Comparative Study of Lipid Profile, Oxidant, Antioxidant Levels and Thyroid Hormone Status in Normal and Abnormal 2 Pregnancy 3 $Uma.T^1$

¹ RIMS/Dr.NTRUHS

Received: 11 December 2013 Accepted: 1 January 2014 Published: 15 January 2014

Abstract 8

Δ

5

6

The aim of the study is to evaluate thyroid status, Oxidative stress lipid profile levels in abnormal and normal pregnancies. Material and Methods: The study includes 25 abnormal 10 pregnant women, 25 normally healthy pregnant women. Ages ranged from 20 to 30 years. 11 Thyroid status was estimated by ELISA method. For determination of oxidative stress 12 Malandialdehyde (MDA) is estimated as oxidants by TBARS, Uric acid is estimated by pap 13 method, cholesterol HDL-C by CHOD-POD method, triglycerides by GPO-POD 14 method.Results: In the present study, we observed that TSH is significantly increased 15 compared to normal pregnancies TSH and FT4 are slightly decreased compared to normal 16 pregnancies. In the present study cholesterol levels were increased in abnormal pregnancies 17 compared to normal pregnancies and decrease in HDL cholesterol were observed in 18 preeclamptic and eclamptic pregnant women. 19

20

Index terms—lipid profile, oxidant, antioxidant, abnormal pregnancy. 21

1 Introduction 22

he thyroid gland secretes hormones having important role in embryogenesis and fetal development during normal 23 pregnancy. (1) Thyroid function tests change during pregnancy due to the influence of two main hormones: 24 human chorionic gonadotropin (hCG), the hormone that is measured in the pregnancy test and estrogen, 25 the main female hormone. HCG can weakly turn on the thyroid and the high circulating hCG levels in the 26 first trimester may result in a slightly low TSH. When this occurs, the TSH Estrogen increases the amount 27 of thyroid hormone binding proteins in the serum which increases the total thyroid hormone levels in the 28 blood. (2) The most common cause of hyperthyroidism in pregnancy is Grave's disease and hypothyroidism 29 30 is autoimmune thyroid disease (Hashimoto's thyroiditis). The complications associated with hyperthyroidism 31 during pregnancy are first trimester spontaneous abortions, preterm deliveries, low birth weight infants, still 32 births, neonatal deaths, fetal, neonatal hyperthyroidism and intrauterine growth retardation; in hypothyroidism are preeclampsia, preterm labour, placental abruption, threatened abortions, neonatal hypothyroidism, neonatal 33 hyperbilirubinemia, postpartum hemorrhage and increased frequency of low birth weight infants. (2) The aim of 34 the study is to evaluate thyroid status in abnormal and normal pregnancies by the estimation of FT3, FT4, and 35 TSH, Oxidative stress is estimated by estimating MDA as oxidant & uric acid as antioxidant & lipid profile levels 36 include cholesterol, triglyceride, HDL. Hb%. Abnormal pregnancies includes pregnancy induced hypertension 37 (preeclampsia, eclampsia), gestational diabetes, hydroamnios, and abortions. 38

³⁹ 2 II.

40 3 Materials and Methods

41 Study Population: The study populations were investigated consisted of 50 women divided into two groups 42 consisted of 25 abnormal pregnant women, 25 normally healthy pregnant women. Ages ranged from 20 to 30

43 years. The prospective study was carried out at the RIMS Hospital, Srikakulam, Andhra Pradesh, India, between

January 2013 to June 2013. The study was approved by the Institutional Human Ethical Committee (IHEC).
 Informed verbal consent was obtained from all subjects. The objectives of the study were explained and a written

45 Informed verbal consent was obtained from an subjects. The objectives of the study were explained and a written
 46 concept was taken from each subject. The data on family history and personal history of diabetes, jaundice,
 47 hypertension and miscarriages were collected through standard questionnaire.

Biochemical Investigation: Serum FT3 (3) , FT4 (4) , and TSH (5) were assessed quantitatively using ELISA

⁴⁹ kits. The FT3 and FT4 were expressed as pmol/L, however the TSH was expressed as mIU/L. Total Material and

50 Methods: The study includes 25 abnormal pregnant women, 25 normally healthy pregnant women. Ages ranged 51 from 20 to 30 years. Thyroid status was estimated by ELISA method. For determination of oxidative stress

⁵² Malandialdehyde (MDA) is estimated as oxidants by TBARS, Uric acid is estimated by pap method, cholesterol ⁵³ k HDL-C by CHOD-POD method triglycerides by CPO-POD method

 $\scriptstyle 53$ $\,$ kDL-C by CHOD-POD method, triglycerides by GPO-POD method.

Results: In the present study, we observed that TSH is significantly increased compared to normal pregnancies TSH and FT4 are slightly decreased compared to normal pregnancies. In the present study cholesterol levels were increased in abnormal pregnancies compared to normal pregnancies and decrease in HDL cholesterol were observed in preeclamptic and eclamptic pregnant women. In the present study Hb levels were decreased in abnormal pregnancies compared to normal pregnancies. The levels of MDA and uric acid were significantly increased in abnormal pregnancies. The glucose (RBS) levels were increased compared to normal pregnancies.

59 increased in abnormal pregnancies. The glucose (RDS) levels were increased compared to normal pregnancies

Conclusions: Altered Thyroid status which leads to hypothyroidism may be due to the loss of protein and protein bound hormones in urine in abnormal pregnancies compared with normals. Increased oxidative stress is due to increase in TSH levels which directly induces the pro-inflammatory return to normal throughout the duration of pregnancy.

cytokines. Study suggests that all abnormal pregnancies will be slightly decreased in the first trimester and
 then studied are associated with hypothyroidism.

Comparative Study of Lipid Profile, Oxidant, method (6,7) and Tgl by GPO -POD method (8). For determination of oxidative stress Malandialdehyde (MDA) is estimated as oxidant by thiobarbituric acid reactive substance (TBARS) (9), Uric acid is estimated by pap method (10), Hb is estimated by Drabkin's method (11), Glucose is estimated by glucose oxidase and peroxidase method (12). Statistical Analysis: All data were expressed as mean \pm S.D. The 'P' value was used to compare mean values of abnormal pregnant women with normal pregnant women.

72 **4** III.

73 5 Results

In the present study, observed that TSH (7.5 ± 3.93) is significantly increased compared to normal pregnancies 74 TSH (4.0 ± 2.82) . FT3 (1.1 ± 0.34) and FT4 (0.7 ± 0.24) are slightly decreased compared to normal pregnancies 75 (2.0 ± 0.95) and (1.2 ± 0.26) . In the present study cholesterol levels were increased in abnormal pregnancies 76 (228.9 ± 47.12) compared to normal pregnancies (212.2 ± 38.95) and decrease in HDL cholesterol were observed 77 in preeclamptic and eclamptic pregnant women. In the present study Hb levels were decreased in abnormal 78 pregnancies (8.8 ± 1.7) compared to normal pregnancies (7.9 ± 1.9) . The levels of MDA and uric acid were 79 significantly increased in abnormal pregnancies. The glucose (RBS) levels were increased compared to normal 80 pregnancies [Table 1]. 81

⁸² 6 COMPARATIVE STUDY VALUES OF NORMAL AND ⁸³ ABNORMAL PREGNANT WOMEN

84 Graph-2

85 IV.

86 7 Discussion

The present study was conducted to find out the thyroid status in abnormal pregnancies. Pregnancy is associated with substantial but reversible changes in thyroid function. The metabolic demands and hormonal changes which occur as a result of pregnancy induce a complex alteration in thyroid function. Many authors earlier reported the hypothyroidism in abnormal pregnancy ??13, ??4,15,16). In the present study which included a group of 25 abnormal pregnant women, it is observed that TSH (7.5 \pm 3.93) is significantly increased compared to normal pregnancies, TSH (4.0 \pm 2.82). FT3 (1.1 \pm 0.34) and FT4 (0.7 \pm 0.24) are slightly decreased compared to normal pregnancies (FT3 2.0 \pm 0.95 and FT4 1.2 \pm 0.26).

94 Preeclampsia women have high incidence of hypothyroidism that might correlate with the severity of 95 preeclampsia .It has been suggested that reduced serum concentrations of thyroid hormones in toxemia may be due to the loss of protein and protein bound hormones in urine. Modest decrease in thyroid hormones with concomitant increase in TSH levels in maternal serum correlated with severity of preeclampsia (or) eclampsia and high levels of endothelia. The endothelial cell dysfunction plays an important role in the pathogenesis of preeclampsia. Nitric oxide, a vasodilator released from the endothelial cells, regulates secretion of thyroid hormones by modulating regional blood flow. The decrease in thyroid hormones with significant increase in TSH has been found to be correlated with the severity of preeclampsia.

In the present study cholesterol levels were increased in abnormal pregnancies compared to normal pregnancies 102 and decrease in HDL cholesterol were observed in preeclamptic and eclamptic pregnant women. Oestrogen is 103 responsible for induction of TG and HDL and suppression of serum LDL and oestrogen levels falls in preeclampsia. 104 The low level of HDL in preeclampsia is however not only because of hypoestrogenaemia but also due to insulin 105 resistance. In the present study Hb levels were decreased in subjects compared to normal pregnancies. Pregnant 106 women are often iron deficient, and iron deficiency had adverse effects on thyroid metabolism. Poor maternal iron 107 status predicts both higher TSH and lower TT4 concentration during pregnancy. Many authors reported increased 108 MDA levels in abnormal pregnancies (17). In the present study also MDA levels is significantly increased 109 (9.2 ± 8.28) . The elevated levels of free radicals as evidence by elevated TBRS (thiobarbutric acid reactive 110 substance) they are products of lipid per oxidation effecting cell membrane. Uncontrolled lipid peroxidation is a 111 112 key contributing factor to pathophysiological condition of preeclampsia.

113 Uric acid levels were increased in abnormal pregnancies compared to normal, uric acid is one of the important 114 redox systems in the body. Uric acid acts as an extracellular antioxidant because of its ability to remove singlet oxygen hydroxyl and peroxyl radical. The current study assessed the effects of TSH on endothelial function, and 115 its possible involvement of inflammation and oxidative stress. Raised TSH levels directly induces TNF ? secretion 116 by bone marrow cells and IL 6 by adipocytes TNF ? is a pivotal NO-controlling cytokine, and elevated TNF? 117 levels may promote the expression of inducible NO synthase, leading to increased oxidative stress. So raised 118 serum TSH levels leads to impairment of endothelium dependent vasodilatation, possibly through the induction 119 of low grade inflammation and reduced NO availability by oxidative stress. 120

In some pregnant women oxidative stress may lead to dysmorphogenesis, abortions and intrauterine growth restriction. The study summarized the role of free radicals (or) reactive oxygen species mediate their action through many of the pro-inflammatory cytokines and this mechanism has been proposed as a common underlying factor for endometriosis, ovarian cancer, polycystic ovary disease, and various other pathologies affecting the female reproductive process.

126 V.

127 8 Conclusion

TSH is significantly elevated and FT3, FT4 were decreased slightly in abnormal pregnancies compared with normals.TBRS (malandialdehyde) as marker of lipid peroxidation was elevated significantly. Antioxidant uric acid is elevated significantly. The rise in antioxidant is probably to compensate the increased peroxide load in abnormal pregnancies. Hb levels are decreased and RBS levels are significantly increased in abnormal pregnancies. Cholesterol, TGL levels were increased and HDL levels were significantly decreased in abnormal pregnancies.

133 Study suggests that all abnormal pregnancies studied are associated with hypothyroidism. ¹

¹Comparative Study of Lipid Profile, Oxidant, Antioxidant Levels and Thyroid Hormone Status in Normal and Abnormal Pregnancy



Figure 1: T

1

		Normal pregnant women		Abnormal			
Sl.	Parameters			pregnant women		Z-value	P-value
No.							
		mean	S.D	mean	S.D		
1	FT3	2.0	0.95	1.1	0.34	0.3409	0.3669
2	FT4	1.2	0.26	0.7	0.24	0.223	0.4129
3	TSH	4.0	2.82	7.5	3.93	3.6231	0.001
4	MDA	6.9	3.27	9.2	8.28	1.292	0.001
5	Uric acid	4.1	0.69	6.9	1.59	8.092	0.001
6	RBS	85.7	12.27	165.1	68.82	5.68	0.001
7	Cholesterol	212.2	38.95	228.9	47.12	1.366	0.0869
8	HDL	35.9	11.62	28.7	6.68	2.680	0.0037
9	TGL	201.4	60.47	222.1	45.80	1.364	0.0869
10	$\mathrm{Hb}\%$	7.9	1.98	8.8	1.70	0.344	0.3669

Figure 2: Table 1 :

- [Hubel et al.], C A Hubel, J M Roberts, R N Taylor, T J Musci, G M Rogers, M K Mclaughin. (Lipid
 peroxidation in pregnancy: new perspectives on preeclampsia)
- 136 [Drabkin ()], D L Drabkin . J. boils. Chem 1932. 98 p. 719.
- 137 [Jacobs and Vandemark ()], N J Jacobs, P J Vandemark. Arch. Biochem. Biophys 1960. 88 p. .
- 138 [Barham and Trinder ()] , D Barham , P Trinder . Analyst 1972. 97 p. 142.
- 139 [Schettler and Nussel ()], G Schettler, E Nussel. Arb. Med, soz. Med 1975. 10 p. 25.
- 140 [Wosilait ()], W D Wosilait. Res. Com. Chem. Pathol. Pharmcal 1977. 16 p. 541.
- 141 [Gordon ()] , T Gordon . Am. J. Med 1977. 62 p. 707.
- 142 [American ()] , J American . Obstet Gynecol 1989. 161 p. .
- [Wild and Immunoassay Handbook ()], D Wild, Immunoassay Handbook. 2001. Stockton Press. p. 551. (nd
 edition)
- [Amino et al. ()] 'Autoimmune thyroid disease in pregnancy'. N Amino , H Tada , Y Hidaka . J Endocrinol
 Investig 1996. 19 p. .
- [Weetman and Mcgregor ()] 'Autoimmune thyroid disease: developments in our understanding'. A P Weetman ,
 A M Mcgregor . *Endocr Rev* 1984. 5 p. .
- [Trinder ()] 'Bishnoi A, Sachmechi I. Thyroid disease in pregnancy'. P Trinder . American Family Physician
 1969. 1996. 14 p. . (Annals. Clin. Bio Chem)
- [Banker ()] 'Determination of Protein Bound Iodine'. S B Banker . Journal Biological Chemistry 1948. 173 p.
 175.
- 153 [Kunyoji ()] 'Estimation of lipid peroxide'. Kunyoji . Methods in enzymology 1984. 105 p. .
- [Khandakar and Ali (2002)] 'Kahtun M Thyroid status of normal pregnant women in Dhaka City'. M A
 Khandakar , M S Ali . Mymensingh Med J 2002 Jan. 11 (1) p. .
- [Thyroid Disease and Pregnancy. The American Thyroid Association website at www] Thyroid Disease and
 Pregnancy. The American Thyroid Association website at www, .thyroid.org