

Frequency and Association of Subclinical Thyroid Disorders with Poor Obstetric Outcome

Subclinical
Thyroid
Disorders with
Poor Obstetric
Outcome

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ABSTRACT

Objective: To determine the frequency and association of subclinical thyroid disorders in patients with poor obstetric outcomes.

Study Design: Cross sectional study

Place and Duration of Study: This study was conducted at the Department of Obstetrics and Gynecology, Lady Reading hospital Peshawar from August 2019 till July 2020.

Materials and Methods: Women of reproductive age with a bad obstetrical history during her visit in the outpatient department of the hospital were enrolled. Obstetrical details including previous miscarriages, stillbirths, neonatal deaths and congenital anomalies were recorded by interviewing the patient as per questionnaire. patients fulfilling the inclusion criteria were subjected to thyroid function tests. Data was analyzed on SPSS version 18.

Results: Out of the 549 women presenting with a poor obstetric outcome, 103 women were recruited for the study. subclinical thyroid disorders were found to be 24.27%, with subclinical hypothyroidism more prevalent (21.35 %) in patients with poor obstetric outcome. significant weak positive correlation was reported between neonatal deaths and TSH levels ($r= 0.200$; $p= 0.043$).

Conclusion: Subclinical thyroid dysfunction is more prevalent in patients with poor obstetric outcome. various adverse fetal and neonatal outcomes are expected to be encountered in patients with subclinical thyroid dysfunction.

Key Words: Subclinical hypothyroidism, thyroid stimulating hormone, miscarriage, stillbirth

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INTRODUCTION

Thyroid hormones have an important role in neurophysiologic development of the fetus. Subclinical thyroid dysfunction implicates an abnormal serum thyroid-stimulating hormone level (normal TSH reference range: 0.45 to 4.5 μ U per mL) in clinically asymptomatic patient. Serum TSH 0.1-0.45 mIU/L will denote subclinical hyperthyroidism and serum TSH 4.5-10.0 mIU/L will demarcate as subclinical hypothyroidism. Some studies recommend against routine treatment of patients and free thyroxine(T4) and triiodothyronine(T3) levels within their normal ranges¹. thyroid disease represents the second most recurrent endocrine condition that might affect females in the reproductive age².

The risk of hypertension, miscarriages, fetal growth restriction, abruption and preterm births has been increased if thyroid disorder is not treated in pregnancy³. In order to reduce these risks, Screening has to be performed during the first trimester of pregnancy. It has been recommended by American Association of Endocrinologist, but European and American Thyroid Association experts do not recommend the universal screening of all pregnant population, though testing in case of high risk is suggested⁴. Females with the history of stillbirth, neonatal death, three or more abortions are considered as poor obstetrical history⁵. The causes of poor obstetric outcome are varied including diabetes, hypertension anaemias, uterine factors, cervical incompetence, prelabour ruptured membranes, antepartum hemorrhage or intrapartum adverse incidents. Some studies revealed an association of these poor obstetric events with subclinical hypothyroidism or thyroid autoimmunity⁶. Others have reported a relationship with a higher rate of stillbirth in the second trimester of pregnancy with subclinical hypothyroidism⁷.

The present study is, therefore, being conducted to assess the frequency and association of subclinical thyroid dysfunction with obstetric outcomes in women of reproductive age group and to evaluate patients with poor obstetrics outcome for any underlying subclinical thyroid dysfunction.

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MATERIALS AND METHODS

This cross sectional study was done at the department of obstetrics and gynecology, Lady Reading hospital Peshawar over one year from August 2019 till July 2020.

Patients were selected by non-probability convenient sampling after taking ethical approval from the institutional review board of the hospital. Women of reproductive age who had a previous history of poor obstetric outcome were included in the study. Patients with any underlying comorbidities such as diabetes, hypertension which per se would increase likelihood of increased fetal and perinatal mortality and in turn lead to poor obstetric outcome were excluded from the study. clinically proven thyroid disorders, previous history of thyroid surgery was also excluded from the study.

After informed consent detailed history was taken regarding maternal age, parity, previous miscarriage, still births and early neonatal deaths etc., as per attached questionnaire. Blood samples were then taken for thyroid hormones i.e. thyroxin(T4) and thyroid stimulating hormone (TSH) levels on their visit in the outpatient department. Patients were also inquired about personal and family history of thyroid disease; those patients who come up with deranged TFTS (thyroid function tests were thus retrospectively diagnosed to be have subclinical thyroid dysfunction which led to a poor obstetric outcome.

Data analysis: The collected data was entered into SPSS version 18.0 and analyzed. Descriptive statistics were presented as frequency and percentages and were calculated for qualitative variables. Pearson correlation was applied to assess the association. P-values less than 0.05 were considered as a level of significance.

RESULTS

Out of the 549 women presenting with a poor obstetric outcome, 446 women had known risk factors for their poor obstetric outcome. These patients were thus excluded. Rest of 103 without any identifiable cause for their bad obstetric history were enrolled for the study. The mean age was noted to be 28.94±4.07years. Mean parity was 3.47±1.70 in which 7(6.8%) were nulliparous (although they had recurrent miscarriages) whereas 96 (93.2%) were multiparous women, (as shown in Table-1).

48(46.6%) had 2 miscarriages. 38 (36.9%) of them had previous 1 still birth. 10 (9.7%) women had a history of abnormal babies, 6 (5.82%) had babies with cerebral palsy (CP), 2(1.94%) women had 2(1.94%) growth retarded child births, whereas 2 (1.94%) had congenitally abnormal baby with one having skeletal dysplasia and other having polydactyly. 27 (26.2%) had previous 1 neonatal death.

78(75.5%) patients had normal thyroid function tests. Frequency of subclinical thyroid disorders was found to be 24.27% out of which subclinical hypothyroidism was found in 22 (21.35 %) who have normal T4 levels but elevated TSH, whereas subclinical hyperthyroidism in 3 (2.9%) patients with poor obstetric outcome. (As shown in Table-2)

Table No.1: Maternal demographic characteristics and variables of poor obstetric outcome

Variable	Mean±SD
Age (years)	28.94±4.07
Parity	3.47±1.70
Miscarriages	2.37±1.2051
Stillbirths	0.65±0.73
Neonatal Deaths	0.32±0.52

Table No.2: Adverse fetal outcomes of maternal thyroid dysfunction.

Variables of poor obstetric outcome		Number of patients	%tage
Previous Miscarriages	1 miscarriage	10	9.7
	2 miscarriage	48	46.6
	3 miscarriage	26	25.2
	4 miscarriage	7	6.8
	5 miscarriage	3	2.9
	6 miscarriage	3	2.9
	No miscarriage	6	5.8
Previous Still Births	1 Still Birth	38	36.9
	2 Still Births	13	12.6
	3 Still Births	1	1.0
	No Still Births	51	49.5
Abnormal Babies	No abnormality	93	90.29
	Cerebral Palsy.	6	5.82
	Growth Restriction	2	1.94
	Congenital anomaly	2	1.94
Previous Neonatal Deaths	1 Neonatal Death	27	26.2
	2 Neonatal Deaths	3	2.9
	Nil	73	70.9

Table No.3: Correlation of demographic variables and obstetric outcomes with TSH levels.

Variable	TSH Levels	
	P	p-value
Age (years)	-.057	0.564
Parity	0.058	0.559
Miscarriages	0.016	0.874
Still Births	0.037	0.713
Neonatal Deaths	0.200	0.043

An insignificant correlation for TSH levels was reported with age ($r = -0.057$; $p = 0.564$), an insignificant correlation was reported with parity ($r = 0.058$; $p = 0.559$), with miscarriages ($r = 0.016$; $p = 0.874$), with stillbirths ($r = 0.037$; $p = 0.713$) whereas a significant weak

positive correlation was reported between neonatal deaths and TSH levels ($r= 0.200$; $p= 0.043$), as shown in Table 3.

Table No.4: - TSH Levels of patients presenting with poor obstetric outcome

TSH Levels	Number of patients =N	Percentage (%)
Reduced(hyperthyroid)	3	2.9
Normal (euthyroid)	78	75.7
Increased(hypothyroid)	22	21.4
MeanTSH Values	3.34±2.26	

DISCUSSION

Thyroid dysfunction is the second most common recurrent endocrine disorder faced by 2 to 3% of women during pregnancy⁸. Adverse or poor obstetrics outcome as detected in overt hypothyroidism may also be encountered in subclinical hypothyroidism. Our study showed prevalence of subclinical hypothyroidism in patients with poor obstetric outcome to be 21.4% whereas subclinical hyperthyroidism was observed in 2.9% cases which shows a relatively higher percentage of subclinical hypothyroidism in patients with poor obstetric outcome. Subclinical thyroid dysfunction in females in reproductive age group fluctuates from 2%-10%^{9,10}. A study conducted in Jordan showed 3% pregnant women to have hypothyroidism during pregnancy¹¹. A much higher prevalence of subclinical hypothyroidism 65.2% in pregnant women with a bad obstetrical history was revealed by a study¹². The reason could be the iodine deficiency in the population which is unmasked and potentiated in a physiologically hyperactive thyroid gland in pregnancy.

The effects of Subclinical (SC) thyroid dysfunction on poor obstetrics outcome have still not been clearly known. Certain studies¹³ reported that SC Hypothyroidism did not lead to any consistent adverse maternal and prenatal outcomes, while other showed that SC Hypothyroidism is associated with a number of obstetric complications, which include abruption of placenta, GDM, and IUGR¹⁴. Our results showed a weak positive significant correlation between TSH levels with neonatal deaths ($r= 0.200$; $p=0.043$) while insignificant association was observed in terms of miscarriages, parity and still births.

Women with hypothyroidism were more prone to miscarriages during their first trimester, which is comparable to the outcomes of earlier studies. Our study results were in agreement with the above mentioned studies and revealed that 48(46.6%) had 2 miscarriages during pregnancy whereas 26(25.2%) had 3 miscarriages but there was an insignificant association between miscarriages and TSH level ($p=0.874$).

A study by Su and colleagues¹⁵ showed greater risks of deformities of circulatory system (11.1%) and

musculoskeletal (4.7%) in fetuses of women with hypothyroidism. As far as our study is concerned, it is in compliance with the above mentioned studies in way that maternal thyroid disorder led to deliver abnormal babies as skeletal dysplasia and polydactyly were noted in 1.94%. Moreover, our results have shown Cerebral Palsy in 5.82% and growth retarded babies were reported in 1.9% cases of subclinical thyroid dysfunction

Study by Casey as mentioned earlier demonstrated that the rate of stillbirth was more among those women with SCH than women with euthyroid in the second trimester, but no statistically significant differences were seen. Though higher rate of stillbirth in women with hypothyroid with TSH levels 0.10mU/L has been earlier stated, SCH (described by much lesser amount of TSH levels) appears to have small impact on stillbirth¹⁶. Our findings are in agreement with those reported in the literature and indicated that stillbirth rate was higher in women hypothyroidism 38(36.9%) but there was an insignificant correlation observed between stillbirth and TSH levels. ($r= 0.037$; $p= 0.713$). Therefore, thyroid screening in women with poor obstetric outcome may be useful to predict its prior incidence. Further, timely management of subclinical thyroid dysfunction might reduce the morbidity and mortality related to poor obstetric outcome.

CONCLUSION

Various adverse fetal outcomes can be correlated with subclinical thyroid dysfunction in patients with poor obstetrical history. Subclinical hypothyroidism in reproductive age group may increase the risks of miscarriages and neonatal deaths.

Recommendations: Routine screening of women with poor obstetric outcome is essential to rule out subclinical thyroid dysfunction to improve fetal and perinatal outcome.

Author's Contribution:

Concept & Design of Study: Sumaira Yasmin
 Drafting: Farnaz Zahoor
 Data Analysis: Farnaz Zahoor
 Revisiting Critically: Sumaira Yasmin, Farnaz Zahoor
 Final Approval of version: Sumaira Yasmin

Conflict of Interest: The study has no conflict of interest to declare by any author.

REFERENCES

1. OR S. Subclinical thyroid disease. JAMA 2004; 291:228-38.
2. Haddow JE, McClain MR, Lambert-Messerlian G. Variability in thyroid - stimulating hormone suppression by human chorionic gonadotrophin

- during early pregnancy. *J Clin Endocrinol Metab* 2008;93(9):3341-3347.
3. Gluud C, Gluud LL. Evidence based diagnostics. *BMJ* 2005;330(7493):724-6.
 4. Stagnaro-Green A, Abalovich M, Alexander E, Azizi F, Mestman J, Negro R, et al. Guidelines of the American Thyroid Association for the diagnosis and management of thyroid disease during pregnancy and postpartum. *Thyroid* 2011; 21(10):1081-125.
 5. Elahi S, Rizvi NB, Nagra SA. Iodine deficiency in pregnant women of Lahore. *J Pak Med Assoc* 2009;59(11):741-3.
 6. Dal Lago A, Vaquero E, Pasqualetti P, Lazzarin N, De Carolis C, Perricone R, et al. Prediction of early pregnancy maternal thyroid impairment in women affected with unexplained recurrent miscarriage. *Human Reproduction* 2011;26(6):1324-30.
 7. Casey BM, Dashe JS, Wells CE, McIntire DD, Byrd W, Leveno KJ, Cunningham FG. Subclinical hypothyroidism and pregnancy outcomes. *Obstet Gynecol* 2005;105(2):239-45.
 8. Wang W, Teng W, Shan Z, Wang S, Li J, Zhu L, et al. The prevalence of thyroid disorders during early pregnancy in China: the benefits of universal screening in the first trimester of pregnancy. *Eur J Endocrinol* 2011;164(2):263-8.
 9. Alkafajei A, Amarin Z, Alazaizeh W, Khader Y, Marji M. Prevalence and risk factors for hypothyroidism in Jordanian women: comparison between different reference ranges. *East Med Health J* 2012;18:2.
 10. Maraka S, Singh Ospina NM, Mastorakos G, O'Keefe DT. Subclinical Hypothyroidism in Women Planning Conception and During Pregnancy: Who Should Be Treated and How? *J Endocr Soc* 2018;2(6):533-546.
 11. Lambert-Messerlian G, Sullivan L, Canick J, Porter TF, Luthy D, Gross S, et al. Maternal thyroid hypofunction and pregnancy outcome. *Obstet Gynecol* 2008;112:85-92.
 12. Butt F, Chohan A. Subclinical thyroid dysfunction among pregnant women with bad obstetrical history. *MOJ Womens Health* 2017;4(1):11-14.
 13. Ashoor G, Maiz N, Rotas M, Jawdat F, Nicolaidis KH. Maternal thyroid function at 11 to 13 weeks of gestation and subsequent fetal death. *Thyroid* 2010; 20(9):989-93.
 14. Ezzeddine D, Hamadi C, Abbas HA, Nassar A, Abiad M, Ghazeeri G. Prevalence and correlation of hypothyroidism with pregnancy outcomes among lebanese women. *J Endocrine Society* 2017; 1(5):415-22.
 15. Su PY, Huang K, Hao JH, Xu YQ, Yan SQ, Li T, Xu YH, et al. Maternal thyroid function in the first twenty weeks of pregnancy and subsequent fetal and infant development: a prospective population-based cohort study in China. *J Clin Endocrinol Metabolism* 2011;96(10):3234-41.
 16. Tudela CM, Casey BM, McIntire DD, Cunningham FG. Relationship of subclinical thyroid disease to the incidence of gestational diabetes. *Obstet Gynecol* 2012;119(5):983-8.