

## **Prevention and nutritional therapy of metabolic syndrome**

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### **ABSTRACT**

The term metabolic syndrome (MetS) defines the co-occurrence of the related risk factors of metabolic origin that promote the development of cardiovascular diseases with atherosclerotic background and type 2 diabetes. The diagnostic criteria of MetS have undergone modifications for years. Until now no clear definition of MetS has been established. The latest diagnostic criteria of MetS published in 2009 by a group of IDF (International Diabetes Federation) and AHA/NHLBI (American Heart Association/ National Heart, Lung and Blood Institute) experts discern three out of five risk factors: abdominal

obesity (taking into consideration population differences), elevated level of triglycerides, reduced HDL cholesterol, hypertension and fasting hyperglycemia. Genetic predispositions and environmental factors, such as lack of physical activity and improper diet are considered to be responsible for MetS development. Therefore, prevention and treatment of MetS should be based first of all on a change in modifiable lifestyle factors, among which proper diet is of essential importance.

**Keywords:** metabolic syndrome, obesity, blood pressure, dyslipidemia, hyperglycemia

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## DEFINITION AND DIAGNOSTIC CRITERIA OF METABOLIC SYNDROME

In medicine, reports on the so called X syndrome – a metabolic syndrome (MetS) appeared already in the middle of the 20th century. However, it is Gerald Reaven who was the first to define in 1988 the main MetS components to be insulin resistance, hyperinsulinemia, impaired glucose tolerance, hypertension and dyslipidemia [1].

In 1998, WHO (World Health Organization) issued a definition [2], according to which MetS can be identified only when one of the following risk factors occurs:

- Diabetes mellitus type 2 (fasting glucose  $\geq 126$  mg/dl or glycaemia two hours after glucose load  $\geq 200$  mg/dl);
- Impaired glucose tolerance (IGT) (fasting glucose 110-125 mg/dl and glycaemia 2 hours after glucose load  $\geq 140$  mg/dl);
- Impaired fasting glucose (IFG) (fasting glucose 110-125 mg/dl and glycaemia 2 hours after glucose load  $\geq 140$  mg/dl);
- Insulin resistance (IR) (glucose uptake below the lowest quartile for background population investigated under hyperinsulinemic-euglycaemic conditions).
- Two out of four risk factors were considered an additional criterion:
- Abdominal obesity (waist-hip ratio WHR  $\geq 0.9$  for men and  $\geq 0.85$  for women and/or body mass index BMI  $\geq 30$  kg/m<sup>2</sup>);
- Dyslipidaemia (triglycerides  $\geq 150$  mg/dl and/or HDL cholesterol  $< 35$  mg/dl for men and  $< 39$  mg/dl for women);
- Arterial blood pressure ( $\geq 140/90$  mm Hg and/or controlled hypertension);
- Microalbuminuria (urinary albumin excretion  $\geq 20$   $\mu$ g/min or albumin/creatinine ratio  $\geq 30$  mg/g).

WHO definition took into account MetS pathogenesis, i.e. insulin resistance. However, MetS diagnosis should be simple and commonly available, whereas WHO criteria required specialist diagnostic examinations, thus limiting their application.

The European Group for the Study of Insulin Resistance (EGIR) [3] modified the WHO criteria; however, the indispensable condition was still insulin resistance (plasma insulin  $> 75$  percentile).

Additional criteria (2 out of 4) included:

- Abdominal obesity ( $\geq 94$  cm for men and  $\geq 80$  cm for women);
- Dyslipidaemia (triglycerides  $\geq 150$  mg/dl and/or HDL cholesterol  $< 39$  mg/dl for men and women);

- Arterial blood pressure ( $\geq 140/90$  mm Hg and/or controlled hypertension);
- Fasting glycaemia ( $\geq 110$  mg/dl, IGF, IGT, but not diabetes).

The modification of MetS diagnostic criteria was issued in 2001 in the Third Report of National Program of Cholesterol Education NCEP-ATP III (National Cholesterol Education Program, Adult Treatment Panel III) [4,5]. These new criteria differed in the lack of insulin resistance as an indicator of MetS, which made them more suitable for every day clinical practice. In the NCEP-ATP III definition, all the risk factors of MetS were equivalent and to diagnose MetS three out of five factors had to be met:

- Abdominal obesity ( $\geq 102$  cm for men and  $\geq 88$  cm for women);
- Level of triglycerides ( $\geq 150$  mg/dl);
- Level of HDL cholesterol ( $< 40$  mg/dl for men and  $< 50$  mg/dl for women);
- Arterial blood pressure ( $\geq 130/85$  mm Hg);
- Fasting glucose ( $\geq 110$  mg/dl or diabetes).

However, NCEP-ATP III criteria in the diagnosis of MetS also had some limitations. The threshold values of abdominal obesity in the NCEP-ATP III criteria were established on the basis of the National Institute of Health as the main quartile for the USA population and did not take into account race or ethnic differences.

In 2003, the American Society of Clinical Endocrinologists (AACE) [6] again proposed the criteria with insulin resistance as the leading factor. The prerequisite for MetS diagnosis was:

- IGT or IGF, but not diabetes.
- Moreover, an additional factor was one of the following:
- Overweight or obesity (BMI  $\geq 25$  kg/m<sup>2</sup>);
- Dyslipidaemia (triglycerides  $\geq 150$  mg/dl and/or HDL cholesterol  $< 40$  mg/dl for men and  $< 50$  mg/dl for women);
- Arterial blood pressure ( $\geq 130/85$  mm Hg);
- - Other risk factors: positive family history of type 2 diabetes, polycystic ovarian syndrome, sedentary lifestyle, old age, being part of the ethnic group exhibiting high risk for developing type 2 diabetes.

However, diagnostic recommendations of AACE are not commonly applied due to the lack of precise criteria. In 2005, the International Diabetes Federation (IDF) presented new diagnostic criteria of MetS [7]:

- Abdominal obesity (waist measurement  $\geq 94$  cm for men and  $\geq 80$  cm for women in the European population or BMI  $\geq 30$  kg/m<sup>2</sup>);
- Additionally, at least two out of four factors have to occur:
- Triglycerides ( $\geq 150$  mg/dl) or management of hypertriglyceridemia;

- HDL cholesterol (<40 mg/dl for men and <50 mg/dl for women) or treatment of low HDL-C;
- Arterial blood pressure (systolic  $\geq 130$  and/or diastolic  $\geq 85$  mm Hg, or treatment of arterial hypertension);
- Fasting glucose ( $\geq 100$  mg/dl or treatment of diabetes).

Insulin resistance, which is difficult to identify in everyday medical practice, was not taken into consideration. However, abdominal obesity by ethnic group, which is closely associated with the occurrence of insulin resistance, was accepted as a criterion. In the IDF definition, the lower limit of abnormal fasting glucose level was reduced from 110 mg/dl to 100 mg/dl. Due to simplicity of the

diagnostic criteria presented by IDF, the definition is frequently used in epidemiological studies [8].

In 2009, IDF and AHA/NHLBI (American Heart Association/ National Heart, Lung and Blood Institute) [9] accepted common criteria for the diagnosis of MetS, according to which the metabolic syndrome is diagnosed when at least three out of five risk factors are identified (table 1). In the presented criteria, abdominal obesity is not a prerequisite for the diagnosis of the metabolic syndrome. Abdominal obesity depending on the different population and ethnic group, for example in the European population recommendations for the waist circumference are  $\geq 94$  cm for men and  $\geq 80$  cm for women; in the USA population  $\geq 102$  cm for men and  $\geq 88$  cm for women; in the Asian population  $\geq 90$  cm for men and  $\geq 80$  cm for women.

**Table 1.** Current criteria for clinical diagnosis of the Metabolic Syndrome [9].

Measure	Categorical cut points
Elevated waist circumference	Population- and country-specific definitions
Elevated triglycerides or drug treatment for elevated triglycerides	$\geq 150$ mg/dl (1,7 mmol/l)
Reduced HDL cholesterol or drug treatment for reduced HDL cholesterol	<40 mg/dl (1,0 mmol/l) for men, <50 mg/dl (1,3 mmol/l) for women
Elevated blood pressure or antihypertensive drug treatment	systolic $\geq 130$ and/or diastolic $\geq 85$ mm Hg
Elevated fasting glucose or drug treatment of elevated glucose	$\geq 100$ mg/dl

Currently, no definition can be totally rejected. Further prospective, randomized epidemiological and clinical studies are needed. Additionally, other symptoms and disorders that go beyond the scope of the definition are observed in MetS, which for practical reasons should be individually identified. However, the finding of any MetS component is a prerequisite for diagnostic performance to diagnose or exclude all other disorders.

## EPIDEMIOLOGY

The incidence of MetS in Poland and worldwide increases each year, despite taking into account different diagnostic criteria. Research conducted in Poland shows that approximately 20% of the adult population suffer from MetS, and that more frequently women are affected. Worldwide, this problem affects even 33% of the USA population, 27% of the population of China and in Europe - 15% of the population of France and up to 34% of the Italian or Finland population [10].

The major cause of MetS is abdominal obesity, which subsequently leads to type 2 diabetes and cardiovascular disorders, and contributes to the increased mortality due to complications. In the USA, over 25% of adult inhabitants are affected

[11]. Worldwide, in 2014 overweight was diagnosed in 39% of the population (38% of men and 40% of women) above the age of 18, whereas obesity in 13% (11% of men and 15% of women). Moreover, 41 million of children under 5 years of age had overweight or obesity. Between 1980 and 2014 the world incidence of obesity increased twice [12].

## PREVENTION AND NUTRITIONAL THERAPY OF THE METABOLIC SYNDROME

The metabolic syndrome is a complex disease. Therefore, prevention and treatment also have to be multifaceted and involve not only the institution of dietetic management and increased physical activity, but also pharmacotherapy to rebalance lipid parameters, glycaemia and arterial blood pressure [13].

Body mass normalization plays a major role [14]. Body mass reduction should be gradual and should involve a 10% drop within a year, resulting in a decrease in dietary energy value by 500-1000 kcal/day. The diet should be based on easily digestible carbohydrates with low or moderate glycaemic index (<70) and low glycaemic

load (<10), accounting for 50% of daily dietary energy value. The intake of simple sugars contained in sweets, cakes and honey has to be limited (below 10% of dietary energy value). Dietary carbohydrates are obtained from wholegrain cereal products, vegetables and fruit, legume seeds and nuts, which also provide fibre, group B vitamins, polyphenols and mineral components. The content of dietary fibre in a reducing diet should oscillate around 30-40 g/day. Clinical examinations showed that a low-glycaemic index diet improves glycaemia control in diabetes [15]. The glycaemic index of food products is determined by the degree of grinding and boiling (the higher the degree, the higher the glycaemic index), the presence of other nutrients in food (fibre, protein and fats reduce the glycaemic effect of carbohydrates), the effect of the preceding food product (low glycaemic index meal may reduce glycaemia after the consumption of a subsequent meal) [16].

The quantitative dietary content of fat should account for 30% of energy intake available from food; however, also the quality of fats has to be considered. The recommended consumption of monosaturated fatty acids should constitute 10-15% of energy intake, polyunsaturated fatty acids 6-10%, including omega-6 fatty acids (5-8%) and omega-3 fatty acids (1-2%), whereas saturated fatty acids should be limited to 7%. Moreover, the intake of trans type fatty acids should be as low as possible, not exceeding 1% of dietary energy value. Dietary cholesterol content should be no more than 200-300 mg/day. The best food sources of fat for patients with MetS include vegetable virgin oils, cold-pressed and eaten raw, as well as nuts (especially walnuts) and saltwater fish rich in omega-3 fatty acids. The intake of animal fats and hard margarines has to be limited. It has been demonstrated that the use of energy-low diet with modified proportion of polyunsaturated fatty acids (n-3/n-6; 1:4-6) for 12 weeks reduces lipid metabolism disorders and arterial blood pressure in obese patients with clinical symptoms of the metabolic syndrome [17].

The dietary protein intake should account for 20% of dietary energy, with the 50/50% of animal to vegetable protein ratio. Low-fat poultry, low-fat dairy products, especially fermented milk drinks, fish, eggs, legume seeds and wholegrain cereal products are a good source of dietary protein.

Dietary salt consumption should not exceed 5g, as excessive salt intake causes hypertension and leads to treatment-resistant hypertension. Therefore, all highly-processed salt-rich products, such as salty snacks, fast food, powdered soups and sauces, spice mixtures, smoked cheese and meat have to be eliminated from the diet [18,19].

Plant polyphenols are an important dietary component, which in the Polish diet is mainly

available from coffee, tea, vegetables and fruit. It has been demonstrated that the consumption of polyphenols in the Polish diet is inversely correlated with the presence of MetS and the risk of hypertension [20,21].

The consumption of 5-6 small-volume meals daily are recommended, taking into account food preferences of patients, and 1.5 litre of low-sodium water. Regular physical activity (30-45 min daily) is of utmost importance [22].

## CONCLUSIONS

MetS prevention and therapy in the first place should involve non pharmacological methods, such as proper diet and physical activity, which lead to body mass reduction and restrict the occurrence of risk factors.

## Conflicts of interest

The authors declare that they have no conflict of interest.

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