

## Profile of Metabolic Syndrome in Newly Detected Hypertensive Patients in India: An Hospital-Based Study

### Abstract

**Background:** Metabolic syndrome (MetS) is recognized as an emerging threat and interest of public health because the factors defining syndrome are associated with increased risk of mortality and morbidity. Hypertension further adds to risk factor leading to target organ damage. Recognizing the MetS in patients with hypertension provides a great opportunity for more aggressive treatment. **Objective:** The objective of the study is to estimate the prevalence and evaluate the metabolic profile of MetS in newly diagnosed adult hypertensive participants. **Methodology:** This is an hospital-based cross-sectional study. A total of 400 participants with newly detected essential hypertension were included in the study after detailed medical and laboratory investigations done to exclude the secondary hypertension. After the informed consent, a structured proforma was filled containing the demographic data; medical history; physiological parameters such as blood pressure, height, weight, and waist circumference; some laboratory investigations such as lipid profile and fasting blood glucose. Descriptive statistics such as mean and proportion were used. To compare the proportions, Chi-square test was used. Kappa agreement was utilized to know the level of agreement between various criteria defining MetS. **Results:** The prevalence of MetS according to the International Diabetes Federation criteria in the new hypertensive study participants was 50.5% more common in females. One-fourth of young hypertensives was having MetS. Hypertensive patients with MetS show risk factors at significantly higher range than their counterparts. High-density lipoprotein was the most common risk factor present apart from increased waist circumference. There was a wide variation in the prevalence of MetS in the Indian population by different criteria. **Conclusion:** Half of the new hypertensive patients had MetS and thus it becomes very important to screen all the hypertensives at the onset for MetS and treat them aggressively to decrease the cardiovascular events.

**Keywords:** Hospital-based, India, metabolic syndrome, new detected hypertensive patient, risk factor

### Introduction

Metabolic syndrome (MetS) is characterized by an array of cardiovascular risk factors such as diabetes and raised fasting plasma glucose, abdominal obesity, high cholesterol, and high blood pressure (BP). Although detected very early in the year 1923, five decades later, i.e., 1980s Reaven coined the term “syndrome X” for this conglomeration of various metabolic abnormalities with insulin resistance being the basic underlying pathophysiologic problem.<sup>[1]</sup> Since then, the definition and terminology of MetS are continuously evolving. The WHO (1999) given a first working definition of MetS with the presence of diabetes, impaired glucose tolerance, or insulin resistance as mandatory criteria.<sup>[2]</sup> It was later commented by “the

European Group for the Study of Insulin Resistance” which coined the term insulin resistance syndrome.<sup>[3]</sup> Both were criticized to be highly technical and impracticable. The first clinical definition was released by the National Cholesterol Education Program Adult Treatment Panel III (NCEP ATP III) with clear cutoff point for each risk factor, differentiated as per sex for waist circumference (WC) and high-density lipoprotein (HDL). Insulin resistance was not considered mandatory but the presence of three or more risk factor.<sup>[4]</sup> This was further modified by the International Diabetes Federation that identified central obesity (WC) as an essential component, given with race- and gender-specific WC cutoff and lowers the threshold for several parameters such as WC, BP, and fasting plasma glucose.<sup>[5]</sup> However, optimum definition is still unknown.

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According to the global burden of diseases 2016 nearly in India, 28.09% of total deaths are attributed to CVD.<sup>[6]</sup> This increase is attributed to the increased prevalence of coronary risk factors in Indians. It is estimated that around 20%–25% of the world's adult population has the MetS. MetS is recognized as an emerging threat and interest of public health because the factors defining syndrome are associated twice as likely to die from and three times as likely to have a heart attack or stroke compared with people without the syndrome.<sup>[5,7,8]</sup> National representative study on the prevalence of MetS is not available in India. A recent study in India shows that at least one-third of urban Indians have MetS.<sup>[9,10]</sup>

BP levels are strongly associated with insulin levels and the degree of insulin resistance. It has been reported that insulin resistance might be involved in the pathogenesis of primary hypertension in up to 40%–50% of cases.<sup>[11]</sup> MetS is present in up to one-third of hypertensive patients. This not only exposed them to a higher risk of target organ damage but also increased the incidence of cardiovascular complications.<sup>[12,13]</sup> Hypertension tends to cluster with other metabolic risk factors. Early recognition of MetS in hypertensive patients, therefore, acquires greater clinical relevance to improving prognosis. Recognizing the MetS in patients with hypertension provides a great opportunity for more aggressive treatment, including lifestyle, dietary modification, weight management, and treatment of comorbid factors to attain cardiovascular risk reduction. To our knowledge, there are no studies conducted in India, highlighting the prevalence and metabolic profile of MetS in newly diagnosed adult hypertensive participants.

### Objective

The objective is to study the prevalence of MetS in newly detected hypertensive patients, to find the association between risk factors and presence of MetS in newly detected hypertensive patients, and to find the prevalence of risk factors for diagnosed MetS in newly detected hypertensive patients.

### Methodology

This was a hospital-based cross-sectional study.

Considering the 52%<sup>[14]</sup> prevalence of MetS in hypertensive patient, 5% precision, and 95% confidence level, sample size comes to be 384, we recruited 400 newly detected hypertensive patients enrolled from June 16 to December 2017 on the basis of below criteria.

#### Inclusion criteria

Age >18 years newly detected hypertensive patients (BP >140/90) were included in the study.

#### Exclusion criteria

Patients who have secondary hypertension, endocrinopathies, long-term medications causing dyslipidemia such as beta-blockers, and steroid use were excluded from the study.

Informed consent was obtained from each patient as per the approved format by the IEC. After obtaining demographic (age, sex, ethnicity, and residential address) medical history was recorded, height, weight, and WC were measured. Standing height was measured nearest to 0.1 cm and weight nearest to 0.1 kg using a calibrated digital weighing machine. The WC measured at the level between the iliac crest and the lower margin of the ribs at the end of expiration. Body mass index (BMI) was calculated and categorized as per the WHO.<sup>[15]</sup>

Seated BP was measured after at least 10 min of rest, two BP recordings were obtained with the help of aneroid sphygmomanometer after a gap of at least 10 min interval. Physical activity was assessed as per the questionnaire blood sample was obtained for the estimation of fasting blood sugar (FBS) and serum for the estimation of HDL and triglycerides (TGs). All analyses were done in biochemistry laboratory by the automated analyzer in a biochemistry laboratory, with commercially available kits. The International Diabetes Federation (IDF) criteria were used for the diagnosis of MetS as given below in Table 1.<sup>[5]</sup>

Index of central obesity (ICO)<sup>[16]</sup> was calculated as ratio WC to height.

**Table 1: International Diabetes Federation criteria for the diagnosis of metabolic syndrome**

| Criteria for diagnosis of MetS  | Cut off values   |
|---|--|
| Central obesity (defined as waist circumference × with ethnicity-specific values) for Asian | Male >90 cm<br>Female >80 cm   |
| Plus any two of the following four factors  |  |
| Raised triglycerides  | ≥150 mg/dL (1.7 mmol/L) or specific treatment for this lipid abnormality   |
| Reduced HDL cholesterol   | <40 mg/dL (1.03 mmol/L) in males <50 mg/dL (1.29 mmol/L) in females or specific treatment for this lipid abnormality |
| Raised blood pressure   | Systolic BP ≥130 or diastolic BP ≥85 mm Hg or treatment of previously diagnosed hypertension                         |
| Raised fasting plasma glucose   | FPG ≥100 mg/dL (5.6 mmol/L), or previously diagnosed type 2 diabetes   |

HDL: High-density lipoprotein; BP: Blood pressure; FPG: Fasting plasma glucose

## Data analysis

Data were analyzed in MS Excel 2013 and SPSS 23 by proportion IBM Corp. Released 2015. IBM SPSS Statistics for Windows, Version 23.0. (Armonk, NY: IBM Corp.), mean, and standard deviation. ANOVA and Chi-square test were used to compare means and proportion, respectively. Kappa coefficient was used to find the agreement of IDF with various other criteria for diagnosing MetS.

## Results

A total of 400 newly detected cases of hypertension were enrolled in the study. The prevalence of MetS according to the IDF criteria in the new hypertensive study participants was 50.5%. Among those having MetS, 72.27% were females. Compared to study participants not having MetS, cases were 5 years younger, significantly more WC, ICO, TG, BMI, and lower HDL level [Table 2]. Table 3 shows a maximum number of cases belong to middle aged (52.47%) and old aged (32.67%) but 75% of young adults had MetS followed by 60% of middle aged. As WC is obligatory criteria for diagnosis of metabolic syndrome as per IDF criteria, we found 100% of study participants to be having central obesity. The prevalence of HDL as a risk factor

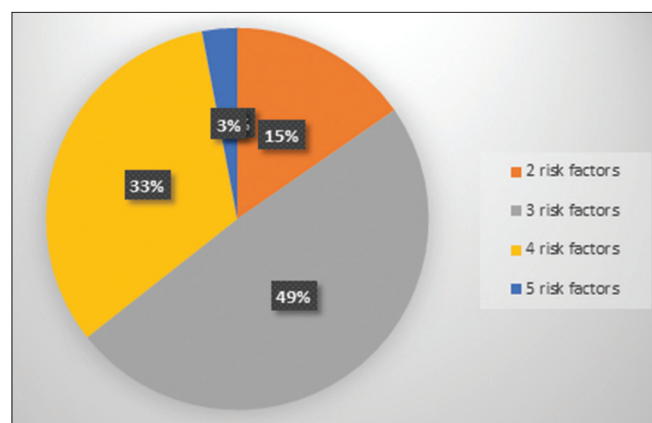


Figure 1: Number of risk factors in patient with metabolic syndrome

was found to be significantly more in females (64.28% vs. 90.41%), whereas FBS was more in males (60.7% vs. 45.2%) [Table 4]. Figure 1 shows that 49%, 33%, and 3% had three, four, and five risk factors, respectively.

In the study participants, the prevalence of MetS by NCEP ATP III was 64.5%, joint interim IDF task force was 60.5%, and as per Parikh and Mohan criteria, considering ICO as obligatory criteria was 68%. Fair agreement (0.519) was found between NCEP ATP III and IDF. Moderate agreement in the diagnosis of MetS was seen between criteria of joint interim IDF task force (0.789) and Parikh and Mohan (0.635) with IDF criteria [Table 5].

## Discussion

In this study, 50.5% of the newly detected hypertensive patient had MetS according to the IDF criteria. However, according to the modified IDF criteria for Asian Indian<sup>[18]</sup> with WC as nonobligatory criteria and presence of three or more factors, the prevalence was much higher, i.e., 60.5%. When NCEP ATP III guidelines were considered the prevalence was 64.5%, and according to Parikh and Mohan criteria, it was 68%. The earlier studies in 2004 showed the prevalence of 30%–35%, but recent studies show more than half of hypertensive patient had MetS.<sup>[10,19,20]</sup> This prevalence in hypertensive is twice more than the general population.<sup>[21]</sup> PRESCOT study done in Spain on 12,000 hypertensives shows the prevalence of 52% and 75.5% as per the ATP III and IDF guidelines, respectively.<sup>[14]</sup> According to the IDF guidelines, 72% of cases of MetS were females and this gender difference was statistically significant which is in line many studies showing comparatively higher prevalence in females, but the gap was only 10%–15% documented till now. Apart from daily household chores, Indian women hardly have access or aware of physical activity exposing them to central obesity. Those having a MetS in study participants were found to be 6 years younger. Though the overall prevalence of MetS was more in middle aged, but proportionately higher number

Table 2: Characteristic of patient with newly detected hypertension by metabolic syndrome

| Variables                    | New hypertensive with MetS (n=202) | New hypertensive without MetS (n=198) | Difference (CI, P)          |
|------------------------------|------------------------------------|---------------------------------------|-----------------------------|
| Sex (male/female percentage) | 27.72/72.27                        | 78.78/21.21                           | 48.9, (40.1-57.2, 0.0001)   |
| Age                          | 52±11                              | 57±12                                 | 6.14 (3.4-6.5, <0.0001)     |
| Waist circumference          | 36.57±3.78                         | 31.89±2.73                            | -20 (-5.1--4.2, <0.0001)    |
| ICO                          | 0.59±0.06                          | 0.50±0.04                             | -58 (-58.5--58.49, <0.0001) |
| Systolic BP                  | 159±18                             | 160±25                                | 0.64 (-2-4, NS)             |
| Diastolic BP                 | 92±11                              | 93±11                                 | 1.28 (-0.5-2.5, NS)         |
| FBS                          | 110±30                             | 107±47                                | -1.07 (-0.8-2.4, NS)        |
| HDL                          | 40±7                               | 43±14                                 | 3.8 (1.4-4.5, 0.0001)       |
| Triglyceride                 | 182±89                             | 148±88                                | -5.4 (-46--21.7, <0.0001)   |
| BMI                          | 28.74±4.39                         | 23.22±4.08                            | -18.4 (-6.1--4.9, <0.0001)  |

ICO: Index of central obesity; HDL: High-density lipoprotein; BP: Blood pressure; FPS: Fasting blood sugar; BMI: Body mass index; CI: Confidence interval; MetS: Metabolic syndrome; NS: Not significant

of young adults from total young adults were found to have MetS. This suggests that younger patient detected with hypertension should be meticulously screened for MetS.

Except for FBS, all the risk factors were significantly more in MetS cases. Only one participant was taking the anti-diabetic drug at time of survey. This suggest that inspite being a new case of hypertension and not previously being diagnosed as diabetes those with MetS have comparatively higher level of TG, BMI and lower level of HDL than without MetS group. Therefore it can be concluded that hypertension may be an one of the early marker for MetS. All patient with newly detected hypertension should be screen for MetS in primary care setting.

As WC was obligatory criteria so it was 100% and ICO was 99% and as only hypertensive was considered so it

**Table 3: Age and sex distribution of newly detected hypertension by metabolic syndrome**

| Age                                   | Metabolic syndrome, n (%) | P       |
|---------------------------------------|---------------------------|---------|
| Young adult (25-39 years) (n=52)      | 30 (75)                   | <0.05   |
| Middle aged (40-59 years) (n=176)     | 106 (60.2)                |         |
| Old aged (60 years and above) (n=172) | 66 (38.37)                |         |
| Sex                                   |                           |         |
| Male (212)                            | 56 (26.4)                 | <0.0001 |
| Female (188)                          | 146 (77.65)               |         |
| Total (400)                           | 202 (50.5)                |         |

was again 100% in MetS cases. HDL and BMI were the most common risk factor (83.16%) followed by TG and FBS. A similar finding was reported by Makwana *et al.*,<sup>[19]</sup> while the most common risk factors in Salagre *et al.*<sup>[20]</sup> and PRESCOT study<sup>[14]</sup> were fasting blood glucose/DM followed by TG. This difference may be due to strict criteria for selecting an only new hypertensive patient. The study done on normal participants shows hypertension followed by HDL to be the most common risk factor.<sup>[21]</sup> HDL as a risk factor was profoundly seen in females while BMI and FBS more in males. No correlation was found between the BP and a number of metabolic risk factors might be due to selection criteria and hypertension in the earlier stage if not treated optimally will further lead to the development of risk factors as demonstrated in another study.<sup>[14]</sup> About 85 % of participants with MetS had three or more risk factors at the time of study. It has been noted in various studies<sup>[12,14]</sup> that more the number of risk factors, worse is the prognosis.

Considering various criteria for defining MetS, there was the very varying prevalence of MetS found in our study from 50.5% to 68%. When we considered WC as the sole criteria then 18 participants (4%) were categorised as not having MetS. Although these participants were overweight and had more than three risk factors for MetS. Therefore considering only WC as sole criteria for MetS will give comparatively lower prevalence of MetS than other

**Table 4: Prevalence of risk factors for metabolic syndrome in newly detected hypertensive patient by sex**

| Risk factors         | Total (%)   | Male (n=56), n (%) | Female (n=146), n (%) | P       |
|----------------------|-------------|--------------------|-----------------------|---------|
| Waist circumference  | 202 (100)   | 56 (100)           | 146 (100)             |         |
| ICO                  | 200 (99)    | 56 (100)           | 144 (98.6)            | NS      |
| Hypertension Stage 1 | 94 (46.53)  | 24 (42.85)         | 70 (47.9)             | NS      |
| Hypertension Stage 2 | 108 (53.46) | 32 (57.14)         | 76 (52.05)            | NS      |
| FBS                  | 100 (49.5)  | 34 (60.7)          | 66 (45.2)             | 0.002   |
| HDL                  | 168 (83.16) | 36 (64.28)         | 132 (90.41)           | <0.0001 |
| TG                   | 128 (63.36) | 38 (67.85)         | 90 (61.6)             | NS      |
| BMI                  | 168 (83.16) | 50 (89.3)          | 118 (80.8)            | 0.01    |

ICO: Index of central obesity; HDL: High-density lipoprotein; BP: Blood pressure; FPS: Fasting blood sugar; BMI: Body mass index; CI: Confidence interval; NS: Not significant

**Table 5: Agreement between International Diabetes Federation with National Cholesterol Education Program Adult Treatment Panel III, Joint interim International Diabetes Federation Task Force; National Heart, Lung, and Blood Institute; American Heart Association, World Heart Federation; International Atherosclerosis Society; and International Association for the Study of Obesity and Parikh and Mohan**

| Criteria for classification of MetS  | MetS(+/-) | IDF <sup>[5]</sup> |          | Observed agreement (%) | Kappa coefficient   |
|--|-----------|--------------------|----------|------------------------|---------------------|
|  |           | MetS (+)           | MetS (-) |                        |                     |
| NCEP ATP III <sup>[4]</sup>  | MetS (+)  | 92                 | 37       | 76.24                  | 0.519 (0.405-0.634) |
|  | MetS (-)  | 10                 | 61       |                        |                     |
| Joint interim IDF task force; national heart, lung, and blood institute; AHA, WHF; International Atherosclerosis Society; and International Association for the Study of Obesity <sup>[17]</sup> | MetS (+)  | 101                | 20       | 89.5                   | 0.789 (705-0.873)   |
|  | MetS (-)  | 1                  | 78       |                        |                     |
| Parikh and Mohan <sup>[6]</sup>  | MetS (+)  | 101                | 35       | 81.91                  | 0.635 (0.533-0.736) |
|  | MetS (-)  | 1                  | 62       |                        |                     |

IDF: International Diabetes Federation; NCEP ATP III: National Cholesterol Education Program Adult Treatment Panel III; AHA: American Heart Association; WHF: World Heart Federation; MetS: Metabolic syndrome

definitions. The prevalence found by IDF in our study was lowest compared to other criteria while other studies done in Mexican American showed maximum prevalence by IDF.<sup>[14,22]</sup> No such comparison was found in any study with Indian participants as per our knowledge.

It has been demonstrated by Barrios *et al.*<sup>[14]</sup> that general practitioners failed to diagnose 43.7% of MetS cases in hypertensive. This will further aggravate the cardiovascular risk if MetS is not aggressively treated timely. We need to choose the best feasible criteria for screening MetS. Among the other three criteria, best (moderate) agreement was found between IDF and joint interim (three or more risk factor).

However, for screening, it is better to use ICO over WC as WC cutoff given for Asians is still under consideration and it is easiest to calculate and interpret. Moderate agreement was found with IDF and yield a maximum number of cases. If the ratio of WC and height (ICO) is more than half then it is recommended to screen further for MetS.

## Conclusion

Half of the newly detected hypertensive study participants had MetS, 2.6 times more in females than males. The MetS cases had risk factors such as WC, ICO, HDL, TG, and BMI at a higher level than non-MetS cases. HDL and BMI were the most common risk factor followed by TG. This finding can be generalized to hypertensive population due to sufficient sample size.

Thus, considering high prevalence and increased cardiovascular risk, all the newly detected hypertensive should be screened for MetS preferably by ICO. There is a need for online medical education for focused management of MetS in hypertensive.

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## Conflicts of interest

There are no conflicts of interest.

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