

## SHORT COMMUNICATION

# URINARY EXCRETION OF 17-KETOSTEROIDS AND 17-HYDROXYCORTICOIDS IN THE OVULATORY AND ANOVULATORY MENSTRUAL CYCLES

U. MATHUR, P. BHARGAVA AND B.B.L. MATHUR

*Department of Physiology, G.R. Medical College, Gwalior*

**Summary:** Excretion of 17-ketosteroids and 17-hydroxycorticoids in three consecutive ovulatory menstrual cycles in pre and post ovulatory phases was investigated in 22 normal medical students. There was a consistent and statistically significant increase in the excretion of 17-hydroxycorticoids during the post ovulatory phase in all the subjects. Except for one subject, the excretion of 17-ketosteroids was also increased in the post ovulatory period. To ascertain if ovulation has any relationship with the increase in the excretion of these metabolites, five an-ovulating subjects were compared with the five ovulating ones. In the anovulating cycles too there was an increase in the 17-ketosteroids and 17-hydroxycorticoids in the premenstrual period.

**Key words:** 17-hydroxycorticoids      17-ketosteroids      urinary excretion      ovulating cycles  
anovulating cycles

## INTRODUCTION

Menstrual cycle is controlled by the cyclic variation in the production of gonadotropins and ovarian hormones. Adrenal cortical and thyroid hormones are also known to exert influence on it. Variations in adrenocortical functions are associated with disturbances in menstrual cycle. There are numerous reports on the excretion of 17-ketosteroids and 17-hydroxycorticoids in normal women (3, 5, 6, 7) but information regarding their variation during the follicular and luteal phases in the normal menstrual cycle and in the anovulatory cycles is negligible. The present study was undertaken to investigate if there are any cyclic variation in the excretion of these metabolites and if it has any relationship with ovulation in normal adult females.

## MATERIALS AND METHODS

The study was carried out in two stages. In the first stage, variations in the excretion of 17-ketosteroids and 17-hydroxycorticoids between the pre and post ovulatory period were observed in 22 normally ovulating medical students who kept a record of their basal body temperature during the entire menstrual cycle. Two 24 hr urine samples were collected: (i) in the first week of the menstrual cycle and (ii) in the last week of the cycle. Thus 17-ketosteroids and 17-hydroxycorticoids were estimated in the pre and post ovulatory phases of three consecutive ovulatory cycles.

In the second part of the study, 5 ovulating and 5 non-ovulating females in the same age group as in the first part *i.e.* 17-20 yrs were investigated. The criterion for anovulation was the failure of elevation of the basal body temperature in the later half of the menstrual cycle. Their 17-ketosteroids and 17-hydroxycorticoids were estimated every 3 or 4 days throughout the menstrual cycle to see if ovulation has any relation with the alteration in the excretion of these metabolites. The 17-ketosteroids were estimated by the method of Callow *et al.* (4) as modified by Vestergaard *et al.* (15), and 17-hydroxycorticoids were estimated by the method of Norymberski *et al.* (9) as modified by Brike *et al.*(2).

## RESULTS

The average excretion of 17-ketosteroids in the pre and post ovulatory phases of the menstrual cycle was 7.45 and 8.72 mg respectively. All the 22 subjects except one showed post ovulatory rise in the 17-ketosteroids excretion.

TABLE I

Sl. No.	Excretion of 17-Ketosteroids in 24 hr. urine in mg. (Average of 3 cycles)			Excretion of 17-Hydroxycorticoids in 24 hr. urine in mg. (Average of three cycles).		
	Preovu	Postovu	Average difference in 17-K.S. excretion	Preovu	Postovu	Average difference in 17-Hydroxycorticoid excretion
1.	9.08	10.01	+0.93±0.14	10.50	13.60	+3.09±2.54
2.	5.89	8.18	+2.29±2.24	10.04	14.67	+4.62±3.73
3.	6.99	6.83	-0.16±1.02	11.83	14.12	+2.29±0.25
4.	5.65	6.77	+1.14±3.08	8.82	12.92	+4.10±0.51
5.	7.67	7.95	+0.28±0.42	5.88	8.59	+2.70±0.22
6.	11.92	12.50	+0.58±9.46	9.90	12.72	+2.82±1.61
7.	7.68	9.81	+2.13±3.48	5.47	9.94	+4.46±3.88
8.	7.68	9.69	+2.01±1.18	5.95	7.99	+2.14±0.84
9.	10.77	11.43	+0.66±0.69	4.69	5.91	+1.32±0.74
10.	5.13	5.92	+0.79±1.20	7.93	9.18	+1.25±0.49
11.	5.76	7.06	+1.29±0.75	9.19	10.67	+1.47±1.11
12.	3.65	6.82	+3.17±1.46	6.65	8.52	+1.86±0.66
13.	5.26	6.39	+1.13±0.83	10.28	12.00	+1.72±0.28
14.	5.50	5.53	+0.03±0.62	4.41	8.17	+3.76±0.96
15.	7.07	7.86	+0.79±2.05	7.03	12.86	+5.83±5.97
16.	6.93	7.21	+0.29±1.57	11.59	13.24	+1.65±0.87
17.	16.43	17.55	+1.14±4.82	15.88	20.32	+4.44±3.92
18.	7.28	9.44	+2.16±1.44	5.64	14.18	+8.64±4.12
19.	5.50	8.61	+3.11±1.18	5.76	10.40	+4.65±1.65
20.	9.02	9.79	+0.77±.55	13.59	15.66	+2.07±1.11
21.	6.10	6.53	+0.43±1.06	8.22	17.20	+8.98±5.32
22.	7.83	9.87	+2.87±3.07	7.00	8.28	+1.28±0.99

The exception *i.e.* subject No. 3 (Table I) showed an increase (average of 0.16 mg) in the pre-ovulatory excretion of 17-ketosteroids in all the three cycles. Statistically, however, this was not significant. The average difference in the 24 hr 17-ketosteroid urinary excretion in the two phases of the menstrual cycle was 1.26 mg (S.D. = 0.99). The excretion of 17-hydroxycorticoids in all the 22 subjects was more in the post-ovulatory period. The average excretion in pre and post ovulatory period was 8.47 and 11.91 mg respectively, with an average difference of 3.84 (S.D. = 2.197) between the two phases.

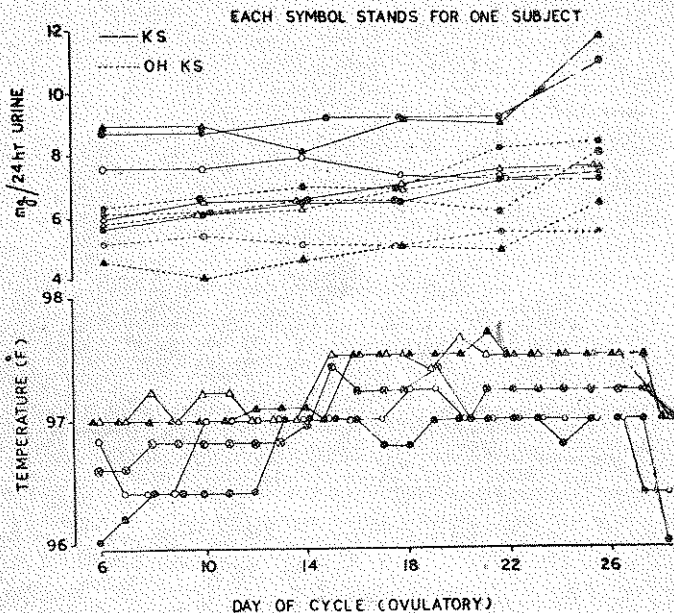


Fig. 1: Urinary excretion of 17-ketosteroids and 17-hydroxycorticoids in the ovulatory cycle.

In the second series of experiments (Fig 1 and 2), there was a premenstrual rise in the excretion of 17-ketosteroids and 17-hydroxycorticoids in all subjects both in ovulatory and anovulatory cycles. The average rise in the 17-ketosteroids excretion in the anovulatory cycles during the premenstrual period was 1.78 mg and that in the 17-hydroxycorticoids excretion during the same period was 1.52 mg. All five ovulating cases studied in this series showed premenstrual rise in the excretion of 17-ketosteroids and 17-hydroxycorticoids, with an average difference of 2.5 mg and 2.22 mg respectively. The premenstrual increase in the excretion of 17-hydroxycorticoids was more marked in the ovulatory than in the anovulatory cycles.

### DISCUSSION

Smith *et al.* (14) using formaldehydogenic method observed 25 to 50% higher excretion of 17-hydroxycorticoids during the period within 36 hrs preceding the onset of menstruation. Pathak and Kahali (11) using indirect method of eosinophil count for the assessment of the adrenocortical

function, observed that in the 18 cycles out of 20 cycles studied, the lowest eosinophil counts fell in the luteal phase of the menstrual cycle.

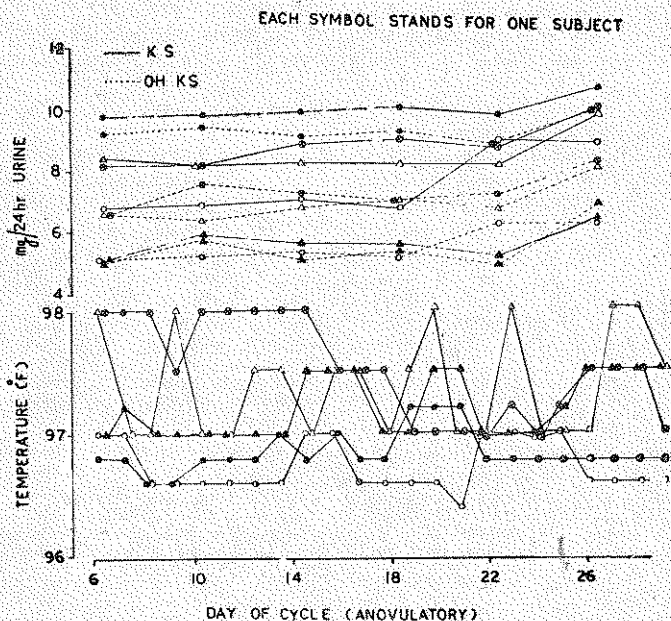


Fig. 2: Urinary excretion of 17-ketosteroids and 17-hydroxycorticoids in the anovulatory cycle.

Smith suggested that the onset of menstruation is preceded by a sudden oxidative inactivation of estrogen(13). A fall in plasma estrogen level in the late luteal phase has also been reported by Mishell Jr. *et al.* (8). The present study shows that the increase in the 17-ketosteroids and 17-hydroxycorticoids in the premenstrual period does not have any relationship with ovulation since it occurs both in ovulating and anovulating females. These results signify that the premenstrual stress acts at the level of central nervous system, possibly by stimulating the hypothalamic pituitary axis thus bringing about increased release of corticotrophin during the premenstrual period (1, 10 & 12).

## REFERENCES

1. Brewer, J. A. Textbook of Gynaecology. ed. 4., 106-108 Baltimore, 1967. The Williams and Wilkins Co.
2. Brike, G., E. Diczfalussy and L. O. Plantin. Assessment of the functional capacity of the adrenal cortex 1. Establishment of normal values. *J. Clin. Endocrinol and Metab.*, **18** : 736-754, 1958.
3. Bulbrook, R. D., B. S. Thomas, J. Utsunomiya and E. Hamauchi. The urinary excretion of 11-Deoxy-17-Oxosteroids and 17-Hydroxycorticosteroids by normal Japanese and British women. *J. Endocrinol.*, **38** : 401-406, 1967.
4. Callow, N. H., R. K. Callow., C. W. Emmens. Colorimetric determination of substances containing grouping  $\text{CH}_{300}$ — in urine extracts as indication of androgen content. *Biochem. J.*, **32** : 1312-1331, 1938.
5. Deshpande, N., J. L. Hayward and R. D. Bulbrook. Plasma 17-Hydroxycorticosteroids and 17-Oxosteroids in patients with breast cancer and normal women. *J. Endocrinol.*, **32** : 167-177, 1965.

6. Hanrotte, J. G., S. Subrahmanyam, A. N. Ramnathan and M. P. Satyanarayanan. Urinary excretion of 17-Ketosteroids in South Indian population. *Lancet.*, 1 : 84-85, 1965.
7. Migdalska, B. Urinary excretion of 17-Hydroxycorticosteroids in normal men and women from 14-80 yr. of age. *Endokr. (Pol)* 15 : 301-308, 1966.
8. Mishell, (Jr) D. R., J. D. Robert, M. Nakamura, P. G. Crosigani, S. Stone, K. Kharama, Y. Nagata and I. H. Thorneycroft. Serum gonadotrophins and steroids pattern during the normal menstrual cycle. *J. Obst. Gyn.*, 111 : 60-65, 1971.
9. Norymberski, J. K., R. D. Stabb, and H. F. West. Assessment of adrenocortical activity by assay of 17-ketogenic steroids in urine. *Lancet.*, I : 1276-1281, 1953.
10. Novak, E. R. and G. S. Jones. Novak's textbook of Gynaecology, ed. 7 : 736-738, Baltimore 1965. The Williams and Wilkins Co.
11. Pathak, C. L. and B. S. Kahali. Cyclic variation in the eosinophil count during the phases of menstrual cycle. *J. Clin. Endocrinol. and Metab.*, 17 : 862-869, 1957.
12. Reeves, B. D., J. E. Garvin and T. W. Incclin. Premenstrual tension: symptoms and weight changes related to potassium therapy. *Am. J. Obst. Gyn.*, 109 : 1036-1041, 1971.
13. Smith, O. W. Menstrual toxin. *Am. J. Obst. Gyn.* 54 : 201-211, 1947.
14. Smith, O. W., V. S. Smith. Rec. Progress in Hormone Research., 8 : 209-245.
15. Vestergaard, P. Rapid micro-modification of Zimmermann/Callow procedure for determination of 17-Ketosteroids in urine. *Acta Endocrinol.*, 8 : 193-214, 1951.