Association between metabolic syndrome and thyroid stimulating hormone - A cross sectional study

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<u>Abstract</u>

Background: Thyroid dysfunction especially hypothyroidism and Metabolic Syndrome (MeS) are two entities which affect blood pressure, fasting plasma glucose, serum triglycerides and HDL-Cholesterol. Both are associated with increased chance of cardiovascular disease. **Methodology:** This was a cross sectional study done among 98 patients with metabolic syndrome in a tertiary care hospital in South India with an aim to find prevalence of hypothyroidism and its association with components of metabolic syndrome. **Results:** Majority 92(92.9%) were females in our study. 89.2% had history of Diabetes Mellitus (DM) and 62.2% had history of Hypertension (HTN). Among the subjects 39(40%) had subclinical hypothyroid state and 11(11%) had overt hypothyroidism. In our study we found that thyroid dysfunction that is hypothyroidism is significantly present in middle aged people (40-49 years), over weight patients and female gender. In our study we found that mean Waist circumference, diastolic BP and fasting blood glucose were more among thyroid dysfunction group compared to that of Euthyroid state **Conclusion:** Thyroid dysfunction particularly hypothyroidism is associated with metabolic syndrome and has to be seriously viewed as a threat in long term.

Key Word: Hypothyroidism, Metabolic Syndrome, Thyroid Stimulating Hormone, Cardiovascular disorders

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Received Date: 16/11/2018 Revised Date: 12/12/2018 Accepted Date: 13/01/2019 DOI: https://doi.org/10.26611/1021922

Access this article online			
Quick Response Code:	Website:		
	www.medpulse.in		
	Accessed Date: 01 February 2019		

INTRODUCTION

Metabolic syndrome (MeS) – A complex mechanism of insulin resistance (IR) and compensatory release of more insulin leading to constellation of obesity (particularly central adiposity), hyperglycaemia, elevated blood pressure, dyslipidaemias (hypertriglyceridemia, decreased High Density-Lipoprotein Cholesterol (HDL-C) and

increased Low Density Lipoprotein cholesterol (HDL-C)), and multiple pro-inflammatory conditions.(1-3) The prevalence of metabolic syndrome is increasing worldwide with an 2.5-fold increase in cardiovascular mortality and a 5-fold higher risk of developing diabetes.^{2,4,5,6} It is also associated with increased risk of kidney disease and mortality due to all causes.⁷ Subclinical hypothyroidism is an asymptomatic condition with high serum thyroid stimulating hormone (TSH) and normal free thyroid hormone levels. It is a prevalent condition among adult population which is frequently overlooked. Subclinical hypothyroidism has also been suggested as a risk factor for atherosclerosis, hyperlipidaemia, hypertension, low grade inflammation and hypercoagulability.8 Thyroid dysfunction especially hypothyroidism is associated with all parameters of metabolic syndrome except increase in fasting blood glucose.⁹ More than 60% of hypothyroid patients have obesity; there is decrease in basal metabolic rate and

How to cite this article: Karanam Madhuri *et al.* Association between metabolic syndrome and thyroid stimulating hormone - A cross sectional study. *MedPulse International Journal of Medicine*. February 2019; 9(2): 87-90. <u>https://www.medpulse.in/Medicine/</u>

metabolism.¹⁰ hypothyroidism, energy In the hemodynamic alterations cause narrowing of pulse pressure, prolongation of circulatory time and decrease in blood flow to the tissues leading to increased systemic vascular resistance and hypertension.11,12 Both the synthesis and degradation of lipids are affected in hypothyroidism and the activity of cholesterol ester transfer protein is decreased thus HDL cholesterol level is reduced.¹³ As metabolic syndrome and hypothyroidism are independent risk factors for the same disease process namely cardiovascular disease, it is possible that patients suffering from both these disease entities may have a compounded risk. Hence this study was conducted to explore the association between thyroid dysfunction and components of metabolic syndrome.

OBJECTIVES

- 1. To study the prevalence of hypothyroidism among metabolic syndrome patients attending a tertiary care hospital.
- 2. To know the association of hypothyroidism and components of metabolic syndrome among patients attending a tertiary care hospital.

METHODOLOGY

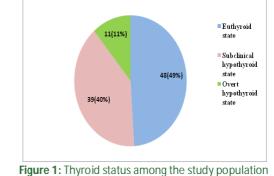
This was a cross sectional study done among 98 patients with metabolic syndrome in a tertiary care hospital in South India. All the patients were interviewed and physical examination was done. Informed consent was obtained from all the participants prior to start the study. Blood pressure (BP) was measured using an automated sphygmomanometer, with the patient in the sitting position before the blood test. Body mass index (BMI) was calculated as weight (kg) divided by height squared (m²) and BMI >25 kg/m² was considered as overweight and >30 kg/m² as obese. Waist circumference (WC) was measured with the measuring tape positioned midway between the lowest rib and the superior border of the iliac crest while the patient exhaled normally. The blood sample was collected in the morning after an 8-12-hour fast. Levels of glucose, Total cholesterol (TC), High density lipoprotein cholesterol (HDL-C), Low-density lipoprotein cholesterol (LDL-C), and Triglyceride (TG), Very Low-density lipoprotein cholesterol (VLDL-C), were determined in the hospital laboratory using standard methods. The components that contribute to metabolic syndrome were defined as high BP ($\geq 130/85$ mmHg), Ttruncal obesity (WC > 90 cm for men, > 80 cm for women). Hypertriglyceridemia (>150mg/dL or 1.7mmol/L), Low HDL-C (<40mg/ dL or 1.0mmol/L for men, < 50mg/ dL or 1.3mmol/L for women) and Hyperglycaemia (fasting blood glucose level ≥ 110 mg/ dL or 6.1 mmol/L).^{14,15} Thyroid stimulating hormone (TSH) was estimated by the electro-chemiluminescence immune assay (ECLIA). Normal range for TSH was $0.27-4.2 \mu$ IU/ml. A high serum TSH level (4.2–10 μ IU/ml) was required for the diagnosis of subclinical hypothyroidism. Patients with high TSH (>10 μ IU/ml) was classified as being overt or clinical hypothyroid. Patients with normal TSH, T3, and T4 were considered euthyroid.¹⁶ Demographic data was summarised descriptively. Continuous variables were expressed as mean \pm standard deviation. Categorical variables were presented as percentages.

RESULTS

A total of 98 subjects were included in the study with mean (SD) age of 50.09 (10.14) years. Among the 98 subjects, 30(30.6%) belonged to the age group of 40 to 49 years. Majority 92 (93.9%) were females in the study. Among the subjects 88(89.8%) had history of Diabetes Mellitus (DM) and 61 (62.2%) had history of Hypertension (HTN). Majority 75(76.5%) were overweight with a Body Mass Index (BMI) between 25-29.9 kg/m². The mean (SD) BMI was 27.96(2.23) with a minimum of 23 and maximum of 35 kg/m².

 Table 1: Descriptive statistics of variables among the study

Variable	Category	Frequency (%)
Age(years)	<40	21(21.4%)
	40-49	30(30.6%)
	50-59	19(19.4%)
	>60	28(28.6%)
Gender	Male	6(6.1%)
	Female	92(93.9%)
Presence of	Yes	88(89.8%)
Diabetes Mellitus	No	10(10.2%)
Presence of	Yes	61(62.2%)
Hypertension	No	37(37.8%)
Body Mass Index (Kg/m ²)	Normal (18.5-24.9)	4(4.1%)
	Overweight(25-29.9)	75(76.5%)
	Obese(>30)	19(19.4%)



Among the subjects 39(40%) had subclinical hypothyroid state and 11(11%) had overt hypothyroidism. Mean TSH

level was $6.89\pm7.67 \mu$ IU/ml with a minimum of 1.00 and maximum of 55 μ IU/ml. In our study we found that thyroid dysfunction that is hypothyroidism is significantly present in middle aged people (40-49 years), over weight patients and female gender. Thyroid dysfunction was present more in hypertensives and diabetics. But the association is not statistically significant. (Table2). In our study we found that mean Waist circumference, diastolic BP and fasting blood glucose were more among thyroid dysfunction group compared to that of Euthyroid state. (table 3)

Table 2: Association between TSH levels and different variables

Variable	Category	Euthyroid state	Thyroid dysfunction	Chi square value	p value
	<40	11(52.4%)	10(47.6%)		
Age	40-49	8(26.7%)	22(73.3%)	17.01	0.001ª*
(years)	50-59	7(36.8%)	12(63.2%)	17.01	0.0014
	>60	22(78.6%)	6(21.4%)		
BMI	18.5-24.9	4(100%)	0		<0.001b
	25-29.9	43(57.3%)	32(42.7%)	22.32	<0.001*
(Kg/ m²)	>30	1(5.3%)	18(94.7%)		
Gender	Male	6(100%)	0		
Gender	Female	42(45.65 %)	50(54.35%)	7.50	0.006 ^{b*}
HTN	Yes	32(52.46 %)	29(47.54%)	1.83	0.32 ^b
	No	16(43.2%)	21(56.8%)		
DM	Yes	44(50%)	44(50%)	1.32	0.42 ^b
	No	4(40%)	6(60%)	1.52	0.42

* p value,0.05 is significant a- chi-square test, b-Fischer's exact test

Table 3: Components of metabolic syndrome among Euthyroid and thyroid dysfunction group

Parameters	Euthyroid state	Thyroid dysfuction
Systolic BP(mmHg)	145.17 ±9.41	140.72±2.01
Diastolic BP(mm Hg)	89.17±8.23	91.44±5.69
Waist Circumference(cm)	100.58±8.55	109.48±8.81
Triglyceride(mg/dl)	180.71±16.90	180.70±18.87
Fasting blood glucose(mg/dl)	148.15±27.29	156.22±28.69
HDL cholesterol(mg/dl)	37.19±9.25	36.40±4.45

DISCUSSION

In our study we aimed to identify thyroid dysfunction patients particularly hypothyroid among metabolic syndrome patients attending tertiary care hospital. Metabolic syndrome by itself is related to may endocrine and non endocrine disease and can predict the development of those diseases. In our study among the subjects 39(40%) had subclinical hypothyroid state and 11(11%) had overt hypothyroidism. This finding is

similar to many studies.^{17,19} In a study¹⁷ among metabolic syndrome patients in Nepal identified thyroid dysfunction as a common endocrine disorder in metabolic syndrome patients; subclinical hypothyroidism (26.6 %) was the commonest followed by overt hypothyroidism (3.5 %) and subclinical hyperthyroidism (1.7 %). In our study we found that thyroid dysfunction that is hypothyroidism is significantly present in middle aged people (40-49 years), over weight patients and female gender. The mean age of our study population was 50.09±10.14 years. Similar mean age and peak incidence among >40 years is found in many studies.^{20,21} Increased BMI and thyroid dysfunction is a common feature which is demonstrated in many studies.^{22,23} As previously demonstrated overweight and obesity can accelerate the development of endocrine disorders. In our study we found that mean Waist circumference, diastolic BP and fasting blood glucose were more among thyroid dysfunction group compared to that of Euthyroid state. The link between waist circumference and thyroid dysfunction was demonstrated by other studies^{19,20} also. Increased waist circumference denotes abdominal obesity which eventually leads to many endocrine disorders. Demonstration of raised diastolic BP among thyroid dysfunction patients denotes a increased probability of cardiovascular disorders. It has been reported that 95 % of newly diagnosed hypothyroid patients have increased levels of cholesterol and 5 % of have hyper triglyceridemia. Hypothyroidism also leads to increased level of LDL cholesterol. All these factors directly contribute to accelerated atherosclerosis.²⁴ Positive association of metabolic syndrome and thyroid dysfunction is demonstrated by many studies^{17,18,20} whereas one study²⁵ particularly rejects the link between the two.

CONCLUSION

In our study we found that majority of metabolic syndrome patients had hypothyroidism. Also we found that thyroid dysfunction that is hypothyroidism is significantly present in middle aged people (40-49 years), over weight patients and female gender. High prevalence of overt and subclinical hypothyroidism in metabolic syndrome as seen in our study may have harmful effect on cardiovascular health. Hypothyroidism will lead to increased lipid levels and hypertension leading to increased risk for CVD. Our study lacks the presence of a control group to get a better view about the presence of hypothyroid state in metabolic syndrome patients. Also inclusion of dietary and exercise factors would have added up more value.

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Source of Support: None Declared Conflict of Interest: None Declared