

## FREQUENCY OF METABOLIC SYNDROME IN PATIENTS WITH TYPE-2 DIABETES

Nasir Ahmed, Tauqir Ahmad, Syed Javed Hussain\*, Muhammad Javed

Department of Medicine, \*Endocrinology, Ayub Medical College, Abbottabad, Pakistan

**Background:** Diabetes, Hypertension, Obesity and Ischaemic Heart Disease have become a problem of public health magnitude with substantial economic burden both in the developed as well as the developing countries. Obesity is quite frequent in Type 2 diabetics and also plays a central role in causing Metabolic Syndrome (MetS). Metabolic Syndrome significantly increases the incidence of cardiovascular complications. This study was done to determine the frequency of MetS in our Type 2 diabetic patients as most of the components of MetS can be modified and identifying/managing these at an early stage might be of considerable help in reducing cardiovascular complications. **Methods:** This cross-sectional study was done in Medical 'B' and Medical 'A' wards of Ayub Teaching Hospital, Abbottabad from Nov, 08 to April, 09. Type 2 Diabetic patients aged above 40 years who gave informed consent were included in the study. Data was collected through a structured proforma. Frequency of Metabolic Syndrome was estimated according to the IDF consensus worldwide definition of the MetS. **Results:** Of the 100 patients enrolled in this study 56 were females and 44 were males with a mean age of 59.9 years. Out of these 100 participants seventy six (76%) were diagnosed to have metabolic syndrome. Of the 56 females, forty eight (85.71%) were having metabolic syndrome while twenty eight (63.63%) of the 44 male participants were having the syndrome. The difference was statistically significant ( $p < 0.05$ ). **Conclusion:** Frequency of MetS was found to be significantly high in this study with female preponderance. All the components, except Hypertension were more frequent in females. Diabetic patients with metabolic syndrome need more aggressive approach in management so as to decrease the incidence of cardiovascular complications.

**Keywords:** Metabolic Syndrome, Type 2 Diabetes Mellitus, Cardiovascular Complications

### INTRODUCTION

Type 2 Diabetes Mellitus is increasing at an alarming rate especially in the developing countries<sup>1</sup> and is almost an epidemic now-a-days<sup>2</sup>. There is a high prevalence of long-term diabetic complications in our patients due to multiple factors (e.g. poor control, obesity etc.) responsible for substantial economic burden.<sup>3</sup> It is also established that there is a worldwide association between Type 2 Diabetes Mellitus, Obesity and Physical inactivity.<sup>4</sup> Obesity in turn plays a central role in causing Metabolic Syndrome<sup>5</sup> which has been defined as a constellation of variables that predicts premature coronary artery disease and type 2 diabetes mellitus and these are the principal causes of long-term morbidity, mortality as well as excess health care cost<sup>6,7</sup>. The syndrome is said to be present when three or more of the following are present according to the NCEP-ATP III definition:<sup>8</sup>

- Abdominal obesity (waist circumference >102 cm in men and >88 cm in women).
- Triglycerides (TGs)  $\geq 1.7$  mmol/l.
- HDL Cholesterol <1.03 mmol/l in men and <1.29 mmol/l in women.
- Systolic BP  $\geq 130$  mmHg and/or Diastolic BP  $\geq 85$  mmHg.
- Fasting Plasma Glucose  $\geq 6.1$  mmol/l.

Although the need to diagnose MetS in diabetes is controversial, we believe it is worth considering because diabetic patients frequently visit

health-care providers. A diagnosis of MetS in them will necessitate a more aggressive approach in management thus improving their life-style and decreasing the incidence of cardiovascular complications.<sup>9</sup> The importance of Diabetes Mellitus and its association with MetS is evident from the WHO definition of MetS<sup>10</sup> which states:

- Diabetes (Fasting Plasma Glucose  $\geq 7.0$  mmol/l and/or 2-hour plasma glucose  $\geq 11.1$  mmol/l, or impaired glucose regulation (FPG 6.1–6.9 mmol/l and/or 2-hour plasma glucose 7.8–11.0 mmol/l and/or insulin resistance (below lowest quartile of glucose uptake in the euglycaemic clamp) and two or more of the following.
- Raised triglycerides ( $\geq 1.7$  mmol/l) and/or low HDL Cholesterol (<0.9 mmol/l in men, <1.0 mmol/l in women).
- Central Obesity (waist to hip ratio >0.90 in men, >0.85 in women) and/or body mass index (BMI) >30 kg/m<sup>2</sup>.
- Raised blood pressure (Systolic blood pressure  $\geq 140$  mmHg and/or Diastolic blood pressure  $\geq 90$  mmHg).
- Microalbuminuria (urinary albumin excretion  $\geq 20$   $\mu$ g/min or albumin/creatinine ratio  $\geq 30$  mg/g)

Almost similar criteria are considered in the IDF consensus worldwide definition of the metabolic syndrome<sup>11,12</sup> which states:

Central Obesity as defined for Asian population (Waist Circumference  $\geq 90$  cm in men,  $\geq 80$  cm in women), together with 2 of the following:

- Blood Pressure (Systolic blood pressure  $\geq 130$  mmHg, Diastolic BP  $\geq 85$  mmHg) or treatment of previously diagnosed hypertension
- Fasting Plasma Glucose  $\geq 5.6$  mmol/l ( $\geq 100$  mg/dl) or previously diagnosed type 2 diabetes
- HDL Cholesterol  $< 1.03$  mmol/l ( $< 40$  mg/dl) in men,  $< 1.29$  mmol/l ( $< 50$  mg/dl) in women
- Triglycerides  $\geq 1.7$  mmol/l ( $\geq 150$  mg/dl)

It is clear from the definitions stated above of metabolic syndrome that almost all the variables of MetS are preventable/modifiable. Public awareness can be started from diabetic patients because of their frequent encounter with health-care providers and general public can then be targeted to have a good quality life and excess health-care cost saved by decreasing cardiovascular complications.

## MATERIAL AND METHODS

This cross-sectional study was conducted at Ayub Teaching Hospital, Abbottabad in Medical 'B' and 'A' units. The study period was 06 months from 1<sup>st</sup> Nov 2008 to 30<sup>th</sup> April 2009. Non-probability convenience sampling technique was used and 100 type 2 diabetics of either gender aged  $\geq 40$  years completed the study. Informed consent was taken from all the patients. Already diagnosed cases of Hypothyroidism, Nephrotic syndrome and patients with renal failure were excluded. Specifically designed forms were used to collect the data. Samples for blood glucose, TGs and HDL Cholesterol were taken after an overnight fast. Sitting blood pressure was measured with mercury sphygmomanometer after 10 minutes of rest. Two readings were taken and the mean was recorded for data. Waist circumference was measured at the level of the mid point between the high point of the iliac crest and the last rib. Metabolic syndrome was diagnosed using the IDF consensus worldwide definition as described above. This definition of MetS is simple, more practical and uniformly agreed upon.

Data was analysed using SPSS Version 10. Descriptive statistics are presented as a percentage of the total no. of the participants. Cross tabulation was used to determine the relationship of different variables of the metabolic syndrome. Chi-square test was applied for determination of *p*-values, and *p*  $< 0.05$  was considered as significant.

## RESULTS

A total of 100 known type 2 diabetics were enrolled in this study. Fifty-six (56%) were females and 44 (44%) were males. The mean age of the study participants was 59.9 years ranging from 41 to 80 years (Table-1). Out of 100 participants 76 (76%) were diagnosed to have

metabolic syndrome by applying the IDF consensus worldwide definition of MetS (Table-2). Of the 56 females 48 (85.71%) were found to have MetS while 28 (63.63%) of the 44 male participants were having the syndrome. The results are shown in Table-3 along with age group of patients. By applying IDF consensus worldwide definition of metabolic syndrome the following results obtained:

Twelve (12%) participants fulfilled all 5 criteria for metabolic syndrome. Twenty-eight (28%) of all the participants had four criteria of metabolic syndrome. Three criteria were fulfilled by 36 (36%) of all the participants.

Of the 44 males, 4 (9.09%) had all 5 components of MetS, 8 (18.18%) had 4 criteria while 3 criteria were present in 16 (36.36%). Of the 56 females, 8 (14.28%) had all 5 criteria while 4 and 3 criteria of metabolic syndrome were met by 20 (35.71%) participants each (Table-4).

Among the patients having metabolic syndrome the facts and figures were as follows:

The total number was 76 (76%). Out of 76 participants having the syndrome males were 28 (36.84%) and females were 48 (63.15%). All the patients were known diabetics. Increased waist circumference was found in 56 (73.68%), 16 (28.57%) were males and 40 (71.43%) were females. Hypertension was seen in 48 (63.16%) participants. Among these 24 (50%) were males and 24 (50%) were females. Triglycerides were increased in 36 (47.37%) participants with 12 (33.33%) males and 24 (66.67%) females having raised TGs. Decreased HDL Cholesterol levels were found in 64 (84.21%) patients of metabolic syndrome. Out of these 20 (31.25%) were males and 44 (68.75%) were females (Table-5). A comparison of participants having metabolic syndrome with those without the syndrome is shown (Table-6).

Frequency of metabolic syndrome increased with age. It increased from 66.66% in the age group 41–49 to 81.81% in the age group  $\geq 70$  years. The most common component of the metabolic syndrome was low HDL Cholesterol. The most common component of the metabolic syndrome in males was Hypertension while in females it was low HDL Cholesterol. Obesity (increased waist circumference) was more common in females as compared to males.

**Table-1: Age distribution of the participants**

Age Group (years)	Male	Female	Total
40–49	9	18	27
50–59	13	23	36
60–69	14	12	26
$\geq 70$	8	3	11
<b>Total</b>	<b>44</b>	<b>56</b>	<b>100</b>

**Table-2: Frequency of metabolic syndrome**

Gender	Total (n=100)	MetS (n=76)	% with MetS
Male	44	28	36.84
Female	56	48	63.16

**Table-3: Metabolic Syndrome in Different Age Groups (n=76)**

Age Group (in years)	Male	Female	Total	Percentage
41-49	3	15	18	23.68
50-59	8	20	28	36.84
60-69	10	11	21	27.63
≥70	7	2	9	11.84
Total	28	48	76	100

**Table-4: No of the patients fulfilling different criteria of metabolic syndrome among patients having the syndrome (n=76)**

	5 Criteria		4 Criteria		3 Criteria	
	No	%	No	%	No	%
Men (n=28)	4	9.09	8	18.18	16	36.36
Women (n=48)	8	14.28	20	35.71	20	35.71
Total (n=76)	12	15.78	28	36.84	36	47.38

**Table-5: Frequency of the components of the metabolic syndrome in patients diagnosed to have MetS (n=76)**

	Total (n=76)		Male (n=28)		Female (n=48)		p
	No	%	No	%	No	%	
Type 2 Diabetes	76	100	28	36.84	48	63.16	>0.05
Increased Waist Circumference	56	73.6	16	28.57	40	71.43	<0.05
Hypertension	48	63.2	24	50	24	50	>0.05
Raised Triglycerides	36	47.4	12	33.33	24	66.67	>0.05
Low HDL-C	64	84.2	20	31.25	44	68.75	<0.05

**Table-6: Comparison of participants having metabolic syndrome (n=76) with those without the syndrome (n=24)**

	Metabolic Syndrome	Mean±SD	p
Waist Circumference (in cm)	Yes	101.26±16.36	0.012*
	No	82.25±5.91	
Systolic Blood Pressure (in mmHg)	Yes	150.33±27.53	0.000*
	No	123.54±18.73	
Diastolic Blood Pressure (in mmHg)	Yes	89.28±11.68	0.000*
	No	75.21±13.14	
Triglycerides (in mg/dl)	Yes	133.47±65.15	0.001*
	No	95.75±38.52	
High Density Lipoprotein (in mg/dl)	Yes	36.16±11.58	0.001*
	No	46.62±12.50	

\*Statistically significant

## DISCUSSION

The diagnosis of metabolic syndrome in patients might hold promise for better control of diabetes and enhanced prevention of cardiovascular complications. According to IDF consensus worldwide definition of the syndrome<sup>11,12</sup>, it is said to be present when central obesity together with any 2 among hypertension, diabetes mellitus or raised fasting blood glucose, raised triglycerides or decreased HDL-C are present in an individual.

In this study 100 type 2 diabetics were studied for the presence of metabolic syndrome. The frequency of metabolic syndrome was 76% which is in accordance with other studies, i.e., prevalence of 70-80% among Caucasian type 2 diabetics<sup>13</sup> and 75.6% among Chinese

population with type 2 diabetes mellitus<sup>14</sup> were estimated. In our study metabolic syndrome was found to be more common in female type 2 diabetics as compared to their male counterparts. Different studies showed quite varied effects of gender on the metabolic syndrome in different populations. In USA, metabolic syndrome is more prevalent in white males.<sup>15</sup> In American blacks, Mexicans Americans, Korea, Kinmen, Iran, India and Oman, women had higher prevalence of the syndrome than men.<sup>16-19</sup> Nigerian women also have higher percentage than men.<sup>20</sup>

Prevalence of metabolic syndrome tends to increase with age.<sup>21</sup> This study also showed similar trend with 66.66% in age group 40-49 and increasing to 81.81% in age group ≥70 years. The prevalence of different risk factors in patients of both sexes was looked at. As all of them were diabetics, comparison was made for the presence of abdominal obesity, hypertension, low HDL and high triglycerides levels. Low HDL was most frequent while hypertriglyceridaemia was least frequent. Abdominal obesity was the second most prevalent factor. The combination of abdominal obesity and low HDL was also reported as the most common combination among Chinese type-2 diabetics with metabolic syndrome.<sup>22</sup> In Greece the most prevalent risk factor was abdominal obesity.<sup>23</sup>

WHO estimates the prevalence of obesity to 4.8% in less affluent countries, 17.1% in countries undergoing economic transition and 20% in the developed world.<sup>24</sup> In our study 56% of the participants were obese but the sample population was type-2 diabetics. This high incidence of obesity contributes to a very high frequency of metabolic syndrome in our patients, especially women. Physical inactivity and excess weight have been shown to be the main underlying contributors to the development of metabolic syndrome.<sup>25</sup> The goal of identifying metabolic risk factors is to prevent morbidity and mortality due to type 2 diabetes and cardiovascular disease. The role of family physicians is important as they are the first line of health care providers to encounter patients and to provide preventive measures besides taking steps to reverse metabolic syndrome in those who have a combination of risk factors.<sup>26</sup> Since physical inactivity and excessive weight gain are the main underlying contributors to the development of metabolic syndrome, getting more exercise and losing weight can help reduce or prevent the complications associated with this condition. Most of the risk factors are preventable and identifying these at an early stage at screening clinics and managing them accordingly might be of significant help as done elsewhere.<sup>27</sup>

It is evident from this study that metabolic syndrome is quite prevalent in our type 2 diabetic patients like in other parts of the world. It is very important to diagnose it early to prevent the life

threatening cardiovascular complications. Counselling and education on life-style modification, exercise and diet are paramount for diabetics in particular and obese people in general to decrease the risks associated with the metabolic syndrome. The limitations of our study are in large part, related to the cross-sectional design of the study. The study is limited to reporting disease prevalence data and the incidence of disease occurrence in this population cannot be inferred from this study.

## CONCLUSION

Frequency of MetS was found to be significantly high in this study with female preponderance. All the components, except Hypertension were more frequent in females. Diabetic patients with metabolic syndrome need more aggressive approach in management so as to decrease the incidence of cardiovascular complications.

## ACKNOWLEDGEMENT

Special thanks to all colleagues of Medical 'B' for their help. We are also grateful to laboratory staff of Department of Pathology, Ayub Teaching Hospital, Abbottabad, for carrying out the tests.

## REFERENCES

1. Alberti KG, Zimmet PZ. Definition, diagnosis and classification of diabetes mellitus and its complications. Part 1; Diagnosis and classification of diabetes mellitus; Provisional report of a WHO consultation. *Diabet Med* 1998;15:539-47.
2. David MN, John BB, Mayer BD, Robert JH, Rury RH, Robert S, Bernard Z. Management of Hyperglycemia in Type 2 Diabetes: A consensus algorithm for initiation and adjustment of therapy; A consensus statement from the American Diabetes Association and the European Association for the Study of Diabetes. *Diabetes Care* 2006;29:1963-72.
3. Basit A, Hydrie MZ, Hakeem R, Ahmedani MY, Masood Q. Frequency of chronic complications of type II diabetes. *J Coll Physicians Surg Pak* 2004;14:79-83
4. Chaudhary GM. Metabolic syndrome X in diabetic patients. Experience in 3275 diabetic patients at Jinnah Hospital, Lahore. *J Coll Physicians Surg Pak* 2000;10:278-80.
5. Scott CL. Diagnosis, prevention and intervention for the metabolic syndrome. *Am J Cardiol* 2003;92:35i-42i.
6. Wilson PWF, Kannel WB, Silbershatz H, D'Agostino RB. Clustering of metabolic factors and coronary heart disease. *Arch Intern Med* 1999;159:1104-9.
7. Khwaja AK, Rafique G, White F. Macrovascular complications and their associated factors among persons with type 2 diabetes in Karachi, Pakistan. *J Pak Med Assoc* 2004;54:60-6.
8. Executive Summary of the Third Report of the National Cholesterol Education Program (NCEP), Expert Panel on Detection, Evaluation and Treatment of High Blood Cholesterol in Adults (Adult Treatment Panel III). *JAMA* 2001;285:2486-97.
9. Shera A, Rafique G, Khawaja I. Pakistan National Diabetes Survey. Prevalence of glucose intolerance and associated factors in North West Frontier Province (NWFP) of Pakistan. *J Pak Med Assoc* 1999;49:206-11.
10. World Health Organization. Definition, diagnosis and classification of diabetes mellitus and its complications. Report of a WHO consultation. Part 1; Diagnosis and classification of diabetes mellitus. Geneva: World Health Organization. 1999. (Publ. No. WHO/NCD/NCS/00.2).
11. International Diabetes Federation. The IDF consensus worldwide definition of the metabolic syndrome. Part 1: Worldwide definition for use in clinical practice. Berlin 2005. Available at: [http://www.idf.org/webdata/docs/IDF\\_Metasyndrome\\_definition.pdf](http://www.idf.org/webdata/docs/IDF_Metasyndrome_definition.pdf).
12. Grundy SM, Cleeman JI, Daniels SR, Donato KA, Eckel RH, Franklin BA, *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.
13. Abdul-Rahim HF, Husseini A, Bjertness E, Gordon NH, Jervell J. The metabolic syndrome in the West Bank population: an urban-rural comparison. *Diabetes Care* 2001;24:275-9.
14. Bruno G, Merletti F, Biggeri A, Bargreo G, Ferrero S, Runzo C, *et al.* Metabolic syndrome as a predictor of all-cause and cardiovascular mortality in type 2 diabetes: the Casale Monferrato Study. *Diabetes Care* 2004;27:2689-94.
15. Park YW, Zhu S, Palaniappan L, Heshka S, Carnethon MR, Heymsfield SB. The metabolic syndrome: prevalence and associated risk factor findings in the US population from the Third National Health and Nutrition Examination Survey, 1988-1994. *Arch Intern Med* 2003;163:427-36.
16. Chuang SY, Chen CH, Tsai ST, Chou P. Clinical identification of the metabolic syndrome in Kinmen. *Acta Cardiol Sin* 2002;18:16-23.
17. Azizi F, Salehi P, Eternadi A, Zahedi AS. Prevalence of metabolic syndrome in an urban population; Tehran Lipid and Glucose Study. *Diabetes Res Clin Pract* 2003;61:29-37.
18. Gupta A, Gupta R, Sarna M, Rastogi S, Gupta VP, Kothari K. Prevalence of diabetes, impaired fasting glucose and insulin resistance syndrome in an urban Indian population. *Diabetes Res Clin Pract* 2003;61:69-76.
19. Park JS, Park HD, Yun JW, Jung CH, Lee WY, Kim SW. Prevalence of the metabolic syndrome as defined by NCEP-ATP III among the urban Korean population. *Korean J Med* 2002;63:290-8.
20. Isezuo SA, Ezunu E. Demographic and clinical correlates of metabolic syndrome in Native African type 2 diabetic patients. *J Natl Med Assoc* 2005;97:557-63.
21. Thomas GN, Ho SY, Janus ED, Lam KS, Hedly AJ, Lam TH. The US National Cholesterol Education Programme Adult Treatment Panel III (NCEP-ATP III) prevalence of the metabolic syndrome in a Chinese population. *Diabetes Res Clin Pract* 2005;67:251-7.
22. Lee YJ, Tsai JC. ACE gene insertion/deletion polymorphism associated with 1998 World Health Organization definition of metabolic syndrome in a Chinese type 2 diabetic patients. *Diabetes Care* 2002;25:1002-8.
23. Athyros VG, Bouloukos VI, Pehlivanidis AN, Dionysopoulou SG, Symeonidis AN. The prevalence of the metabolic syndrome in Greece: The MetS-Greece Multicentre Study. *Diabetes Obes Metab* 2005;7:397-405.
24. Islam N. Obesity: An epidemic of 21<sup>st</sup> century. *J Pak Med Ass* 2005;55:118-22.
25. Lopez-Candales A. Metabolic syndrome X. A comprehensive review of the pathophysiology and recommended therapy. *J Med* 2001;32:283-300.
26. Grundy SM. Obesity, metabolic syndrome and coronary atherosclerosis. *Circulation* 2002;105:2696-8.
27. Kevin E, Oscar C Kip. Clinical importance of obesity versus the metabolic syndrome in cardiovascular risk in women. *Circulation* 2004;109:706-13.

## Address for correspondence:

**Dr. Nasir Ahmed**, Assistant Professor of Medicine, Ayub Medical College, Abbottabad, Pakistan. **Cell:** +92-300-9111583  
**Email:** drnaseertanoli61@yahoo.com