Thyroid Dysfunction in Women during First Trimester of Pregnancy:
Correlation with Anti-Thyroid Peroxidase Antibodies

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Abstract

**Background:** During pregnancy, thyroid dysfunction has been associated with a number of adverse outcomes. The presence of anti-Thyroid Peroxidase Antibody (anti-TPO) also results in post-partum complications. Hypothyroidism is closely associated with the presence of anti-TPO. The study aimed to evaluate anti-TPO and thyroid function tests in first trimester of pregnancy. **Materials and Methods:** The study was carried out at Department of Biochemistry, Medical College and SSG Hospital, Vadodara, Gujarat, India over a period of one year after prior approval from institutional ethics committee. Total 200 normal pregnant women in their first trimester were randomly selected after informed written consent. Women having known thyroid dysfunction, other endocrinopathies, undergone thyroid surgery or taking thyroid medications were excluded. Overnight fasting blood samples from participants were analyzed for anti-TPO, thyroid stimulating hormone (TSH), total T3 and total T4. **Results:** Mean maternal age was 25.56 ± 3.32 years. Out of 200, total 30 (15%) pregnant women of first trimester had TSH level >2.5 mIU/l. Out of these 30 females, 9 (4.5%) had overt hypothyroidism & 21 (10.5%) had subclinical hypothyroidism. Among these 30 hypothyroid women, 26 (13%) had anti-TPO positive (anti-TPO >35 IU/ml) which also showed a significant positive correlation with TSH level and negative correlation with T3 and T4 level. **Conclusion:** Hypothyroidism especially sub-clinical hypothyroidism is prevalent among pregnant women with positive correlation with anti-TPO. It is suggested that anti-TPO should be added as a screening test in first trimester of pregnancy which can be useful to identify early thyroid dysfunction.

**Keywords:** Anti-Thyroid Peroxidase Antibody, Thyroid stimulating hormone, T3, T4.

**INTRODUCTION**

Pregnancy is a physiological state associated with many significant changes in thyroid function. The role of thyroid hormone in embryogenesis and fetal development during pregnancy is well known. During the first trimester of pregnancy, the fetus is reliant on trans-placental passage of maternal thyroid hormone, as the fetal thyroid is not fully functional until about 16 weeks of gestation [1]. It has been seen that thyroid dysfunction is associated with many complications during pregnancy and delivery like miscarriage, preterm birth, placental abruption and the child may have low IQ, cretinism and developmental disorders. Many studies have shown that hypothyroid state in pregnancy is associated with maternal and fetal complications [2, 3]. During the first trimester, the fetus is completely dependent upon thyroid hormone produced by the mother. Even a small unnoticed malfunction of the thyroid gland, which doesn’t endanger the course of pregnancy, can affect the psychomotor development of the child [4]. Thyroid function tests (TFT) are frequently assessed during pregnancy, both to diagnose suspected thyroid abnormalities and to monitor the status of pre-existing thyroid disease. In addition to conventional TFT (T3, T4 & TSH) tests, many studies have suggested testing of anti-thyroid peroxidase antibodies (anti-TPO) [5]. Anti-TPO works against thyroid peroxidase, an enzyme which catalyzes iodide oxidation and iodination of tyrosyl residues of thyroglobulin. It also causes tissue destruction in other forms of thyroiditis such as postpartum thyroiditis [6]. Anti-TPO may be considered as a marker of generalized autoimmune dysfunction in the body. Anti-TPO positive women have a risk for post partum thyroid dysfunction, hypothyroidism, miscarriage, preterm delivery and perinatal death. Screening for anti-TPO level in early pregnancy may help to diagnose women at risk of hypothyroidism and thereby prevent adverse outcome of pregnancy [7]. Many cases of hypothyroidism especially subclinical hypothyroidism remain undiagnosed during pregnancy leading to complications. Early treatment significantly reduces these complications [8]. Hence, early diagnosis of hypothyroidism and initiation of treatment is
necessary. Thus, the present study was designed to correlate the anti-TPO levels with thyroid function tests in first trimester of pregnancy.

**Materials & Methods**

This cross sectional study was carried out at Sir Sayajirao General Hospital and Medical College, Vadodara over a period of one year. Before starting this study, approval from the Institutional Ethics Committee for Human Research, Medical College and S.S.G. Hospital, Vadodara was obtained (ECR/85/Inst/GJ/2013). Total 200 pregnant women visited antenatal clinic of Obstetrics and Gynecology Department in 1st trimester were enrolled in study after written informed consent. Women having known thyroid dysfunction, thyroid autoimmunity, other endocrinopathies, undergone thyroid surgery or taking thyroid medications were excluded. Detailed medical history and physical examination findings were recorded in a predesigned proforma. 3 ml of fasting blood sample was collected from participants under aseptic precautions and was analyzed for TFT i.e. total T3, total T4, TSH and anti-TPO. The analysis was done by immunoassay technique ELISA. Reference values used for these parameters were: T3=0.5-2.0 ng/ml, T4=53-121 ng/ml, TSH= 0.4-2.5 mU/l and anti-TPO= <35 IU/ml. Women with T3, T4 below the reference range along with elevated TSH were classified as having overt hypothyroidism while those having T3, T4 in normal range with TSH more than 2.5 mIU/l were diagnosed as having Subclinical Hypothyroidism. Statistical analysis was done by using GraphPad Prism software. Regression analysis was used to find out statistical correlation between two variables and its significance.

**Results**

Total 200 pregnant women of first trimester were enrolled. Mean maternal age was 25.56 ± 3.32 years. Table-1 showed Thyroid dysfunction found in first trimester of pregnancy. Out of 200, 30 (15%) pregnant women had hypothyroidism (TSH >2.5mIU/l) Out of these 30 hypothyroid women, 9 (4.5%) had overt hypothyroidism & 21 (10.5%) had subclinical hypothyroidism. Among these 30 hypothyroid women, 26 (13%) had found anti-TPO positive (TPO-Ab >35 IU/mL, p<0.001). Table-2 showed anti-TPO status in hypothyroid women. There was a significant positive correlation between positive TPO-Ab and serum TSH level (r=0.6330) of study subjects and there was a negative correlation between anti-TPO and serum T_{3}, T_{4} level (r=-0.4871, r=-0.3037 respectively) in study subjects. Correlation of anti-TPO with thyroid function tests are shown in Figure 1, 2 & 3.

**Table-1: Thyroid dysfunction found in first trimester of pregnancy (n=200)**

<table>
<thead>
<tr>
<th>Frequency</th>
<th>Percentage</th>
</tr>
</thead>
<tbody>
<tr>
<td>Overt hypothyroidism</td>
<td>9</td>
</tr>
<tr>
<td>Sub-clinical hypothyroidism</td>
<td>21</td>
</tr>
<tr>
<td>Euthyroid</td>
<td>170</td>
</tr>
<tr>
<td><strong>Total</strong></td>
<td><strong>200</strong></td>
</tr>
</tbody>
</table>

**Fig-1: Correlation of serum TSH with TPO-Ab**
DISCUSSION

Thyroid dysfunctions are common during pregnancy and it can affect the outcome also. Our study revealed 15% prevalence of hypothyroidism in first trimester of pregnancy. Out of this 15%, 4.5% had overt hypothyroidism and 10.5% had sub-clinical hypothyroidism. There are only few reports on prevalence of hypothyroidism during pregnancy from India with prevalence rates ranging from 4.8% to 24%. Dhanwal et al., reported 14.3% prevalence of hypothyroidism in the first trimester of pregnancy and majority of them had sub-clinical hypothyroidism [9]. High prevalence of hypothyroidism is due to high intake of dietary iodine, goitrogens and minerals like iron, selenium deficiency [10-12]. The prevalence rates may be found difference in different geographic areas and it is also due to the selection of TSH upper cut-off value. Sailakshmi et al., showed a prevalence of 7.5% [6]. Sahu et al., reported 6.4% prevalence of hypothyroidism [13]. Lata et al., have found 24% prevalence of hypothyroidism among pregnant females with history of two or more consecutive miscarriages [14]. Thyroid autoimmunity can also alter the progression of pregnancy and thus affect its outcome. Western studies showed 10-15% prevalence of thyroid autoimmunity in pregnancy [15]. Teng et al., found anti-TPO positivity in 9.6% of first trimester pregnant Chinese women [11]. Dhanwal et al., [9] reported an anti-TPO prevalence of 6.82% in pregnant women of North India. In the current study, anti-TPO was elevated in 26 (13%) women who had hypothyroidism which also suggest that mean TSH level was high with anti-TPO positivity. The possible reason for effect of anti-TPO on pregnancy outcome is subclinical deficiency of the thyroid hormones in women and they are unable to meet the increased requirements of thyroid hormones during pregnancy. The presence of anti-TPO adversely affects the foetus and result into spontaneous abortions, preterm delivery, IUGR etc [16, 17].

CONCLUSION

Our study concluded that Hypothyroidism especially subclinical hypothyroidism is prevalent among pregnant females. Anti-TPO positivity is associated with high level of TSH suggestive of hypothyroidism. Since thyroid dysfunction during pregnancy is associated with many fetal and maternal complications, anti-TPO status of pregnant females should be tested in early pregnancy so that in case of thyroid dysfunction, early treatment can be started to prevent complications.
REFERENCES