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#### ORIGINAL RESEARCH

# Hypertriglyceridemic waist phenotype in primary health care: comparison of two cutoff points

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Objective: We aimed to evaluate the prevalence of hypertriglyceridemic waist (HTGW) phenotype among users of primary health care using two different cutoff points used in the literature. Methods: We evaluated adults and elderly individuals of both sexes who attended the same level of primary health care. HTGW phenotype was determined with measurements of waist circumference (WC) and triglyceride levels and compared using cutoff points proposed by the National Cholesterol Education Program - NCEP/ATP III (WC ≥102 cm for men and ≥88 cm for women; triglyceride levels  $\geq$ 150 mg/dL for both sexes) and by Lemieux et al (WC  $\geq$ 90 cm for men and  $\geq$ 85 cm for women; triglyceride levels  $\geq$ 177 mg/dL for both).

Results: Within the sample of 437 individuals, 73.7% was female. The prevalence of HTGW phenotype was high and statistically different with the use of different cutoff points from the literature. The prevalence was higher using the NCEP/ATP III criteria compared to those proposed by Lemieux et al (36.2% and 32.5%, respectively, p<0.05). Individuals with the presence of the phenotype also presented alterations in other traditional cardiovascular risk markers.

Conclusion: The HTGW phenotype identified high prevalence of cardiovascular risk in the population, with higher cutoff points from the NCEP/ATP III criteria. The difference in frequency of risk alerts us to the need to establish cutoff points for the Brazilian population.

Keywords: abdominal obesity, cardiovascular disease, dyslipidemia, cardiovascular risk

## Introduction

According to the World Health Organization (WHO), the prevalence of cardiovascular disease (CVD) is increasing in the world, and it is the leading cause of death and disability over the past years.<sup>1</sup> In this context, it is important to have criteria for the screening of CVD and cardiovascular events for different populations.<sup>2</sup>

In order to make the cardiovascular risk assessment a simple and accessible task at all levels of health care,<sup>2</sup> Lemieux et al<sup>3</sup> proposed a cardiometabolic marker called hypertriglyceridemic waist (HTGW) phenotype. The HTGW phenotype is characterized by the simultaneous presence of increased waist circumference (WC) and high levels of triglycerides, and it is a better predictor for individuals with cardiovascular risk when compared to the components in isolation.<sup>4</sup> Therefore, the authors propose that the HTGW phenotype may be used as a first marker for tracing and identifying patients with risk of developing metabolic syndrome (MS).<sup>5</sup> This association is justified by the presence of hyperinsulinemia, high levels of apolipoprotein B and high levels of low-density lipoprotein-cholesterol (LDL-C), due to the levels of visceral adipose tissue.6

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There are two cutoff points available in the literature to identify the presence of HTGW phenotype. The Canadian pioneer group suggested a cutoff point for men<sup>3</sup> and, later, a different cutoff point for women, with the only difference being the WC.<sup>5</sup> The cutoff points of WC and triglyceride levels from this group are different from those proposed by the National Cholesterol Education Program (NCEP-ATP III).<sup>7</sup> Both cutoff points have been used in studies evaluating the HTGW phenotype. Thus, different prevalence rates can be found in the literature and, sometimes, within the same population, generating confusion among various authors.

In Brazil, there are few studies available about the prevalence of the HTGW phenotype, and no study has proposed a specific cutoff point. Recognizing the lack of studies, the objective of this study was to evaluate the prevalence of the HTGW phenotype in a population of the state of Rio Grande do Norte, Brazil, as well as to compare the results obtained using the two cutoff points available in the literature.

#### Methods

An observational study with a cross-sectional design was conducted including individuals of both sexes older than 20 years of age, living in the state of Rio Grande do Norte (Brazil) and who attended a primary health care unit. The participants of the study were selected for convenience, and they were approached at the health service unit which they attended. Only those patients who had biochemical test results, including fasting triglyceride levels, within <30 days of the date of assessment were included. The following subjects were excluded: pregnant women, adults and elderly individuals with hepatic diseases, patients with AIDS or malnutrition, or patients who were undergoing renal replacement therapy. All patients signed the form of informed consent in duplicate, and the research project was approved by the Research Ethics Committee of the Federal University of Rio Grande do Norte (protocol 284.437, CAAE 13148313.1.0000.5568).

For the sample size calculation, the HTGW prevalence rate of 10.9% among the inhabitants of Minas Gerais was considered.<sup>2</sup> Using an absolute precision of 3% and a significance level of 5%, it would be necessary to evaluate at least 384 individuals.

Data collection was performed at the primary health care units by properly trained nutrition professionals and academics, in a room reserved for the purpose. First, the patients answered a structured questionnaire with information about their housing, socioeconomic condition and lifestyle. Economic class was evaluated using the Brazilian Association of Research Companies (ABEP) methodology,<sup>8</sup> which takes into account the number of material goods and monthly-paid employees in the household, the educational level of the main breadwinner, and clustering within the economic classes A1/ A2, B1/B2, C1/C2, D and E, which were later on regrouped as A/B [better condition], C [intermediary condition] and D/E [worse condition]). After this interview, the anthropometric nutritional evaluation was performed to characterize the nutritional status and to obtain the WC measurement.

The measurement of body mass was obtained using a properly calibrated mechanical professional balance (Welmy, Sao Paulo, Brazil), supporting up to 150 kg; the individual was asked to remove shoes and heavy clothing and to remain with minimal clothes. To measure the height, the individual remained barefoot in the center of the equipment, erect, with feet together and arms extended along the body, using the measurement technique according to the Frankfurt plan.<sup>9</sup> To determine the nutritional status, the BMI was calculated, and the nutritional status was ranked according to the cutoff points recommended by WHO.<sup>10</sup> The nutritional status of people aged >60 years was classified according to the cutoff points of Lipschitz.<sup>11</sup>

The WC measurement was performed using inelastic tape at the midpoint between the lower costal margin and the iliac crest. The presence of cardiovascular risk was determined by considering a WC of 102 cm for men and 88 cm for women.<sup>7</sup> The hip circumference (HC) was measured using a measuring tape positioned at the point of greatest circumference of the gluteal region. The waist-to-hip ratio (WHR) was calculated by dividing the WC by the HC, and the cutoff points used were those established by the WHO,<sup>12</sup> where the values of WHR >1 and WHR >0.85 represent risk of developing CVD in men and women, respectively.

The waist-to-height ratio was calculated by dividing the WC (in centimeters) by the height (in centimeters); the cutoff point adopted for abdominal obesity was 0.53.<sup>13</sup> The sagittal abdominal diameter (SAD) was measured with the mobile rod of an anthropometer having subdivisions of 0.1 cm. During the evaluation, the volunteer remained lying on a firm-surface examination table in the supine position with knees flexed. The measurement was taken at the umbilical level.<sup>14</sup> For females and males, the cutoff points were 19.3 and 20.5 cm, respectively.<sup>15</sup>

The results of the biochemical tests for the lipid profile (total cholesterol, LDL-C, high-density lipoproteincholesterol [HDL-C] and triglycerides) and glycemia were collected from the medical records or provided by the patients. The results for the blood pressure (BP) were also collected from the medical records. The cutoff point to identify changes was BP  $\geq$ 140/90 mmHg, established by the Brazilian Hypertension Guideline.<sup>16</sup>

The HTGW prevalence was evaluated through two cutoff points available in the literature: 1) that proposed by Lemieux et al,<sup>3,5</sup> which considers the HTGW phenotype to be present when the triglyceride levels are  $\geq$ 177 mg/dL and WC  $\geq$ 90 cm in men, and WC  $\geq$ 85 cm for women; and 2) that proposed by the NCEP-ATP III, which establishes an increased risk of CVDs for measurements of WC  $\geq$ 102 cm for men and  $\geq$ 88 cm for women, and triglyceride levels  $\geq$ 150 mg/dL for both sexes.<sup>7</sup>

Statistical analyses were performed using SPSS software (version 22.0; IBM Corporation, Armonk, NY, USA). The normality of the data was tested with the Shapiro–Wilk test. Descriptive statistics were used to characterize the sample, with measures of central tendency, dispersion and frequencies. The chi-square test was used to compare the prevalences of the HTGW phenotype, and Cohen's kappa was used as a measure of agreement between the cutoff points. Comparison of the quantitative variables between individuals with or without the HTGW phenotype was performed with independent *t*-test or the Mann–Whitney *U*-test for parametric and nonparametric variables, respectively. The significance level adopted for all analyses was  $p \leq 0.05$ .

## Results

A total of 437 individuals of both sexes (73.7% female), with a mean age of  $48.4\pm15.9$  years, ranging from 20 years to 78 years, were evaluated. Table 1 shows the characteristics of the sample, demonstrating that the users were mostly married, nonsmokers and sedentary. No statistically significant association was observed between the sociodemographic data and the presence of the HTGW phenotype.

Table 2 presents data on the frequency of the HTGW phenotype, agreement between cutoff points and comparison of the prevalence of the phenotype between the sexes, with the different cutoff points. Significant differences between the sexes were observed only when the cutoff points of NCEP-ATP III<sup>7</sup> were used. The frequency of the HTGW phenotype was elevated at both cutoff points, with a statistically significant difference (p<0.01) between them. A higher prevalence (36.2%) was observed on using the cutoff point suggested by NCEP-ATP III<sup>7</sup> when compared to the cutoff points of Lemieux et al (32.5%).<sup>3,5</sup> The agreement between the cutoff points showed that, although the prevalence of the HTGW phenotype was different, agreement was good (kappa =0.756; SE =0.03).

Characteristics	Number of patients	Percentage	
Profession			
Houseworker	69	15.8	
Farmer	47	10.8	
Retired	93	21.3	
Teacher	11	2.5	
Housekeeper	19	4.3	
Autonomous	37	8.5	
Other	155	35.5	
Marital status			
Single	118	27	
Married	275	62.9	
Widower	26	5.9	
Divorced	18	4.1	
Economic class <sup>a</sup>			
A/B	0	0	
С	198	45.3	
D/E	239	54.7	
Smoker			
Yes	67	15.6	
No	312	71.6	
Former smoker	57	13	
Active physically			
Yes	141	31.3	
No	296	67.7	
Alcohol consumption			
Never	190	43.5	
Rarely	47	10.7	
Once or twice weekly	92	21.1	
, Daily	108	24.7	

**Note:** <sup>a</sup>This definition is based, overall, on the households gross monthly income: Class A, above Brazilian Real (BRL) 10.200; Class B, above BRL 5.100; Class C, above BRL 2.040; Class D, above BRL 1.020; Class E, below BRL 1.020.

Tables 3 and 4 present comparisons between metabolic, cardiovascular and anthropometric variables among individuals with the presence or absence of the HTGW phenotype. In general, for both cutoff points, it is observed that the phenotype is present in individuals with higher age, worse lipid profile, higher glycemia level, overweight and excessive abdominal fat, reinforcing its usefulness to track people with high cardiovascular risk.

# Discussion

The HTGW phenotype stands out because of its high concordance with MS in predicting the risk of developing metabolic diseases<sup>17</sup> in a simpler way, being easy to apply and inexpensive as well as the involvement of only two parameters. For the diagnosis of MS, at least three of five proposed criteria are needed.<sup>18</sup> Although simple, it remains little known and is rarely used by health professionals in routine patient evaluation.<sup>19</sup>

The characteristics of the sample in the present study show the main profile found in basic health care units in 
 Table 2 Differences between sexes, frequency of the HTGW according to comparative classifications and agreement between cutoff points

Classifications	Sex		Total	p-value <sup>a</sup>	<b>p-value</b> ⁵	p-value <sup>c</sup>
	Male	Female				
HTGW/NCEP						
Presence	33 (28.7%)	125 (38.8%)	158 (36.2%)	0.05	<0.01	<0.01
Absence	82 (71.3%)	197 (61.2%)	279 (63.8%)			
Total	115 (100%)	322 (100%)	437 (100%)			
HTGW/Lemieux						
Presence	41 (35.7%)	101 (31.4%)	142 (32.5%)	0.4	-	-
Absence	74 (64.3%)	221 (68.6%)	295 (67.5%)			
Total	115 (100%)	322 (100%)	437 (100%)			

**Notes:** <sup>3</sup>*p*-value for chi-square test between the sexes. <sup>5</sup>*p*-value for chi-square test between cutoff points. <sup>c</sup>*p*-value for Cohen's kappa between cutoff points. HTGW/ NCEP, HTGW according to references of NCEP<sup>7</sup> for waist circumference and triglycerides. HTGW/Lemieux, HTGW according to references of Lemieux et al<sup>3.5</sup> for waist circumference and triglycerides.

Abbreviations: HTGW, hypertriglyceridemic waist; NCEP, National Cholesterol Education Program.

Table 3 Characteristics of the sample according to the presence or absence of HTGW<sup>a</sup>

Variables	Presence of HTGW	Absence of HTGW	p-value
Age, years (n=437)	51.03±14.73	47.2±16.29	< 0.01 <sup>b</sup>
Triglycerides, mg/dL (n=437)	281.43±121.20	I26±60.80	<0.01°
Cholesterol, mg/dL (n=433)	233.32±51.03	188.89±40.62	<0.01 <sup>b</sup>
HDL-cholesterol, mg/dL (n=365)	45.16±10.76	50.93±14.52	<0.01
DL-cholesterol, mg/dL (n=344)	136.90±41.93	113.17±37.55	<0.01
Glucose, mg/dL (n=425)	I 18.20±52.70	95.4±32.80	<0.01
BP, mmHg (n=393)	127.26±14.46	123.01±18.74	<0.01
DBP, mmHg( n=393)	76.84±17.01	77.90±14.35	<0.01
SMI, kg/m² (n=437)	30.05±4.92	28.05±6.48	<0.01
Vaist circumference, cm (n=437)	96.08±12.65	88.90±15.20	<0.01
lip circumference, cm (n=315)	104.58±9.16	101.48±10.77	<0.01
WHR (n=315)	0.89±0.83	0.84±0.09	<0.01
Vaist-to-height ratio (n=437)	0.60±0.07	0.55±0.10	<0.01°
SAD, cm (n=209)	21.67±3.58	20.36±3.90	<0.01

Notes: \*As proposed by Lemieux et al (2000/2007).<sup>3.5</sup> b-value obtained using the chi-square test; \*p-value obtained using independent *t*-test.

Abbreviations: BMI, body mass index; DBP, diastolic blood pressure; HDL, high-density lipoprotein; HTGW, hypertriglyceridemic waist; LDL, low-density lipoprotein; SAD, sagittal abdominal diameter; SBP, systolic blood pressure; WHR, waist-to-hip ratio.

Variables	Presence of HTGW	Absence of HTGW	p-value	
Age, years (n=437)	51.01±14.99	46.99±16.22	< 0.01b	
Triglycerides, mg/dL (n=437)	250.26±116.27	I 35.28±84.87	<0.01°	
Cholesterol, mg/dL (n=433)	223.94±51.66	191.60±43.10	<0.01 <sup>b</sup>	
HDL-cholesterol, mg/dL (n=365)	45.46±10.80	51.09±14.68	<0.01 <sup>b</sup>	
LDL-cholesterol, mg/dL (n=344)	133.14±42.48	113.71±37.65	<0.01	
Glucose, mg/dL (n=425)	I16±50.50	100.06±39.63	<0.01	
SBP, mmHg (n=393)	125.68±13.92	123.66±19.41	<0.01	
DBP, mmHg (n=393)	77.77±16.05	77.4I±I4.8I	<0.01	
BMI, kg/m² (n=437)	31.05±5.18	27.36±6.16	<0.01 <sup>b</sup>	
Waist circumference, cm (n=437)	97.92±12.66	87.44± 14.59	<0.01	
Hip circumference, cm (n=315)	106.44±9.20	100.16±10.26	<0.01	
WHR (n=315)	0.89±0.08	0.84±0.09	<0.01	
Waist-to-height ratio (n=437)	0.61±0.07	0.54±0.09	<0.01°	
SAD, cm (n=209)	22.24±3.67	20.03±3.71	<0.01 <sup>b</sup>	

Notes: "Based on NCEP criteria (2000).7 bp-value obtained using the chi-square test; cp-value obtained using independent t-test.

Abbreviations: BMI, body mass index; DBP, diastolic blood pressure; HDL, high-density lipoprotein; HTGW, hypertriglyceridemic waist; LDL, low-density lipoprotein; NCEP, National Cholesterol Education Program; SAD, sagittal abdominal diameter; SBP, systolic blood pressure; WHR, waist-to-hip ratio.

Brazil: the majority were female, married and sedentary, with no usual alcohol consumption and belonging to the less-favored socioeconomic class. These characteristics were also found in other studies in the country, which indicate that women are the main users of basic health care units.<sup>20,21</sup> The predominance of the population in the less-favored economic classes, as observed by Haack et al,<sup>21</sup> can be explained by the data collection site, which is a public health service. The low frequency of smokers, physically active persons or frequent consumers of alcohol explains the fact that these characteristics are not associated with the HTGW phenotype.

Our results showed differences in the presence of the phenotype according to the cutoff points, as well as a good agreement between the cutoff points, with the highest frequency observed when the cutoff point suggested by the NCEP/ATP III<sup>7</sup> was used. Until now, there have been few studies to report appropriate cutoffs of HTGW among Brazilian adults or the association between HTGW and risk for chronic diseases. Unfortunately, in the present study, it was not possible to predict the best cutoff point for the study population because we did not have access to the medical diagnoses of diabetes or hypertension (many subjects were not sure if they had these diseases). Recognizing that the Brazilian population is characterized by different ethnicities and does not have a specific cutoff point for indicators of abdominal obesity, we suggest the use of NCEP/ATP III cutoff point because it can recognize more patients at risk and, using this, we anticipate the prevention of CVDs.

In addition to the increased incapacitation and mortality associated with overweight and obesity, there is numerous evidence associating visceral fat accumulation with increased cardiovascular risk of metabolic origin. Thus, visceral adipose tissue becomes a major cardiovascular risk factor correlating well with proatherogenic, prodiabetogenic, prothrombogenic and proinflammatory status.<sup>22</sup> It is also established that the accumulation of abdominal fat may be the result of an increase in visceral or subcutaneous adipose tissue and is an indicator for the prediction of cardiovascular risk.<sup>15</sup> Thus, it is proven that the HTGW phenotype is a useful marker for the identification of cardiovascular risk by recognizing the association between WC and triglyceride levels.

Among the few available studies on the prevalence of the HTGW phenotype in Brazil, a study conducted by Guattini et al<sup>4</sup> in the city of Porto Alegre, Rio Grande do Sul, with individuals aged  $\geq$ 40 years, showed a 48.9% prevalence of individuals with the phenotype and 55.6% of individuals with increased WC. The cutoff point adopted for that study was suggested by Brazilian guidelines (WC  $\geq$ 94 cm for

men and  $\geq$ 80 cm for women and TG  $\geq$ 150 mg/dL for both sexes).<sup>23</sup> Another study carried out by Cabral et al<sup>20</sup> found a prevalence of 33% in hypertensive women  $\geq$ 50 years old in the city of São Luís, Maranhão, using the cutoff points of the NCEP.<sup>7</sup> However, in the southern region of Brazil, smaller values were found in a study conducted with young adults between 22 and 23 years of age, with a prevalence of the HTGW phenotype of 5.9% in men and 4.5% in women.<sup>21</sup> The cutoff point adopted in that study was that proposed by Lemieux et al.<sup>3,5</sup>

Mendes and Melendez<sup>2</sup> also performed a population-based study in Brazil with the purpose of describing the HTGW phenotype. A total of 506 individuals, 42.13±15.94 years old, were evaluated in the city of Belo Horizonte, Minas Gerais, and a 11.1% prevalence of individuals with HTGW was found. The cutoff point adopted in that study was calculated from the receiver operating characteristic (ROC) curve obtained with the values found in the sample, wherein the following cutoff values were obtained: WC ≥85.30 cm and triglyceride levels ≥114.8 mg/dL for men; and WC  $\geq$ 87.50 cm and triglyceride levels  $\geq$ 124.5 mg/dL for women. Although the proposed cutoff point presented lower values, the prevalence of HGTW was lower than that found in the present study, which used cutoff points with higher values commonly found in the literature. This can be explained by the fact that the population in that study was younger and because it was evaluated at random. Our study was carried out with older individuals who seek primary health care, which thus is characterized as a higher-risk group among the public. The low value of the proposed cutoff points were unsuitable for use in the present study because it would be an underestimation of the result found for the phenotype.

In other studies performed in populations from different countries, different prevalence rates of HTGW were observed. In Spain, in a study carried out by Cortés et al,<sup>17</sup> it was possible to evaluate the presence of the phenotype from two cutoff points: NCEP ATP III and the International Diabetes Federation (IDF) (WC ≥94 cm and ≥80 cm for men and women, respectively; and triglyceride levels ≥150 mg for both sexes).<sup>24</sup> The study observed a prevalence of 49.2% of individuals with HTGW using the first cutoff point and a prevalence of 79.7% using the second. This difference can be explained by the fact that the cutoff point suggested by the IDF presents lower values for WC. In another study, also carried out in Spain and evaluating 2270 individuals between 18 and 80 years old, a variation of 14.5% in individuals with the presence of the HTGW phenotype was observed through the cutoff point suggested by NCEP-ATP III.25

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Amini et al<sup>26</sup> evaluated 1323 Iranians with a clinical history of type II diabetes and age between 30 and 55 years, and they found a prevalence of HTGW of 23.6% among women and 9.6% among men using the cutoff point of NCEP-ATP III. Gasevic et al<sup>27</sup> evaluated 809 adults of different ethnicities living in Canada and aged 30-65 years old, and detected a prevalence of 22.3% in men and 17% in women, using the cutoff point of Lemieux et al.<sup>3,5</sup> In Malaysia, Zainuddin et al<sup>28</sup> evaluated 298 people aged 18-70 years using two different cutoffs for the HTGW phenotype: IDF<sup>24</sup> and NCEP/ATPIII.<sup>7</sup> Similar to the results of the present study, those authors found good concordance of these cutoff points, and the prevalence of the HTGW phenotype was 19.7% (with the IDF cutoff point). Thus, we can see that there is no consensus among authors about the best cutoff point for identifying the HTGW phenotype in different countries.

The prevalence of MS in adults identified with the HTGW phenotype is high. A study observed that 97.5% of individuals with the phenotype also had MS.<sup>28</sup> Thus, the correlation in the diagnosis of cardiovascular risk among these evaluation criteria can be considered high. Similar results were observed in a study of 254 women undergoing angiography to verify coronary artery disease.<sup>29</sup> The cutoff point used was the one proposed by NCEP-ATP III,<sup>7</sup> in which a higher cardiovascular risk was detected among participants with MS, the HTGW phenotype and a high Framingham risk score.<sup>29</sup>

The present study also observed that individuals with the HTGW phenotype show different changes in cardiometabolic markers, regardless of the cutoff point. People who do not have the phenotype have fewer changes in these markers as well as in the factors considered protective against the development of CVDs, such as higher HDL-C. Similar results were found by Rocha et al<sup>30</sup> in a study carried out in Viçosa, Minas Gerais, evaluating 976 individuals aged 20–59 years, wherein the authors identified that the highest averages for all cardiometabolic risk factors analyzed were observed in the HTGW phenotype group and lower values of HDL were also found only in this group.

One limitation of the present study was that a cardiovascular event was not adopted to test the prediction of the phenotype. Studies in the literature have investigated the association of the phenotype with subclinical atherosclerosis,<sup>27</sup> cholesterol and C-reactive protein<sup>23</sup> as well as overweight and glycemia levels.<sup>22</sup> It is suggested that future investigations study the frequency of the phenotype and its association with events associated with metabolic risk in order to improve the knowledge about these interrelations and to potentiate the clinical use of this indicator.

#### Conclusion

There was a high frequency of the HTGW phenotype in the population, regardless of the cutoff point used, as well as a high frequency of cardiovascular risk factors, especially in participants with HTGW. The frequency was higher when using the cutoff points suggested by NCEP-ATPIII. Thus, it is reaffirmed that the HTGW phenotype is a simple and inexpensive marker that should be used in clinical practice because, through this research and other reports available in the literature, it was observed that there is a high prevalence of the HTGW phenotype in the population. Further prospective investigations are necessary to understand the predictive usefulness of the HTGW phenotype in a population attended to in a primary health care setting, to screen for patients at a high risk for diabetes and other cardiometabolic diseases.

#### Author contributions

All authors contributed toward data analysis, drafting and critically revising the paper and agree to be accountable for all aspects of the work.

## Disclosure

The authors report no conflicts of interest in this work.

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