EFFECTS OF DISRUPTION OF DIURNAL RHYTHMS IN FOOD INTAKE OF RATS

B. S. RAO

Department of Physiology, St. John's Medical College, Bangalore–560 034

and

K. N. SHARMA

Department of Physiology, University College of Medical Sciences, Ring Road, New Delhi–110 016

(Received on April 28, 1980)

Summary : The calorie intake, body weight (bw) mainenance and oral intake of *ad lib* fed (Gr I) and feeding rhythm-disrupted (Gr II) adult rats of either sexes are investigated. Gr I calorie intake $(19.2\pm0.6 \text{ cal}/100 \text{ } gm \text{ bw})$ and bw $(214\pm5.6 \text{ } gms)$ were stabilized after 130-140 days of age. Rats of similar age (130-140 days) on feeding rhythm disruption (Gr II) showed initial decrease in calorie intake and bw but 136 days later their bw $(224\pm6.5 \text{ } gms)$ was similar to Gr I bw $(226\pm9.2 \text{ } gms)$ though calorie intake $(17.5\pm0.4 \text{ cal } /100 \text{ } gm \text{ bw})$ was still less as compared Gr I intake. This disparity in calorie intake and bw in Gr II appears to be linked to the adaptive metabolic changes.

The solution and mixed-diets tests showed that oral intake is taste-dependant in Gr II but not in Gr I. Gr II intake was increased on sweet taste and decreased on salt and bitter tastes indicating that feeding rhythm-disruption enhances sensory regulation of intake in contrast to calorie regulation found predominant on free feeding.

Key words: feeding rhythm

calorie-intake

body weight,

, intake regulation

INTRODUCTION

Rats are nocturnal animals and ingest nearly 60 to 80% of their total daily for intake during the night (11 and 12). The feeding patterns are shown to parallel th general activity (22 and 23) and blood glucose levels (8). The feeding rhythms ma easily be disrupted by restricting the time of availability of food to a period which fall outside the normal rhythm of the animal. A few such studies have indicated change in metabolic activities (6) as a corollary to the disruption of feeding rhythms. Howeve the long-term effects of feeding rhythm disruption on calorie intake, body weight main tainance, responses to sensory and metabolic properties of the diet are largely unknown The present investigation is intended to meet this lacuna in the knowledge.

s ha s - 1 e b rat: sts ven ir 1 the ilph ige on

) :

4

me

Volume 24 Number 3

MATERIALS AND METHODS

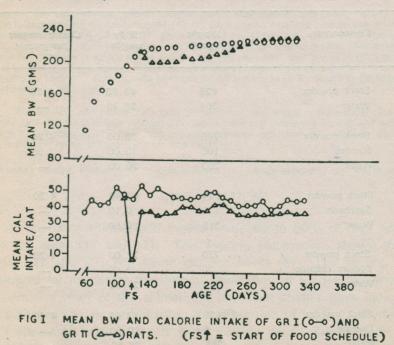
Adult Wistar rats of either sexes and housed in individual cages were used. Group I rats had access to *ad lib* diet whereas Gr II rats were adapted to 3 hr food schedule (09:15 hrs - 12:15 hrs) for a period of 3-4 months. Water was available *ad lib* for both groups. The body weight (bw), food intake and water intake, were measured daily. Both groups of rats were investigated for gustatory responses to 1 hr (08:00 hrs - 9:00 hrs) single bottle tests and to mixed diet intake tests administered randomly. The stock or mixed diet was given to animals immediately after the solution test either for 24 hours (Gr I) or for 3 hours (Gr II) depending on the food schedule. The test solution or mixed diet used, contained either sweet (glucose, sucrose and saccharin) or salty (sodium chloride) or bitter (quinine sulphate) substances in concentrations as shown in Table I. During solution tests, the cage was cleared of food and water, whereas during mixed diet tests the water was available along with the diet. The intake of solution was observed at regular intervals of 5, 15, 30 and 60 min whereas the intake of diet was measured only after the scheduled time of 24 hours (Gr I) or 3 hrs (Gr II).

SI. No.	Diet	Components	Weight (gms)	% by weight	Caloric content per gram	Total cal/ 1 gram of diet
1.	Stock	Stock powder	636	63.60	4.20	2.62
		Water	364	36.40	-	
2.	Sucrose	Stock powder	700	70.00	4.20	3.3
		Sucrose	_ 100	10.00	3.98	
		Water	200	20.00		
3.	Saccharin	Stock powder	634	63.40	4.20	
		Saccharin	1	0.10	S-10-3	2.62
		Water	365	36.50	56-201	
4.	Salt	Stock powder	720	72.00	4.20	
		Sodium chloride	30	3.00	A-10-53	3.0
		Water	250	25.00	- 60	
5. (Quinine	Stock powder	630	63.00	4.20	
		Quinine	5	0.50	1 93 -	2.62
		Water	365	36.50	-	

TABLE I : Components of diets and their caloric density.

RESULTS

The calorie intake of Gr I and Gr II rats and their bws are shown in Fig. 1 The Gr I rats showed a rapid bw growth upto the age of 130-140 days after which the bw wa almost stabilized at 214+5.6 gm. The bw of Gr. II rats (216+4.5 gm) at the age d 130 days was comparable to that of Gr I rats when the 3 hr-food schedule was initiated However, in the first 36 day-period following the initiation of 3-hr food schedule the bu of Gr II rat steadily decreased to the lowest level of 200+8.7 gm. After this 36 da period, the Gr II rats showed a steady increase in their bw and at the end of further 100 day period the bw of Gr II rats $(224\pm6.5 \text{ gm})$ was similar to the bw of Gr I rats (226 ± 9.2) am) of comparable age (260 days). At 320 days of age the bw of Gr II rats (236.2 \pm 73 am) even surpassed by of Gr I rats (234.5±9.8 gm.) Parellel to the changes in by, the calorie intake of Gr I rats showed in general an increase during the period of steep growth in bw and evidenced lowered calorie intake (19.2 ± 0.6) cal 100 gm bw) when the growth reached an asymptote. On the other hand, the Gr II rats reduced calorie intake for a period of 8-10 days immediately after they were put on 3-hr food schedule. Later they stabilized food intake at 17.5 ± 0.4 cal/100 gm bw (Table II) throughout the period which was significantly less as compared to the intake of Gr I rats (P<0.05).



The results of oral intake of solutions and diets of both Gr I and Gr II rats computed

Volume 24 Number 3

per 100 gm bw are shown in Table II and Table III respectively. Intra group analysis of solution intake showed that the Gr I rats ingested higher amounts of glucose solution $(45\pm0.2 \text{ ml})$ as compared to their intake of the other three (Saccharin, sodium chloride and quinine sulphate) solutions whereas Gr II rats intake of glucose $(4.8\pm0.1 \text{ ml})$ was similar to the intake of saccharin $(4.3\pm0.3 \text{ ml})$ and was higher as compared to intake of salt $(3.6\pm0.4 \text{ ml})$ and of quinine $(0.8\pm0.1 \text{ ml})$ solutions. The intergroup analysis showed that the intake of the Gr II rats increased slightly on glucose $(4.8\pm0.1 \text{ ml})$ and saccharin $(4.3\pm0.3 \text{ ml})$, and significantly on sodium chloride $(3.6\pm0.4 \text{ ml})$, but decressed slightly on quinine $(0.8\pm0.1 \text{ ml})$ as compared to respective solution intakes of Gr I rats. It is note worthy that the intake of Gr I rats on glucose solution was significantly higher as compared to their intake on saccharin solution, though both glucose and saccharin are sweet in taste. In contrast, the Gr II rats ingested similar amounts on glucose and saccharin solutions.

TABLE II : Solution intake (mean $mI \pm SE/100 \ gm$ bw) of rats.

Solution	Gr / Million de dist	Gr II
THE OF BRIDER	te ald gostanon contract the of	
Glucose	4.5±0.2	4.8±0.1
Saccharin	3.7±0.2	4.3±0.3
Sodium chloride	2.5 <u>+</u> 0.1	3.6±9.4
Quinine sulphate	0.9±0.2	0.8±0.1

The calorie intake of both Gr I and Gr II rats on stock diet and on the mixed diets Table III) showed some notable changes. The calorie intake of Gr I rats on sucrose $(19.1\pm0.9 \text{ cal})$, saccharin $(20.5\pm0.5 \text{ cal})$ and on salt diet $(17.7\pm1.2 \text{ cal})$ was comparable to their intake on stock diet $(19.2\pm0.6 \text{ cal})$. The Gr II rats on the other hand increased their calorie intake on sucrose $(22.1\pm0.9 \text{ cal})$ and saccharin diet $(19.2\pm0.7 \text{ cal})$ but showed a decrease on the salt diet (14.7±0.9 cal) as compared to their intake on stock diet $(17.5\pm0.4 \text{ cal})$. Both Gr I and Gr II rats decreased significantly on the quinine diet the intakes being 7.6 ± 1.3 cal and 6.8 ± 0.6 cal respectively. Intergroup analysis showed that the calorie intake of Gr. I rats on stock diet (19.2 ± 0.61) was significantly higher as compared to the intake of Gr II rats $(17.5\pm0.4 \text{ cal})$ on the similar stock diet. The Gr I rats intake on salt diet $(17.7 \pm 1.2 \text{ cal})$ and on quinine diet $(7.6 \pm 1.3 \text{ cal})$ was also higher as compared to Gr II rats intake on salt (14.7 \pm 0.9 cal) and quinine (6.8 \pm 0.6 (al) diets, though the increase on quinine diet was only slight. This trend was almost reversed on the addition of sweet taste to diet. The Gr II rats calorie intake on sucrose det (22.1 \pm 0.9 cal) was higher than Gr I calorie intake on sucrose diet (19.1 \pm 0.85 cal). In saccharin diet the Gr II intake (19.2 ± 0.7) was similar to Gr I intake $(20.5 \pm 0.5 \text{ cal})$.

174 Rao and Sharma

July-September 1980 Ind. J. Physiol, Pharmac

TABLE III : Calorie intake (mean ± SE/100 gm bw) of Group ! and Group !I rats and on stock and test diets.

Diet	Group 1	Group II
Stock	19.2 <u>+</u> 0.6	17.5±0.4
Sucrose	19.1 <u>+</u> 0.9	22.1±0.9
Saccharin	20.5±0.5	19.2 <u>±</u> 0.7
Salt	17.7±1.2	14.7±0.9
Ouinine	7.6 <u>+</u> 1.3*	6.8±0.6

*P < 0 05 Stock diet intake of the group taken as control.

DISCUSSION

The diurnal eating rhythm of rats was disrupted by restricting the availability of food between 09.15 and 12.15 hrs. The effects of eating rhythm disruption on by maintenance, calorie, intake, and gustatory responses were investigated. The earlier investigations have shown that the bw of the animals is proportional to its calorie intake (3). However, the Gr II rats, adapted to the 3-hour food schedule for 136 days, had bws (224±6.5 gms) similar to the bws of Gr I rats (226±9.2 gms) though the calore intake of Gr II rats (17.5±0.4 cal/100 gm bw) was less as compared to the intake of GrI rats (19.2±0.6 cal/100 gm bw). Identical bws on different calorie intakes appear to be due to the changes in the metabolic adaptive changes induced by the disruption in rhythms of eating (2) or reduction in energy expenditure by "meal eaters" as the compared to "nibblers" (4 and 5) or perhaps due to both. As an adaptation to the nonavailability of food during species-specific rhythms of eating, the animals are shown to develop efficiency in the conversion of ingested food into bodily stores of lipids and glycogen (9, 17, 18). It has been shown by several investigations (1, 2, 6, 15, 16, 21, 24) that the increase in bw of animals on food schedule is due to the deposition of fat and to a lesser extent to increased glycogen. The increase in bodily stores (lipids and glycogen) of rats on food schedule may be an anticipatory response because the increase in the deposition of fat has also been observed in migrating birds, in hybernants, in early pregnancy of female mammals and in long distance swimmers (7). In addition, anticipatory systemic responses are also demonstrated for several physiological early processes (19 and 20). The fat deposits of animals starved intermittenty may be used by them during periods of starvation. However the mechanisms triggering the changes in metabolic activity favouring the fat deposition are still not clear though ventromedial hypothalamus (VMH) is speculated to lacate such mechanisms (7 and 13). Further, Lepkovsky (14) suggested that the "set point" for energy stores may be shifted to a higher Volume 24 Number 3

level during intermittent starvation, which in turn leads to increased deposition of fats. By such a shift in the "set point" the rats on 3-hr food schedule (Gr II) may behave as animals in energy-deficit favouring the conservation of energy.

One of the behaviours of energy-deficit animals as suggested by the model of Jacobs and Sharma (10) is that they show prepotent sensory regulation of intake. The Gr II rats ingested similar amounts of glucose and saccharin solutions (Table I). As both the solutions are sweet for taste though calorically different the ingestive behaviour of Gr II rats could be interpreted to be sensory regulated. This conclusion was further substantiated by mixed diet tests (Table II). The Gr II rats ingested significantly higher amounts of calories on sweet taste and significantly lower amounts on salty and bitter taste as compared to their intake on stock diet. In contrast the Gr I rats calorie intake on sucrose, saccharin and salt mixed diets was similar to their intake on stock diet. Thus both on solution as well as on mixed diet tests the Gr I rats showed predominent calorie regulation of intake.

In short, the present investigation shows that on the disruption of diurnal rhythms of eating in rat, the calorie intake fells sharply and later stabilized at lower level as compared to intake on free feeding. The bw usually followed the fall and rise in calorie intake. However, after adaptation to rhythm disruption the bw is disproportionately increased on lowered calorie intake. Further the intake of rhythm-disrupted rats was sensory regulated in contrast to calorie-regulated intake of *ad lib* fed rats though the bws of both the rhythmdisrupted and the *ad lib* fed rats were similar.

REFERENCES

- 1. Cohn, C., E. Shrago and D. Joseph. The effect of food administration on weight gain on body composition of normal and adrenalectomised rats. Am. J. Physiol., **18**: 503-507, 1955.
- 2. Cohn, C. and D. Joseph. Role of rats of ingestion of diet on regulation of intermediate metabolism (mealeating vs nibbling). Metab. Clin. Exp., 9: 492-500, 1960.
- 3. Collier, G. Body weight loss as a measure of motivation in hunger and thirst. Ann. N.Y. Acad. Sci., 157 : 594-609, 1969.
- 4. Fabry, P., R. Petrasek and L. Krulich. Oxygen consumption of rats adapted to intermediate starvation. *Physiol. Bohemoslov.*, **10** : 362-369, 1961.
- 5. Fabry, P., R. Petrasek, E. Horakova, E., Konapesek, and T. Braun. Energy metabolism and growth in rats adapted to intermittent starvation. *Brit. J. Nutr.*, **17**: 295-301, 1963.
- 6. Fabry, P. Metabolic consequences of the pattern of the food intake. In : Hand Book of Physiology, Sec 6, Alimentary Canal, Vol I, (Ed) C.F. Code., American Physiological Society, Washington, 1967.
- 7. Harvey, G. In : Hunger : Basic mechanism and clinical implications. (Ed) D. Novin, W. Wyrwicka and G. Bray, Raven Press, New York, 1976.
- 8. Higgins, G.M., J. Berkson and E. Flock. The diurnal cycle in the liver. Am. J. Physiol., 102: 673, 1932.
- 9. Hollifield, G. and W. Parson. Metabolic adaptation to a "stuff and starve" feeding program, I, studies of adipose tissue and liver glycogen in rats limited to a short daily feeding period. J. Clin. Invest., 41: 245-249, 1962.
- 10. Jacobs, H.L. and K.N. Sharma. Taste versus calories : sensory and metabolic signals in the control of food intake. Ann. N.Y. Acad. Sci., 157 : 1084-1125, 1969.

176 Rao and Sharma

July-September 19 Ind. J. Physiol. Pharma

- 11. Le Magnen, J. and S. Tallon. La periodicite spontanee de la prise dialiments ad libitum du rat blanc. J. Physical Science of the second seco
- 12. Le Magnen, J. Habits and food intake. In : Hand Book of Physiology, Sec 6, Alimentary Canal, Vol I, B C.F. Code, American Physiological Society, Washington, D.C., 1967.
- 13. Le Magnen, J. Gluco lipostatic mechanisms and feeding. In : Hunger : Basic mechanisms and clinical imple tions. (Ed) D. Novin, W. Wyrwicka and G. Bray, Raven Press, New York, 1976.
- 14. Lepkovsky, S. Control mechinary in the regulation of food intake and body weight. In : Regulation and Com in Physiological systems. (Ed) A.S. iberall, and A.C. Guyton, 1973.
- 15. Leveille, G.A. Lipogenesis in adipose tissue of meal-fed rats. A possible regulatory role of alpha-glycerophyper
- formation. Canad. J. Physiol., 45: 201-214, 1966a. 16. Leveille, G.A. Glycogen metabolism in meal-fed rats and chicks and the time sequence of lipogenic a enzymatic adaptive changes. J. Nutr., 90: 449-460, 1966b.
- 17. Leveille, G.A. In vivo fatty acid systhesis in adipose tissue and liver of meal-fed rats. Proc. Soc. Exp. But
- 18. Leveille, G.A. and K. Chakrabarty. Absorption and utilization of glucose by meal-fed and nibbling rats. J. Nat
- 19. Nicolaidis, S. Early systemic responses to orogastric stimulation in the regulation of food and water balance functional and electrophysiological data. Ann. N.Y. Acad. Sci., 157 : 1176-1203, 1969.
- 20. Nicolaidis, S. Sensory-neuroendocrine reflexes and their anticipatory and optimizing role on metabolism in The chemical senses and nutrition., (Ed) M.R. Kare and O. Maller, Academic Press, New York, 1977.
- 21. Putten, L.M., D.W. Van, Bekkum Van and A. Quericd. Influence of hypothalamic lesions producing hyperplay and of feeding regimens on carcass composition in the rat. Metabolism, 4: 68-74, 1955.
- Pichter, C.P. A behaviouristic study of the activity of the rat. Comp. Psychol. Monogr., 1: 55, 1922. 23. Richter, C.P. Animal behaviour and internal drives. Quart. Rev. Biol., 2: 307-343, 1927.
- 24. Tepperman, J. and H. N. Tepperman, The hexose mono phosphate shunt and adaptive hyper lipogenes J. Diabetes, 7: 478-485, 1958.