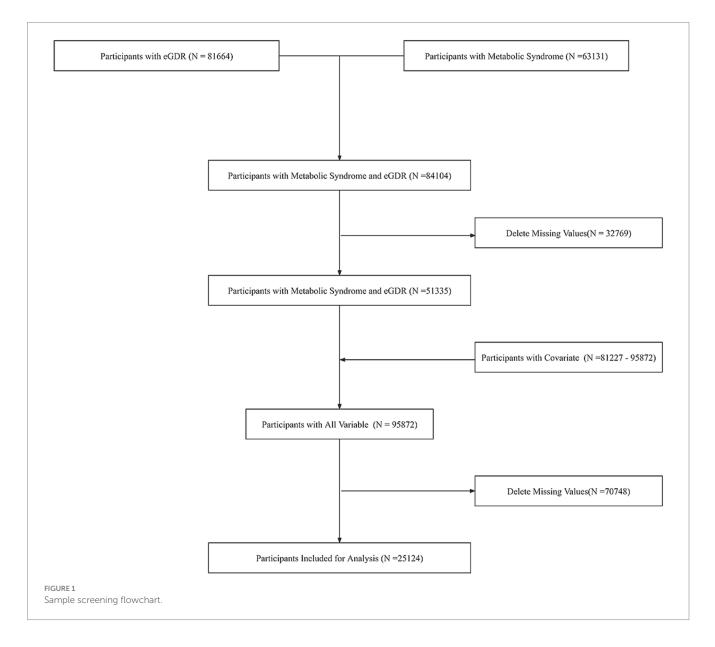


TABLE 1 Differential analysis of baseline characteristics in weighted eGDR quartiles.

Characters	Total	Q1	Q2	Q3	P Value	
Sex, %					< 0.001	
Male	51.84	59.13	54.58	43.47		
Female	48.16	40.87	45.42	56.53		
Age, mean (SD)	45.74 (0.24)	54.40 (0.28)	46.54 (0.28)	38.11 (0.28)	< 0.001	
BMI, mean (SD)	28.72 (0.08)	33.39 (0.12)	29.92 (0.08)	23.88 (0.05)	< 0.001	
ce/Ethnicity, %						
White	70.62	71.18	70.49	70.29		
Mexican	7.58	6.42	9.12	7.06		
Black individuals	9.63	12.42	8.77	8.22		
Other	12.17	9.98	11.62	14.44		
Marital status, %		<u> </u>	<u> </u>		< 0.001	
Married	57.16	62.11	59.59	50.95		
Living with partner	7.45	4.89	7.91	9.06		
Separated	1.92	2.12	1.89	1.79		
Divorced	9.00	11.28	9.07	7.12		
Widowed	5.45	8.96	5.52	2.61		
Never married	19.02	10.65	16.01	28.47		
Education, %					< 0.001	
Under high school	11.82	13.52	12.12	10.19		
High school or	22.67	25.79	24.11	18.85		
equivalent	65.51	60.69	63.77	70.97		
Above high school					0.46	
Poverty, median (IQR)	3.34 (1.69,5.00)	3.23 (1.65,5.00)	3.29 (1.74,5.00)	3.45 (1.69,5.00)	0.46	
Physical Activity, median (IQR)	1920.00 (720.00,5300.00)	1680.00 (567.00,4800.00)	1920.00 (720.00,5760.00)	2280.00 (840.00,5760.00)	< 0.001	
Smoking, %	I				< 0.001	
Never	55.62	50.60	54.42	60.72		
Former	24.79	33.28	24.28	18.52		
Now	19.59	16.12	21.30	20.76		
Alcohol intake, %	1	1	1		< 0.001	
Never	8.93	9.42	8.50	8.94		
Former	10.06	13.56	10.70	6.69		
Mild	38.85	41.22	39.09	36.75		
Moderate	18.97	16.33	18.06	21.92		
Heavy	23.19	19.47	23.66	25.70		
Diabetes, %					< 0.001	
No	91.59	78.11	94.77	99.35		
Yes	8.41	21.89	5.23	0.65		
					< 0.001	
Hypertension, %						
Hypertension, %	65.80	7.68	78.67	100.00		
	65.80 34.20	7.68 92.32	78.67 21.33	0.00		
No Yes					< 0.001	
No Yes					< 0.001	
No Yes Cancer, %	34.20	92.32	21.33	0.00	< 0.001	
Yes Cancer, % No Yes	34.20 90.78	92.32 86.27	21.33 91.13	0.00 94.04	< 0.001	
No Yes Cancer, % No	34.20 90.78	92.32 86.27	21.33 91.13	0.00 94.04		

Exposure	Model 1		Model 2		Model 3				
	OR (95%CI)	P value	OR (95%CI)	P value	OR (95%CI)	P value			
eGDR (mg/kg/min)	0.56 (0.55, 0.57)	< 0.001	0.56 (0.55, 0.58)	<0.001	0.48 (0.44, 0.52)	<0.001			
Quartile of eGDR (mg/kg/min)									
Q1	Ref	Ref	Ref	Ref	Ref	Ref			
Q2	0.18 (0.17, 0.20)	<0.001	0.19 (0.17, 0.20)	<0.001	0.35 (0.30, 0.42)	<0.001			
Q3	0.02 (0.01, 0.02)	<0.001	0.02 (0.01, 0.02)	<0.001	0.07 (0.05, 0.09)	<0.001			
P for trend		< 0.001		<0.001		<0.001			

TABLE 2 Weighted ORs for the associations between eGDR and MS across three models.

Model 1: Add eDGR. Model 2: Add eDGR, Age, Sex, and Race/Ethnicity. Model 3: All the covariates. eGDR: Estimated glucose disposal rate; MS: Metabolic syndrome.

smoked, and were light drinkers. The majority of study participants did not have diabetes or cancer, 34% had hypertension, and 21.26% had metabolic syndrome.

In the eGDR tertile subgroups, concerning gender, the proportion of males was higher in the Q1 and Q2 groups, while the proportion of females was higher in the Q3 group. Age and BMI tended to decrease as the eGDR quartile increased. In terms of race, the Q3 group had a higher percentage of other races, while the Q1 group had a higher percentage of Black individuals. In terms of marital status, the married group had the largest percentage in the Q1 group, while those who had never been married had the highest percentage in the Q3 group. The group with higher educational level has the largest percentage in group Q3. Individuals with higher poverty index and more physical activity tended to be located in the higher eGDR quartile groups. Smoking prevalence declines with increasing eGDR quartiles, while the proportion of moderate and higher level drinkers increases with higher eGDR. The proportion of people with diabetes, hypertension and cancer decreased with higher eGDR quartiles. The prevalence of metabolic syndrome followed a similar trend, with a higher proportion in the Q3 group not suffering from MS.

3.2 Relationship between eGDR level and MS prevalence

The logistic regression analysis results showed that eGDR was negatively associated with MS prevalence in Model 1, Model 2 and Model 3. Both in the model containing only eGDR and in the logistic regression model with different covariates, eGDR was found to be a protective factor for MS prevalence. According to the results of eGDR tertile analysis, in model 1, with Q1 as the control group, both Q2 and Q3 groups showed a protective effect. After covariate correction, the results in models 2 and 3 were consistent with model 1. The *p*-values of the trend tests were all less than 0.01, indicating that there was indeed a linear relationship between eGDR and MS prevalence after using eGDR as an ordered categorical variable (Table 2).

3.3 Subgroup analysis

Table 3 presents the results of the analysis of eGDR levels and their tertiles across different subgroups. The findings revealed a significant negative correlation between eGDR levels and MS prevalence in all covariate subgroups. Specifically, higher eGDR levels were associated

with a lower prevalence of MS, indicating a protective effect. The odds ratios (ORs) were all less than 0.6 in every subgroup, except in the subgroup with higher BMI, suggesting that the protective effect of eGDR levels was significant in most subgroups. In the subgroup analyses by eGDR tertiles, using the Q1 group as a reference, the ORs for the Q2 group were all below 0.5 (p < 0.001), while the ORs for the Q3 group were all below 0.2 (p < 0.001). The higher the eGDR level, the more pronounced the protective effect, which was consistent with the observed negative correlation between eGDR levels and MS prevalence.

3.4 RCS of eDGR and MS prevalence

The RCS analysis revealed a significant nonlinear association between eGDR levels and the log odds ratio of MS events (Figure 2A), which remained consistent across genders (Figure 2B). Specifically, the log probability of an MS event was significantly reduced with increasing eGDR levels, indicating that an increase in eGDR levels is effective in reducing the risk of an MS event. However, this relationship is nonlinear. At low eGDR levels (eGDR<8), the log odds ratio decreases more slowly, suggesting that low eGDR levels have a limited effect on reducing the risk of MS events. As eGDR levels increased further (\geq 8), the rate of decrease in the log odds ratio increased, indicating that the effect of eGDR on reducing the risk of MS events became more pronounced and stronger at higher eGDR levels. This non-linear change suggests that eGDR may have a stronger protective effect after some cut points.

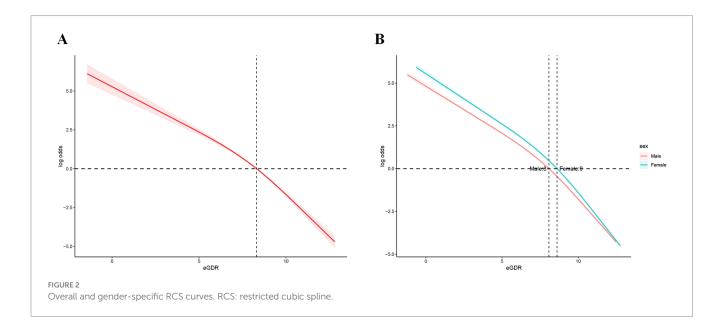
In addition, we further verified the stability of the curve fit over the entire range of eGDR levels by confidence interval analysis. This means that the non-linear relationship between eGDR and risk of MS events is reliable and consistent across different eGDR levels. These results further highlight the potential role of eGDR as an important indicator in MS risk assessment, especially at high eGDR levels, which may have greater clinical significance.

4 Discussion

In this epidemiologic study, the US population was carefully screened to ensure it was representative of the study cohort, and a significant association was found between higher eGDR levels and a reduced prevalence of MS in a population without MS. This association remained significant even after adjusting for known risk factors for MS development. Furthermore, the negative correlation

Character	eGDR (mg/kg/min)		Quartile of eGDR (mg/kg/min)					
	OR (95%CI)	P value	Q1 Q2		2	Q3		
			OR (95%CI)	OR (95%CI)	P value	OR (95%CI)	P value	
Sex								
Male	0.50 (0.44, 0.57)	< 0.001	Ref	0.45 (0.34, 0.59)	<0.001	0.06 (0.03, 0.10)	< 0.001	
Female	0.46 (0.42, 0.52)	< 0.001	Ref	0.28 (0.22, 0.36)	< 0.001	0.06 (0.04, 0.09)	< 0.001	
Age								
= < 60	0.46 (0.42, 0.50)	< 0.001	Ref	0.35 (0.28, 0.43)	<0.001	0.05 (0.03, 0.07)	< 0.001	
>60	0.44 (0.37, 0.52)	< 0.001	Ref	0.29 (0.22, 0.40)	< 0.001	0.11 (0.06, 0.20)	< 0.001	
BMI								
Normal	0.33 (0.28, 0.39)	< 0.001	Ref	0.28 (0.20, 0.40)	<0.001	0.06 (0.03, 0.09)	< 0.001	
Low	0.27 (0.21, 0.35)	< 0.001	Ref	0.21 (0.13, 0.34)	<0.001	0.02 (0.01, 0.06)	< 0.001	
High	0.74 (0.70, 0.78)	< 0.001	Ref	0.43 (0.32, 0.58)	<0.001	0.18 (0.10, 0.31)	< 0.001	
Smoke								
Never	0.49 (0.44, 0.55)	< 0.001	Ref	0.40 (0.30, 0.52)	< 0.001	0.07 (0.04, 0.11)	< 0.001	
Former	0.48 (0.41, 0.55)	< 0.001	Ref	0.31 (0.22, 0.42)	<0.001	0.05 (0.02, 0.10)	< 0.001	
Now	0.43 (0.37, 0.51)	< 0.001	Ref	0.28 (0.18, 0.43)	<0.001	0.06 (0.03, 0.13)	< 0.001	
Alcohol								
Never	0.52 (0.44, 0.62)	< 0.001	Ref	0.41 (0.28, 0.61)	<0.001	0.11 (0.05, 0.21)	< 0.001	
Former	0.50 (0.40, 0.62)	<0.001	Ref	0.29 (0.17, 0.49)	<0.001	0.07 (0.03, 0.16)	< 0.001	
Mild	0.47 (0.42, 0.53)	< 0.001	Ref	0.37 (0.27, 0.52)	< 0.001	0.05 (0.03, 0.10)	< 0.001	
Moderate	0.36 (0.30, 0.43)	<0.001	Ref	0.33 (0.20, 0.55)	<0.001	0.05 (0.02, 0.11)	< 0.001	
Heavy	0.55 (0.48, 0.64)	< 0.001	Ref	0.34 (0.24, 0.46)	< 0.001	0.08 (0.05, 0.13)	< 0.001	

TABLE 3 Stratified logistic regression analysis.



between eGDR levels and MS risk was consistent across stratified analyses of risk factors and socio-behavioral variables.

The metabolic syndrome contributes to the spread of diseases such as type 2 diabetes, coronary heart disease, stroke, and other disabilities (17, 18). A major driver of this phenomenon is the high degree of overlap between the core pathological features of the metabolic syndrome (e.g., insulin resistance, chronic low-grade inflammation, and abnormalities of lipid metabolism) and the pathogenesis of the aforementioned disorders (19, 20). This further increases the risk of coronary heart disease and stroke (21). In addition, chronic low-grade inflammation, an important feature of metabolic syndrome, leads to elevation of pro-inflammatory cytokines (e.g., TNF- α , IL-6), which induces systemic inflammatory responses and thus plays a key role in the course of many diseases (22, 23). Therefore, the level of eGDR levels may not only serve as a reliable marker of insulin resistance but may also indirectly reflect the chronic low-grade inflammation and abnormal lipid metabolism changes, thus providing a new perspective for early screening of metabolic syndrome and its related complications.

Despite the widespread health risks associated with MS, current diagnostic and management strategies remain limited. The internationally accepted diagnostic criteria for MS are mainly based on the definitions and recommendations of relevant international organizations and authoritative guidelines. For instance, the International Diabetes Federation uses a waist circumference-centered metabolic standard, combined with other metabolic risk factors (24). The National Cholesterol Education Program Adult Treatment Panel III incorporates a comprehensive set of indicators, including waist circumference, blood lipids, blood glucose, and blood pressure (1). Additionally, the World Health Organization emphasizes insulin resistance as a core factor, alongside indicators of blood glucose, blood pressure, and dyslipidemia (25). While these guidelines provide useful frameworks for identifying metabolic syndrome, they still fall short in disease prediction and individualized risk assessment. Existing criteria lack comprehensive and unified metrics for diagnosing MS. eGDR levels, as an insulin resistance-based biomarker, show promise in integrating the core pathological features of metabolic syndrome. Our study found that eGDR levels are not only significantly correlated with the prevalence of MS, but their stability in stratified analyses also suggests their potential application across different subgroups. Given that metabolic syndrome is a complex medical diagnosis involving multiple biomarkers, simple anthropometric definitions (such as waist circumference and blood pressure) have certain limitations for clinical application. In contrast, eGDR levels provide a more straightforward and acceptable alternative, offering an effective screening tool for large populations, particularly in resource-limited settings and high-risk groups (17).

As a marker of insulin resistance, eGDR levels exhibit accuracy comparable to the normoglycemic hyperinsulinemic clamp, making them suitable for clinical practice (26, 27). The normoglycemic hyperinsulinemic clamp, considered the gold standard for identifying and quantifying insulin resistance, is labor-intensive, time-consuming, and invasive, rendering it impractical for routine clinical use (28). Alternative methods, such as the homeostatic model assessment of insulin resistance, which relies on fasting insulin and glucose levels, have been proposed (29). However, simpler and more reliable alternatives are needed. Consequently, eGDR levels have been widely utilized in studies predicting vascular disease mortality (15, 30, 31), adverse nephropathic outcomes (22), adverse nephropathic outcomes (32), demonstrating excellent predictive efficacy. Similarly, eGDR levels have shown high sensitivity in predicting MS-related conditions, such as chronic diabetes mellitus (33) and the onset of type 1 diabetes (34). Notably, eGDR levels have demonstrated superior diagnostic performance for identifying metabolic syndrome in patients with type 1 diabetes, including children and adolescents (13, 35, 36), a finding consistent with the present study. Our study found that eGDR levels exhibited a protective effect across different racial groups, suggesting the potential for incorporating eGDR levels into routine diabetes care (33). However, it has also been shown that, in a population of type 1 diabetic patients from an urban clinic, lower eGDR levels in the Black individuals population were associated with diabetic complications (12). The inconsistency in these findings may be due to differences in the study samples. For example, genetic predispositions, lifestyle factors, or socioeconomic status in Black individuals populations may influence the relationship between eGDR levels and diabetic complications (37). Given the significant association between eGDR and MS risk, low eGDR levels could serve as an early indicator for identifying high-risk individuals, particularly those with existing metabolic risk factors. This can guide personalized interventions, such as lifestyle modifications and, where necessary, pharmacological treatment. Regular monitoring of eGDR levels could also facilitate the assessment of intervention efficacy and enable timely adjustments to treatment plans. However, despite eGDR's significant potential in MS prediction, further research is needed to validate its standardization and application across diverse populations and clinical settings.

Although this study clarified the stable association between eGDR levels and the prevalence of MS, several potential limitations should be acknowledged. First, this was a cross-sectional study, which precludes the establishment of a causal relationship between eGDR levels and MS prevalence, limiting our ability to make causal inferences. Second, NHANES data are primarily based on U.S. populations, which may limit their applicability to other countries or regions. Third, NHANES data reflect health status at a single point in time, which prevents tracking of health changes or the effects of interventions over time. This represents a limitation in assessing the long-term progression of metabolic syndrome and fluctuations in eGDR levels. Fourth, a more detailed exploration of subgroup analysis is needed. For example, the weaker protective effect of eGDR in individuals with high BMI compared to other BMI groups warrants further investigation. Finally, NHANES did not consistently record whether participants received specific health interventions or treatments, such as glucose-lowering medications or insulin therapy, which may influence the relationship between eGDR levels and MS. In conclusion, while our findings provide valuable insights, they should be interpreted with caution. Further studies are needed to confirm the direction and strength of the relationship between eGDR levels and MS prevalence.

5 Conclusion

In conclusion, our findings suggest a significant association between higher eGDR levels and a lower prevalence of MS. However, further large-scale prospective studies are needed to validate these findings. Nonetheless, eGDR levels demonstrate important clinical utility as a valid predictor of MS prevalence and a potential therapeutic target.

Data availability statement

The raw data supporting the conclusions of this article will be made available by the authors, without undue reservation.

Ethics statement

The studies involving humans were approved by the NHANES database has officially been certified for ethics. 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

KX: Conceptualization, Data curation, Formal analysis, Writing – original draft. SJ: Methodology, Resources, Software, Visualization, Writing – original draft. GC: Software, Visualization, Writing – original draft. HZ: Investigation, Validation, Writing – original draft. QZ: Funding acquisition, Project administration, Validation, Writing – review & editing.

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References

1. Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults. Executive summary of the third report of the National Cholesterol Education Program (NCEP) expert panel on detection, evaluation, and treatment of high blood cholesterol in adults (adult treatment panel III). *JAMA*. (2001) 285:2486–97. doi: 10.1001/jama.285.19.2486

2. Alberti KGMM, Eckel RH, Grundy SM, Zimmet PZ, Cleeman JI, Donato KA, et al. Harmonizing the metabolic syndrome: a joint interim statement of the international diabetes federation task force on epidemiology and prevention; National Heart, Lung, and Blood Institute; American Heart Association; world heart federation; international atherosclerosis society; and International Association for the Study of obesity. *Circulation.* (2009) 120:1640–5. doi: 10.1161/CIRCULATIONAHA.109. 192644

3. Ranasinghe P, Mathangasinghe Y, Jayawardena R, Hills AP, Misra A. Prevalence and trends of metabolic syndrome among adults in the asia-pacific region: a systematic review. *BMC Public Health*. (2017) 17:101. doi: 10.1186/s12889-017-4041-1

4. Do Vale Moreira NC, Hussain A, Bhowmik B, Mdala I, Siddiquee T, Fernandes VO, et al. Prevalence of metabolic syndrome by different definitions, and its association with type 2 diabetes, pre-diabetes, and cardiovascular disease risk in Brazil. *Diabetes Metab Syndr.* (2020) 14:1217–24. doi: 10.1016/j.dsx.2020.05.043

5. National Center for Health Statistics and others. Crude and age-adjusted percentage of civilian, noninstitutionalized adults with diagnosed diabetes, United States, 1980–2010. (2012). Available at: https://scholar.google.com/scholar?hl=en&as_sdt=0% 2C5&q=Crude+and+ageadjusted+percentage+of+civilian%2C+noninstitutionalized+ adults+with+diagnosed+diabetes&btnG=

6. Li Y, Zhao L, Yu D, Wang Z, Ding G. Metabolic syndrome prevalence and its risk factors among adults in China: a nationally representative cross-sectional study. *PLoS One.* (2018) 13:e0199293. doi: 10.1371/journal.pone.0199293

7. Eckel RH, Grundy SM, Zimmet PZ. The metabolic syndrome. Lancet. (2005) 365:1415-28. doi: 10.1016/S0140-6736(05)66378-7

 Esposito K, Chiodini P, Colao A, Lenzi A, Giugliano D. Metabolic syndrome and risk of cancer: a systematic review and meta-analysis. *Diabetes Care*. (2012) 35:2402–11. doi: 10.2337/dc12-0336

9. Tahapary DL, Pratisthita LB, Fitri NA, Marcella C, Wafa S, Kurniawan F, et al. Challenges in the diagnosis of insulin resistance: focusing on the role of HOMA-IR and Tryglyceride/glucose index. *Diabetes Metab Syndr.* (2022) 16:102581. doi: 10.1016/j.dsx.2022.102581

10. Wei X, Min Y, Song G, Ye X, Liu L. Association between triglyceride-glucose related indices with the all-cause and cause-specific mortality among the population with metabolic syndrome. *Cardiovasc Diabetol.* (2024) 23:134. doi: 10.1186/s12933-024-02215-0

11. Williams KV, Erbey JR, Becker D, Arslanian S, Orchard TJ. Can clinical factors estimate insulin resistance in type 1 diabetes? *Diabetes*. (2000) 49:626–32. doi: 10.2337/diabetes.49.4.626

12. Epstein EJ, Osman JL, Cohen HW, Rajpathak SN, Lewis O, Crandall JP. Use of the estimated glucose disposal rate as a measure of insulin resistance in an urban multiethnic population with type 1 diabetes. *Diabetes Care*. (2013) 36:2280–5. doi: 10.2337/dc12-1693

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13. Köken ÖY, Kara C, Yılmaz GC, Aydın HM. Utility of estimated glucose disposal rate for predicting metabolic syndrome in children and adolescents with type-1 diabetes. *J Pediatr Endocrinol Metab.* (2020) 33:859–64. doi: 10.1515/jpem-2020-0012

14. Kong X, Wang W. Estimated glucose disposal rate and risk of cardiovascular disease and mortality in U.S. adults with prediabetes: a nationwide cross-sectional and prospective cohort study. *Acta Diabetol.* (2024) 61:1413–21. doi: 10.1007/s00592-024-02305-1

15. Guo R, Tong J, Cao Y, Zhao W. Association between estimated glucose disposal rate and cardiovascular mortality across the spectrum of glucose tolerance in the US population. *Diabetes Obes Metab.* (2024) 26:5827–35. doi: 10.1111/dom.15954

16. Center for Health Statistics. About the National Health and nutrition examination survey. (2023) Available online at: https://www.cdc.gov/nchs/nhanes/about_nhanes.htm

17. Saklayen MG. The global epidemic of the metabolic syndrome. *Curr Hypertens Rep.* (2018) 20:12. doi: 10.1007/s11906-018-0812-z

18. Grundy SM, Brewer HB, Cleeman JI, Smith SC, Lenfant C. American Heart Association, National Heart, Lung, and Blood Institute. Definition of metabolic syndrome: report of the National Heart, Lung, and Blood Institute/American Heart Association conference on scientific issues related to definition. *Circulation*. (2004) 109:433–8. doi: 10.1161/01.CIR.0000111245.75752.C6

19. Kahn SE, Hull RL, Utzschneider KM. Mechanisms linking obesity to insulin resistance and type 2 diabetes. *Nature*. (2006) 444:840–6. doi: 10.1038/nature05482

20. DeFronzo RA, Ferrannini E. Insulin resistance. A multifaceted syndrome responsible for NIDDM, obesity, hypertension, dyslipidemia, and atherosclerotic cardiovascular disease. *Diabetes Care.* (1991) 14:173–94. doi: 10.2337/diacare.14.3.173

21. Zheng Y, Ley SH, Hu FB. Global aetiology and epidemiology of type 2 diabetes mellitus and its complications. *Nat Rev Endocrinol.* (2018) 14:88–98. doi: 10.1038/nrendo.2017.151

22. Peng J, Zhang Y, Zhu Y, Chen W, Chen L, Ma F, et al. Estimated glucose disposal rate for predicting cardiovascular events and mortality in patients with non-diabetic chronic kidney disease: a prospective cohort study. *BMC Med.* (2024) 22:411. doi: 10.1186/s12916-024-03582-x

23. Duncan BB, Schmidt MI, Pankow JS, Ballantyne CM, Couper D, Vigo A, et al. Atherosclerosis risk in communities study. Low-grade systemic inflammation and the development of type 2 diabetes: the atherosclerosis risk in communities study. *Diabetes*. (2003) 52:1799–805. doi: 10.2337/diabetes.52.7.1799

24. International Diabetes Federation (IDF). The IDF consensus worldwide definition of the metabolic syndrome. (2006) Available online at: https://idf.org/media/uploads/2023/05/attachments-31.pdf

25. Alberti KG, Zimmet PZ. Definition, diagnosis and classification of diabetes mellitus and its complications. Part 1: diagnosis and classification of diabetes mellitus provisional report of a WHO consultation. *Diabet Med.* (1998) 15:539–53. doi: 10.1002/(SICI)1096-9136(199807)15:7<539::AID-DIA668>3.0.CO;2-S

26. Shi W, Qin M, Wu S, Xu K, Zheng Q, Liu X. Value of estimated glucose disposal rate to detect prevalent left ventricular hypertrophy: implications from a general population. *Postgrad Med.* (2023) 135:58–66. doi: 10.1080/00325481.2022.2131153

27. Komosinska-Vassev K, Gala O, Olczyk K, Jura-Półtorak A, Olczyk P. The usefulness of diagnostic panels based on circulating Adipocytokines/regulatory peptides, renal function tests, insulin resistance indicators and lipid-carbohydrate metabolism parameters in diagnosis and prognosis of type 2 diabetes mellitus with obesity. *Biomol Ther.* (2020) 10:1304. doi: 10.3390/biom10091304

28. Wang S, Shi J, Peng Y, Fang Q, Mu Q, Gu W, et al. Stronger association of triglyceride glucose index than the HOMA-IR with arterial stiffness in patients with type 2 diabetes: a real-world single-Centre study. *Cardiovasc Diabetol.* (2021) 20:82. doi: 10.1186/s12933-021-01274-x

29. Zheng T, Ge B, Liu H, Chen B, Qin L, Xiao L, et al. Triglyceride-mediated influence of serum angiopoietin-like protein 8 on subclinical atherosclerosis in type 2 diabetic patients: results from the GDMD study in China. *Cardiovasc Diabetol.* (2018) 17:84. doi: 10.1186/s12933-018-0687-y

30. Lu Z, Xiong Y, Feng X, Yang K, Gu H, Zhao X, et al. Insulin resistance estimated by estimated glucose disposal rate predicts outcomes in acute ischemic stroke patients. *Cardiovasc Diabetol.* (2023) 22:225. doi: 10.1186/s12933-023-01925-1

31. Zheng X, Han W, Li Y, Jiang M, Ren X, Yang P, et al. Changes in the estimated glucose disposal rate and incident cardiovascular disease: two large prospective cohorts in Europe and Asia. *Cardiovasc Diabetol.* (2024) 23:403. doi: 10.1186/s12933-024-02485-8

32. Chen Y, Lin H, Xu J, Zhou X. Estimated glucose disposal rate is correlated with increased depression: a population-based study. *BMC Psychiatry*. (2024) 24:786. doi: 10.1186/s12888-024-06257-2

33. Lam-Chung CE, Martínez Zavala N, Ibarra-Salce R, Pozos Varela FJ, Mena Ureta TS, Berumen Hermosillo F, et al. Association of estimated glucose disposal rate and chronic diabetic complications in patients with type 1 diabetes. *Endocrinol Diabetes Metab.* (2021) 4:e00288. doi: 10.1002/edm2.288

34. Chillarón JJ, Goday A, Flores-Le-Roux JA, Benaiges D, Carrera MJ, Puig J, et al. Estimated glucose disposal rate in assessment of the metabolic syndrome and microvascular complications in patients with type 1 diabetes. *J Clin Endocrinol Metab.* (2009) 94:3530–4. doi: 10.1210/jc.2009-0960

35. Mao Y, Zhong W. Changes of insulin resistance status and development of complications in type 1 diabetes mellitus: analysis of DCCT/EDIC study. *Diabetes Res Clin Pract.* (2022) 184:109211. doi: 10.1016/j.diabres.2022.109211

36. Grabia M, Socha K, Bossowski A, Markiewicz-Żukowska R. Metabolic syndrome as a factor of impairment of antioxidant defense system in youth with T1DM. *Int J Mol Sci.* (2023) 24:9428. doi: 10.3390/ijms24119428

37. Tatulashvili S, Fagherazzi G, Dow C, Cohen R, Fosse S, Bihan H. Socioeconomic inequalities and type 2 diabetes complications: a systematic review. *Diabetes Metab.* (2020) 46:89–99. doi: 10.1016/j.diabet.2019.11.001