

remaining testis after unilateral orchidectomy (UO). It is important to realize that, although oxygen tension gives a good picture of circulatory changes over short periods of continuous measurement, it is necessary to compare the results of oxygen tension measurements to a specific method of recording capillary blood flow, since increased circulation would presumably reestablish a normal oxygen tension over a long period of time and thus obscure metabolic changes.

Experimental. Animals. Male rats of the Sprague-Dawley strain (Møllegaard-Hansen, Ejby, Denmark) were kept in a controlled environment 14 days before UO, which was performed as described earlier², and an additional 20 days before blood flow measurements. Food and water were available ad libitum.

Xenon-133 clearance. Xenon-133 dissolved in sterile saline was obtained commercially (AB Atomenergi, Studsvik, Sweden). Approximately 20–40 μ Ci in 30–40 μ l of saline was injected percutaneously into the testis with a gastight Hamilton syringe. A Na-I crystal detector (Friesche-Hoepfner 421 A) was used with a 10 mm exit diameter collimator (Friesche-Hoepfner 417 B) with a distance of approximately 20 cm between the crystal and the testicular surface. The detector was connected to a Friesche-Hoepfner 49 A scaler and recording was made on a Hewlett-Packard 7172 A strip chart recorder.

After background subtraction recordings were plotted on semi-logarithmic paper, $t^{1/2}$ was calculated from the wash-out curve and introduced into the following formula in order to obtain blood flow:

$$F = \frac{0.693 \times K \times 100}{t^{1/2}}$$

The partition coefficient K used was 0.7 as given by Wax⁹. Oxygen tension recording. Oxygen tension was measured polarographically as described by Cross and Silver⁸. Recording electrodes were made of 60 nm platinum wire coated with a single layer of glass. A scrotal incision was made and the electrode was inserted through the tunica albuginea. An indifferent silver/silver chloride electrode was inserted into the scrotal cavity via the incision. A polarizing voltage of 0.6 V was applied via the indifferent electrode, and the resultant minute current at the platinum cathode (proportional to the concentration of dissolved oxygen) was amplified and fed to a galvanometer and a Devices strip chart recorder. The mean value from 10 min of stable recording was introduced into a calibration curve, for conversion to oxygen tension values. Calibrations were made before and after each recording in order to avoid errors due to damage to the electrode. The results are presented in the table.

Conclusion. Statistical analysis revealed no significant differences in blood flow or oxygen tension between control rats and UOR. We conclude that no increase in testicular blood flow is detectable with the method used and furthermore that no major metabolic changes are detectable after UO in rats. It is, however, possible that minute changes in capillary blood flow or metabolic rate in the interstitial tissue may be sufficient to account for the endocrine compensation observed¹ since the Leydig cells of the rat testis occupy a mere 2% of the testicular volume¹⁰, and that such minute changes are not detectable with any of the 2 methods used.

9 S. H. Wax, *Invest. Urol.* 9, 167 (1971).

10 E. C. Roosen-Runge, *Anat. Rec.* 123, 385 (1955).

Cyclic variations in luteinizing and follicle stimulating hormone excretion by male rabbits, indicating a male equivalent to the oestrous cycle

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Summary. Cyclic variations of the urinary concentrations of gonadotropins in the male rabbit have been demonstrated. The mean durations of the cycles, 5–7 days, are close to that of the oestrous cycle of the female rabbit.

Ever since the initial observations by Doggett¹ and Kihlström², a considerable amount of data has accumulated indicating that at least some male sexual functions vary cyclically, often with a cycle duration very close to that of the oestrous cycle of the same species. These aspects of male sexual physiology have been discussed at a recent conference³. Previously, the literature within this field was reviewed by Kihlström^{4,5} and Voss⁶.

There are some observations indicating a hormonal regulation of this male cycle in rabbits^{7–12} and in men^{13–15}. Moreover, long-term studies have revealed cyclic variations in plasma testosterone¹⁶ and testosterone excretion^{17,18} in men. There are as yet no similar studies of gonadotropin levels. Because of the short-term pulsatile variation of gonadotropic hormone concentrations in blood^{19,20}, determinations of the urinary concentrations of these hormones are to be preferred in long-term studies. Therefore, urine was continuously collected from healthy 15 sexually mature male rabbits of different breeds daily for 7 weeks. The animals were individually caged and the urine was secured by means of a plastic foil, formed into a funnel below each cage. Every morning the volume of the past 24 h specimen was measured and 5 ml pipetted

into a plastic tube, which was frozen at once for later analysis. The quantitative analysis of the gonadotropins LH and FSH in the urine was carried out using the radioimmunosorbent technique according to Wide et al.²¹. All specimens from each animal were analyzed simultaneously in duplicate, recording the radioactivity with an automatic gammacounter (Autowell, Pickers, USA). Power spectral analysis was performed using an electronic computer, looking for cyclic durations from 2 to 16 days. As seen from the table, all 15 rabbits show statistically significant cyclic variations in LH and/or FSH excretion. Besides, 6 rabbits also show a longer cycle superimposed upon the shorter one. One possible explanation of this phenomenon is that the shortest cycle, having the most narrow band width, is the primary one, the longer ones being multiples of the former. The mean duration of these shortest cycles is 5.1 (LH) and 6.7 days, respectively. Consequently, the present results strongly indicate a hormonal background to a male sexual cycle, corresponding to the female cycle in the rabbit. A full account of this work will be published elsewhere. Studies of the continuous variation of LH and FSH excretion from day to day in man are also in progress.

Power spectral analysis of LH and FSH excretion in urine from 15 rabbits

Rabbit No.	F-value	Luteinizing hormone		Possible cycle duration, days		F-value	Follicle stimulating hormone		Possible cycle duration, days	
		Band width, days	Midpoint of successive bands				Band width, days	Midpoint of successive bands		
G1	2.11	7.09–8.91	8.0	–	8	–	3.24	8.91–12.37	10.6	11 – –
G2	–	–	–	–	–	–	2.21 1.64	8.91–12.37 2.37–2.55	10.6 2.5	11 – 3
G3	2.07 2.04 2.38	4.23– 4.91 2.37–2.55 2.19–2.37	4.6 2.4	–	5	2	–	–	–	– – –
G4	1.60	1.93–2.07	2.0	–	–	2	2.33 1.64	3.77–4.23 1.93–2.07	4.0 2.0	– 4 2
B5	2.38	8.91–12.37	10.6	11	–	–	3.00 3.29	8.91–12.37 7.09– 8.91	10.6 8.0	11 8 –
B6	2.57 2.56	8.91–12.37 7.09– 8.91	9.3	9	–	–	2.09 2.08	7.09– 8.91 5.73– 7.09	7.2	– 7 –
B7	1.65	2.37– 2.55	2.5	–	–	3	1.78	3.77– 4.23	4.0	– 4 –
B8	1.84 2.28	2.79– 3.03 2.55– 2.79	2.8	–	–	3	1.78	3.35– 3.77	3.6	– 4 –
R9	1.90 3.49	4.91– 5.73 5.73– 7.09	5.9	–	6	–	2.15	8.91–12.37	10.6	11 – –
R10	2.49 2.09	7.09– 8.91 5.73– 7.09	7.2	–	7	–	2.45 1.90	8.91–12.37 5.73– 7.09	10.6 6.4	11 6 –
R11	2.29	8.91–12.37	10.36	11	–	–	2.61	8.91–12.37	10.6	11 – –
R12	1.89 1.96 1.80	2.37– 2.55 2.19– 2.37 2.07– 2.19	2.3	–	–	2	3.22	8.91–12.37	10.6	11 – –
G13	2.41 2.84	2.37– 2.55 2.19– 2.37	2.4	–	–	2	2.11 3.14	2.55– 2.79 2.37– 2.55	2.6	– – 3
G14	2.48 2.20 1.63	3.35– 3.77 3.03– 3.35 1.93– 2.07	3.4 2.0	–	–	3,2	1.80 1.89	3.03– 3.35 1.93– 2.07	3.2 2.0	– – 3,2
G15	1.77 2.19 1.78	3.35– 3.77 3.03– 3.35 2.79– 3.03	3.2	–	–	3	2.50	8.91–12.37	10.6	11 – –

All F-values indicated above: $p < 0.05$.

- V. C. Doggett, *Am. J. Physiol.* 187, 445 (1956).
- J. E. Kihlström, Studies on some activities of the male accessory glands, especially the production of male sperm antagglutin and their relations to fertility (Uppsala 1958).
- M. Ferin, F. Halberg, R. M. Richart and R. L. Vande Wiele, *Biorhythms and human reproduction*. John Wiley & Sons, New York 1974.
- J. E. Kihlström, *Experientia* 22, 630 (1966).
- J. E. Kihlström, in: *Current problems in fertility*. Ed. A. Ingelman-Sundgren and N-O Lunell. Plenum Press, New York 1971.
- H. E. Voss, *Handb. exp. Pharmacol.* 35, 420 (1973).
- J. E. Kihlström and O. Hornstein, *Acta endocr., Copenh.* 46, 597 (1964).
- J. E. Kihlström and O. Hornstein, *Z. vergl. Physiol.* 49, 191 (1964).
- O. Hornstein, J. E. Kihlström and G. Degerman, *Acta endocr., Copenh.* 46, 608 (1964).
- J. E. Kihlström and D. Fjellström, *J. Reprod. Fert.* 19, 375 (1969).
- G. Degerman and J. E. Kihlström, *Acta physiol. scand.* 78, 567 (1970).
- J. E. Kihlström, *Acta endocr., Copenh.* 70, 360 (1972).
- J. C. Månsson, *Life Sci.* 4, 329 (1965).
- J. E. Kihlström, *Life Sci.* 10, part II, 321 (1971).
- G. H. Schmid, O. P. Hornstein, O. Mittmann and M. Münstermann, *Acta Cytol.* 16, 352 (1972).
- C. H. Doering, H. C. Kraemer, H. K. H. Brodie and D. A. Hamburg, *J. clin. Endocr. Metab.* 40, 492 (1975).
- A. A. A. Ismail and R. A. Harkness, *Acta endocr.* 56, 469 (1967).
- R. A. Harkness, in: *Biorhythms and human reproduction*, p. 469. Ed. M. Ferin, F. Halberg, R. M. Richart and R. L. Vande Wiele. John Wiley & Sons, New York 1974.
- H. R. Nankin and P. Troen, in: *Biorhythms and human reproduction*, p. 457. Ed. M. Ferin, F. Halberg, R. M. Richart and R. L. Vande Wiele. John Wiley & Sons, New York 1974.
- A. Vermeulen, L. Verdonck and F. Comhaire, in: *Biorhythms and human reproduction*, p. 427. Ed. M. Ferin, F. Halberg, R. M. Richart and R. L. Vande Wiele. John Wiley & Sons, New York 1974.
- L. Wide, S. J. Nillius, C. Gemzell and P. Roos, *Acta endocr., Copenh., suppl.* 174 (1973).