

Investigational Profile in Case of Subclinical Hypothyroidism and its Effect on Cardiovascular System

Dr. Neha Sharma¹, Dr. R. V. Dosi², Dr. Hardik Rathwa³, Dr. Satish Gurjar⁴

¹Senior Resident

²Professor and Head

³rd Year Resident

⁴rd Year Resident

Abstract: ***Introduction:** Subclinical hypothyroidism (also known as compensated hypothyroidism or mild hypothyroidism) is a condition associated with a raised serum concentration of thyroid stimulating hormone (TSH) but a normal serum free thyroxine (FT4). Hypothyroidism leads to hypercholesterolemia due to increased level of LDL cholesterol. **Aim:** to determine ECG, ECHO and lipid profile changes in case of subclinical hypothyroidism. **Methodology:** This was a cross sectional observational study conducted on 40 patients at outpatient medicine department of SSG hospital, Vadodara. **Results:** In majority of patients, total cholesterol, HDL, LDL and VLDL were deranged. Triglycerides were within normal limits. 24 (60.00%) patients had grade 1 LV diastolic dysfunction. No abnormality was detected in only 16 out of 40 patients (40.00%). **Conclusion:** There is association between Lipid Profile changes and 2D Echo changes. There is no major ECG change in patients with subclinical hypothyroidism. Subclinical Hypothyroidism can be considered as independent risk factor of cardiovascular system morbidity.*

Keywords: Hypothyroidism, lipid profile, cardiac changes,

1. Introduction

Subclinical hypothyroidism (also known as compensated hypothyroidism or mild hypothyroidism) is a condition associated with a raised serum concentration of thyroid stimulating hormone (TSH) but a normal serum free thyroxine (FT4).¹ Prevalence of subclinical hypothyroidism is 6 - 8% in women (10% over 60 year of AGE) and 3% in men.

Though subclinical hypothyroidism can affect various organ system Cardio vascular system is a major target.^{2, 3, 4} Hypothyroidism produces a decrease in myocardial contractility, pericardial effusion, increase in left ventricular mass and prolonged duration of contraction and relaxation. When lasting more than 6 - 12 months, SH may be associated with an atherogenic lipid profile, a hypercoagulable state, a subtle cardiac defect with mainly diastolic dysfunction, impaired vascular function, and reduced submaximal exercise capacity. The deviation from normality usually increases with serum TSH level ('dosage effect' phenomenon).^{5, 6}

Hypothyroidism leads to hypercholesterolemia due to increased level of LDL cholesterol, Total cholesterol and triglycerides. Restoration of euthyroidism by levothyroxine (LT4) treatment may correct the lipid profile and cardiac abnormalities, especially in patients with an initially higher deviation from normality and higher serum TSH levels.^{7, 8} Importantly, a strong association between SH and atherosclerotic cardiovascular disease, independent of the traditional risk factors, has been recently reported in a large cross - sectional survey (the Rotterdam Study). However,

whether SH confers a high risk for cardiovascular disease, and whether LT4 therapy has a long - term benefit that clearly outweighs the risks of overzealous treatment in these individuals, The current study was planned to determine ECG, ECHO and lipid profile changes in case of subclinical hypothyroidism.

2. Methodology

This was a cross sectional observational study conducted on 40 patients at outpatient medicine department of SSG hospital, Vadodara. Following were the inclusion and exclusion criteria.

Inclusion Criteria: Any patients having subclinical hypothyroidism attending OPD of medicine department with age more than 18 years.

Exclusion Criteria: The following patients were excluded from the study

- 1) Patients with known cardiac disorder
- 2) Patients who had hypertension
- 3) Diabetes mellitus,
- 4) Renal failure
- 5) Pregnancy
- 6) Hyperthyroidism on treatment
- 7) Cholesterol lowering agents and on drugs like Amiodrane, lithium & levothyroxine
- 8) Age less than 18 years

3. Results

40 patients with subclinical hypothyroidism were included in the study. ECG findings, ECHO findings and lipid profile were noted. 36 (90.00%) patients were females and 4 (10.00%) patients were males.

Table 1: Distribution of demographic characteristics of study subjects

| Demographic characteristics | Frequency | Percentage |
|-----------------------------|-----------------|------------|
| Gender | | |
| Female | 36 | 90.00% |
| Male | 4 | 10.00% |
| Age(years) | | |
| Mean±SD | 35.9±11.6 | |
| Median(25th-75thpercentile) | 35(27.25-45.25) | |
| Range | 18-65 | |

Table 2: Descriptive statistics of thyroid profile of study subjects

| Thyroid Profile | Mean± SD | Median (25th-75 th percentile) | Range |
|--------------------|------------|---|-----------|
| Serum TSH (mIU/mL) | 7.8±1.18 | 7.7(7.075-8.425) | 5.6-9.8 |
| Free T3 (pg/mL) | 3.83 ±0.36 | 3.8(3.548-4.12) | 3.12-4.43 |
| Free T4 (ng/dL) | 1.3±0.23 | 1.34(1.115-1.462) | 0.86-1.67 |

Mean value of serum TSH (mIU/mL), free T3 (pg/mL) and free T4 (ng/dL) of study subjects was 7.8 ± 1.18, 3.83 ± 0.36 and 1.3 ± 0.23 with median (25th - 75th percentile) of 7.7 (7.075 - 8.425), 3.8 (3.548 - 4.12) and 1.34 (1.115 - 1.462) respectively.

Table 3: Distribution of lipid profile of study subjects

| Lipid Profile | Frequency | Percentage |
|--|----------------------|------------|
| Total cholesterol (mg/dL) | | |
| Deranged | 23 | 57.50% |
| Normal | 17 | 42.50% |
| Mean±SD | 204.32±41.21 | |
| Median (25 th -75 th percentile) | 207.5(167.75-238.25) | |
| Range | 125-278 | |
| Triglyceride(mg/dL) | | |
| Deranged | 19 | 47.50% |
| Normal | 21 | 52.50% |
| Mean±SD | 144.05±50.83 | |
| Median (25 th -75 th percentile) | 142(99.75-169) | |
| Range | 74-278 | |
| HDL(mg/dL) | | |
| Deranged | 26 | 65.00% |
| Normal | 14 | 35.00% |
| Mean±SD | 41.5±12.16 | |
| Median (25 th -75 th percentile) | 37(33.5-48.25) | |

| Range | 28-79 | |
|---|----------------------|--------|
| LDL(mg/dL) | | |
| Deranged | 21 | 52.50% |
| Normal | 19 | 47.50% |
| Mean±SD | 121.33±46.43 | |
| Median(25 th -75 th percentile) | 112.5(89.338-163.55) | |
| Range | 49.4-215.6 | |
| VLDL(mg/dL) | | |
| Deranged | 14 | 35.00% |
| Normal | 26 | 65.00% |
| Mean±SD | 29.38±14 | |
| Median(25 th -75 th percentile) | 23.3(18.75-33.2) | |
| Range | 14.8-59 | |

In majority of patients, total cholesterol, HDL, LDL and VLDL were deranged. Triglycerides were within normal limits.

Table 4: Distribution of 2D Echo of study subjects

| 2DEcho | Frequency | Percentage |
|------------------------------|-----------|------------|
| No abnormality detected | 16 | 40.00% |
| Grade1LVdiastolicdysfunction | 24 | 60.00% |
| Total | 40 | 100.00% |

24 (60.00%) patients had grade 1 LV diastolic dysfunction. No abnormality was detected in only 16 out of 40 patients (40.00%).

Table 5: Distribution of ECG of study subjects

| ECG | Frequency | Percentage |
|-------------------------|-----------|------------|
| No abnormality detected | 35 | 87.50% |
| Low voltage complexes | 1 | 2.50% |
| Sinus brady cardia | 3 | 7.50% |
| Sinus tachy cardia | 1 | 2.50% |
| Total | 40 | 100.00% |

In majority (35 (87.50%)) of patients, no abnormality was detected in ECG findings followed by sinus bradycardia (3 (7.50%)). Low voltage complexes and sinus tachycardia was seen in only 1 out of 40 patients (2.50%) each.

Distribution of grade 1 LV diastolic dysfunction was comparable with lipid profile. (**Total cholesterol (mg/dL):** - Deranged (56.52%) vs Normal (64.71%) (p value=0.601), **Triglyceride (mg/dL):** - Deranged (73.68%) vs Normal (47.62%) (p value=0.093), **HDL (mg/dL):** - Deranged (57.69%) vs Normal (64.29%) (p value=0.685), **LDL (mg/dL):** - Deranged (52.38%) vs Normal (68.42%) (p value=0.301) and **VLDL (mg/dL):** - Deranged (64.29%) vs Normal (57.69%) (p value=0.685).

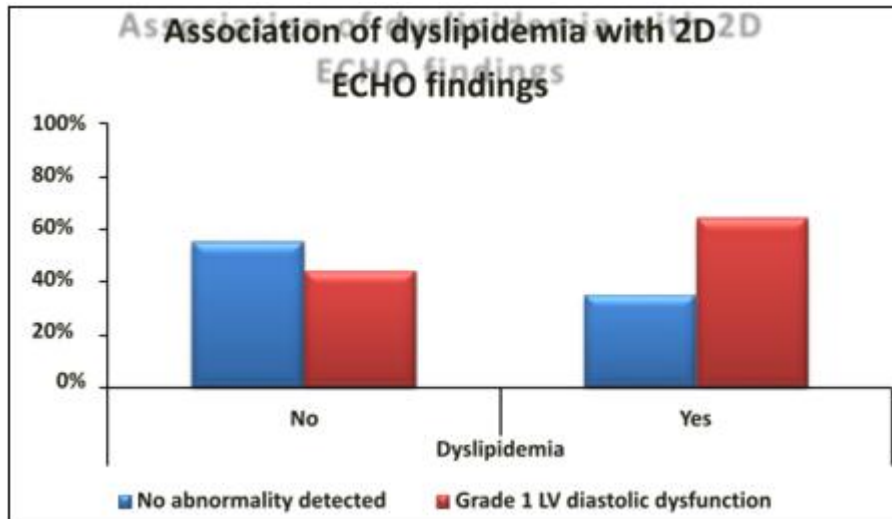


Figure 1: Association of dyslipidemia with 2D ECHO findings

Table 6: Association of lipid profile with ECG findings

| Lipid profile | No abnormality detected (n=35) | Abnormal (n=5) | Total | P value |
|--------------------------|--------------------------------|----------------|----------|---------|
| Total cholesterol(mg/dL) | | | | |
| Deranged | 19(82.61%) | 4(17.39%) | 23(100%) | 0.373* |
| Normal | 16(94.12%) | 1(5.88%) | 17(100%) | |
| Triglyceride(mg/dL) | | | | |
| Deranged | 15(78.95%) | 4(21.05%) | 19(100%) | 0.172* |
| Normal | 20(95.24%) | 1(4.76%) | 21(100%) | |
| HDL(mg/dL) | | | | |
| Deranged | 23(88.46%) | 3(11.54%) | 26(100%) | 1* |
| Normal | 12(85.71%) | 2(14.29%) | 14(100%) | |
| LDL(mg/dL) | | | | |
| Deranged | 18(85.71%) | 3(14.29%) | 21(100%) | 1* |
| Normal | 17(89.47%) | 2(10.53%) | 19(100%) | |
| VLDL(mg/dL) | | | | |
| Deranged | 12(85.71%) | 2(14.29%) | 14(100%) | 1* |
| Normal | 23(88.46%) | 3(11.54%) | 26(100%) | |

*Fisher's exact test

4. Discussion

Dyslipidemia in overt hypothyroidism is clearly established, and studies have shown beneficial effects of treatment with levothyroxine on dyslipidemia in these subjects.

However, studies on the effects of SCH on lipid profile and beneficiary effects of treatment in these patients are controversial. Hence, this topic is still open for further studies. There was female preponderance in the study with 90% female and 10% male which is comparable with study conducted by Mubashir Alam Khan et al.⁹

In study conducted by Manoj Kumar et al¹⁰, 87 patients with subclinical hypothyroidism (SCH) and 101 age and sex matched euthyroid controls were evaluated which showed following results: Total cholesterol (TC) and low density lipoprotein (LDL - C) are higher in patients with subclinical hypothyroidism (SCH) as compared to euthyroid individuals. Other lipid like Triglycerides (TG) and very low density lipoprotein cholesterol (VLDL - C) may be marginally elevated whereas high density lipoprotein cholesterol (HDL - C) may be slightly reduced in these patients as compared to euthyroid individuals which are comparable with my study. 2D Echo findings of my study

showed that 60% of patients have grade 1 LV diastolic dysfunction which was comparable with study conducted by Haridose Sripriya Vasudevan et al¹¹ in which a total of 84 patients with newly diagnosed drug naïve hypothyroidism were selected, of which 80 patients who fit the inclusion criteria of age >18 years of age, with subclinical hypothyroidism thyroid - stimulating hormone (TSH) >5.5 µ IU/ml with normal FT4 and FT3, or overt hypothyroidism were included which also showed similar results

5. Conclusion

Subclinical hypothyroidism have deranged lipid profile in form of raised Total Cholesterol and LDL and low HDL. 2D Echo in patients of subclinical hypothyroidism shows Grade 1 LV diastolic dysfunction. There is association between Lipid Profile changes and 2D Echo changes. There is no major ECG change in patients with subclinical hypothyroidism. Subclinical Hypothyroidism can be considered as independent risk factor of cardiovascular system morbidity.

References

- https://www.jfmpc.com/article.asp?issn=22494863; year=2016; volume=5; issue=4; page=809; epage=816; aulast=Jayasingh
- https://www.ijem.in/article.asp?issn=22308210; year=2012; volume=16; issue=8; page=347; page=349; aulast=Asranna
- https://academic.oup.com/eurheartj/article/39/7/503/3056578
- https://actascientific.com/ASMI/pdf/ASMI - 01 - 0027.pdf
- http://sciaeon.org/articles/Evaluation - of - Cardiac - Functions in Hypothyroidism - and - Subclinical - Hypothyroidism. pdf
- https://www.ahajournals.org/doi/epub/10.1161/CIRCULATIONAHA.106.678326
- GuelKW, van Sluisveld IL, Grobbee DE, et al.1993 The importance of thyroid microsomal antibodies in the development of elevated serum TSH in middleaged women: associations with serum lipids. Clin

Endocrinol (Oxf) 39: 275–280

- [8] <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5509395/>
- [9] Mubashir Alam Khan, Tasnim Ahsan, Urooj Lal Rehman, Rukhshand Jabeen, and Saad Farouq. Subclinical Hypothyroidism: Frequency, clinical presentations and treatment indications. Pak J Med Sci. 2017 Jul - Aug; 33 (4): 818–822. doi: 10.12669/pjms.334.12921.
- [10] *Manoj Kumar, Raveendra S. Rajpoot.* A study of lipid profile in patients of subclinical hypothyroidism in rural area of Western Uttar Pradesh, India. <http://dx.doi.org/10.18203/2320-6012.ijrms20162273>.
- [11] Preethi JJ, Vasudevan HS. Echocardiographic Changes in Overt and Subclinical Primary Hypothyroidism. Int J Sci Stud 2018; 6 (1): 65 - 68.