## Hypothyroidism- A Risk Factor for Menstrual Disorders among Nulliparous Females

Gulab Kanwar<sup>1</sup>, Monika Shekhawat<sup>2</sup>, Nidhi Sharma<sup>3</sup>, Rinki Hada<sup>4</sup>,

Rahul Kabra<sup>5</sup>, Juber Ahmed<sup>6</sup>

1- Professor & Head, Department of Biochemistry, Govt. Medical College, Kota, Rajasthan, India
2,3,5,6- Residents, Department of Biochemistry, Govt. Medical College, Kota, Rajasthan, India
4- Senior Demonstrator, Department of Physiology, SMS Medical College, Jaipur, Rajasthan
Corresponding Author: Dr.Monika Shekhawat Email ID : drmonikashekhawat@gmail.com

**Abstract: Background:** Menstrual disorders are very common among the young females and its incidence decreases as the age advances. Any abnormality in the hypothalamic or pituitary hormone leads to disturbance in menstrual cycle. Hypothyroidism is one of the major endocrine disorder prevalent amongst the females. The increased level of TSH influences the secretion of FSH and LH, leading to menstrual disorders.

The aim of the study was to estimate the TSH levels in the nulliparous females with menstrual cycle disorders and so as to establish hypothyroidism as a risk factor. **Method**: The study was conducted in Govt. Medical College, Kota and attached group of Hospitals, Kota, Rajasthan.Duration of study is from January 2015 to September2015. A total of 102 nulliparous females of ages between 18-28 years presenting with menstrual cycle disorderswere included in the study. Theserum was analysed for TSH by chemiluminescence technique on Cobas e411 in Hormonal Assay Lab, Department of Biochemistry, Govt. Medical College, Kota, Rajasthan. Statistical Analysis was done on Microsoft Excel. Continuous parameters were expressed as Mean $\pm$  SD. The results were obtained by students' Unpaired t-test. P value <0.05 was considered statistically significant. **Conclusion**: By this study we conclude that menstrual disorders are more common in hypothyroidism thus establishing that hypothyroidism is a risk factor for development of menstrual cycle disorders among nulliparous females and has to be ruled out at an early stage.

Keywords: Hypothyroidism, Menstrual disorders, Nullipara, Thyroid Stimulating Hormone(TSH)

## I. Introduction

Menstruation is the periodic change occurring in primates, which results in the flow of blood and endometrium from the uterine cavity, and may be associated with various constitutional disturbances[1]. The normal menstrual functions depend upon the complex interaction between the hypothalamic-pituitary-ovarian axis and endogenous hormones[2]. The menstrual cycle is divided in two phases, the proliferative phase and the secretory phase, which are under the control of ovarian hormones oestrogen and progesterone respectively. Oestrogen and Progesterone are secreted by ovary under the influence of pituitary(FSH and LH), which is further under the control of hypothalamus[3]. The first menstruation usually begins between twelve and fifteen years of age, a point of time known as Menarche[4]. Menstrual cycle is usually of twenty eight days, measured by the time between the first day of one period and first day of next period. But the regular twenty eight days cycle is seen in only a small proportion of women. Bleeding usually occurs for 2 to 7 days[5]. Menstruation stops occurringbetween ages 45 to 55 years, which is termed as Menopause[6]. The most frequent menstrual cycle disorders are polymenorrhoea, oligomenorrhea and dysmenorrhea [7]. Polymenorrhoea is defined as a menstruation interval lasting less than 21 days, oligomenorrhea is defined as menstruation interval of more than 35 days[8,9]; dysmenorrhea as abdominal pain severe enough to interfere with normal activities, or require medication[10]. Menstrual abnormalities are more common in younger girls, becoming less frequent as they grow older[11,12]. Alterations in the hormones of hypothalamic-pituitary-ovarian origin affect menstrual cycle such as cycle length, regularity and bleeding patterns[13].

Thyroid hormones released by the thyroid gland under the influence of hypothalamus(Thyroid Releasing Hormone) modulate the carbohydrates, fats and proteins metabolism, gene expression and also the sexual and reproductive functions[14]. Hypothyroidism is caused by insufficient production of thyroid hormones by the thyroid gland. Hypothyroidism may be caused by immune or non-immune causes, Autoimmune thyroiditis being the most common cause. It has numerous effects on reproductive system development and functions[15].In females hypothyroidism is associated with anovulatory cycles, amenorrhea, polymenorrhoea, menstrual irregularities, menorrhagia, infertility and increased frequency of spontaneous abortions[16]. The

changes in menstrual cycle reflect that thyroid disorders are associated with ovarian hyperactivity like hyperoestrogenemia, hyperprolactinemia, impaired fertility. These effects are to a greater extent thought to be due to changes in TSH levels, whose secretion overlaps with FSH,LH and prolactin[17].

Nullipara/Para 0 is a woman who has never carried a pregnancy beyond 20 weeks[18].

Measurement of plasma TSH concentration provides the cornerstone of biochemical evaluation of hypothyroidism[19]. Normal level of TSH is 0.3-3.5 mU/L and in hypothyroidism, the level is found to be >10mU/L[20].

**AIM :** To estimate the TSH levels in the nulliparous females with menstrual cycle disorders and so as to establish hypothyroidism as a risk factor.

## II. Materials And Methods

**Study population** : The study was carried out in Govt. Medical College and attached group of hospitals, Kota, Rajasthan. The study period was from January 2015 to September 2015. A total of 102 nulliparous females of agesbetween 18-28 years presenting with menstrual cycle disorders were included in the study.

## Exclusion criterion included :

- Pregnancy
- Females using oral contraceptives and Intrauterine Contraceptive Devices(IUCD).
- Previous history of thyroid disorders and its treatment, known cases of hyperthyroidism and diabetes mellitus
- Chronic smokers and alcoholics.
- Patients suffering from any pituitary disorders. Females on treatment for infertility.
- Patients with drug history of corticosteroids, Lithium carbonate, etc. Any gynaecological disorder eg. Ovarian cyst, PCOD, etc.
- The patients who did not give the consent to participate.

Among the total of 102 females presenting with menstrual disorders, 78 were euthyroid, 3 were hyperthyroid and 21 were found to be hypothyroid. The hyperthyroid cases were excluded from the study.

Sample Collection : A sample of<br/>left standing for an hour and serum5ml was collected<br/>was obtained byfollowing the consent of the patient . The sample was then<br/>centrifugation at 3000 rpm for 10 minutes. There after the<br/>chemiluminescence<br/>technique on Cobas e411 in Hormonal Assay Lab,<br/>Department of Biochemistry, Govt. Medical College, Kota.

**Statistical Analysis**: Statistical Analysis was done on Microsoft Excel. Continuous parameters were expressed as Mean  $\pm$  SD. The results were obtained by students' unpaired t-test. P value <0.05 was considered statistically significant.

## III. Results

During the 9 months study period from January 2015 to September 2015, a total of 102 females were studied of which 78 were euthyroid (TSH level between 0.3-3.5 mU/L), 3 were hyperthyroid (TSH level<0.3 mU/L) and 21 were found to be hypothyroid (TSH level >10 mU/L).

# Table 1: Number and percentage of euthyroid, hypothyroid and hyperthyroid nulliparous females with menstrual disorders. The hyperthyroid females were excluded from the study.

CATEGORY	NUMBER OF	PERCENTAGE
	CASES	
EUTHYROID FEMALES	78	76.5%
(TSH level between 0.3-3.5mU/L)		
HYPERTHYROID FEMALES	3	2.9%
(TSH level<0.3 mU/L)		
HYPOTHYROID FEMALES	21	20.6%
(TSH level >10mU/L)		
TOTAL NUMBER OF NULLIPAROUS	102	100%
FEMALES WITH MENSTRUAL DISORDERS		

The Mean $\pm$  SD in case of euthyroid nulliparous females with menstrual disorders is  $2.36\pm 0.96$  and in case of hypothyroid nulliparous females with menstrual disorders, it is  $15.51\pm 4.82$ . P value <0.05 was found to be

statistically significant.				
	PARAMETER	EUTHYROID NULLIPAROUS FEMALES	HYPOTHYROID NULLIPAROUS FEMALES	P VALUE
		WITH MENSTRUAL DISORDER	WITH MENSTRUAL DISORDER	
	TSH LEVEL(mU/L)	$2.36\pm\ 0.96$	15.51± 4.82	<0.05* (significant)
				(significant)

## Table 2: Showing Mean± SD of TSH in cases of Euthyroid and Hypothyroid Nulliparous Females with Menstrual Disorders

#### IV. **Discussion And Conclusion**

Thyroid disorders are one of the most common endocrine problem encountered all over the world since past two decades and India also carries its significant burden. In India, in a population based study done in Cochin, the prevalence of hypothyroidism was found to be 3.9%[21]. Its prevalence is also seen more in females as compared to males. Menstrual irregularities are common among nulliparous females. Abnormalities related to menstruation contribute to the major health problems that adversely affect the lives of not only women but also family, social and national economy as well. Medical management is required in majority of the cases.

The patients of hypothyroidism suffer from menstrual disorders frequently. Females with irregular menstrual cycle are at increased risk of development of anaemia, obesity, infertility, cardiovascular diseases, type 2 diabetes mellitus, osteoporosis, breast and endometrial cancers [22-25]. Hypothyroidism is also closely related to ovulatory problems, leading to reduction in number of ova produced each cycle. Anovulatory cycles are also present in hypothyroid patients which leads to issues related to infertility. It also leads to obesity which itself is associated with two-fold increase in the prevalence of heavy menstrual flow[26]. Excessive blood loss has been associated with ovulatory disorders, which are common among obese females[27].Polymenorrhea (increased bleeding) is more common in hypothyroidism due to the defects in haemostatis. By this study we conclude that menstrual disorders are more common in hypothyroidism thus establishing that hypothyroidism is a risk factor for development of menstrual cycle disorders and has to be ruled out at an early stage. Our results go in favour of study done by Krassas GE et al in 1999, who reported that 23.4% of the hypothyroid females presented with irregular menstrual cycle[28]. It becomes an important issue of concern for the gynaecologists and physicians, so as decrease the burden of both the problems in the society.

## Aknowledgement

Department of Biochemistry, GMC, Kota for their kind cooperation.

### References

- Symonds EM, Symonds LM. Essential Obstetrics and Gynecology.2004. UK. Churchill Livingstone. [1].
- [2]. Harlow Sd, Ephross SA. Epidemiology of menstruation and its relation to women's health. Epidemiol Rev. 1995; 17(2): 265-286.
- [3]. V.G. Padubidri and Shirish N. Daftary. Shaw's Textbook of Gynaecology.12th edition. Churchill Livingstone.
- [4]. Women Gynecologic Health. Jones and Barlett Publishers. 2011. p 94.
- [5]. "Menstruation and menstrual cycle fact sheet". Office of women's health. December 23,2014. Retrieved on 25 June 2015.
- "Menopause : Overview", 2013-06-28. Retrieved on 8-03-2015. [6].
- ACOG Comittee on Adolescent Health Care: Menstruation in girls and adolescents using menstrual cycle as a vital sign. Obstet [7]. Gynecol 2006,108(5):1323-1328.
- [8]. Petrozza J, poley K: Dysfunctional Uterine Bleeding. In Glass's Office gynecology. 5th edition.Edited by Curtis MG, Hopkins MP, Baltimore: Williams and Wilkins: 1999:241-264.
- Speroff L, Glass RH, Kase NG: Dysfunctional Uterine Bleeding. In Clinical Gynecologic Endocrinology and Infertility.5th edition. [9]. Baltimore: Williams and Wilkins, 1994:575-593.
- [10]. World Health Organisation: International Statistical Classification of Diseases and Related Health Problems, Tenth Revision. Geneva. Switzerland: World Health Organisation;1992.
- [11]. Treloar AE, Boynton RE, Behn BG, Brown BW: Variation of the Human menstual cycle through reproductive life. Int J Fertil 1967,12(1 Pt 2):77-126.
- Vollman RF: The Menstrual Cycle. Major Probl Obstet Gynecol. 1977, 7:1-193. [12]
- Jensen JT, Speroff I. Health benefits of oral contraceptives. Obstet Gynecol Clin North Am. 2000;27(4): 705-721. [13].
- [14]. Comprehensive Thyroid Assessment. Geneva Diagnostics. Retrieved on 2007-05-21
- [15]. Longcope C. The male and female reproductive systems in hypothyroidism. In Werner and Ingbars. The Thyroid a Fundamentalclinical Text, eds l braveman and GB Lipincott, Philadelphia-New York1996;p849-852.
- William J Marshall and Stephen K. Bangert. "Thyroid dysfunction-Clinical Biochemistry : Metabolic and Clinical Aspects". 2<sup>nd</sup> [16]. edition. 394-421.
- [17]. Marou T, Hiramatsu et al: Increase in expression of thyroid hormone receptors in porcine granulose cells early in follicular maturation Acta Endocrinal 1992;127:1562-60(3).
- [18].
- [18] F. Gray Cunnigham.2005. Williams Obstetrics. 22<sup>nd</sup> edition. Mc Graw-Hills companies.
   [19] William J Marshall and Stephen K Bangert. "Thyroid Dysfunction" Clinical Biochemistry- Metabolic and Clinical aspects.2<sup>nd</sup> [19]. edition.p416.
- Carl A. Burtis and David E. Bruns Hormones "Tietz Fundamentals of Clinical Chemistry and Molecular Diagnostics". 7th [20]. edition. 806-823
- [21]. Usha Menon V, Sundaram KR, Unnikrishnan AG, Jayakumar RV, Nair V, Kumar H. High prevalence of undetected thyroid disorders in an iodine sufficient adult south Indian population. J Clin Biochem. 2009;24:52-9.

- [22]. Barret-Connor E, Bush TL. Estrogen and coronary heart disease women. JAMA. 1991; 265 (4):705-721.
- [23]. Butler LM, Potishman NA, Newman B, et al. Menstrual risk factors and early onset of breast cancers. Cancer Causes Control.2000;11(5):451-458.
- Titus-Ernstoff L, Perez K, Cramer DW, Harlow BL, Baron JA, Greenberg ER. Menstrual and reproductive factors in relation to [24]. ovarian cancer risk. Br J Cancer. 2001;84(5):714-721.
- [25]. Wise LA, Mikkelsen EM, Rothman KJ, et al. A prospective cohort study of menstrual characteristics and time of pregnancy. Am J Epidemiol.2011;174(6):701-709.
- Santos IS, Minten GC, Valle NC, et al. Menstrual bleeding patterns: a community based cross-sectional study among women aged [26]. between 18-45 years in Southern Brazil. BMC Womens Health.2011;11(1):26.
- [27]. Harlow SD. Menstruation and menstrual disorders: the epidemiology of menstruation and menstrual dysfunction. In: Goldman MB, Hatch MC, editors. Women and Health. San Diego, CA: Academic Press;2000:99-113. Krassas GE, Pontikides N, Kaltsas T, Papabopoulau P, Paunkovic J, Paunkovic N, Duntas LH. Disturbance of menstruation in
- [28]. hypothyroidism. Clin Endocrinol(Oxf).1999 May;50(5):655-9.