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A WAIST-TO-HEIGHT RATIO OF 0.54 IS A GOOD PREDICTOR OF METABOLIC SYNDROME IN 16-YEAR-OLD MALE AND FEMALE ADOLESCENTS

Fabian Vasquez¹, Paulina Correa-Burrows¹, Estela Blanco², Sheila Gahagan², and Raquel Burrows¹

¹Institute of Nutrition and Food Technology, University of Chile. Santiago, Chile.

²Division of Child Development and Community Health, University of California San Diego. La Jolla, California

Abstract

Background: We aimed to determine the sensitivity and specificity of selected anthropometric indicators as predictors of cardiovascular risk in adolescents.

Methods: Cross-sectional study in 678 adolescents ($16.8y \pm 0.3$) from an infancy cohort. Weight, height, waist circumference (WC) and hip circumference were measured. Body-Mass Index (BMI), waist-to-hip ratio (WHR) and waist-to-height ratio (WHtR) were estimated. MetS was diagnosed with IDF/AHA/NHLBI. Optimal cutoffs of BMI, WC, WHR and WHtR for diagnosing MetS were determined using ROC analysis.

Results: In males, WHtR (0.96) had the greatest area under the ROC curve, followed by WC (0.95) and BMI (0.93). In females, BMI (0.84) had the greatest area under the ROC curve (0.84), followed by WHtR (0.83) and WC (0.83). In both sexes, the optimal WHtR cutoff for MetS diagnosis was 0.54. A BMI of 26.9 in males and 26.3 in females were the optimal cutoffs for diagnosing MetS. Finally, WC values of 92 and 81.6 cm in males and females, respectively, were the optimal cutoffs for MetS diagnosis.

Conclusions: In both sexes, a WHtR value of 0.54 was a good predictor of MetS. In males and females, the optimal cutoff of BMI for MetS diagnosis was below the values for diagnosing obesity.

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Contact Information: Prof. Dr. Raquel Burrows. Unidad de Epidemiología Nutricional. Instituto de Nutrición y Tecnología de los Alimentos. Universidad de Chile. Avda. El Líbano 5524, Macul. CP: 7830490. Santiago de Chile, Chile. Tel. +56 2 29781492. Fax: +56 2 22214030. rburrows@inta.uchile.cl.

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INTRODUCTION

Obesity in childhood and adolescence is associated with metabolic and endocrine disorders, which predispose children to early development of cardiovascular disease and type-2 diabetes mellitus (T2DM) (1,2). A clustering of metabolic abnormalities, including obesity, hyperglycemia, dyslipidemia, and hypertension, has been referred to as Metabolic Syndrome (MetS). MetS is an independent predictor of cardiovascular morbidity and mortality, and identifies greater additional biological risk beyond the sum of the individual risk factor related to obesity and insulin resistance (IR)(3,4).

Excessive weight gain has dramatically increased in Chilean children and adolescents over the past two decades. In 2015, 52% of 1st graders and 44% of 9th graders were either overweight or obese, according to a population survey (5). A study conducted in Chilean children and adolescents, ages 10-15, found a high prevalence of obesity (16.1%), MetS (7.3%) and insulin resistance (IR) (26%). Among obese participants, prevalence rates of MetS and IR were especially high (62% and 29%, respectively)(6). Similarly, in a sample of 16-year-old Chilean adolescents of mid- to low socioeconomic status, 16% had obesity, 79% had at least one cardiovascular risk, and 9.5% had MetS. One in three adolescents had abdominal obesity, 70% had low high-density lipoproteins (HDL-cholesterol) and fasting hyperglycaemia prevalence was 8.7% (7).

Anthropometric indicators of obesity such as waist circumference (WC), waist-to-hip ratio (WHR) and waist-to-height ratio (WHtR) are associated with adverse cardiometabolic consequences in children and adolescents (8-11). Abdominal fat is metabolically active and has been linked to insulin hypersecretion and MetS. In obesity and IR, fat tissue increases the plasma concentration of free fatty acids, adipokines and proinflammatory cytokines (12-14). Although WC correlates with the amount of intra-abdominal visceral fat, it is also associated with abdominal subcutaneous fat and total body fat (15-18). However, WHtR best expresses central adiposity and, therefore, is better able to determine the cardiovascular risk associated with MetS in all age groups (11-15). In adults, it was found that WHtR is strongly and positively correlated with BMI ($r=0.85-0.91$) and body fat percentage ($r=0.69-0.76$), assessed with plethysmography (19). Additionally, in adults, WHtR was as reliable as Body-Mass Index (BMI) in predicting endothelial dysfunction compared to WHR and WC (20). Studies suggest that WHtR has a stronger association with cardiovascular risk factors in children, adolescents and adults compared to BMI (8-11,15). However, differences between WHtR and BMI are relatively small (9,19,21). Inexpensive, easy-to-measure anthropometric indicators which have good sensitivity to identify obesity-related cardiometabolic disorders might be useful in the early detection of individuals with increased biological risk. In this study, we aimed to compare the sensitivity and specificity of BMI, WC, WHR and WHtR as predictors of cardiovascular risk in male and female adolescents and to determine the optimal cutoff values for MetS diagnosis.

METHODS

Participants

We studied 16-17-year-old adolescents living in Santiago, Chile, from low-to-middle socioeconomic status (SES), who were part of follow-up study beginning in infancy. Participants were recruited at 4 months from public healthcare facilities in the southeast area of Santiago ($n=1,791$) to participate in a randomized controlled trial of iron supplementation to prevent iron deficiency anemia (IDA). They were born at term of uncomplicated vaginal births, weighted >3.0 kg, and were free of acute or chronic health problems (22). They were assessed for developmental outcomes in infancy, 5, 10 and 15 years. At 16y, those with complete data ($n=678$) were also assessed for obesity/cardiovascular risk (7). The study has been approved by the institutional review boards of the University of Michigan, Institute of Nutrition and Food Technology, University of Chile, and University of California, San Diego. Participants and their primary caregiver provided informed and written consent, which was obtained according to the norms for Human Experimentation, Code of Ethics of the World Medical Association (Declaration of Helsinki, 1995).

Anthropometric assessment

Weight (kg), height (cm), waist and hip circumference (cm) were measured by physician-investigators. Standardized procedures were used to measure weight to the closest 0.1 kg, using a SECA scale (SECA 703, Seca GmbH & co. Hamburg, Germany), and height to the closest 0.1 cm, using a Holtain stadiometer. BMI (Kg/m^2). BMI Z score for age (BAZ) was estimated and weight status was evaluated according to WHO references (23). WC and hip circumference (HC) were measured with non-elastic flexible tape (Seca 201, Seca GmbH & co. Hamburg, Germany) at the highest point of the iliac crest around the abdomen and at the level of the greater trochanter, respectively, and recorded to the nearest 0.1 cm (24). Measurements were taken twice, with a third measurement if the difference between the first two exceeded 0.3 kg for weight, 0.5 cm for height and 1.0 cm for waist and hip. WC, HC and height were used to calculate WHtR and WHR.

Cardiometabolic assessment

Systolic and diastolic blood pressures (SBP and DBP) were measured in the non-dominant arm with a standard mercury sphygmomanometer in the morning after 15 minutes at rest, according to the National High Blood Pressure Education Working Group recommendations; the mean of three measurements was used for analysis (25). Fasting serum total glucose (Gli), total cholesterol (TChol), triglycerides (TG), HDL, and insulin levels were measured after a 12-hour overnight fast. Radioimmunoassay (RIA, DCP Diagnostic Products Corporation, LA) with intra-assay CV of 5.1% and inter-assay CV of 7.1% for 14.4 uUI/ml, and a sensitivity of 1.2 uUI/ml was used for insulin determination. Glucose was measured with enzymatic-colorimetric test (QCA S.A. Amposta, Spain). Baseline insulin sensitivity was calculated using the homeostasis model assessment (HOMA) method [$\text{HOMA} = \text{fasting insulin (uUI/ml)} \times \text{fasting glycaemia (mg/dl)} / 405$]. Cholesterol profile (HDL and TG mg/dl) was determined using the drychemical method (Vitros, Johnson & Johnson, Clinical Diagnostics Inc.).

Definition of Metabolic Syndrome

MetS was diagnosed with the joint IDF/AHA/NHLBI phenotype, which includes having three of five risk factors using the following definitions: abdominal obesity (WC 80 and 90 cm in females and males, respectively); high blood pressure (SBP 130 mmHg and/or DBP 85 mmHg); hypertriglyceridaemia (TG 150 mg/dl); low HDL (HDL 50 and 40 mg/dl in females and males, respectively); and fasting hyperglycaemia (Gli 100 mg/dl) (26). Adolescents in the sample were not taking any antihypertensive, lipid-lowering or hypoglycemic medications.

Statistical analysis

All variables were checked for normality of distribution (Shapiro-Wilk test) before the analysis. Statistical analysis included Student's *t* test and Wilcoxon's rank-sum test for comparison of mean or median values of anthropometric and cardiometabolic variables. The χ^2 test was used for comparison of categorical variables. Receiver operating characteristic (ROC) analysis was used to find the optimal cutoff of BMI, WC, WHR and WHtR for MetS diagnosis in males and females. A test with perfect discrimination has a ROC plot that passes through the upper left corner, indication of 100% sensitivity and 100% specificity. A ROC plot closer to the upper left corner denotes greater accuracy of the test. To determine the optimal cutoffs for MetS diagnosis, the point on the ROC curve with maximum Youden Index [sensitivity-(1-specificity)] was calculated. Next, the values were verified with the likelihood ratio for a positive result (LR+) and the post-test probability (the proportion of participants above cutoffs who truly have the MetS). Data were analyzed using Stata for Windows V.15.0 (Lakeway Drive College Station, Texas).

RESULTS

Anthropometric characteristics and individual components of MetS are presented for males and females in Table 1. A total of $n=678$ adolescents (52% males) were evaluated. Participants' mean age was 16.8y (0.3 *SD*). Males had significantly higher values of height, weight and WHR, and lower values of BMI, HC and WHtR compared to females. As for the cardiometabolic profile, males had significantly higher levels of SBP, DBP and fasting glucose and lower values of insulin, total cholesterol and HDL cholesterol than females. No difference was found in the prevalence of MetS between males (8.5%) and females (8.8%).

In males (Figure 1), WHtR had the greatest area under the ROC curve (0.96), followed by WC (0.95) and BMI (0.93), suggesting a greater accuracy in predicting MetS. In females (Figure 2), BMI was the anthropometric indicator with the greatest area under the ROC curve (0.84), followed by WHtR (0.83) and WC (0.83).

In Table 2, the optimal cutoff points of anthropometric indicators to predict MetS in male and female adolescents are presented. In males, cutoffs of 26.9 for BMI, 92.0 cm for WC, 0.91 for WHR, and 0.54 for WHtR had the best sensitivity and specificity for MetS diagnosing. In females, the optimal cutoffs for MetS diagnosis were 26.3 for BMI, 81.6 cm for WC, 0.87 for WHR and 0.54 for WHtR. In males, the percentage of correctly classified participants was higher than females. Also, the probability of having MetS if the adolescent

was above the optimal cut-off point of BMI, WC, WHR and WHtR was markedly higher in males compared with females.

The cardiometabolic profile of participants by optimal cutoff values of BMI, WC, WHR and WHtR are shown in Table 3. In males and females with BMI ≥ 26.9 and ≥ 26.3 , respectively, we found significantly higher values of all MetS-related biomarkers. In the sample, 18.9% of men and 26.5% of women had BMI values equal to or greater than the optimal cut-off point. In males, participants with WC ≥ 92.0 cm had significantly higher values of SBP, DBP, fasting insulin, TChol and TG. Similarly, in females with WC values ≥ 81.6 cm significantly higher values of SBP, DBP, fasting insulin and TG were observed. Males with WHR ≥ 0.91 and females with values ≥ 0.87 had significantly higher values in all MetS biomarkers, with the exception of fasting glucose and TChol in females. In both sexes, WHtR ≥ 0.54 was associated with significantly higher values of all MetS-related biomarkers, except for fasting glucose and TChol in females.

DISCUSSION

Main findings

This study aimed to compare the sensitivity and specificity of BMI, WC, WHR and WHtR as predictors of cardiometabolic risk in Chilean adolescents between 16-17 years of age, as measured by MetS. We found that WHtR and BMI, along with WC, which is a component of the MetS, had high sensitivity and specificity for the diagnosis of MetS. A study examining the association of total and central adiposity with cardiovascular risk in eutrophic and obese adolescents (13.2y) showed that BMI and WHtR were associated with higher levels of triglycerides and total cholesterol and low HDL cholesterol and apolipoprotein B levels in the obese participants (27). In our sample, WHtR in males and BMI in females had the highest sensitivity and specificity to define an optimum cutoff value for predicting MetS. These results are consistent with those obtained by the Bogalusa Heart Study, in children ages 5-17y ($n=2,498$) (28). WHtR was a good predictor of altered LDL and HDL concentrations whereas BMI performed well in predicting high levels of fasting insulin, SBP and DBP. Similarly, in the Childhood and Adolescence Surveillance and Prevention of Adult Non-communicable Disease-IV Study conducted in a representative sample of children and adolescents ages 6-18 y from Iran ($n=4,811$), BMI, WC and WHtR were the most accurate indicators for predicting cardiovascular risk factors (29). Although, BMI for age and WHtR did not differ in their ability to identify children with adverse risk factors, it is worth noting that the optimal cutoffs value of BMI for MetS diagnosis in our sample were below the cutoff for obesity diagnosis according to WHO 2007 and CDC 2000 standards (23,30). This suggests that biological risk associated with increased body fat is being underestimated in adolescents. In non-Hispanic white US adolescents, Peterson *et al.*, found that the tri-ponderal mass index was a more reliable estimator of body fat than BMI (31). Furthermore, in a sample of $n= 3,091$ US children (ages 7-17) from the Bogalusa Heart Study, Mokha *et al.* validated WHtR as predictors of cardiovascular risk and reported that WHtR performed better than BMI in predicting the cardiovascular risk in children with and without obesity (8). These results are consistent with those obtained by Khoury *et al.* in a representative sample of the US population of children and adolescents (10).

WHtR has been suggested as a useful measure of cardiovascular risk and a cutoff value of 0.50 and 0.54 has been proposed to identify children and adults with increased cardiometabolic risk, which is similar to the cutoff obtained in our study (8,9,32,33). Other studies in children and adolescents report results similar to ours. Using data from the National Health and Nutrition Examination Survey III, Kahn *et al.*, found that WHtR performed better than sex- and age-specific BMI percentiles to identify US children and adolescents with adverse cardiovascular risk factors (34). Similarly, in n=209 Chilean children (ages 6 to 17), WHtR was the best predictor of components aggregation of MetS and the optimal cut-off was 0.55 (35). In the Bogalusa Study, overweight children with WHtR in the top tertile were 2-3 times more likely to have cardiovascular risk factors than those with low WHtR (36). Finally, two studies conducted in US and Japanese children are also in line with our findings (8,37). US children and adolescents with a WHtR ≥ 0.5 were more likely to have significant adverse levels of LDL cholesterol, HDL cholesterol, triglycerides and insulin (8). In Japanese children ages 9-13 years, WHtR performed better in predicting total cholesterol, triglycerides, LDL cholesterol and atherogenic index compared to BMI, WHR and WC (37). In this sample, WHtR was the most sensitive indicator for males and the second most sensitive indicator in female for diagnosing MetS and the optimal cutoff value was the same for both sexes (0.54). This facilitates use and proves its utility at the clinical and population level.

We observed sex differences in the four anthropometric indicators used to identify adolescents with higher biological risk. We found that if the adolescent was above the optimal cut-off point for all anthropometric indicators, the likelihood of having MetS was higher in males than females. Thus, some females may have lower cardiovascular risk as measured by the MetS despite having increased values of BMI, WC, WHR, and WHtR. Several studies show a sexual dimorphism in the cardiometabolic risk associated with the increased body fat mass (38,39). Several studies show that body fat distribution has a greater impact on cardiometabolic risk compared to total adiposity. Males with excess weight usually have visceral fat deposits, whereas in women with excess weight, subcutaneous fat deposits predominate (40,41). Although our research did not identify potential protective factors of cardiometabolic risk in females with higher body fat, our findings confirm the importance of estimating the optimal cut-off point of each anthropometric indicator separately by sex, in order to identify adolescents at greater biological risk.

Limitations and strengths

This study has limitations that should be considered when interpreting its results. First, our sample is not representative of the Chilean adolescent population, as this was comprised of adolescents from low- to middle SES between a narrow range of 16 to 17 years. Our findings, however, may be equally relevant for several of reasons. The prevalence of obesity and cardiometabolic risk is significantly higher in individuals of low- to middle SES according to population-based surveys (44,45). Second, low- to middle-SES adolescents are highly exposed to risk factors that have a direct effect on the development of MetS (6,7,42-45). Third, studies in children and adolescents of all SES levels have found similar values for the WHtR (8,35,37). A further limitation is the cross-sectional nature of the study, which limits the ability to draw conclusions related to the temporality of these associations.

Future studies should aim to longitudinally explore the performance of these indicators in predicting the risk of cardiometabolic disorders in adulthood. Yet, our study contributes with scientific evidence to select the best anthropometric indicators to identify early cardiovascular risk. Also, it provides knowledge that allows nutritional diagnosis using inexpensive, easy-to-measure anthropometric indicators which are based on biological risk and are potentially useful in both clinical and population settings. Last, our study observed sex differences in the effectiveness of these anthropometric indicators to identify adolescents at greater cardiometabolic risk. To the best of our knowledge, this has not been described in adolescents.

Finally, although a controversy exists on how to diagnose the MetS in children and adolescence, in this study we use the joint IDF/AHA/NHLB criteria. This represents the consensus of several major organizations in an attempt to unify criteria to diagnose MS in a population older than 15 years (26).

Conclusion

WHtR, WC and BMI-z are good predictors of MetS in adolescents. Although BMI was one of the three best indicators associated with cardiovascular risk, especially in females, the cut-off point of greater sensitivity and specificity for predicting biological risk was below the cut-off point for obesity diagnosis. The use of WHtR might have advantages over BMI z-score and WC percentile related to the ease of calculation and application at the individual- or population-level, and because the same cut-off value can be used in both sexes.

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References

1. Obesity Aggoun Y., metabolic syndrome, and cardiovascular disease. *Pediatr Res* 2007; 61:653–659. [PubMed: 17426660]
2. Liang Y, Hou D, Zhao X, Wang L, Hu Y, Liu J, et al. Childhood obesity affects adult metabolic syndrome and diabetes. *Endocrine* 2015; 50(1):87–92. [PubMed: 25754912]
3. Weiss R, Dziura J, Burgert TS, Tamborlane WV, Taksali SE, Yeckel CW, et al. Obesity and the metabolic syndrome in children and adolescents. *N Engl J Med* 2004; 350: 2362–2374. [PubMed: 15175438]
4. Burrows R, Correa-Burrows P, Reyes R, Blanco E, Albala C, Gahagan S. *J Diabetes Res.* 2015;2015:783296. doi: 10.1155/2015/783296. [PubMed: 26273675]
5. Informe Mapa Nutricional 2015 Junta Nacional de Auxilio Escolar y Becas. Ministerio de Educación: Santiago, Chile [Online]. Available from: <https://www.junaeb.cl/wp-content/uploads/2016/11/Informe-Mapa-Nutricional-2015-final.pdf>. Accessed Jan. 23, 2015.
6. Mardones F, Amaiz P, Barja S, Giadach C, Villarroel L, Domínguez A, et al. Nutritional status, metabolic syndrome and insulin resistance in children from Santiago (Chile). *Nutr Hosp* 2013; 28: 15871593.

7. Burrows R, Correa-Burrows P, Reyes M, Blanco E, Albala C, Gahagan S. High cardiometabolic risk in healthy Chilean adolescents: Association with anthropometric, biological and life style factors. *Public Health Nutrition*, 2015; Doi.org/10.1017/S136898001500158
8. Mokha J, Srinivasan S, Dasmahapatra P, et al. Utility of waist-to-height ratio in assessing the status of central obesity and related cardiometabolic risk profile among normal weight and overweight/obese children: The Bogalusa Heart Study. *BMC Pediatr* 2010; 10:73. doi: 10.1186/1471-2431-10-73. [PubMed: 20937123]
9. Sijtsma A, Bocca G, L'abée C, Liem E, Sauer P, Corpeleijn E. Waist-to-height ratio, waist circumference and BMI as indicators of percentage fat mass and cardiometabolic risk factors in children aged 3-7 years. *Clin Nutr* 2014; 33(2):311–5. doi: 10.1016/j.clnu.2013.05.010. [PubMed: 23768783]
10. Khoury M, Manlhiot C, McCrindle B. Role of the waist/height ratio in the cardiometabolic risk assessment of children classified by body mass index. *J Am Coll Cardiol* 2013; 62(8):742–51. doi: 10.1016/j.jacc.2013.01.026. [PubMed: 23500256]
11. Zhou D, Yang M, Yuan Z, et al. Waist-to-Height Ratio: a simple, effective and practical screening tool for childhood obesity and metabolic syndrome. *Prev Med* 2014; 67:35–40. doi: 10.1016/j.ypmed.2014.06.025. [PubMed: 24967955]
12. Dain A, Repossi G, Das UN, Eynard A. Role of PUFAs, the precursors of endocannabinoids, in human obesity and type 2 diabetes. *Front Biosci* 2010; 2: 1432–1447.
13. Capurso C, Capurso A. From excess adiposity to insulin resistance: the role of free fatty acids. *Vascul Pharmacol* 2012; 57: 91–97. [PubMed: 22609131]
14. Ouchi N, Parker J, Lugus J, et al. Adipokines in inflammation and metabolic disease. *Nat Rev Immunol*. 2011; 11: 85–97. [PubMed: 21252989]
15. Lee C, Huxley R, Wildman R, Woodward M. Indices of abdominal obesity are better discriminators of cardiovascular risk factors than BMI: A meta-analysis. *J Clin Epidemiol* 2008; 61:646653.
16. Lean M, Han T, Deurenberg P. Predicting body composition by densitometry from simple anthropometric measurements. *Am J Clin Nutr* 1996; 63:4–14. [PubMed: 8604668]
17. Molarius A, Seidell J. Selection of anthropometric indicators for classification of abdominal fatness: A critical review. *Int J Obes Relat Metab Disord* 1998; 22:719–727. [PubMed: 9725630]
18. Hebert P, Rich-Edwards J, Manson J, Ridker P, Cook N, O'Connor G, et al. Height and incidence of cardiovascular disease in male physicians. *Circulation* 1993; 88:1437–1443. [PubMed: 8403290]
19. Bosy-Westphal A, Geisler C, Onur S, Korth O, Selberg O, Schrezenmeir J, et al. Value of body fat mass vs anthropometric obesity indices in the assessment of metabolic risk factors. *Int J Obes (Lond)* 2006; 30:475–483. [PubMed: 16261188]
20. Maher V, O'Dowd M, Carey M, Markham C, Byrne A, Hand E, et al. Association of central obesity with early Carotid intima-media thickening is independent of that from other risk factors. *Int J Obes* 2009; 33(1): 136–143. doi: 10.1038/ijo.2008.254
21. Hsieh S, Yoshinaga H, Muto T. Waist-to-height ratio, a simple and practical index for assessing central fat distribution and metabolic risk in Japanese men and women. *Int J Obes Relat Metab Disord* 2003; 27:610–616. [PubMed: 12704405]
22. Lozoff B, De Andraca I, Castillo M, Smith JB, Walter T, Pino P. Behavioral and developmental effects of preventing iron-deficiency anemia in healthy full-term infants. *Pediatrics* 2003; 112:846–854. [PubMed: 14523176]
23. De Onis M, Onyango A, Borghi E, Siyam A, Nashidaa C, Siekmanna J. Growth reference data for 5-19 years. *Bull World Health Organ* 2007; 85:660–667. [PubMed: 18026621]
24. Fernandez J, Redden D, Pietrobelli A, Allison D. Waist circumference percentiles in nationally representative samples of African-American, European-American, and Mexican-American children and adolescents. *J Pediatric* 2004; 145:439–444.
25. National High Blood Pressure Education Program Working Group on High Blood Pressure in Children and Adolescents. The fourth report on the diagnosis, evaluation, and treatment of high blood pressure in children and adolescents. *Pediatrics* 2004; 114: 555–576. [PubMed: 15286277]

26. Alberti K, Eckel R, Grundy S, Zimmet P, Cleeman J, Donato K, 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: 1640–1645. [PubMed: 19805654]
27. Teixeira P, Sardinha L, Going S, Lohman T. Total and regional fat and serum cardiovascular disease risk factors in lean and obese children and adolescents. *Obes Res* 2001; 9: 432–442. [PubMed: 11500523]
28. Freedman D, Kahn H, Mei Z, Grummer-Strawn L, Dietz W, Srinivasan S, et al. Relation of body mass index and waist-to-height ratio to cardiovascular disease risk factors in children and adolescents: the Bogalusa Heart Study. *Am J Clin Nutr* 2007; 86: 33–40. [PubMed: 17616760]
29. Kelishadi R, Heidari-Beni M, Azizi-Soleiman F, Ardalan G, Khoshhali M, Heshmat R, et al. Reference curves of anthropometric indices in two national studies conducted among Iranian children in 2003-2004 and 2009-2010: The Caspian study. *J Res Med Sci* 2014; 19: 709–714. [PubMed: 25422654]
30. Kuczmarski R, Ogden C, Guo S, Grummer-Strawn L, Flegal K, Mei Z, et al. 2000 CDC Growth Charts for the United States: methods and development. *Vital Health Stat* 11 2002;1–190.
31. Peterson C, Su H, Thomas D, Heo M, Golnabi A, Pietrobelli A, et al. Tri-Ponderal Mass Index vs Body Mass Index in Estimating Body Fat During Adolescence. *JAMA Pediatr*. 2017. doi: 10.1001/jamapediatrics.2017.0460.
32. Ashwell M, Hsieh S. Six reasons why the waist-to-height ratio is a rapid and effective global indicator for health risks of obesity and how its use could simplify the international public health message on obesity. *Int J Food Sci Nutr* 2005; 56: 303–307. [PubMed: 16236591]
33. Nakamura K, Nanri H, Hara M, Higaki Y, Imaizumi T, Taguchi N, et al. Optimal cutoff values of waist circumference and the discriminatory performance of other anthropometric indices to detect the clustering of cardiovascular risk factors for metabolic syndrome in Japanese men and women. *Environ Health Prev Med*. 2011; 16(1): 52–60. doi: 10.1007/s12199-010-0165-y. [PubMed: 21432217]
34. Kahn H, Imperatore G, Cheng Y. A population-based comparison of BMI percentiles and waist-to-height ratio for identifying cardiovascular risk in youth. *J Pediatr* 2005; 146: 482–488. [PubMed: 15812450]
35. Arnaiz P, Marin A, Pino F, Barja S, Aglony M, Navarrete C, et al. Waist height ratio, ultrasensitive c reactive protein and metabolic syndrome in children. *Rev Med Chil* 2010; 138: 13781385.
36. Freedman D, Dietz W, Srinivasan S, Berenson G. Risk factors and adult body mass index among overweight children: the Bogalusa Heart Study. *Pediatrics* 2009; 123: 750–757. [PubMed: 19254998]
37. Hara M, Saitou E, Iwata F, Okada T, Harada K. Waist-to-height ratio is the best predictor of cardiovascular disease risk factors in Japanese schoolchildren. *J Atheroscler Thromb* 2002; 9: 127–132. [PubMed: 12226553]
38. Caranti D, Lazzar S, Damaso A, Agosti F, Zennaro R, De Mello M, et al. Prevalence and risk factors of metabolic syndrome in Brazilian and Italian obese adolescents: a comparison study. *Int J Clin Pract* 2008; 62:1526–1532 [PubMed: 18822022]
39. Schorr M, Dichtel L, Gerweck A, Valera R, Torriani M, Miller KK et al. Sex differences in body composition and association with cardiometabolic risk. *Biol Sex Differ* 2018; 9:28. doi: 10.1186/s13293018-0189-3. [PubMed: 29950175]
40. Bjorntorp P, Rosmond R. Visceral obesity and diabetes. *Drugs* 1999; 58(Suppl 1): 13–18. [PubMed: 10576519]
41. Tchermof A, Despres I. Pathophysiology of human visceral obesity: an update. *Physiol Rev*. 2013; 93:359–404. [PubMed: 23303913]
42. National Health Survey 2016-2017. Department of Epidemiology. Santiago de Chile: Ministry of Health; 2017 Retrieved from: www.minsal.cl/wp-content/uploads/2017/11/ENS-2016-17_PRIMEROS-RESULTADOS.pdf

43. National Food and Consumption Survey 2016-17. Preliminary results. Department of Epidemiology. Santiago de Chile: Ministry of Health; 2017 Retrieved from: http://www.minsal.cl/wpcontent/uploads/2017/11/ENS-2016-17_PRIMEROS-RESULTADOS.pdf
44. Burrows R, Díaz E, Sciaraffia V, Gattas V, Montoya A, Lera L. Intake and physical activity habits among elementary and high school children according to type of school. *Rev Med Chil* 2008; 136: 53–63. [PubMed: 18483654]
45. Manyanga T, Tremblay M, Chaput J, Katzmarzyk P, Fogelholm M, Hu G, et al. Socioeconomic status and dietary patterns in children from around the world: different associations by levels of country human development? *BMC Public Health*. 2017; 17:457. doi: 10.1186/s12889-017-4383-8. [PubMed: 28511721]

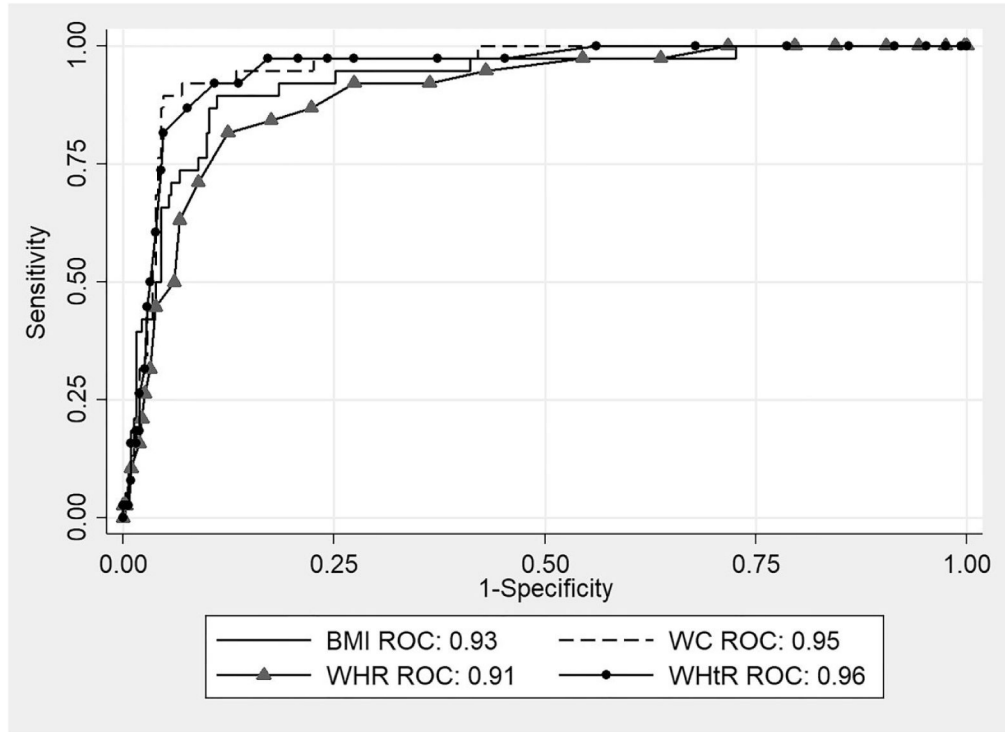


Figure 1. ROC curves of anthropometric indicators as predictors of metabolic syndrome in 16-year-old male adolescents. BMI: Body-Mass Index. WC: Waist circumference. WHR: Waist-to-Hip ratio. WHtR: Waist-to-Height ratio

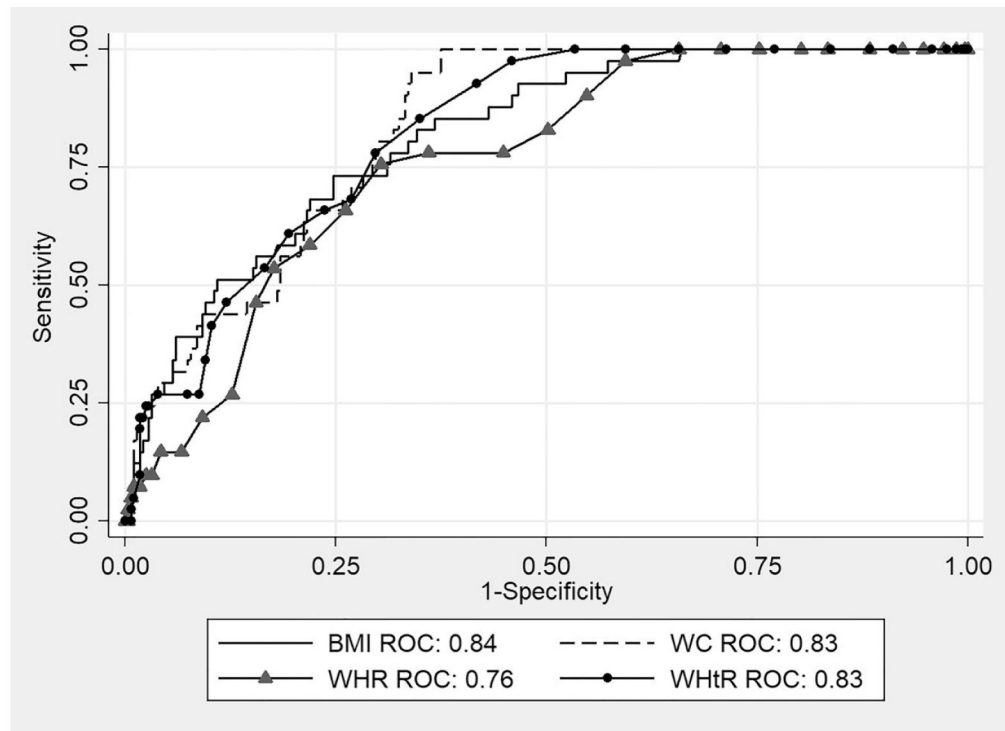


Figure 2. ROC curves of anthropometric indicators as predictors of metabolic syndrome in 16-year-old female adolescents. BMI: Body-Mass Index. WC: Waist circumference. WHR: Waist-to-Hip ratio. WHtR: Waist-to-Height ratio

Anthropometric characteristics and MetS related biomarkers in male and female adolescents in the sample (n=678)

Table 1.

Variables	Males (n=354)	Females (n=324)	P value
Age (years)	16.8(0.3)	16.8(0.4)	NS [§]
Weight (kg)	69.0±13.9	61.8±13.1	<0.001
Height (cm)	171.3±6.1	159.6±6.0	<0.001
Body-Mass Index (kg/m ²)	22.3(5.2)	23.4(5.7)	0.030 [§]
Body-Mass Index z score	0.5(1.6)	0.7(1.5)	NS [§]
Waist circumference (cm)	81.2±11.1	81.3±11.8	NS
Hip circumference (cm)	93.9±8.1	96.3±8.8	<0.001
Waist-to-height ratio	0.46(0.08)	0.50(0.09)	0.001 [§]
Waist-to-hip ratio	0.85(0.07)	0.84(0.09)	0.001 [§]
Systolic Blood Pressure (mm Hg)	115.4±10.4	108.9±9.8	<0.001
Diastolic Blood Pressure (mm Hg)	70.8±7.0	67.7±6.7	<0.001
Fasting glucose (mg/dl)	90.6±9.6	86.5±9.0	<0.001
Fasting insulin (uIU/ml)	6.0(5.1)	7.1(5.0)	0.001 [§]
HOMA-IR	1.3(1.1)	1.5(1.0)	NS [§]
Total cholesterol (mg/dl)	143(29.3)	153.9(32.8)	0.001 [§]
HDL cholesterol (mg/dl)	36.9(11.5)	41.7(14.9)	0.00H
Triglycerides (mg/dl)	71.5(45.5)	76.3(43.3)	NS [§]
Metabolic Syndrome (%)	8.5	8.8	NS [#]

Values are Mean±SD, Median(IQR), and relative frequencies. Two-tailed Student's t test for independent samples, except as indicated.

[§] Wilcoxon rank-sum test.

[#] χ^2 (Pearson).

Table 2. Optimal cutoff values of BMI, WC, WHR and WHtR to predict Metabolic Syndrome in male and female adolescents

	Cutoff	Sensitivity (%)	Specificity (%)	Correctly Classified	LR+	Post test probability	
						(+ test)	(- test)
MALES (n=354)							
Body mass index (BMI)	26.9	96.7	87.4	88.2	7.7	0.935	[35%,48%] [0%,2%]
Waist circumference (WC; cm)	92.0	92.1	92.9	92.9	18.7	0.950	[44%,64%] [0%,3%]
Waist-to-hip ratio (WHR)	0.91	86.7	87.8	87.6	7.0	0.911	[32%,48%] [1%,3%]
Waist-to-height ratio (WHtR)	0.54	93.3	92.6	92.7	12.5	0.960	[44%,64%] [0%,2%]
FEMALES (n=324)							
Body mass index (BMI)	26.3	78.6	78.4	78.6	3.6	0.840	[21%,32%] [1%,5%]
Waist circumference (WC; cm)	81.6	96.4	63.9	69.8	2.8	0.830	[18%,23%] [0%,4%]
Waist-to-hip ratio (WHR)	0.87	95.1	66.1	63.9	2.7	0.760	[18%,24%] [0%,3%]
Waist-to-height ratio (WHtR)	0.54	72.3	72.8	72.6	2.8	0.830	[16%,26%] [2%,6%]

Table 3. Cardiovascular and metabolic profile in males and females according to optimal cutoff values of BMI, WC, WHR and WHtR.

	BMI < 26.9		WC < 92.0		WC 92.0		WHR < 0.91		WHR 0.91		WHR < 0.54		WHR 0.54	
	(n=287)	(n=67)	(n=297)	(n=57)	(n=281)	(n=73)	(n=295)	(n=59)						
Systolic Blood Pressure (mm Hg)	110.7 ± 9.9	118.4 ± 10.9 ¹	113.9 ± 9.7	123.2 ± 10.7 ¹	111.0 ± 9.9	117.5 ± 11.7 ¹	113.7 ± 9.6	123.6 ± 10.3 ¹						
Diastolic Blood Pressure (mm Hg)	68.4 ± 6.8	72.6 ± 7.2 ¹	70.0 ± 6.8	74.6 ± 6.8 ¹	68.4 ± 6.7	72.8 ± 7.3 ¹	70.0 ± 6.6	74.7 ± 7.8 ¹						
Fasting Glucose (mg/dl)	88.1 ± 8.8	90.4 ± 11.6 ¹	90.3 ± 8.6	91.7 ± 13.5	88.5 ± 8.8	89.0 ± 11.9	90.3 ± 8.7	91.9 ± 13.2						
Fasting Insulin (uIU/ml)	6.1 (4.3)	9.9 (8.4) ²	5.6 (4.2)	11.6 (8.1) ²	6.3 (4.4)	9.5 (9.1) ²	5.6 (4.3)	10.2 (8.0) ²						
Total Cholesterol (mg/dl)	145.5 (30.0)	156.2 (40.9) ²	142.6 (27.9)	149.8 (43.0) ²	146.2 (30.9)	152.4 (41.9) ²	142.6 (26.7)	151.8 (46.3) ²						
HDL-cholesterol (mg/dl)	40.3 (14.4)	35.9 (11.3) ²	37.8 (11.8)	31.4 (10.4) ²	40.2 (14.3)	35.4 (12.5) ²	37.7 (12.2)	32.2 (11.2) ²						
Triglycerides (mg/dl)	70.8 (38.1)	98.9 (75.2) ²	67.7 (39.1)	103.8 (71.2) ²	71.4 (40.7)	91.1 (68.3) ²	67.6 (40.0)	103.3 (80.8) ²						
Metabolic Syndrome (%)	2.0	33.8 ³	1.4	59.6 ³	3.5	29.9 ³	0.7	48.3 ³						
Females (n=324)	BMI < 26.3	BMI 26.3	WC < 81.6	WC 81.6	WHR < 0.87	WHR 0.87	WHR < 0.54	WHR 0.54						
	(n=238)	(n=86)	(n=220)	(n=104)	(n=207)	(n=117)	(n=220)	(n=104)						
Systolic Blood Pressure (mm Hg)	110.7 ± 9.9	117.6 ± 11.2 ¹	105.6 ± 8.1	113.6 ± 10.1 ¹	110.1 ± 9.8	115.7 ± 10.9 ¹	106.8 ± 8.6	113.3 ± 10.6 ¹						
Diastolic Blood Pressure (mm Hg)	68.4 ± 6.8	72.2 ± 7.1 ¹	65.7 ± 5.9	70.5 ± 7.0 ¹	67.8 ± 6.5	71.6 ± 7.3 ¹	66.5 ± 6.3	70.2 ± 7.0 ¹						
Fasting Glucose (mg/dl)	88.2 ± 8.9	90.1 ± 11.2 ¹	85.9 ± 8.5	87.4 ± 9.6	88.5 ± 8.8	88.7 ± 10.5	86.4 ± 8.4	86.7 ± 10.1						
Fasting Insulin (uIU/ml)	6.0 (4.2)	9.7 (8.3) ²	6.8 (4.0)	8.6 (5.9) ²	6.1 (4.2)	8.0 (6.2) ²	6.9 (4.0)	8.7 (6.5) ²						
Total Cholesterol (mg/dl)	145.6 (30.4)	153.0 (39.4) ²	152.7 (31.2)	155.2 (36.7)	146.3 (30.1)	148.8 (34.9)	152.9 (32.3)	156.0 (36.6)						
HDL-cholesterol (mg/dl)	40.3 (14.5)	36.0 (11.3) ²	41.7 (17.1)	41.6 (10.7)	40.3 (14.0)	37.4 (13.9) ²	42.3 (17.3)	40.7 (11.2) ²						
Triglycerides (mg/dl)	70.8 (38.1)	97.0 (74.3) ²	72.6 (39.3)	83.3 (46.2) ²	68.4 (37.9)	86.5 (53.1) ²	73.3 (41.9)	83.3 (47.8) ²						
Metabolic Syndrome (%)	1.5	32.1 ³	1.6	28.4 ³	2.2	18.7 ³	3.6	20.2 ³						

¹ Student's t test.

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² Wilcoxon rank-sum test.

³ Chi2 test. BMI: Body-Mass Index. WC: Waist circumference (cm). WHR: Waist-to-Hip Ratio. WHHR: Waist-to-Height Ratio