

# The Relevance of Inflammatory Markers in Metabolic Syndrome

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## ABSTRACT

**Objectives:** To identify how severe is inflammation in metabolic syndrome using as inflammatory markers: C-reactive protein and leukocytes. To assess these markers considering the diversity of metabolic syndrome elements.

**Material and method:** We performed a study that enrolled 258 patients registered to a family physician and diagnosed with metabolic syndrome. The subjects included in the study were divided in two groups: group A-137 subjects diagnosed with metabolic syndrome that was defined by 3 elements: abdominal obesity+arterial hypertension+diabetes mellitus; group B-121 patients diagnosed with metabolic syndrome based on 5 elements: abdominal obesity+arterial hypertension+diabetes mellitus+decreased high density lipoprotein cholesterol (HDL-C)+increased triglycerides.

**Results:** We observed increased values of CRP and leukocytes for group B in comparison to group A:  $0.9 \pm 0.8$  mg/dl vs  $0.79 \pm 0.8$  mg/dl ( $p=0.02$ , significantly statistic). Leukocytes value was higher for group B, but not significantly statistic.

**Conclusions:** Inflammation in patients with metabolic syndrome depends on the number and association of elements that define this entity and it is more accentuated for subjects who associate more elements.

**Keywords:** metabolic syndrome, inflammation, markers

## INTRODUCTION

**M**etabolic syndrome (MetS) is an entity that represents a global health problem; in our country there is an increased prevalence of metabolic syndrome (1,2). It is a multiplex risk factor that arises from insulin resistance accompanying abnormal adipose tissue (3-8). Clinical manifestations of the syndrome may include: abdominal obesity, hy-

per-tension, diabetes mellitus, hypertriglyceridemia, reduced high density lipoprotein cholesterol. According to International Diabetes Federation (IDF) and the National Heart, Lung and Blood Institute metabolic syndrome is diagnosed when a patient has at least 3 of the following 5 conditions (2005): obesity – waist circumference  $\geq 94$  cm in men and  $\geq 80$  cm in women (European population); blood pressure  $\geq 130/85$  mmHg (or receiving drug therapy

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for hypertension); fasting glucose  $\geq 100$  mg/dl (or undergoing therapy); tryglicerides  $\geq 150$  mg/dl (or receiving treatment); HDL – C  $< 40$  mg/dl in men and  $< 50$  mg/dl in women (or receiving drug therapy) (9-13). Abundent data suggests that patients meeting these diagnostic criteria have a greater risk for significant clinical consequences: doubled risk of coronary artery disease, increased risk of stroke, fatty liver disease, diabetes and cancer (14-17). □

**MATERIALS AND METHODS**

The study included 250 people with metabolic syndrome enrolled on the lists of family physician. The group of study included patients from both urban and rural areas. The study was carried on from 2007 to 2009. Patients agreement for participation to the study was obtained. Subjects admitted in the study were divided in two groups: first group consists of 137 subjects diagnosed with metabolic syndrome that was defined by 3 elements: abdominal obesity+arterial hypertension + hyperglycemia; the second group consists of 121 patients diagnosed with metabolic syndrome based on 5 elements: abdominal obesity +arterialhypertension+hyperglycemia+decreased high density lipoprotein+increased triglycerides. There was an even distribution of patients in terms of age, obesity degree and sex in the study groups. We assesed inflammatory status for patients from the two study groups using as inflammatory markers: C-reactive protein and leukocytes. We considered as normal values for leukocytes: 5000-10000/ $\mu$ l. C-reactive protein was evaluated through a quantitative technique and we considered as normal values: 0.1-0.8 mg/dl. Inflammatory markers values obtained were compared for the two study groups. Patients diagnosed with other diseases beside metabolic syndrome were excluded from the study as there was a possibility to interfere with the results obtained for inflammatory markers: chronic pulmonary disease, chronic kidney failure, records of neoplasm or actual neoplasia, pulmonary microembolism (diagnosed by various procedures: chest X-ray, spiral computerized tomography scan, ultrasound, magnetic

resonance imaging), chronic focal infections, recent acute infections, colagenous diseases, nonsteroidian antiinflammatory therapy or cortisone therapy, surgery in the past six months, acute myocardial infarction or stroke.

Statistics used Epilnfo 6 (ANOVA). A value of  $p < 0.05$  was considered with a statistical significance.

Our study complies with Ethical Principles for Medical Research Involving Human Subjects stated by the Declaration of Helsinki. □

**RESULTS**

Females predominate in our study, 55.5% (142) vs. 45.5% (116) male. The average age is higher in women, 61.5 + / -10 versus men 55.1 + / - 11.5 years, with a statistically significant difference ( $p = 0.03$ ).

In the second group C-reactive protein is higher than in the first group with statistical significance ( $p = 0.02$ ).

Leukocytes have a less important value in establishing proinflammatory and cardiovascular risk contribution in patients with metabolic syndrome compared with C-reactive protein. □

**DISCUSSION**

In the present conditions that obesity is increasing at global level metabolic syndrome will always be in the top of medical problems. Our study is very actual as metabolic syndrome is very important for current medical practice due to a progressive increasing frequency and atherogenic risk. Metabolic syndrome may affect most of the population and it may generate both vascular and metabolic complications (1-2, 18-19). The severity of inflammation in the metabolic syndrome measured by determining C-reactive protein and leukocytes is influenced by the number of criteria that make up metabolic syndrome.

Pro-inflammatory mechanisms can be considered as a base of increased cardiovascular risk. Proinflammatory activity is more significant if metabolic syndrome is characterised by more elements (group B is defined by 5 elements and group A by 3 elements). The results we obtained ascertain that inflammatory status is increased in patients diagnosed with metabolic syndrome (significantly statistic in subjects that associate more than 3 elements). Inflammatory injury has different severity depending

	CRP (mg/dl)	Leukocytes (/ $\mu$ l)
First group	0.79 $\pm$ 0.8	12600 $\pm$ 1000
Second group	0.9 $\pm$ 0.8	14100 $\pm$ 1000
p	0.02	0.07

TABLE 1. Leukocytes and CPR values in the two groups.

on the elements that define metabolic syndrome and on their association. Once the inflammation level increases there is a differentiated prognostic impact for cardiovascular events (20-22).

Metabolic syndrome frequency is progressively increasing and evaluation of proinflammatory risk of this entity is valuable, as assessment of some inflammatory biomarkers implies minimum costs and it can be repeated (23). In our study CRP proved to be an accurate indicator of inflammation for patients with metabolic syndrome. In subjects with acute coronary syndrome, stroke, periferic vascular disease and sudden death, recent epidemiological data ascertained a positive association between CRP levels and clinic manifestations of atherothrombosis. Increased values of CRP represent a predictive marker for unfavourable evolution in patients with unstable angina pectoris after myocardial revascularisation, as well as in patients with metabolic syndrome and diabetes – that suggests its role in atherogenesis (24-25).

Leukocytes increase more evidently in acute vascular complications that may occur in these subjects. They seem to be less valuable for chronic inflammatory character (26-30). In our study leukocytes value, even if it was increased for group B in comparison to group A, was not significantly statistic. □

## CONCLUSIONS

Obesity is the central factor of the metabolic syndrome. Patients diagnosed with metabolic syndrome present an activated inflammatory status. Inflammatory syndrome is expressed according to the number of metabolic syndrome components. Bioumoral components of metabolic syndrome have a higher proinflammatory contribution. Proatherogenic impact in those with metabolic syndrome is uneven and increased.

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