Document heading doi: 10.21276/apjhs.2014.1.3.17 Original Article

COMPARISON OF SERUM TESTOSTERONE AND PROGESTERONE LEVELS IN SURGICAL AND NATURAL MENOPAUSE

Raviraj R Naik ¹, Rittu S Chandel ², Sarita D Dakhure ³, Leela G Abichandani ⁴

^{1, 2} Post graduate resident at Grant Government Medical College, Mumbai, India

³Post graduate resident at MIMSR Medical College, Latur, India

⁴ Professor of Biochemistry, Grant Government Medical College, Mumbai, India

ABSTRACT

Background: Natural menopause is a physiological process which occurs around the age of 45 - 55 years. Surgical menopause is not physiological, occurs through an operative procedure which leads to sudden hormonal imbalances. Both the groups whether natural or surgical menopause experience hot flushes and mood swings, but this happens gradually in natural menopause and suddenly in surgical menopause. These symptoms may be correlated to the sudden hormonal imbalances

Aim: To study and compare hormonal levels of Progesterone and Testosterone in Surgical and Natural menopause

Settings and Design: Prospective Study

Methods and Material: 50 women (cases) belonging to surgical menopausal group and 50 women (controls) were belonging to Natural menopausal group.

The hormonal parameters (Progesterone and Testosterone) were measured by Solid Phase Competitive Chemilumniscent Enzyme Immunoassay.

Statistical analysis used: Student's t-test was used to compare mean serum concentrations of Progesterone. Mann - Whitney U Rank Test was used to compare mean testosterone values

Diagramatic Representation: 1] Dot – Plot graphs were used to show significance as per results of t – test. 2] Bar diagrams were used to show mean values of various parameters of this study in surgical and natural menopause

Results: The mean level of progesterone hormone in study group was found to be 0.71 ± 0.57 while that in control group was 2.074 ± 0.42 . Mann- Whitney U Rank Test was used to compare mean testosterone values which showed significant difference between both the groups of surgical and natural menopause.

Conclusion: In natural menopause the hormonal imbalance occurs gradually over a period of time but in surgical menopause the hormonal levels change abruptly. These sudden changes in the hormonal level may be responsible for severe hot flushes, headaches, mood swings and atherosclerosis.

Key words: Serum testosterone, serum progesterone, surgical menopause, natural menopause, chemiluminiscence

INTRODUCTION

Progesterone is secreted mainly by corpus luteum. Partly metabolized to estrogen and partly to testosterone .It also have many significant actions such as secretory hypertrophy of endometrium, myohyperplasia of uterus, inhibit production of FSH

*Correspondence

Raviraj Naik

Grant Government Medical College, Mumbai

Email: raviraj 40@yahoo.com

and causes sodium retention and relaxation of smooth muscles[1]. Fifty percent testosterone is secreted by adrenal gland and remaining secreted in ovaries by all three types of cells i.e stroma, theca & granulosa but mainly by the theca interna cells of ovarian follicles. 80-85% testosterone are bound to sex – hormone binding protein, 10-15 % to albumin and remaining 1-2% remains free which responsible for its action at peripheral targets mainly hair growth and acne[1]. Menopause is defined as that point in time when permanent cessation of menstruation occurs following loss of ovarian activity.[2] It takes 12 months of amenorrhoea to

ISSN: 2349-0659

confirm that menopause has set in and so it is retrospective diagnosis.[3] *Surgical menopause* is the cessation of menses resulting from surgical removal of the uterus, leaving one or both ovaries, or the removal of both ovaries.[4]

MATERIALS AND METHODS

Cases : Fifty women belonging to surgical menopausal group

Control : Fifty women belonging to Natural menopausal group.

Inclusion Criteria

- 1. Women aging between 44 to 50 years who have undergone Total Hysterectomy in past one to two years.
- 2. Women aging between 44 to 52 years who are experiencing natural menopause since past one to two years.

1. Hormonal intake in any form

- 2. Endocrine disorders
- 3. Testosterone secreting tumors.

We used fully automated enzyme amplified chemiluminescent immunoassay based Immulite 1000 analyzer. The solid phase(bead) is coated with rabbit anti-hormonal polyclonal antibody. The reagent contains alkaline phosphatase conjugated to respective hormone. This hormone-enzyme conjugate competes with respective hormone in patients' blood sample for limited antibody binding sites on bead. The excess sample and reagent are removed by centrifugal wash. Finally chemiluminiscent substrate is added to the bead and signal is generated in proportion to the bound enzyme.

ISSN: 2349-0659

Exclusion Criteria

RESULTS

Table 1: Reference Values as per Siemens diagnostic kit insert

Total Testosterone	ND – 43 ng/dl
Progesterone	ND – 11 ng/ml

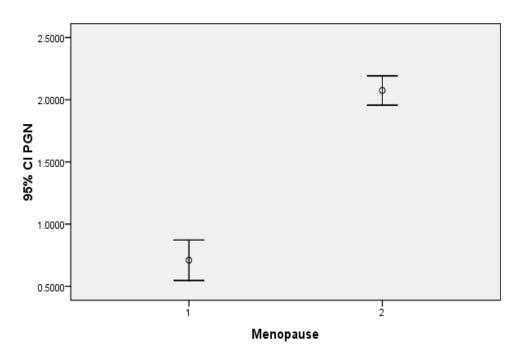
Progesterone: On applying independent (Unpaired) sample t – test, the difference in mean progesterone level between both the groups was found to be very significant with p value < 0.001. (Table – 2 & Figure – 1)

Table 2: Serum mean Progesterone Level of surgical and natural menopausal females

Menopause	N N	sterone Level of surgical a Mean Progesterone [ng/ml]	Standard Deviation [ng/ml]	Standard Error [ng/ml]	
Surgical	50	0.71	0.57	0.08	
Natural	50	2.074	0.42	0.06	
Mean Difference= -1.364 [ng/ml] Independent Sample t test:- t value -13.			63, df-98, p value <0.00)1	

ISSN: 2349-0659

Dot Plot of Serum Progesterone level by Type of menopause



1- Surgical Menopause 2- Natural Menopause

Figure 1: Dot Plot of serum progesterone level by type of menopause

Testosterone: In this study Mann- Whitney U Rank Test was used to compare mean testosterone values which showed significant difference between both the groups of surgical and natural menopause. (Table 3 & Figure 2)

Table 3: Serum mean Testosterone level of surgical and natural menopausal females

Serum mean Testosterone level of surgical and natural menopausal females									
Ranks				Test Statistics					
	Menopause	N	Mean Rank	Sum of Ranks	Mann- Whitn ey U	p Value	Inference		
Serum Testosterone	Surgical	50	43.05	2152.5	877.5	0.01	Significant Difference.		
Level	Natural	50	57.95	2897.5			Difference.		

Naik et al www.apjhs.com

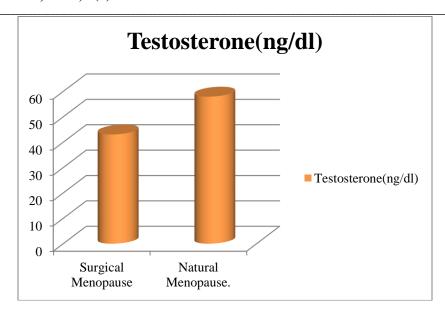


Figure 2: Testosterone levels in surgical and natural menopause

DISCUSSION

In the present study, the serum progesterone levels in surgical menopausal women from 1 to 2 years after surgery was lower than that in the post-menopausal women at similar periods after natural menopause. This indicates that some ovarian progesterone secretion may be sustained for a few years after natural menopause. Similar finding were suggested by Nobuaki Furuhashi et al., [5] who reported significantly decreased levels of progesterone within 2 years after surgical menopause as compared to their levels in natural menopause. As ovary is the major site of synthesis and secretion of estradiol and progesterone in the mammals.[6,7] This might be the reason of significant decrease in progesterone secretion in surgical menopause than that of natural menopause as surgical menopause characterized by faster onset of ovarian dysfunction.[8,9] Present study also reported significant decrease in testosterone levels of surgical menopausal women as compared to its level in natural menopausal women. This view was supported by S. L. Davison et al., [10] who reported that surgical significantly menopausal women had testosterone levels than women in the reference group of natural menopause. Similar findings were also suggested by A. Vermeulen et al., who reported significant decrease in testosterone levels in surgical menopause as compared to natural menopause.[11] Gail A. Laughlin et al., reported 40% decrease while Hughes.[12] CL Jr. et al., reported 50% decrease in testosterone levels in surgical menopause.[13] Significant decrease in testosterone levels in surgical menopause was also revealed by Taylor . M et al., who concluded that this decrease is associated with destabilization of psychiatric axis in women with surgical menopause.[14] In women, testosterone is produced primarily through peripheral conversion of androstenedione (50 percent) with the remainder of production concentrated in the ovary (25 percent) and adrenal cortex (25 percent). [15] Women with surgical menopause have decreased levels of testosterone because production in this condition relies primarily on the adrenal cortex and peripheral conversion of androstenedione[16] while in natural menopause ovary still appears to be source of testosterone.[10] It has also been shown that testosterone levels do not abruptly in women undergoing fall natural menopause due to the preservation of androgen producing theca cells.[17,18] Evidence that a simple hysterectomy hampers ovarian function is proved in various studies.[19-23] Increased bone loss hysterectomy [24] as well premenopausal cardiovascular disease,[25] both conditions associated with ovarian failure has been reported. By blocking the circulation of the uterine vessels, hysterectomy can reduce ovarian blood flow,[26] which may result in of follicular reserve and menopause.[20,22]

ISSN: 2349-0659

CONCLUSION

In the present study, Progesterone and Testosterone were significantly decreased in Surgical menopause

ISSN: 2349-0659

as compared to natural menopause. The sudden change in these hormone levels may be responsible for the increased severity of hot flushes or mood swings seen in the surgical menopausal women as compared to natural menopausal women.

Acknowledgement

To the technical Staff of Special Investigation Laboratory.

REFERENCES

- 1. Padubidri VG, Daftary SN. Shaws Textbook of Gynaecology. 14 ed. New Delhi: Elsevier India Pvt.Ltd; 2008. Physiology. Chapter 3; p:36-41.
- Marc A.Fritz and Leon Speroff. Clinical Gynaecologic Endocrinology and Infertility. 8th ed.New Delhi; Wolter Kluwer India Pvt.Ltd ;2008. Menopausal Perimenopausal and Transition. Chapter 17; p:681.
- Padubidri VG, Daftary SN. Shaws Textbook of Gynaecology. 14 ed. New Delhi: Elsevier India Pvt.Ltd; 2008. Perimenopause, Menopause, Premature Menopause and Postmenopausal bleeding. Chapter 5; p:52-55.
- Kate M. Brett .Can Hysterectomy Be Considered a Risk Factor for Cardiovascular Disease; In the circulation Journal of American heart association. 2005; 111: 1456 - 1458.
- 5. Nobuaki Furuhashi, Tetsuro Abe, Masakuni Suzuki et al : Changes in Hypophysio-ovarian Endocrinological Function of Post-menopausal and Surgical menopausal women: In Tohoku J Med 1976; 120;19-24.
- Abraham, G.E. & Chakmakjian, Z.H. Serum steroid levels during the menstrual cycle in a bilaterally adrenalectomized women. J. clin. Endocr., (1973); 37: 581-587.
- 7. J. Dinny Graham, Christine L. Clarke.Physiological Action of Progesterone in Target Tissues. Endocrine Reviews Aug 1997; 18(4) 502-519.
- 8. Vuorento T, Maenpaa J, Huhtaniemi I. Followup of ovarian endocrine function in premenopausal women after hysterectomy by daily measurements of salivary progesterone.J Clin Endocrinol 1992;36:505–10.
- Riedel HH, Lehmann-Willenbrock E, Semm K. Ovarian failure fenomena after hysterectomy. J Reprod Med 1986;31:597-600.
- 10. S. L. Davison, R. Bell, S. Donath; Androgen Levels in Adult Females: Changes with Age, Menopause, and Oophorectomy: In The Journal of Clinical Endocrinology & Metabolism 2005; 90(7) : 3847-3853.

- 11. A. Vermeulen. The Hormonal Activity of the Postmenopausal Ovary. The Journal of Clinical Endocrinology & Metabolism. 1976; 42(2): 2247-253.
- 12. Laughlin GA, Barrett-Connor E, Kritz-Silverstein D, von Mühlen D. Hysterectomy, oophorectomy, and endogenous sex hormone levels in older women: the Rancho Bernardo Study. J Clin Endocrinol Metab 2000;85:645-51
- 13. Hughes CL Jr, Wall .LL, Creaseman; Reproductive Hormone levels in gynaecologic oncology patients undergoing castration after menopause: In Journal spontaneous gynaecologic oncology . 1991; 40: 42-45.
- **14.** Taylor . M ; Psychological consequences of surgical menopause: In Journal of Reprod Medicine2001; 46(3): 317-24.
- 15. Speroff L, Glass RH, Kase NG. Clinical gynecologic endocrinology and infertility. 5th ed. Baltimore, MD: Williams and Wilkins, 1994:457-
- 16. Norman AW, Litwack G, Hormones, 2nd ed, San Diego, CA: Academic Press, 1997:9-86.
- 17. Braunstein GD. Androgen insufficiency in women: summary of critical issues. J Fertil Steril 2002;77 (Suppl. 4):S94-S99.
- 18. Fogle R, Stanczyk F, Zhang X, Paulson R. Ovarian androgen production in postmenopausal women. J Clin Endocrinol Metab 2007; 92(8): 3040-3043.
- 19. Stone SC, Dickey RP, Mickal A. The acute effect of hysterectomy on ovarian blood flow. Am J Obstet Gynecol 1975;121:193-197.
- 20. Siddle N, Sarrell P, Whitehead M. The effect of hysterectomy on the age at ovarian failure: identification of a subgroup of women with premature loss of ovarian function and literature review. Fertil Steril 1987;47:94-100.
- 21. Kaiser R, Kusche M, Wurz H. Hormone levels in women after hysterectomy. Arch Gynecol Obstet 1989;244:169-73.
- 22. Souza AZ, Fonseca AM, Izzo VM, Clauzet RM, Salvatore CA. Ovarian histology and function after total abdominal hysterectomy. Obstet Gynecol 1986;68:847.
- 23. Oldenhave A, Jaszmann LJ, Everaerd WT, Haspelss AA. Hysterectomized women with ovarian conservation, report more severe climacteric complaints than do normal climacteric women of similar age. Am J Obstet Gynecol 1993;168:765-71.
- 24. Watson NR, Studd JWW, Garnett T, Savas M, Milligan P. Boneloss after hysterectomy with

ovarian conservation. Obstet Gynecol 1995;86:72–7.

- **25.** Ritterband AB, Jaffe IA, Denson PM, Magagna PF, Reed E. Gonadal function and the
- developement of coronary heart disease. Circulation 1963;27:237–51.

ISSN: 2349-0659

26. Janson PO, Janson I. The acute effect on ovarian blood supply. Am J Obstet Gynecol 1977;127:349–352.

Source of Support: NIL Conflict of Interest: None