

OR 1889

**Visceral obesity, skeletal muscle mass and resistin in metabolic syndrome development**

*Obesidad visceral, masa músculo-esquelética y resistina en el desarrollo de síndrome metabólico*

Carmen Paulina Rodríguez-López<sup>1</sup>, María Cristina González-Torres<sup>1</sup>, Ivette Cruz-Bautista<sup>2</sup> and Oralia Nájera-Medina<sup>3</sup>

<sup>1</sup>Departamento de Ciencias de la Salud, CBS. Universidad Autónoma Metropolitana-Iztapalapa. Ciudad de México, México. <sup>2</sup>Departamento de Endocrinología y Metabolismo. Instituto Nacional de Ciencias Médicas y Nutrición Salvador Zubirán. Ciudad de México, México. <sup>3</sup>Departamento de Atención a la Salud, CBS. Universidad Autónoma Metropolitana-Xochimilco. Ciudad de México, México

**Received:** 07/03/2018

**Accepted:** 06/07/2018

**Correspondence:** Oralia Nájera Medina. Universidad Autónoma Metropolitana-Xochimilco. Departamento de Atención a la Salud. Calzada del Hueso, 1100. Col. Villa Quietud A. P. 23-18. 04960 Coyoacán, México

e-mail: onajera@correo.xoc.uam.mx

DOI: 10.20960/nh.1889

**ABSTRACT**

**Background:** obesity implies an increase in the visceral adipose tissue (VAT), which is a risk factor for various metabolic diseases. VAT releases proinflammatory mediators, like resistin. In addition, it has been noted that the skeletal muscle mass (SMM) is involved in the development of metabolic syndrome (MS).

**Objective:** this study was designed to determine the relationship of body components (VAT and SMM) with MS and resistin in patients with obesity.

**Methods:** body composition and anthropometric and biochemical measurements to assess MS, and ELISA tests for resistin were carried out in 40 patients aged 18-40 years.

**Results:** overweight and obesity were observed in 72% of patients; visceral obesity was found in 53% and 35% had MS. A positive correlation between VAT and SMM in patients with MS was detected. In the entire population, an increase of 1 kg of SMM was found to be associated with an increase of 3 cm<sup>2</sup> of VAT, and an increase of 4 cm<sup>2</sup> of VAT was observed in individuals with MS. According to resistin, people with increased VAT had higher concentration than persons with normal VAT. Furthermore, an increase of 1 cm<sup>2</sup> of VAT accounted for a person entertaining a 3.3 fold greater risk of MS for different values of SMM and resistin.

**Conclusion:** the transcendence and significance of VAT as a main factor in triggering the chronic inflammatory process and MS, the SMM and resistin were also related.

**Key words:** Obesity. Visceral adipose tissue. Skeletal muscle mass. Metabolic syndrome. Resistin.

## RESUMEN

**Introducción:** la obesidad implica un aumento del tejido adiposo visceral (TAV), el cual es un factor de riesgo para varias enfermedades metabólicas. El VAT se relaciona con mediadores proinflamatorios, como la resistina. Además, se ha observado que la masa musculoesquelética (MME) interviene en el desarrollo del síndrome metabólico (SM).

**Objetivo:** este estudio fue diseñado para determinar la relación de la composición corporal (TAV y MME) con el SM y la resistina en pacientes con obesidad.

**Métodos:** se realizaron medidas antropométricas, de composición corporal y bioquímica para determinar el SM y prueba de ELISA para resistina en 40 pacientes de 18 a 40 años de edad.

**Resultados:** se observó sobrepeso y obesidad en el 72% de los participantes, obesidad visceral en el 53% y el 35% presentó SM. Se detectó una correlación positiva entre el TAV y la MME en pacientes con SM. En el grupo de estudio encontramos que un aumento de un 1 kg de MME se asociaba con un incremento de 3 cm<sup>2</sup> de TAV y en individuos con SM, con un incremento de 4 cm<sup>2</sup> de TAV. En relación a la resistina, las personas con TAV incrementado

presentan concentraciones más altas que las personas con TAV normal. Además, se observó que un aumento de 1 cm<sup>2</sup> de TAV representa un riesgo 3,3 veces mayor que para las personas de padecer SM para diferentes valores de MME y de resistina.

**Conclusión:** además de la trascendencia y la importancia del TAV como factor principal para desencadenar el proceso inflamatorio crónico y el SM, se observó que la MME y la resistina también están relacionados.

**Palabras clave:** Obesidad. Tejido adiposo visceral. Masa músculo-esquelética. Síndrome metabólico. Resistina.

## INTRODUCTION

Obesity is a growing public health problem at a national and global level. The most recent analysis (2010) conducted by the International Association of the Study of Obesity (IASO) and the International Obesity Task Force (IOTF) reported that one billion adults worldwide are overweight and 475 million are obese (1). In Mexico, 72% of the adult population is overweight or obese (2,3). This problem has been classified as a disease that is caused by multiple factors (physiological, psychological, metabolic, genetic, socioeconomic and cultural). However, the imbalance resulting from an increase in energy intake and diminished usage is the most common cause of the accumulation of adipose body tissue (4-6).

Obesity is not only an accumulation of energy in the form of triglycerides that increases body fat (both subcutaneous and visceral), but also this accumulation of energy in itself, and especially at the level of the visceral adipose tissue (VAT), renders obesity a risk factor for various metabolic diseases and disease progression and mortality (6,7).

VAT is the most bioactive component that is related with the release of inflammatory mediators (resistin, tumor necrosis factor alpha [TNF- $\alpha$ ], interleukin 1beta [IL-1 $\beta$ ], and interleukin 6 [IL-6]), which gives rise to insulin resistance (IR), mainly local, and subsequently in liver and skeletal muscle (8,9). Resistin is a proinflammatory adipocytokine that, in humans with obesity with MS, has been reported to express contradictory results, manifesting both increased and reduced levels of concentration in blood (10).

On the other hand, it has been noted that the skeletal muscle mass (SMM) could play a central role in the development of systemic IR in obesity by reducing the effect of insulin on this, contributing to the development of the metabolic syndrome (MS) (8,11-13).

According to the above, in the present study we decided to address the relationship of body components with the concentration of resistin as an inflammatory marker in patients with overweight and obesity.

## **MATERIALS AND METHODS**

### **Study population**

An observational, cross-sectional and clinical study was performed with undergraduate students and adult workers from the Metropolitan Autonomous University (UAM) of Xochimilco, Mexico City, Mexico. Patients' ages ranged between 18 and 40 years, of both genders, and all were submitted to anthropometric measurements, body composition, and biochemical tests. Exclusion criteria comprised metabolic disease, infections and autoimmune disease, cancer, heart disease, and those who were pregnant or taking medications. Participants were previously informed concerning the objectives of the study and were asked to sign a letter of informed consent. All of the procedures adhered to were reviewed and approved by the Ethics Committee of the Metropolitan Autonomous University of Xochimilco.

### **Anthropometric assessments**

The anthropometric measurements evaluated included weight, height and waist circumference (WC), following the standardized protocol of the International Society for the Advancement of Kinanthropometry (ISAK). A Seca® 213 stadiometer set at 0.1 cm precision was used for measurement of height. A Seca® Cursa Model 818 electronic scale was utilized, with an accuracy of 0.1 kg for measuring weight. WC was measured with a Seca® 201 fiberglass tape.

### **Definition of nutritional status**

The formula employed to calculate body mass index (BMI) was the following:  $BMI (kg/m^2) = \text{weight (kg)}/\text{height (m}^2\text{)}$ , for the first classification of the study participants' nutritional

status, according to World Health Organization (WHO) criteria for adults. Additionally, the waist-height index (WHI) was calculated ( $WHI = \text{waist circumference [cm]} / \text{height [cm]}$ ).

### **Body composition**

Analysis of body composition was evaluated by electrical bioimpedance with InBody720 equipment to obtain VAT in square meters, in which  $\geq 100 \text{ cm}^2$  of fat was diagnosed in persons with visceral obesity, in addition to the percentage of subcutaneous adipose tissue (ST) and kilograms (kg) of SMM.

### **Laboratory analysis and definition of metabolic syndrome**

For biochemical tests, the participants were fasted during the previous 12 hours. Automated Institute for Clinical Experimental Medicine (IKEM) clinical chemistry was utilized for the measurement of triglycerides (TG), high-density cholesterol (HDL-c), and glucose (Glu) from a sample of peripheral blood. Blood pressure measurement was performed according to the guidelines of Mexican Official Norm (NOM-030-SSA2-1999) for the prevention, treatment, and control of hypertension, twice in each patient (14).

The definition of the Cholesterol Education National Program (ATP III), modified for Hispanics, was taken into account for the diagnosis of MS. Accordingly, the presence of three or more of the following conditions is sufficient to diagnose MS: blood pressure  $\geq 130/85 \text{ mmHg}$ ; fasting glucose  $\geq 100 \text{ mg/dl}$ ; triglycerides  $\geq 150 \text{ mg/dl}$ ; HDL-c  $< 40 \text{ mg/dl}$  for men and women; and waist circumferences  $\geq 80 \text{ cm}$  for women and  $\geq 90 \text{ cm}$  for men.

### **Assay**

From each participant, a second blood sample was obtained and it was centrifuged at 1,500 rpm. Plasma was obtained and the resistin concentration was determined using a commercial Enzyme-Linked Immunosorbent Assay (ELISA) kit (Peprotech, Inc., Rocky Hill, NJ, USA).

### **Statistical analysis**

The Kolmogorov-Smirnov test was employed to explore the distribution of each variable. Logarithmic transformation was performed to approximate normality in variables exhibiting

a nonparametric distribution. The results were analyzed by calculating the mean  $\pm$  standard deviation (SD) and the median using confidence intervals (CI) for the values that were transformed. Comparison between two groups was performed using the Student's t-test. The one-way analysis of variance (ANOVA) and Bonferroni post-hoc test were applied to determine differences among > 2 groups. Correlation coefficients by performing step-wise forward linear and logistic regressions were obtained among variables utilizing the Pearson test. A p value of < 0.05 was considered as significant, employing SPSS statistical software program (version 21).

We do not have a large number of study subjects, however, it is enough to detect differences between increased and normal VAT according to resistin, with a power of 93.3%.

## **RESULTS**

### **General information on the study population**

The study-group population comprised 40 persons, with an average age of  $27.3 \pm 6.8$  years. Overall, 60% (n = 24) of participants were female and 40% (n = 16) were male.

### **Association between nutritional status and VAT with MS and body composition**

Normal BMI was observed in 28% (n = 11) of individuals, 35% demonstrated overweight (n = 14), and 37% had obesity (n = 15). On the other hand, 53% of participants had visceral obesity and 47% did not. It was observed that an increase in BMI was related with an increase in VAT; high VAT was observed in 9% of individuals with normal BMI, in 36% of people with overweight, and in 100% of participants with obesity.

An increase in anthropometric measurements (WC and WHI), body composition (ST, VAT, and SMM), biochemical indicators (Glu, TG) and systolic (SBP) and diastolic blood pressure (DBP) was observed in persons with obesity in relation to individuals with normal weight. HDL-c did not exhibit this behavior, because it decreased as BMI values increased. Regarding resistin as an inflammatory parameter, blood concentration demonstrated no statistical difference in relation to BMI (Table I).

When anthropometric measurements, body composition, and biochemical parameters were analyzed in relation to the presence of visceral obesity, it was found that in persons with normal VAT all variables were lower than in individuals with increased VAT. HDL-c

concentration was again the exception, because persons with increased VAT had lower values compared with those with normal VAT. No modifications were found in SBP and DPB in relation to VAT. According to blood levels of resistin, people with increased VAT had higher concentrations ( $405.2 \pm 56.6$ ,  $p < 0.022$ ) than persons with normal VAT ( $257.6 \pm 23.2$ ) (Table II).

### **Relation between metabolic syndrome, BMI and visceral obesity**

According to the ATP III criteria for Hispanics, 35% ( $n = 13$ ) of the individuals studied had MS, and 65% ( $n = 27$ ) did not. On the other hand, according to BMI, a higher percentage of individuals with MS had obesity (60%,  $p < 0.012$ ). Individuals without MS were mostly persons who demonstrated normal BMI (91%) (Fig. 1).

With respect to VAT, it was observed that a higher percentage (77%) of individuals with MS exhibited increased VAT and a lower proportion of normal VAT (23%,  $p < 0.031$ ). The higher percentage of adults without MS demonstrated normal VAT (59%) (data not shown).

### **Relation between MS, anthropometric assessments and body composition**

The relationship between MS diagnosis and anthropometric measurements and body composition was analyzed. This analysis revealed that MS is related to increases in all measurements; thus, individuals without MS had lowest values and persons with MS, highest values. Nevertheless, no changes were found in blood levels of resistin according to the MS (Table III).

Additionally, increased VAT was related to changes in anthropometric variables and body composition (WC, WHI, ST, and SMM) in patients with or without MS. All of these differences were statistically significant, except for SMM in persons who did not have MS, and for ST in persons who did present MS (Table IV).

When the resistin concentration was analyzed with respect to the presence or absence of MS and according to VAT, we observed that resistin showed a significant increase in individuals without MS and with increased VAT ( $p < 0.012$ ) as compared to those with normal VAT (Table IV).

### **Lineal regression between VAT and SMM considering metabolic syndrome**

Based on the aforementioned results, higher SMM was perceived in individuals with MS and increased VAT, compared with individuals with MS but normal VAT. We decided to conduct a correlation between VAT and SMM in general population, where we observed that increased VAT correlated with greater SMM. A positive correlation between VAT and SMM in patients with MS was also found; however, when these same parameters were analyzed in individuals without MS, no correlation was found (Fig. 2).

In addition, to observe the relationship between VAT and SMM, a linear regression in general population (n = 40) and in participants with MS (n = 12) was performed, and a statistical significance was observed in both regressions. Taking the entire population into account, we perceived that an increase of 1 kg of SMM was associated with an increase of 3 (range, 1.1-5) cm<sup>2</sup> of VAT (p < 0.002) and, in persons with MS, an increase of 1 kg of SMM was associated with an increase in VAT of 4 (range, 1-7) cm<sup>2</sup> (data not shown).

#### **Logistic regression between MS and SMM with VAT and resistin**

A logistic regression was also estimated to establish the relationship between MS (dependent variable) and SMM, and VAT and resistin (independent variables). We observed, in a first model, that a 1-cm<sup>2</sup> increase of VAT was related with higher odds (2.7, range: 0.4-5) for acquiring MS (p < 0.023) for different values of SMM. In a second model, VAT, SMM and resistin were analyzed, and the results showed that a 1-cm<sup>2</sup> increase of VAT contributed higher odds (3.3, range: 0.7-5.9) for developing MS (p < 0.013) for different values of SMM and resistin (data not shown).

When examining this adipocytokine in terms of anthropometric measurements in individuals with and without MS, we found that the blood concentration of resistin in individuals without MS had a statistically significant positive correlation with WHI. Additionally, resistin had a statistically significant negative correlation with SMM in patients with MS (Fig. 3).

#### **DISCUSSION**

Obesity is the main risk factor for developing comorbidities such as dyslipidemia, hyperglycemia, hypertension and IR, among others, as well as presenting anthropometric



and body composition changes, mainly VAT distribution. The latter is categorized as the trigger factor of the aforementioned diseases (15-17).

According to our results, we observed that, as BMI and VAT increased, anthropometric measurements, body composition, and biochemical indicators were altered. It was also noteworthy that persons with obesity and increased VAT had the highest proportions of MS. These data were, to a certain degree, expected, as it has been reported that obesity involves increased VAT, negatively affecting biochemical, anthropometric, and body composition indicators, which are all-important for diagnosing a person with MS. This demonstrated an association between visceral obesity and the presence of MS (17).

In the present investigation, important findings were obtained in which an increase in SMM is related with increased BMI and with the presence of increased VAT. When SMM was analyzed according to the presence or absence of MS and, in turn, it was stratified according to normal and increased VAT, it was observed that individuals with MS and increased VAT demonstrated more kg of SMM, this indicating that increased VAT is an important factor for the presence of SMM in persons with MS.

Some authors support the notion that not only abdominal obesity plays an important role in the development of MS and that, in general terms, obesity may perhaps not be expressed in terms of BMI, but that SMM also comprises an important component (11). SMM is the main contributor to glucose uptake (about 75%) from insulin (18), displaying the importance of this tissue in the presence of IR and subsequent morbidities. In this regard, it has been proposed that the proportion of skeletal muscle could possess an important role in the progression of MS (19).

In the population of the present study, we found that a 1-kg increase of SMM is associated with an increase of 3 (range, 1.1-5) cm<sup>2</sup> of VAT, and that in individuals with MS, a 1-kg increase of SMM is associated with an increase of 4 (range, 1-7) cm<sup>2</sup> of VAT. This demonstrated the close relationship between SMM and VAT and the development of MS, considering SMM as a factor that goes hand-in-hand with VAT for the development of MS. In this respect, it has been observed that inflammatory mediators such as the TNF- $\alpha$ , IL-1 $\beta$ , and IL-6 that are released by VAT, and other cells, as well as free fatty acids, contribute to local IR, but can escape from the circulation and contribute to systemic IR through reducing the insulin signal in various organs such as skeletal muscle (12,19,20). The latter is damage

that precedes the development of MS, and these two tissues could represent a risk factor for metabolic diseases.

It was mentioned that muscle is a site of cytokine release in obesity, also called miokines (13,20). In a proteomic *in vitro* study in muscle fibers (C2C12 myotubes) in which IR was induced, it was found that the muscle is capable of releasing about 1,073 putative proteins (32 growth factors, 25 cytokines, and 29 metalloproteinases), of which approximately 40% were regulated under conditions of IR (13).

In addition, in this study, we decided to study the blood concentration of resistin. A tendency toward increased concentrations related to the BMI was detected, but this was significantly elevated in persons with increased VAT. Data analysis, according to the presence or absence of MS and stratified with normal and increased VAT, revealed a significant increase in resistin concentration in individuals without MS and increased VAT compared to those without MS and normal VAT. Resistin is a proinflammatory cytokine associated with inflammatory markers (TNF- $\alpha$  and IL-6) that affect metabolism (21-27). Several authors have suggested that increased levels of serum resistin are linked with visceral fat, IR, inflammatory markers (TNF- $\alpha$  and IL-6), myocardial infarction, and atherosclerosis. These authors suggest that resistin levels could serve as a surrogate marker for metabolic diseases in humans (22,27,28). However, in the present study this was associated in individuals with VAT and increased VAT without MS. This could mean that in our patients, resistin should be taken as a predictor of metabolic damage. Further studies are needed for monitoring and to confirm its role.

In addition, a significant positive correlation between WHI and resistin in individuals without MS was observed. Some authors have linked this index with cardiovascular and metabolic risk (29-31). It is probable that this finding could also be a predictor of metabolic disorders and cardiovascular disease.

Some studies have suggested a proinflammatory role of resistin, and a positive relationship has been found between blood concentrations of resistin with body weight, fat mass, blood lipids, as well as inflammation markers, both in obese patients and in obese morbid patients (32,33). However, there are others who do not relate this adipocytokine with the presence of metabolic disorders, or obesity, or IR, or the MS (25,26,34-36). In the present study, a high concentration of resistin in persons without MS, and a positive correlation with

increased VAT and WHI were found. This, in a certain manner, confirms the previous results of several studies, indicating the presence of controversy among levels of resistin in obesity, IR and MS (21,26,34,37).

## **CONCLUSION**

With these results, we can state that increased VAT in patients with obesity is probably the most important factor for the development MS, moreover highlighting the importance of SMM in the presence of MS.

We also observed an association between resistin and increased VAT, SMM and MS. These results support the notion that resistin might be acting as an inflammatory adipocytokine, contributing to the increase in frequency of MS in those persons who present increased levels of this cytokine.

However, this finding emphasizes the need to conduct further studies related to the presence of SMM, resistin, and other inflammation markers in a greater number of individuals with obesity in order to corroborate the results presented herein.

## **ACKNOWLEDGEMENTS**

We would like to thank Ms. María Magdalena Rodríguez-Magallanes, at the Unit of Nutrition, Body Composition and Energy Expenditure of the UAM-Xochimilco, for facilitating the installation and the equipment to perform the measurements and the biochemical analysis, as well as CONACYT for the scholarship awarded to MBE Carmen Paulina Rodríguez-López (302016).

## **ETHICS APPROVAL**

Ethical approval for the study was obtained from the Committee of the Metropolitan Autonomous University-Xochimilco. Written informed consent was obtained from each participant and archived.

## **REFERENCES**

1. Patel P, Abate N. Body fat distribution and insulin resistance. *Nutrients* 2013;5(6):2019-27.

2. Gutiérrez J, Rivera-Dommarco J, Shamah-Levy T, Villalpando-Hernández S, Franco A, Cuevas-Nasu L, et al. Encuesta Nacional de Salud y Nutrición 2012. Resultados nacionales. México: Instituto Nacional de Salud Pública; 2012.
3. Secretaría de Salud del Distrito Federal (SSDF). Sobrepeso y obesidad. 2014. Accessed on Jan 1<sup>st</sup>, 2016. Available from: [http://www.salud.df.gob.mx/ssdf/index.php?option=com\\_content&task=view&id=4034](http://www.salud.df.gob.mx/ssdf/index.php?option=com_content&task=view&id=4034)
4. Haro M, Antonio M, Ponce y Ponce G, Andrés Núñez A, Ruiz-Esparza J, Soria C, et al. Hipertrofia ventricular en el paciente con obesidad. *RESPYN* 2012;13(1):1-8.
5. Lee MJ, Wu Y, Fried SK. Adipose tissue heterogeneity: implication of depot differences in adipose tissue for obesity complications. *Mol Aspects Med* 2013;34(1):1-11.
6. Sarvottam K, Yadav RK. Obesity-related inflammation and cardiovascular disease: efficacy of a yoga-based lifestyle intervention. *Indian J Med Res* 2014;139(6):822-34.
7. Trachta P, Drapalova J, Kavalkova P, Touskova V, Cinkajzlova A, Lacinova Z, et al. Three months of regular aerobic exercise in patients with obesity improve systemic subclinical inflammation without major influence on blood pressure and endocrine production of subcutaneous fat. *Physiol Res* 2014;63(Suppl 2):S299-308.
8. Esser N, Legrand-Poels S, Piette J, Scheen AJ, Paquot N. Inflammation as a link between obesity, metabolic syndrome and type 2 diabetes. *Diabetes Res Clin Pract* 2014;105(2):141-50.
9. Marette A, Liu Y, Sweeney G. Skeletal muscle glucose metabolism and inflammation in the development of the metabolic syndrome. *Rev Endocr Metab Disord* 2014;15(4):299-305.
10. Procaccini C, De Rosa V, Galgani M, Corbone F, La Rocca C, Formisano L, et al. Role of adipokines signaling in the modulation of T cells function. *Front Immunol* 2013;4(332):1-12.
11. Namwongprom S, Rerkasem K, Wongthanee A, Pruenglampoo S, Mangklabruks A. Relationship between body composition parameters and metabolic syndrome in young thai adults. *J Clin Res Pediatr Endocrinol* 2014;6(4):227-32.
12. Winer DA, Winer S, Chng MH, Shen L, Engleman EG. B lymphocytes in obesity-related adipose tissue inflammation and insulin resistance. *Cell Mol Life Sci* 2014;71(6):1033-43.

13. Deshmukh AS, Cox J, Jensen LJ, Meissner F, Mann M. Secretome analysis of lipid-induced insulin resistance in skeletal muscle cells by a combined experimental and bioinformatics workflow. *J Proteome Res* 2015;14(11):4885-95.
14. Norma Oficial Mexicana NOM-030-SSA2-2009, para la prevención, detección, diagnóstico, tratamiento y control de la hipertensión arterial 2009. Accessed on Nov 20<sup>th</sup>, 2015. Available from: [http://dof.gob.mx/nota\\_detalle.php?codigo=5144642&fecha=31/05/2010](http://dof.gob.mx/nota_detalle.php?codigo=5144642&fecha=31/05/2010)
15. Ryder E, Díez-Ewald M, Mosquera J, Fernández E, Pedreanez A, Vargas R, et al. Association of obesity with leukocyte count in obese individuals without metabolic syndrome. *Diabetes Metab Syndr* 2014;8(4):197-204.
16. López-López J, López-Jaramillo P, Camacho P, Gómez-Arbelaez D, Cohen D. The link between fetal programming, inflammation, muscular strength and blood pressure. *Mediators Inflamm* 2015;2015:1-8.
17. Maurya SK, Periasamy M. Sarcolipin is a novel regulator of muscle metabolism and obesity. *Pharmacol Res* 2015;102:270-5.
18. Funai K, Lodhi IJ, Spears LD, Yin L, Song H, Klein S, et al. Skeletal muscle phospholipid metabolism regulates insulin sensitivity and contractile function. *Diabetes* 2015;65:358-70.
19. Park BS, Yoon JS. Relative skeletal muscle mass is associated with development of metabolic syndrome. *Diabetes Metab J* 2013;37(6):458-64.
20. Gregor MF, Hotamisligil GS. Inflammatory mechanisms in obesity. *Annu Rev Immunol* 2011;29:415-45.
21. Al-Suhaimi EA, Shehzad A. Leptin, resistin and visfatin: the missing link between endocrine metabolic disorders and immunity. *Eur J Med Res* 2013;18:12.
22. DeFuria J, Belkina AC, Jagannathan-Bogdan M, Snyder-Cappione J, Carr JD, Nersesova YR, et al. B cells promote inflammation in obesity and type 2 diabetes through regulation of T-cell function and an inflammatory cytokine profile. *Proc Natl Acad Sci USA* 2013;110(13):5133-8.
23. Carvalho AF, Rocha DQ, McIntyre RS, Mesquita LM, Kohler CA, Hyphantis TN, et al. Adipokines as emerging depression biomarkers: a systematic review and meta-analysis. *J Psychiatr Res* 2014;59:28-37.

24. Giby VG, Ajith TA. Role of adipokines and peroxisome proliferator-activated receptors in nonalcoholic fatty liver disease. *World J Hepatol* 2014;6(8):570-9.
25. Hsieh YY, Shen CH, Huang WS, Chin CC, Kuo YH, Hsieh MC, et al. Resistin-induced stromal cell-derived factor-1 expression through Toll-like receptor 4 and activation of p38 MAPK/ NFkappaB signaling pathway in gastric cancer cells. *J Biomed Sci* 2014;21:59.
26. Barnes MA, Carson MJ, Nair MG. Non-traditional cytokines: how catecholamines and adipokines influence macrophages in immunity, metabolism and the central nervous system. *Cytokine* 2015;72(2):210-9.
27. Muse ED, Feldman DI, Blaha MJ, Dardari ZA, Blumenthal RS, Budoff MJ, et al. The association of resistin with cardiovascular disease in the Multi-Ethnic Study of Atherosclerosis. *Atherosclerosis* 2015;239(1):101-8.
28. Cao H. Adipocytokines in obesity and metabolic disease. *J Endocrinol* 2014;220(2):T47-59.
29. Moreira M. Which of the anthropometric measures to define excess weight is the best discriminator of cardiovascular risk? *Med Clin (Barc)* 2010;134(9):396-8.
30. Remón P, González S, Arpa G. Waist-to-height ratio as a fat accumulation variable to assess cardiovascular risk. *Rev Cub Med Mil* 2013;42(4):444-50.
31. Hernández J, Duchi P. Waist-to-height ratio and its usefulness in detection of the cardiovascular and metabolic risk. *Rev Cubana Endocrinol* 2015;26(1):66-76.
32. De Luis D, González Sagrado M, Conde R, Aller R, Izaola O. Resistin levels and inflammatory markers in patients with morbid obesity. *Nutr Hosp* 2010;25(4):630-4.
33. De Luis D, González M, Conde R, Aller R, Izaola O, Primo D. Lack of association of serum resistin levels with metabolic syndrome criteria in obese female patients. *Clin Biochem* 2011;44:1280-3.
34. Zulet A, Puchau B, Navarro C, Martí A, Martínez J. Biomarcadores del estado inflamatorio: nexo de unión con la obesidad y complicaciones asociadas. *Nutr Hosp* 2007;22(5):511-27.
35. Gómez R, Conde J, Lago F, Gualillo O. Las adipocinas: mediadores emergentes de la respuesta inmune y de la inflamación. *Reumatol Clin* 2009;05(51):6-12.
36. Codoñer-Franch P, Alonso-Iglesias E. Resistin: insulin resistance to malignancy. *Clin Chim Acta* 2015;438(1):46-54.

37. Samsam-Shariat SZ, Bolhasani M, Sarrafzadegan N, Najafi S, Asgary S. Relationship between blood peroxidases activity and visfatin levels in metabolic syndrome patients. *ARYA Atheroscler* 2014;10(4):218-26.



**Table I. Distribution of the anthropometric variables, body composition, biochemical parameters and resistin of the study groups according to BMI**

<i>Variable</i> ( <i>n</i> = 40)	<i>Normal</i> ( <i>n</i> = 11)	<i>Overweight</i> ( <i>n</i> = 14)	<i>Obesity</i> ( <i>n</i> = 15)	<i>p</i>	<i>p (post hoc)</i>
WC (cm)	81.7 (79.5-86)	90.1 (82.8-99.8)*	107 (102-114)* <sup>†</sup>	0.000	0.000
WHI	0.49 (0.47-0.51)	0.56 (0.53-0.58)*	0.64 (0.61-0.70)* <sup>†</sup>	0.000	0.000
ST (%)	29 ± 6.9	33.7 ± 7	41.7 ± 6.1* <sup>†</sup>	0.000	0.000
VAT (cm <sup>2</sup> )	78.9 (40-84.5)	90.3 (76.6-110.3)*	152.8 (131-168.1)* <sup>†</sup>	0.000	0.000
SMM (kg)	24 ± 5.2	26.8 ± 7.5	30.5 ± 5.3*	0.035	0.033
TG (mg/dl)	137.2 (92-170)	113.2 (48-166.2)	205 (123-221)	0.058	
HDL-c (mg/dl)	59.2 ± 13.2	47.2 ± 18.5	36.7 ± 15.1*	0.004	0.003
Glu (mg/dl)	87.5 ± 8.6	92 ± 6.6	97.8 ± 13.9	0.053	
SBP (mmHg)	110 (100-120)	120 (110-120)	120 (120-130)*	0.009	0.008
DBP (mmHg)	70 (60-80)	80 (70-80)	80 (70-90)*	0.021	0.028
Resistin (pg/ml)	240.0 ± 27.5	345.8 ± 46.2	393.3 ± 73.9	0.165	

BMI: body mass index; WC: waist circumference; WHI: waist-height index; ST: subcutaneous adipose tissue; VAT: visceral adipose tissue; SMM: skeletal muscle mass; TG: triglycerides; HDL-c: high-density lipoprotein cholesterol; Glu: glucose; SBP: systolic blood pressure; DBP: diastolic blood pressure. p (post hoc): p value adjusted with Bonferroni test. Data are presented in media ± SD or median and CI. \*Statistically significant difference vs normal. <sup>†</sup>Statistically significant difference vs overweight.



**Table II. Characteristics of the anthropometric media, body composition, biochemical indicators and resistin of the groups according to visceral obesity**

<i>Variable</i> ( <i>n</i> = 40)	<i>Normal VAT</i> ( <i>n</i> = 19)	<i>Increased VAT</i> ( <i>n</i> = 21)	<i>p</i>
WC (cm)	83 (80-88.5)	103.4 (100-111.2)*	0.000
WHI	0.53 (0.49-0.55)	0.63 (0.59-0.69)*	0.000
ST (%)	30.7 ± 5.9	39.6 ± 8.1*	0.000
SMM (kg)	22.6 (21.3-27.2)	31 (24.8-35.3)*	0.003
TG (mg/dl)	128.5 (56-170)	178 (101-194.5)	0.175
c-HDL (mg/dl)	56.4 ± 17.3	37.7 ± 13.7*	0.001
Glu (mg/dl)	89.1 ± 7.9	96.5 ± 12.2*	0.031
SBP (mmHg)	110 (110-120)	120 (110-130)	0.068
DBP (mmHg)	80 (70-80)	80 (70-90)	0.213
Resistin (pg/ml)	257.6 ± 23.2	405.2 ± 56.6*	0.022

WC: waist circumference; WHI: waist-height index; ST: subcutaneous adipose tissue; VAT: visceral adipose tissue; SMM: skeletal muscle mass; TG: triglycerides; HDL-c: high-density lipoprotein cholesterol, Glu: glucose; SBP: systolic blood pressure; DBP: diastolic blood pressure. p (post hoc): p value adjusted with Bonferroni test. Data are presented in media ± SD or median CI. \*Statistically significant difference vs normal VAT (p < 0.05).

**Table III. Characteristics of anthropometric media, body composition and resistin in the study groups according to metabolic syndrome**

<i>Variable</i> ( <i>n</i> = 40)	<i>Without MS</i> ( <i>n</i> = 27)	<i>With MS</i> ( <i>n</i> = 13)	<i>p</i>
WC (cm)	86.5 (82-101)	103.5 (94.2-113.7)*	0.002
WHI	0.54 (0.50-0.62)	0.60 (0.57-0.68)*	0.005
ST (%)	34 ± 8.8	38.3 ± 6.8	0.130
VAT (cm <sup>2</sup> )	88.5 (73.1-124.1)	150.3 (90.3-165.1)*	0.001
SMM (kg)	26.1 ± 6.5	30.2 ± 6	0.059
Resistin (pg/ml)	342.6 ± 47	308 ± 26.5	0.528

MS: metabolic syndrome; WC: waist circumference; WHI: waist-height index; ST: subcutaneous tissue; VAT: visceral adipose tissue; SMM: skeletal muscle mass. Data are presented in media ± SD or median and CI. \*Statistically significant difference ( $p < 0.05$ ).

**Nutrición  
Hospitalaria**

**Table IV. Distribution of anthropometric variables, body composition and resistin in presence or absence of metabolic syndrome and the normal and increased VAT in the study groups**

Variable (%) (n = 40)	Without MS (n = 27)			MS (n = 13)		
	N VAT (n = 15)	I VAT (n = 12)	p	N VAT (n = 4)	I VAT (n = 9)	p
WC (cm)	82.5 (79.8-84)	101.5 (93.7-107.2)*	0.000	90.6 (85.7-95.4)	109 (103-115.5)*	0.000
WHI	0.51 (0.48-0.54)	0.62 (0.55-0.67)*	0.000	0.56 (0.53-0.57)	0.65 (0.60-0.70)*	0.001
ST (%)	29.6 ± 5.8	39.5 ± 8.8*	0.002	34.9 ± 4.6	39.8 ± 7.2	0.251
SMM (kg)	24.3 ± 4.8	28.3 ± 7.8	0.118	24.1 ± 5	32.9 ± 4.3*	0.008
Resistin (pg/ml)	226.9 ± 22.5	493 ± 84.2*	0.012	357.5 ± 36.1	280 ± 33.3	0.172

MS: metabolic syndrome; N VAT: normal visceral adipose tissue; I VAT: increased visceral adipose tissue; WC: waist circumference; WHI: waist-height index; ST: subcutaneous tissue; SMM: skeletal muscle mass. Values are presented in media ± SD or median and CI.

\*Statistically significant difference among normal vs increased VAT p < 0.05.

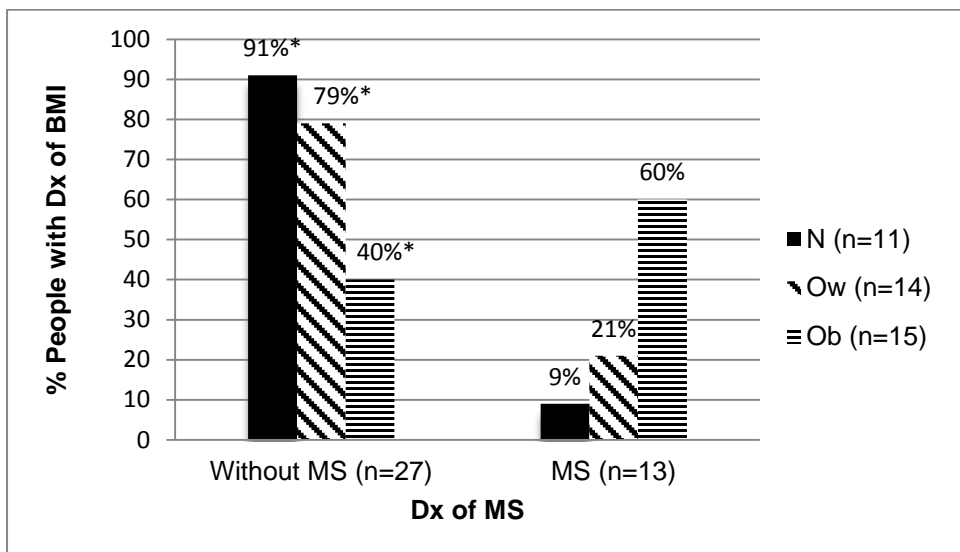


Fig. 1. Metabolic syndrome in relation to BMI. Dx of BMI: diagnosis according to body mass index; Dx of MS: diagnosis according to metabolic syndrome; N: normal; Ow: overweight; Ob: obesity. \*Statistical difference between presence and absence of MS according to BMI.  $p < 0.012$ .

**Nutrición  
Hospitalaria**

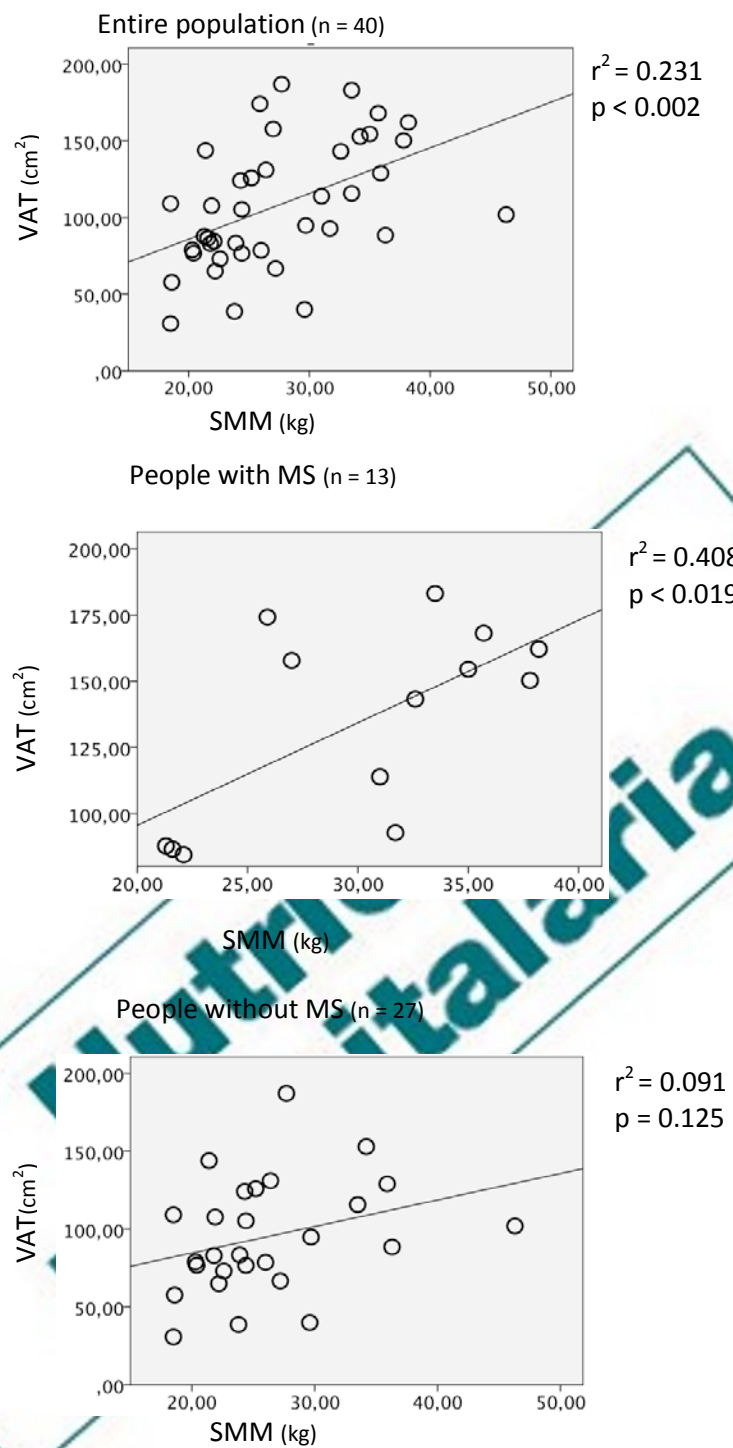


Fig. 2. Correlation between VAT and SMM in the study groups. VAT: visceral adipose tissue; MS: metabolic syndrome; SMM: skeletal muscle mass.

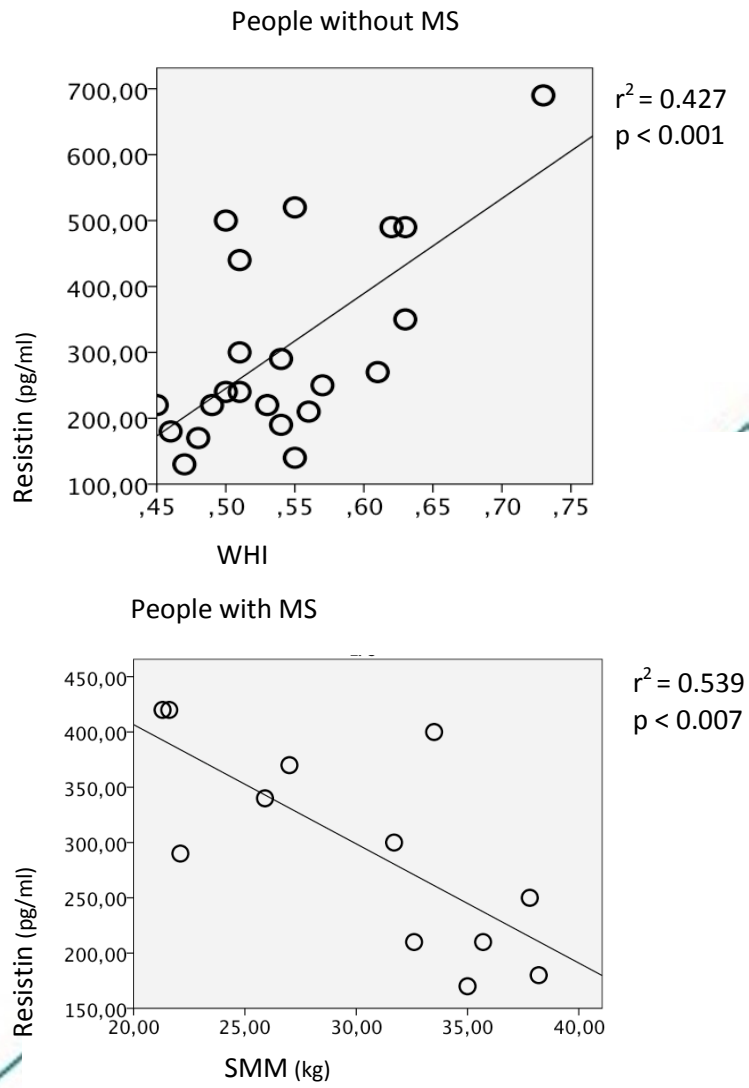


Fig. 3. Correlation of resistin with WHI and SMM. WHI: waist-height index; SMM: skeletal muscle mass.