SUBCLINICAL THYROID DISORDERS AND ITS ASSOCIATION WITH DYSLIPIDAEMIA AT A TERTIARY CARE INSTITUTE

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ABSTRACT

BACKGROUND
Subclinical thyroid disorders with altered Thyroid Stimulating Hormone (TSH) level, yet normal circulating levels of free thyroid hormones (FT4) include subclinical hypothyroidism and subclinical hyperthyroidism. The prevalence of subclinical thyroid disorders is found to be increasing in our country and subclinical hypothyroidism is known to be linked with hypercholesterolemia. Hence, we intend to find the prevalence of subclinical thyroid diseases and its association with lipid profile.

MATERIALS AND METHODS
This study was done on 253 female medical students of PES institute of medical Sciences and research who were willing to participate in the study. Free T4 (FT4) and Thyroid Stimulating Hormone (TSH) levels were estimated by Enzyme Linked Immuno-Fluorescent Assay (ELFA). Lipid Profile of the subjects including total cholesterol, serum triglyceride, HDL, LDL, VLDL was estimated using VITROS instrument. Based on the levels of TSH and FT4, the students were categorized as euthyroid, subclinical hypothyroid and subclinical hyperthyroid.

RESULTS
The prevalence of subclinical hypothyroidism and subclinical hyperthyroidism was found to be 8.3% and 1.58% respectively among the study population. A statistically significant decrease in FT4 levels as the TSH levels increased was seen. As the TSH value increased, HDL levels significantly decreased, and the LDL levels were found to be increased significantly.

CONCLUSION
Prevalence of subclinical hypothyroidism is found to be high in our study. The students with subclinical hypothyroidism were found to have significantly increased levels of LDL and decreased levels of HDL, which is an important risk factor for atherosclerosis, hypertension and coronary heart disease. Hence early detection of subclinical thyroid disorders would help us prevent the individual from developing the overt disease and its complications.

KEYWORDS
Subclinical Hypothyroidism, Subclinical Hyperthyroidism, Atherosclerosis, Coronary Heart Disease.


BACKGROUND
Subclinical Hypothyroid and subclinical Hyperthyroid individuals are asymptomatic. These thyroid disorders can be diagnosed only by measuring the levels of TSH and FT4.

Individuals with elevated TSH and normal FT4 levels are diagnosed as subclinical Hypothyroid and those with decreased TSH and FT4 within the normal range as subclinical hyperthyroid.1,2 Advancing age, female gender and excess intake of dietary iodine are the risk factors for developing subclinical hypothyroidism.3 Several studies have established that subclinical hypothyroidism is the most prevalent thyroid disorder, affecting 3-15% of the adult population.4

Thyroid hormones play an important role in regulating lipid metabolism. It has been thought that subclinical hypothyroidism could be associated with altered lipid profile.5 Several studies have also reported that subclinical hypothyroidism leads to hyperlipidemia.6-8
Recent studies show that subclinical hypothyroidism is linked with an increased risk of coronary heart disease (CHD) events and CHD mortality in those with higher TSH levels.\(^\text{10}\) As subclinical hypothyroidism has been associated with hypercholesterolemia\(^\text{11}\) and atherosclerosis,\(^\text{12}\) screening of subclinical hypothyroid individuals for LDL, VLDL, Total cholesterol, Triglycerides and HDL and treatment has to be advocated to prevent cardiovascular disease.\(^\text{13}\) The changes in lipid metabolism and cardiovascular physiology are alarming, not only in the context of untreated cases, but also among patients with L-thyroxine substitution therapy.\(^\text{14}\)

As the individuals with subclinical thyroid disorders do not present with symptoms, screening the general population, especially the female population for their TSH, FT\(_4\) and lipid profile would greatly reduce the incidence of overt thyroid disorders and the complications associated with it. Hence, we intend to find the prevalence of subclinical thyroid diseases and its association with lipid profile.

**Aims and Objectives**
1. To determine the prevalence of subclinical hypothyroidism and subclinical hyperthyroidism among female medical students.
2. To determine the association of subclinical hypothyroidism with Dyslipidaemia.

**MATERIALS AND METHODS**
This study was done on 253 female medical students within the age group of 18-25 years of PES Institute of medical Sciences and Research, Kuppam who were willing to participate in the study. After obtaining institutional ethical clearance, written informed consent was obtained from the participants. All the individuals participating in the study were instructed to give 5 ml of venous blood after overnight fasting. TSH and FT\(_4\) levels were estimated by Enzyme linked immuno-fluorescent assay (ELFA) using VIDAS FT\(_4\) and TSH kit. The normal ranges of TSH and FT\(_4\) were 0.25-5.5 \(\mu\)IU/ml and 9-24 pmol/L respectively. Asymptomatic individuals with TSH more than 5.5 \(\mu\)IU/ml and FT\(_4\) within the normal range were diagnosed as subclinical hypothyroidism and asymptomatic individuals with TSH less than 0.25 \(\mu\)IU/ml and FT\(_4\) within the normal range were diagnosed as subclinical Hyperthyroid.

Lipid Profile of the subjects including total cholesterol, serum triglyceride, HDL, LDL, VLDL were estimated using VITROS instrument.

**Inclusion Criteria**
Students willing to participate in the study within the age group 18-25 years.

**Exclusion Criteria**
Subjects with ischemic heart disease, cerebrovascular and neurological diseases, Diabetes mellitus, chronic renal impairment, known psychological illnesses, previous H/O thyroid disease or previous thyroxine therapy were excluded.

The data was entered into MS Excel 2007 version and the values of the FT\(_4\), TSH, Total cholesterol, serum triglyceride, HDL, LDL, VLDL was expressed as Mean± SD and prevalence of subclinical Hypothyroid and subclinical Hyperthyroid individuals was expressed in percentage. Correlation of TSH with lipid profile was performed by Pearson’s correlation and simple regression analysis.

**RESULTS**
This study was carried out among 253 female medical students. Table 1 shows the Mean ± SD values of Age, TSH, FT\(_4\) and lipid profile. The prevalence of subclinical hypothyroidism and subclinical hyperthyroidism were found to be 8.3% and 1.58% respectively as shown in table 2. Pearson’s correlation (table 3) and simple regression analysis (Figure 1 - Figure 6) were done. A statistically significant decrease in FT\(_4\) levels as the TSH levels increased was seen. As the TSH value increased, HDL levels significantly decreased, and the LDL levels were found to be increased significantly.

### Table 1. Anthropometric and Baseline Biochemical Parameters of the Study Population

<table>
<thead>
<tr>
<th>Parameters</th>
<th>Total Number</th>
<th>Prevalence among Study Population</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sub-clinical Hyperthyroidism</td>
<td>4</td>
<td>1.58%</td>
</tr>
<tr>
<td>Euthyroid</td>
<td>228</td>
<td>90.12%</td>
</tr>
<tr>
<td>Sub-Clinical Hypothyroidism</td>
<td>21</td>
<td>8.3%</td>
</tr>
<tr>
<td>Total Study Population</td>
<td>253</td>
<td></td>
</tr>
</tbody>
</table>

### Table 2. Prevalence of Sub-Clinical Hyperthyroidism, Euthyroid and Sub-Clinical Hypothyroidism among the Study Population

<table>
<thead>
<tr>
<th>Parameters</th>
<th>r-Value</th>
<th>p-Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>FT(_4) (pmol/L)</td>
<td>-0.31</td>
<td>&lt;0.001**</td>
</tr>
<tr>
<td>Total Cholesterol (mg/dl)</td>
<td>0.10</td>
<td>0.08</td>
</tr>
<tr>
<td>Serum TGL (mg/dl)</td>
<td>0.08</td>
<td>0.19</td>
</tr>
<tr>
<td>HDL (mg/dl)</td>
<td>-0.14</td>
<td>0.02*</td>
</tr>
<tr>
<td>LDL (mg/dl)</td>
<td>0.14</td>
<td>0.02*</td>
</tr>
<tr>
<td>VLDL (mg/dl)</td>
<td>0.06</td>
<td>0.28</td>
</tr>
</tbody>
</table>

* ** P value <0.05 is Statistically Significant.
Figure 1. Correlation between TSH and FT₄ among the Study Population
Correlation Coefficient = -0.28, p = <0.001**

As the TSH value increases the FT₄ levels decreases and it is found to be statistically significant among the study population.

Figure 2. Correlation between TSH and Total Cholesterol among the Study Population
Correlation Coefficient = 0.009, p = 0.08

As the TSH value increases the Total Cholesterol levels increases among the study population, but not found to be statistically significant.

Figure 3. Correlation between TSH and Serum Triglyceride among the Study Population
Correlation Coefficient = 0.005, p = 0.19

As the TSH value increases the serum Triglyceride levels increases among the study population, but not found to be statistically significant.

Figure 4. Correlation between TSH and HDL among the Study Population
Correlation Coefficient = -0.037, p = 0.02*

As the TSH value increases the HDL levels decreases and it is found to be statistically significant among the study population.

Figure 5. Correlation between TSH and LDL among the Study Population
Correlation Coefficient = 0.01, p = 0.02*

As the TSH value increases the LDL levels increases and it is found to be statistically significant among the study population.

Figure 6. Correlation between TSH and VLDL among the Study Population
Correlation Coefficient = 0.02, p = 0.28

As the TSH value increases the VLDL levels increases among the study population but not found to be statistically significant.
DISCUSSION
The prevalence of subclinical hypothyroidism and subclinical hyperthyroidism were found to be 8.3% and 1.58% respectively in our study showing that subclinical hypothyroidism is more common than subclinical hyperthyroidism. It is also established that subclinical hypothyroidism is more common than overt hypothyroidism.\(^1\),\(^\text{14,15,16}\) 8.02% of the study population was found to have subclinical hypothyroidism in an epidemiological study in eight cities of India.\(^1\) Another study reports 9.4% prevalence of subclinical hypothyroidism and it was noted that women were affected more than men. It was also found that the prevalence of subclinical hypothyroidism increased with age.\(^1\) The progression of subclinical hypothyroidism to overt hypothyroidism is definite among 33- 55% of cases in 10 years.\(^1\)

Thyroid hormones play a very important role in the regulation, absorption, and metabolism of lipid synthesis. Several studies have reported the link between subclinical hypothyroidism and alteration in the lipid profile. Abnormal lipid profile is considered as an independent risk factor for cardiovascular diseases. Hence its association with subclinical hypothyroidism may imply an increase in the risk for cardiovascular diseases like coronary heart disease. It has been established that total cholesterol, LDL levels will be higher in subclinical Hypothyroid individuals.\(^3\) In our study, a linear increase in LDL levels and a linear decrease in HDL levels with increasing TSH was seen which is similar to other studies.\(^20,21\) Rise of serum LDL levels is mainly due to impaired clearance of LDL, probably reflecting decreased LDL receptor expression.\(^22\)

Subclinical hypothyroidism has been associated with increased systemic vascular resistance, arterial stiffness, altered endothelial function, increased atherosclerosis and altered coagulability which in turn may accelerate development of coronary heart disease.\(^10\) Therefore screening the young adult females for TSH, FT\(_4\) and lipid profile would greatly reduce the burden of overt hypothyroidism and the complications associated with it like atherosclerosis, hypertension, coronary heart disease.

Further study can be undertaken to know the relationship between dietary habits like excessive intake of iodized salt and its relationship to development of subclinical Hypothyroidism.

CONCLUSION
Subclinical hypothyroidism is found to be more common than overt hypothyroidism or subclinical hyperthyroidism. The individuals with subclinical hypothyroidism were found to have significantly increased levels of LDL and significantly decreased levels of HDL. Subclinical hypothyroidism may be considered as potentially modifiable risk factor for cardiovascular diseases. Hence its early detection would prevent the individual from developing the overt disease and its complications.

REFERENCES


