### **ORIGINAL RESEARCH**

industrial workers

# Evaluation of serum adipokines with the development of metabolic syndrome among

Ravi Prakash Jamalpur, Vinay Kumar Adepu, Surender Jakkam, Ravibabu Kalahasthi Regional Occupational Health Centre Southern, Karnataka, India

#### ABSTRACT

**Background:** Most of the studies were reported serum adipokines with the presence of metabolic syndrome (MetS) in population-based studies.

**Objective:** The present study assessed the levels of serum adipokines, i.e., leptin, adiponectin, leptin:adiponectin ratio (LAR), and adiponectin:leptin ratio (ALR), and the development of MetS among industrial workers.

**Methods:** Eighty-eight workers were recruited in the study. MetS was diagnosed using the definition of the International Diabetes Federation. Serum leptin and adiponectin levels were done using enzyme linked immunosorbent assay methods. LAR and ALR values were calculated using the serum leptin and adiponectin. The data was analyzed using statistical package for the social sciences software. Mann–Whitney U test was used to find out the differences of adipokines among workers with or without MetS. Kruskal–Wallis test was used to find out the effect of the number of metabolic components (0, 1, 2, 3, and >4) on serum adipokines among workers.

**Results:** 37.5% of workers had MetS. The levels of serum leptin (p < 0.01) and LAR (p < 0.01) were significantly increased and serum adiponectin (p = 0.104) and ALR (p < 0.01) were decreased in workers with the presence of MetS. The effect of the number of metabolic components was demonstrated a significant increase of serum leptin (p = 0.018) and LAR (p = 0.026) and significant decrease of ALR (p = 0.046).

**Conclusion:** The levels of serum leptin and LAR were increased and adiponectin and ALR were decreased in workers with the presence of MetS and the number of metabolic components. The data of the present study support the role of adipokines and development of MetS. It could be used as specific indicators for the diagnosis and management of MetS.

#### **ARTICLE HISTORY**

Received September 04, 2019 Accepted December 27, 2019 Published February 29, 2020

#### **KEYWORDS**

Metabolic syndrome; industrial workers; leptin; adiponectin; leptin:adiponectin ratio; adiponectin:leptin ratios

#### Introduction

Metabolic syndrome (MetS) is the risk of obesity, reduced high-density lipoprotein-cholesterol (HDL-C), hypertriglyceridemia, hypertension, and elevated glucose. It is characterized by proinflammatory, prothrombotic state, and atherogenic dyslipidemia [1]. It is a global public health problem and linked with a three-fold risk of type 2 diabetes mellitus and a two-fold risk of cardiovascular disease (CVD) [2]. The potential health problems such as insulin resistance, obesity, and CVD were positively associated with the presence of MetS [3]. Industrial workers with privileged access to healthcare facilities typically reported a high prevalence of MetS [4]. Physical inactivity, worse lifestyle factors, unhealthy dietary intake, and overweight were positively related to the risk of MetS [5]. A high prevalence of MetS was reported in industrial workers from Poland [6] and adults from Jordan [7]. A significant number of the metabolic component score was reported in gas refinery workers as compared to non-industrial workers [8]. Work places of manufacturing, transport, finance, and cooperative association reported a high prevalence of MetS [9]. The self-reporting health complaints were positively associated with the risk of MetS [10]. The cost of medical treatment is 3.6 times higher for those with MetS and disease [11]. Employees with



**∂** Open Access

Contact Ravibabu Kalahasthi 🖾 kalahasthi20012002@yahoo.co.in 🗔 Regional Occupational Health Centre Southern, Karnataka, India.

abdominal obesity [12], high workload [13], short duration of sleep and shift [14], and high job stress [15] are the potential predictors of MetS.

Adipose tissue is an endocrine gland, which produces different kinds of biologically active compounds, namely  $-\alpha$  Tumor necrosis factor (TNF- $\alpha$ ), IL-6, leptin, adiponectin, and retinal binding protein-4 [16]. Adipokines, in particular, leptin and adiponectin modulate the MetS [17]. Adiponectin regulates the metabolism of glucose and lipids and has the property of antiatherogenic and anti-inflammatory. It is served as an eradicating marker of MetS [18]. A negative tendency was noted between adiponectin and the presence of MetS [19]. Subjects with the existence of MetS observed lower levels of adiponectin [20]. Leptin regulates the appetite, carbohydrate, lipid, energy metabolism, arterial pressure, neuroendocrine, and etiopathogenesis of MetS [21]. Leptin was positively correlated with MetS [22]. The existence of MetS in older adults has noted higher leptin and lower adiponectin [23,24]. Jung et al. [25] reported a significant relationship between ALR and the presence of MetS and the number of metabolic components in the population. A recent study also established that the ALR was a better marker for the assessment of acute metabolic stress [26]. LAR was served as a sensitive marker for the early detection of MetS [27,28]. LAR is a better predictor for MetS as compared to alone measurements of adiponectin and leptin [29].

Most of the studies were reported serum adipokines and the presence of Mets in population-based studies. However, the interactions between adipokines (adiponectin, leptin, LAR, and ALR) and the presence of MetS among industrial workers were not explored. This study was undertaken to find out the interaction between adipokines and the presence of MetS among industrial workers.

# **Material and methods**

This study enrolled eighty-eight industrial workers from establishments located in Karnataka, India. The study subjects were enrolled in the study by visiting the industry. These subjects were chosen from the industries which manufactured bearings components and food flavors and fragrances. Subjects with a history of diabetes, thyroid, pituitary, hypogonadism, chronic liver, or chronic renal diseases were excluded from the study to avoid the influences of adipokine secretions on metabolic syndrome among industrial workers. The Institutional Ethics Committee has approved the study with protocol number: 142 dated 13-12-2018. A diagnosis of MetS among industrial workers was done using the definition of the International Diabetes Federation (IDF). Serum adiponectin and leptin concentration among subjects were analyzed using enzyme linked immunosorbent assay (ELISA) methods.

# Clinical examination

Clinical examination and lifestyle factors data were collected from industrial workers using a pre-validated questionnaire, which was validated using a small sample population from the center. Subjects' height in centimeters and weight in kilograms were measured using measuring tape and weighing machine with lightly clothed and without shoes. For waist circumference (WC), we measured in accordance with the World Health Organization diagnostic criteria for abdominal obesity. Blood pressure [systolic blood pressure (SBP) and diastolic blood pressure (DBP)] was monitored in subjects using the HEM-7112 monitor after resting for 4 minutes while sitting.

## **Blood collection**

Four milliliters of whole blood was collected into Vacuurate easy clot activator tubes (manufactured by Labtech Disposables, India) and centrifuged at 4,000 rpm for 10 minutes at 4°C to separate the serum and used for the measurement of fasting serum glucose, triglyceride, and HDL-C among subjects.

## Fasting serum glucose (FSG)

The FSG estimation was performed using glucose oxidase and peroxidase (GOD-POD), an endpoint kit method developed by Trinder [30]. In this method, the GOD enzyme converts the glucose into gluconate. The hydrogen peroxide is produced in the reaction, is degraded by POD, and gives a colored product phenol and 4-aminoantipyrine, which is measured using the Trinder indicator reaction at 505 nm. The absorbance increase was correlated with the glucose concentration of the sample. The limit of detection (LOD) of the method is 1 mg/dl.

## Serum triglycerides

The level of serum triglyceride estimation was done using enzymatic (glycerol 3-phosphate oxidase peroxidase) kit method developed by Fossati et al. [31] and coupled the reaction of classical Trinder [30]. In this approach, triglycerides are hydrolyzed by lipase into glycerol and fatty acids. Glycerol is phosphorylated by ATP in the presence of glycerol kinase to glycerol-3-phosphate, which is oxidized by the glycerol-3-phosphate oxidase and produces hydrogen peroxide. It reacts with 4-aminoantipyrine and 4-chlorophenol in the presence of POD to produce red quinoneimine. The intensity of color developed is proportional to the triglyceride concentration. The LOD of the method is 1 mg/dl.

#### Serum HDL-cholesterol

The levels of serum HDL-C were measured using phosphotungstic acid and magnesium ion precipitate method as described by Lopes et al. [32]. In this approach, chylomicrons, very low density lipoprotein, and low density lipoprotein are precipitated by phosphotungstic acid and magnesium ions. After centrifugation, HDLs are present in the supernatant. Cholesterol is measured by an enzymatic kit method. In which, the determination of cholesterol after enzymatic hydrolysis and oxidation in the above, the colorimetric indicator quinoneimine is formed, which is generated from 4-aminoantipyrine and phenol by hydrogen peroxide under the catalytic action of POD. This is measured using Trinder indicator reaction at 510 nm. The increase in absorbance correlates with the HDL-C concentration of the sample. The LOD of the method is 4 mg/dl.

## Serum adiponectin

The levels of serum adiponectin were measured using the sandwich ELISA kit method (Diagnostics Biochem Canada Inc., CAN-APN-5000). The LOD of the method is 0.06 ng/ml and detection range is 0-50 ng/ml.

### Serum leptin

The level of serum leptin was measured using sandwich ELISA kit method (Diagnostics Biochem Canada Inc., CAN-L-4260). The LOD of the method is 0.50 ng/ml and detection range is 0–100 ng/ml.

## Leptin:adiponectin ratio (LAR)

It was a calculated value from serum leptin and serum adiponectin of individual subjects.

### Adiponectin:leptin ratio (ALR)

It was a calculated value from serum adiponectin and serum leptin of individual subjects.

### Statistical analysis

The Statistical package for the social sciences version 20 was used for data analysis. Shapiro–Wilk test was used to find the normality of serum adipokine levels (adiponectin, leptin, LAR, and ALR) among industrial workers. The distribution of serum adipokine levels was found in non-normal (skewed). The data of serum adipokines (adiponectin, leptin, LAR, and ALR) were presented in median levels and compared using the Mann–Whitney U test in industrial workers with and without MetS. Kruskal–Wallis test was used to find the effect of the number of metabolic components (0, 1, 2, 3, and >4) on serum adipokines (adiponectin, leptin, LAR, and ALR) among industrial workers. Probability of less than 0.05 was considered as significant.

### Result

Table 1 represents the criteria for MetS using the IDF that is WC plus the presence of any two of the risk factors such as elevated triglyceride, low HDL-C, increased blood pressure(SBP or DBP) and elevated levels of fasting glucose. The baseline data of the industrial workers are presented in Table 2. Individuals with a mean and standard deviation of age, height, weight, BMI, WC, blood pressure (SBP and DBP), and lipid profile (serum triglycerides and HDL-C); serum fasting glucose; and adipokines (leptin, adiponectin, LAR, and ALR) were reported.

Table 3 shows the median levels of serum adipokines (adiponectin, leptin, LAR, and ALR) among industrial workers with and without MetS. The non-parametric test such as the Mann–Whitney U test was used to find the differences of adipokine levels among industrial workers with and without MetS. The levels of serum leptin (p < 0.01) and LAR (p < 0.01) were significantly increased and ALR (p < 0.01) was significantly decreased in industrial workers with the presence of MetS. The level of serum adiponectin was decreased (p = 0.104) in workers with MetS, but it was not significantly altered.

Table 1.	IDF	criterion	for	MetS
----------	-----	-----------	-----	------

Parameters	Defining level
Serum triglycerides	≥150 mg/dl
Serum HDL-C	<40 mg/dl in males <50 mg/dl in females
Blood pressure (SBP or DBP)	Systolic BP ≥ 130 or Diastolic BP ≥ 85 mm Hg
Serum fasting glucose	≥ 100 mg/dl
Waist circumference	Male ≥ 90 cm Female ≥ 80 cm

Diagnosis of MetS: WC plus any two of four parameters.

Table 2	<b>Baseline</b>	data o	f the	industrial	workers
	Daseinie	υαια υ	i uic	muustiiai	WUIKEIS.

Parameters	Mean ± SEM (n = 88)
Age (years)	33.65 ± 1.0
Height (cm)	162.76 ± 0.9
Weight (kg)	63.50 ± 1.1
BMI (kg/m²)	$24.03 \pm 0.4$
WC (Cm)	86.04 ± 1.0
SBP (mm Hg)	125.22 ± 1.8
DBP (mm Hg)	74.84 ± 1.2
Glucose (mg/dl)	93.91 ± 3.5
Triglyceride (TG) (mg/dl)	167.42 ± 9.9
HDL-C (mg/dl)	34.51 ± 0.8
Leptin (ng/ml)	$1.30 \pm 0.05$
Adiponectin (µg/ml)	$4.76 \pm 0.4$
Leptin:adiponectin ratio (LAR)	0.52 ± 0.2
Adiponectin:leptin ratio (ALR)	$6.01 \pm 0.8$

**Table 3.** Median levels of serum adipokine among workerswith and without MetS.

(MetS)	n = 88	Leptin (ng/ml)	Adiponectin (µg/ml)	LAR	ALR
(-)	55	1.01	4.51	0.19	5.02
(+)	33	1.49**	3.16	0.48**	2.27**

\*\*p < 0.01.

The prevalence of MetS among industrial workers is presented in Figure 1. The prevalence of MetS among industrial workers was assessed using the definition of IDF, i.e., WC plus any two of the following four factors: increased triglycerides, reduced HDL-C, increased blood pressure (SBP or DBP), and elevated levels of fasting glucose. In this study, it was noted that 37.5% of workers had MetS and 62.5% of workers without MetS.

Figure 2 shows the effect of metabolic components on serum adipokines (adiponectin, leptin, LAR, and ALR) among industrial workers. Kruskal–Wallis test was used to find the effect of the number of metabolic components (0, 1, 2, 3, and >4) on serum adipokines (adiponectin, leptin, LAR, and ALR) among industrial workers. The results of the study shown that the serum leptin (p = 0.018) and LAR (p = 0.026) were significantly increased and ALR (P=0.046) was significantly decreased with the increase of the number of metabolic components.

### Discussion

This study assessed the interaction between serum adipokines and the presence of MetS among indus-



**Figure 1.** Prevalence of metabolic syndrome among industrial workers.



**Figure 2.** Effect of the number of metabolic components on serum adipokines among workers.

trial workers. Nair et al. [33] reported the presence of MetS ranged from 11.11% to 30.7% as per the International Standard Classification of Occupations (ISCO-88); the highest prevalence of MetS was reported in managers and senior officials. Sarang et al. [34] identified the MetS ranged from 9.4% to 52.9% in mining workers. During the present study, it was noted that 37.5% of workers had the MetS. A similar prevalence of MetS was noted in adults from Jordan [7] and gas refinery workers [8]. A study from India reported a high prevalence of MetS among industrial workers, who have good access to health care [4]. Review of the literature indicated that the presence of MetS was associated with unhealthy diet patterns, worse lifestyle factors, and low physical activity [35].

MetS was accompanied by increased visceral tissue and adipokine secretion [36]. An inverse association was noted between adiponectin levels and the presence of MetS in population-based studies [37,38]. During the present study, it was observed a similar trend between adiponectin levels and the presence of Mets among industrial workers. The measurement of adiponectin was considered as a therapeutic marker for MetS [39]. Leptin is a hormone derived from adipose tissue and reported a positive correlation with the MetS in population-based studies [40,41]. In this study, we also noted a similar tendency between serum leptin and the presence of MetS among industrial workers. The existence of MetS in older adults reported higher leptin and lower adiponectin levels [23,24]. In the present study, it was observed higher leptin and lower adiponectin levels with the existence of MetS among workers. Leptin was considered as a predictive marker of MetS [42].

LAR was associated with low-grade inflammation [43] and atherosclerotic risk [44]. It is used as a predictive marker for MetS as compared to individual measurements of adiponectin and leptin [45]. Population-based studies have reported a positive association between LAR and the presence of MetS [46,47]. The findings of this study found a similar relationship between LAR and the presence of MetS among workers.

ALR is a marker of adipose tissue inflammation [48]. It is a useful estimator for obesity-associated cardiometabolic risk [49]. Population studies have found the negative link between ALR and the presence of MetS [50,51]. The findings of this study also indicated a decreased ALR with the presence of MetS. Wang et al. [40] reported a positive tendency between leptin and number of metabolic components. This study also evaluated the effect of the number of metabolic components (0, 1, 2, 3, and >4) on serum adipokines (leptin, adiponectin, LAR, and ALR) among industrial workers. The findings of this study were indicated significantly higher serum leptin and LAR concentration with the presence of MetS. Santamiema et al. [52] reported a negative inclination between adiponectin levels with the components of MetS defined by IDF. Jung et al. [16] reported a negative trend between ALR and the presence of MetS and the number of metabolic components in Korean subjects. In this study, it was noted a similar trend between adiponectin and ALR concentrations with the number of MetS components.

### Conclusion

The levels of serum leptin and LAR were significantly increased and ALR were significantly decreased in workers with the presence of MetS and the number of metabolic components. The data of this study support the role of adipokines and development of MetS. It could be used as a specific indicator for the diagnosis and management of MetS.

#### References

- [1] Grundy SM, Clemens JI, Daniels SR, Donato KA, Eckel RH, Franklin B, et al. Diagnosis and management of the metabolic syndrome: an American Heart Association/National Heart, Lung, and Blood Institute scientific statement. Circulation 2005; 112:2735–52.
- [2] Zimmet P, Magliano D, Matsuzawa Y, Alberti G, Shaw J. The metabolic syndrome: a global public health problem and a new definition. J Atheroscler Thromb 2005; 12(6):295–300.
- [3] Rochlani Y, Pothineni NV, Kovelamudi S, Mehta JL. Metabolic syndrome: pathophysiology, management, and modulation by natural compounds. Ther Adv Cardiovasc Dis 2017; 11(8):215–25.
- [4] Mini GK, Sarma PS, Thankappan KR. Overweight, the major determinant of metabolic syndrome among industrial workers in Kerala, India: Results of a cross-sectional study. Diabetes Metab Syndr 2019; 13(5):3025–30.
- [5] Strasser B. Physical activity in obesity and metabolic syndrome. Ann N Y Acad Sci 2013; 1281:141–59.
- [6] Kwella N, Szczubelek M, Zielecki P, Harazny J, Stompor T. Prevalence of physical activity and metabolic syndrome in population of industrial workers in Olsztyn, Poland. J Hypertension 2017; 35:e257.
- [7] Obeidat AA, Ahmad MN, Haddad FH, Azzeh FS. Alarming high prevalence of metabolic syndrome among Jordanian adults. Pak J Med Sci 2015; 31(6):1377–82.
- [8] Baghshini MR, Nikbakht-Jam I, Mohaddes-Ardabili H, Pasdar A, Avan A, Tayefi M, et al. Higher prevalence of metabolic syndrome among male employees of a gas refinery than in their counterparts in nonindustrial environments. Asian Biomed 2017; 11:227–34.
- [9] Hidaka T, Hayakawa T, Kakamu T, Kumagai T, Hiruta Y, Hata J, et al. Prevalence of metabolic syndrome and its components among Japanese workers by clustered business category. PLoS One 2016; 11(4):e0153368.
- [10] Liu Y, Ozodiegwu ID, Nickel JC, Wang K, Iwasaki LR. Self-reported health and behavioral factors are associated with metabolic syndrome in Americans aged 40 and over. Prev Med Rep 2017; 7:193–7.
- [11] Schultz AB, Edington DW. Analysis of the association between metabolic syndrome and disease in a workplace population over time. Value Health. 2010; 13(2):258–64.
- [12] Ayogu RN, Nwajuaku C, Udenta EA. Components and risk factors of metabolic syndrome among rural Nigerian workers. Niger Med J 2019; 60:53–61.
- [13] Nisa MA, Martiana T, Wahyudiono YD. Analysis of relation between life style, workload, and work stress with metabolic syndrome. Indian J Public Health Res Dev 2018; 9(5):53–8.

- [14] Itani O, Kaneita Y, Tokiya M, Jike M, Murata A, Nakagome S, et al. Short sleep duration, shift work, and actual days taken off work are predictive life-style risk factors for new-onset metabolic syndrome: a 7-year cohort study of 40,000 male workers, Sleep Med 2017. doi:10.1016/j.sleep.2017.07.027
- [15] Kang MG, Koh SB, Cha BS, Park JK, Woo JM, Chang SJ. Association between job stress on heart rate variability and metabolic syndrome in shipyard male workers. Yonsei Med J 2004; 45:838–46.
- [16] Jung U, Choi MS. Obesity and its metabolic complications: the role of adipokines and the relationship between obesity, inflammation, insulin resistance, dyslipidemia and nonalcoholic fatty liver disease. Int J Mol Sci 2014; 15:6184–223.
- [17] Francisco V, Ruiz-Fernández C, Pino J, Mera A, Gonzalez-Gay MA, Gómez R, et al. Adipokines: linking metabolic syndrome, the immune system, and arthritic diseases. Biochem Pharmacol 2019; 165:196–206.
- [18] Adhikar P, Nagtilak S, Parashar P. Association of adiponectin with components of metabolic syndrome in Western U.P. Int J Med Sci Innov Res 2018; 6(3):215–21.
- [19] Von Frankenberg AD, do Nascimento FV, Gatelli LE, Nedel BL, Garcia SP, de Oliveira CS, et al. Major components of metabolic syndrome and adiponectin levels: a cross-sectional study. Diabetol Metab Syndr 2014; 6:26.
- [20] Ntzouvani A, Fragopoulou E, Panagiotakos D, Pitsavos C, Antonopoulou S. Reduced circulating adiponectin levels are associated with the metabolic syndrome independently of obesity, lipid indices and serum insulin levels: a cross-sectional study. Lipids Health Dis 2016; 15:140.
- [21] Li R, Chen LZ, Zhao SP, Huang XS. Inflammation activation contributes to adipokines imbalance in patients with acute coronary syndrome. PLoS One 2016; 11:e0151916.
- [22] Tseng PW, Wu DA, Hou JS, Hsu BG. Leptin is an independent marker of metabolic syndrome in elderly adults with type 2 diabetes. Tzu Chi Med J 2017; 29:109.
- [23] You T, Nicklas BJ, Ding J, Penninx BW, Goodpaster BH, Bauer DC, et al. The metabolic syndrome is associated with circulating adipokines in older adults across a wide range of adiposity. J Gerontol Ser A Biol Sci Med Sci 2008; 63(4):414–9.
- [24] Stenholm S, Koster A, Alley DE, Visser M, Maggio M, Harris TB, et al. Adipocytokines and the metabolic syndrome among older persons with and without obesity: the InCHIANTI study. Clin Endocrinol 2010; 73(1):55–65.
- [25] Jung CH, Rhee EJ, Choi JH, Bae JC, Yoo SH, Kim WJ, et al. The relationship of adiponectin/leptin ratio with homeostasis model assessment insulin resistance index and metabolic syndrome in apparently

healthy Korean male adults. Korean Diabetes J 2010; 34:237–43.

- [26] Elshaari FA, Alshaari AA, Sara A, Hassan SA, Sheriff DS. Adiponectin/leptin ratio as a biomarker of acute metabolic stress. Int J Biol Med Res 2013; 4(3):3278–83.
- [27] Larsen MA, Isaksen VT, Moen OS, Wilsgaard L, Remijn M, Paulssen EJ, et al. Leptin to adiponectin ratio—a surrogate biomarker for early detection of metabolic disturbances in obesity. Nutr Metab Cardiovasc Dis 2018; 28(11):1114–21.
- [28] Ayina CN, Endomba FT, Mandengue SH, Noubiap JJ, Ngoa LS, Boudou P, et al. Association of the leptin-to-adiponectin ratio with metabolic syndrome in a sub-Saharan African population. Diabetol Metab Syndr 2017;9:66.
- [29] Chen VC, Chen CH, Chiu YH, Lin TY, Li FC, Lu ML. Leptin/adiponectin ratio as a potential biomarker for metabolic syndrome in patients with schizophrenia. Psychoneuroendocrinology 2018; 92:34–40.
- [30] Trinder P. Determination of blood glucose using an oxidase-peroxidase system with a non-carcinogenic chromogen. J Clin Pathol 1969; 22:158–61.
- [31] Fossati P, Prencipe L. Serum triglycerides determined colorimetrically with an enzyme that produces hydrogen peroxide. Clin Chem 1982; 28:2077–80.
- [32] Lopes-Virella MF, Stone P, Ellis S, Colwell JA. Cholesterol determination in high-density lipoproteins separated by three different methods. Clin Chem 1977; 23:882–4.
- [33] Nair CV. Metabolic syndrome: an occupational perspective. Indian J Community Med 2010; 35(1):122–4.
- [34] Sarang VD, Subroto SN, Umesh LD. Metabolic syndrome in different sub occupations among mine workers. Indian J Occup Environ Med 2015; 19:76.
- [35] Huang JH, Li RH, Huang SL, Sia HK, Chen YL, Tang FC. Lifestyle factors and metabolic syndrome among workers: the role of interactions between smoking and alcohol to nutrition and exercise. Int J Environ Res Public Health 2015; 12(12):15967–78.
- [36] Zhang Y, Yang P, Cui R, Zhang M, Li H, Qian C, et al., Eosinophils reduce chronic inflammation in adipose tissue by secreting Th2 cytokines and promoting M2 macrophages polarization. Int J Endocrinol 2015; 2015:565760. doi:10.1155/2015/565760
- [37] Sparrenberger K, Sbaraini M, Cureau FV, Teló GH, Bahia L, Schaan BD. Higher adiponectin concentrations are associated with reduced metabolic syndrome risk independently of weight status in Brazilian adolescents. Diabetol Metab Syndr 2019; 11(1):40.
- [38] Kim JY, Ahn SV, Yoon JH, Koh SB, Yoon J, Yoo BS, et al. Prospective study of serum adiponectin and incident metabolic syndrome: the ARIRANG study. Diabetes Care 2013; 36(6):1547–53.

- [39] Ghadge AA, Khaire AA, Kuvalekar AA. Adiponectin: a potential therapeutic target for metabolic syndrome. Cytokine Growth Factor Rev 2018; 39:151–8.
- [40] Wang LH, Liu YC, Wang JH, Lee CJ, Hsu BG. Serum leptin level positively correlates with metabolic syndrome among elderly Taiwanese. Tzu-Chi Med J 2017; 29(3):159.
- [41] García-Jiménez S, Bernal Fernandez G, Martínez Salazar MF, Monroy Noyola A, Toledano Jaimes C, Meneses Acosta A, et al. Serum leptin is associated with metabolic syndrome in obese Mexican subjects. J Clin Lab Anal 2015; 29(1):5–9.
- [42] Ghadge AA, Khaire AA. Leptin as a predictive marker for metabolic syndrome. Cytokine. 2019; 121:154735.
- [43] Chou HH, Hsu LA, Wu S, Teng MS, Sun YC, Ko YL. Leptin-to-adiponectin ratio is related to low grade inflammation and insulin resistance independent of obesity in non-diabetic Taiwanese: a cross-sectional cohort study. Acta Cardiol Sin 2014; 30(3):204.
- [44] Selthofer-Relatic K, Radic R, Stupin A, Sisljagic V, Bosnjak I, Bulj N, et al. Leptin/adiponectin ratio in overweight patients-gender differences. DiabVasc Dis Res 2018; 15(3):260–2.
- [45] Li G, Xu L, Zhao Y, Li L, Fu J, Zhang Q, et al. Leptin-adiponectin imbalance as a marker of metabolic syndrome among Chinese children and adolescents: the BCAMS study. PLoS One 2017; 12(10):e0186222.

- [46] Konsoulova PS, Nyagolova PV, Orbetzova MM, Simitchiev KK, Terzieva DD, Kaleva NN. Leptin, adiponectin, and leptin/adiponectin ratio in adolescents with metabolic syndrome. Int J Pharm Med Res 2016; (5):1–6.
- [47] Gupta V, Mishra S, Mishra S, Kumar S, Gupta V. Association of leptin: adiponectin ratio and metabolic risk markers in postmenopausal women. Immunol Lett 2018; 196:63–7.
- [48] Fruhbeck G, Catalan V, Rodriguez A, Ramírez B, Becerril S, Salvador J, et al. Adiponectin-leptin ratio is a functional biomarker of adipose tissue inflammation. Nutrients 2019; 11(2):454.
- [49] Frühbeck G, Catalán V, Rodríguez A, Gómez-Ambrosi J. Adiponectin-leptin ratio: a promising index to estimate adipose tissue dysfunction. Relation with obesity-associated cardiometabolic risk. Adipocyte 2018; 7(1):57–62.
- [50] Mirza S, Qu HQ, Li Q, Martinez PJ, Rentfro AR, McCormick JB, et al. Adiponectin/leptin ratio and metabolic syndrome in a Mexican American population. Clin Invest Med 2011; 34(5):E290.
- [51] Ko SH. The adiponectin/leptin ratio and metabolic syndrome in healthy korean adult males. Korean Diabetes J 2010; 34(4):220–1.
- [52] Santaniemi M, Kesäniemi YA, Ukkola O. Low plasma adiponectin concentration is an indicator of the metabolic syndrome. Eur J Endocrinol 2006; 155(5):745–50.