

**UNIVERSIDADE FEDERAL DE MINAS GERAIS**  
**ESCOLA DE EDUCAÇÃO FÍSICA, FISIOTERAPIA E TERAPIA**  
**OCUPACIONAL**

**TERMORREGULAÇÃO E PERFORMANCE EM**  
**RATOS DURANTE EXERCÍCIO PROGRESSIVO**  
**EM DIFERENTES FASES DO CICLO**  
**CIRCADIANO**

**Frederico Sander Mansur Machado**

**BELO HORIZONTE**

**JULHO DE 2010**

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Trabalho de conclusão de curso elaborado pelo aluno de graduação em Educação Física **Frederico Sander Mansur Machado**, sob orientação do Prof. Dr. Cândido Celso Coimbra (ICB – UFMG).

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**JULHO DE 2010**

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**Trabalho de Conclusão de Curso**

***TERMORREGULAÇÃO E PERFORMANCE EM RATOS DURANTE EXERCÍCIO  
PROGRESSIVO EM DIFERENTES FASES DO CICLO CIRCADIANO***

Trabalho de Conclusão de Curso aprovado pela Escola de Educação Física, Fisioterapia e Terapia Ocupacional (EEFTO) da Universidade Federal de Minas Gerais (UFMG) como requisito parcial para obtenção do grau de Graduado em Educação Física, em \_\_\_\_ de Julho de 2010.

RESULTADO: \_\_\_\_\_

NOTA: \_\_\_\_\_

CONCEITO: \_\_\_\_\_

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## RESUMO

O presente trabalho tem o objetivo de analisar a temperatura corporal interna ( $T_b$ ) e a atividade locomotora espontânea (ALE) além da performance durante o exercício incremental até a exaustão em ratos machos em diferentes fases do ritmo circadiano.

Os ritmos circadianos da atividade locomotora e temperatura corporal de ratos Wistar adultos ( $n = 11$ ) foram registradas continuamente através de um sensor de telemetria (G2 E-Mitter Mitter Mini-Empresa, Oregon, EUA) implantado intraperitonealmente. Após a avaliação do ritmo circadiano, os animais foram familiarizados ao exercício em esteira motorizada durante cinco dias consecutivos e, em seguida, submetidos ao protocolo de teste incremental de exercício até a fadiga no início da fase clara e escura do ritmo circadiano claro/escuro. Cada teste foi separado por três dias. Os animais começavam correndo a uma velocidade inicial de  $10 \text{ m} \cdot \text{min}^{-1}$  e aumentos de  $1 \text{ m} \cdot \text{min}^{-1}$  a cada 3 min eram realizados até a fadiga. A inclinação da esteira foi mantida constante em  $5^\circ$ . As temperaturas pré-exercício e pós-exercício foram monitoradas. O tempo total de exercício (TTE), carga de trabalho realizado (W) e a temperatura interna ( $\Delta T_b$ ) foram registrados.

Como esperado, as médias circadianas de ALE e  $T_b$  observadas durante escuras foram maiores quando comparadas as da fase clara. A ALE durante a fase escura foi aproximadamente 70% superior ao correspondente da fase clara ( $p < 0,0001$ ). A  $T_b$  observada durante a fase escura foi maior do que a observada durante a fase clara ( $p < 0,0001$ ). Por outro lado, o exercício realizado no início da fase clara apresentou um tempo total de exercício (TTE), hipertermia ( $\Delta T_b$ ), taxa de aumento de temperatura e taxa de acúmulo de calor maiores do que os observados durante a fase escura ( $p < 0,05$ ). A TTE durante a fase clara foi aproximadamente 10% maior ( $p < 0,05$ ) do que os observados durante a fase escura. O trabalho realizado foi 30% maior ( $p < 0,01$ ) durante a fase clara do que o observado em relação à fase escura. Todas as variáveis de temperatura relacionadas ao exercício apresentaram diferenças significativas entre as fases. A taxa de aumento de temperatura ( $^\circ\text{C} \cdot \text{min}^{-1}$ ) da fase

escura foi superior a observada durante a fase clara ( $p<0,05$ ). A taxa de armazenamento de calor foi 80% ( $p<0,01$ ) maior do que a da fase escura correspondente. Os animais com maior carga de trabalho durante a fase escura apresentou os melhores resultados de carga de trabalho também durante a fase clara ( $r=0,89$ ,  $p<0,001$ ). Uma relação direta da atividade locomotora espontânea circadiana e a capacidade de exercício foi observada apenas durante a fase clara mostrando uma relação positiva entre ALE (específica à fase clara) e o W realizado ( $r = 0,75$ ,  $p<0,01$ ).

Em conclusão, os resultados mostraram que ratos em condições ambientais normais de temperatura apresentaram melhor desempenho em protocolos de exercício progressivo durante a fase inicial do período claro e esta foi acompanhada por diferenças na termorregulação pré e pós-exercício. Além disso, apesar das diferenças entre a capacidade de exercício e termorregulação em duas fases circadiano, atividade locomotora espontânea circadiana estão relacionadas com a capacidade de exercício apenas durante a fase clara.

## ABSTRACT

The aims of the present study were to analyze responses to incremental exercise in male rats in different phases of light/dark circadian rhythm of body temperature ( $T_b$ ) and spontaneous locomotor activity (LA).

Circadian locomotor activity and body temperature of adult male Wistar rats ( $n=11$ ), were continuously recorded using an previously implanted (i.p.) telemetric sensor (G2 E-Mitter Mini-Mitter Company, Oregon, EUA). After circadian rhythm recording, animals were familiarized to a motor-driven treadmill during five consecutive days and then submitted to an incremental exercise test protocol until fatigue during early light and dark-phases of activity. Each test was separated for 3 days. During the graded test, animals started running at an initial speed of  $10 \text{ m} \cdot \text{min}^{-1}$  and increased by  $1 \text{ m} \cdot \text{min}^{-1}$  every 3 min until fatigue. Treadmill inclination was held constant at  $5^\circ$ . Pre-exercise and post-exercise temperature was monitored. Time to fatigue (TTF), workload (W) and body temperature ( $\Delta T_b$ ) were registered.

As expected, circadian dark-phase LA and  $T_b$  were higher when compared to light-phase. Dark-phase LA was 70% higher than light-phase correspondent ( $p<0.0001$ ). Dark-phase  $T_b$  was higher than light-phase ( $p<0.0001$ ). In contrast, light-phase exercise time to fatigue (TTF), hyperthermia ( $\Delta T_b$ ), body heat rate (BHR) and heat storage rate (HSR) were higher than dark-phase ( $p<0.05$ ). Light-phase TTF was 10% higher ( $p<0.05$ ) than dark-phase. It was observed a 30% higher ( $p<0.01$ ) light-phase W compared to dark-phase. Temperature exercise-related variables showed significant differences between phases. Dark-phase body heat rate was higher than light-phase correlate ( $p<0.05$ ). Light-phase heat storage rate was 80% higher ( $p<0.01$ ) than dark-phase correspondent. Animals with higher workload during dark-phase also showed better workload results during light-phase ( $r=0.89$ ,  $p<0.001$ ). There was a direct relationship of circadian activity and exercise capacity at light phase with a positive relationship between LA and W ( $r=0.75$ ,  $p<0.01$ ).

In conclusion, results demonstrated that rats under normal environmental conditions showed better graded exercise performance during early light-phase and this was accompanied by differences in thermoregulation pre and post-exercise. Besides, despite the differences between exercise capacity and thermoregulation at two circadian phases, circadian locomotor activity was related to exercise capacity only during light-phase.

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## 1 INTRODUÇÃO





## 2 OBJETIVOS

## **3 MATERIAIS E MÉTODOS**

### **3.1 AMOSTRA**

### **3.2 PROCEDIMENTOS CIRÚRGICOS**

### 3.3 REGISTRO DO RITMO CIRCADIANO

### **3.4 PROTOCOLO DE EXERCÍCIO PROGRESSIVO**

### **3.5 DELINEAMENTO EXPERIMENTAL**

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## **4 RESULTADOS**

### **4.1 RITMO CIRCADIANO**





## 4.2 EXERCÍCIO PROGRESSIVO



## 5 CONCLUSÃO

## 6 REFERÊNCIAS BIBLIOGRÁFICAS

ALDEMIR H, ATKINSON G, CABLE NT *et al.* Comparison of the immediate effects of moderate exercise in the early morning and late afternoon on core temperature and cutaneous thermoregulatory mechanisms. *Chronobiol Int* 2000 17: 197–207.

ARMSTRONG RB, LAUGHLIN MH, ROME L, TAYLOR CR Metabolism of rats running up and down an incline. *J Appl Physiol Respirat Environ Exercise Physiol* 1983; 55(2): 518-521.

ATKINSON G, EDWARDS B, REILLY T, WATERHOUSE J. Exercise as a synchronizer of human circadian rhythms: an update and discussion of the methodological problems. *Eur J Appl Physiol* 2007; 99:331–341.

BALTHAZAR CH, LEITE LH, RIBEIRO RM, SOARES DD, COIMBRA CC. Effects of blockade of central dopamine D1 and D2 receptors on thermoregulation, metabolic rate and running performance. *Pharmacol Rep* 2010; 62(1): 54-61.

BENSTAALI C, MAILLOUX A, BOGDAN A, AUZÉBY A, TOUITOU Y. Circadian rhythms of body temperature and motor activity in rodents: Their relationships with the light-dark cycle. *Life Sciences* 2001; 68: 2645–2656.

BROOKS GA, DONOVAN CM, WHITE TP. Estimation of anaerobic energy production and efficiency in rats during exercise. *J Appl Physiol* 1984; 56 (2): 520-525.

\_\_\_\_\_, WHITE TP. Determination of metabolic and heart rate responses of rats to treadmill exercise. *J Appl Physiol* 1978; 45 (6): 1009-1015.

BUTLER MP, KRIEGSFELD LJ, SILVER R. Circadian Regulation of Endocrine Functions. In: *Hormones, Brain and Behavior*. 2<sup>nd</sup> ed. Edited by PFAFF DW, ARNOLD AP, ETGEN AM, PURPURA DP, FAHRBACH SE, RUBIN RT. USA: Elsevier science, 2009.

BUTTNER D, WOLLNIK F. Spontaneous short-term fluctuations in the daily pattern of heart rate, body temperature and locomotor activity in the laboratory rat. *Laboratory Animals* 1982; 16: 319-326.

CONTI B, SANCHEZ-ALAVEZ M, WINSKY-SOMMERER R, MORALE MC, LUCERO J, BROWNELL S, FABRE V, HUITRON-RESENDIZ S, HENRIKSEN S, ZORRILLA EP, LECEA L, BARTFAI T. Transgenic mice with a reduced core body temperature have an increased life span. *Science* 2006; 314: 3.

DIAZ EO, PRENTICE AM, GOLDBERG GR, MURGATROYD PR, COWARD WA. Metabolic response to experimental overfeeding in lean and overweight healthy volunteers. *Am J Clin Nutr* 1992; 56 (4): 641-655.

GORDON, CJ. 24-Hour control of body temperature in rats - Integration of behavioral and autonomic effectors. *Am J Physiol Regulatory Integrative Comp Physiol* 1994; 267 (36): R71-R77.

HASEGAWA H, YAZAWA T, TANAKA H, YASUMATSU M, OTOKAWA M, AIHARA Y. Effects of ambient light on body temperature regulation in resting and exercising rats. *Neuroscience Letters* 288 (2000) 17-20.

KOTZ CM, TESKE JA, BILLINGTON CJ. Neuroregulation of nonexercise activity thermogenesis and obesity resistance. *Am J Physiol Regul Integr Comp Physiol* 2008; 294 (3): R699-R710.

KRIEGSFELD LJ, LESAUTER J, HAMADA T, PITTS SM., SILVER R. Circadian Rhythms in the Endocrine System. in: *Hormones, Brain and Behavior* volume two. USA: Elsevier Science, 2002.

LACERDA ACR, MARUBAYASHI U, BALTHAZAR CH, LEITE LHR, COIMBRA CC. Central nitric oxide inhibition modifies metabolic adjustments induced by exercise in rats. *Neuroscience Letters* 2006; 410: 152–156.

LEITE LHR, LACERDA ACR, BALTHAZAR CH, MARUBAYASHI U, COIMBRA CC. Central angiotensin AT1 receptors are involved in metabolic adjustments in response to graded exercise in rats. *Peptides* 2009; 30: 1931–1935.

LEVINE JA, EBERHARDT NL, JENSEN MD. Role of nonexercise activity thermogenesis in resistance to fat gain in humans. *Science* 1999; 283 (5399): 212-214.

\_\_\_\_\_. Nonexercise activity thermogenesis (NEAT): environment and biology. *Am J Physiol Endocrinol Metab* 2004; 286: E675–E685.

\_\_\_\_\_, VANDER WEG MW, HILL JO, KLESGES RC. Non-exercise activity thermogenesis: the crouching tiger hidden dragon of societal weight gain. *Arterioscler Thromb Vasc Biol* 2006; 26 (4): 729-736.

LI H, SATINOFF E. Changes in circadian rhythms of body temperature and sleep in old rats. *Am J Physiol* 1995; 269(1 Pt 2): R208-14.

LIMA NR, COIMBRA CC, MARUBAYASHI U. Effect of intracerebroventricular injection of atropine on metabolic responses during exercise in untrained rats. *Physiol Behav* 1998; 64 (1): 69-74.

MOUNT LE, WILLMOTT JV. The relation between spontaneous activity, metabolic rate and the 24 hour cycle in mice at different environmental temperatures. *J Physiol* 1967; 190: 371-380.

NOVAK CM, LEVINE JA. Central neural and endocrine mechanisms of non-exercise activity thermogenesis and their potential impact on obesity. *J Neuroendocrinol* 2007; 19 (12): 923-940.

PRIMOLA-GOMES TN, PIRES W, RODRIGUES LO, COIMBRA CC, MARUBAYASHI U, LIMA NR. Activation of the central cholinergic pathway increases post-exercise tail heat loss in rats. *Neurosci Lett* 2007; 413 (1): 1-5.

\_\_\_\_\_, CAMPOS LA, LAUTON-SANTOS S, BALTHAZAR CH, GUATIMOSIM S, CAPETTINI LS, LEMOS VS, COIMBRA CC, SOARES DD, CARNEIRO-JÚNIOR MA, QUINTÃO-JÚNIOR JF, SOUZA MO, CRUZ JS, NATALI AJ. Exercise capacity is related to calcium transients in ventricular cardiomyocytes. *J Appl Physiol* 2009; 107: 593–598.

REFINETTI R, MA H, SATINOFF E. Body temperature rhythms, cold tolerance, and fever in young and old rats of both genders. *Exp Gerontol* 1990; 25(6): 533-43.

\_\_\_\_\_, Relationship between the daily rhythms of locomotor activity and body temperature in eight mammalian species. *Am J Physiol Regulatory Integrative Comp Physiol* 1999; 277 (46): R1493–R1500.

\_\_\_\_\_. The circadian rhythm of body temperature. *Frontiers in Bioscience* 2010; 15: 564-594.

REILLY T, WATERHOUSE J. Sports performance: is there evidence that the body clock plays a role? *Eur J Appl Physiol* 2009; 106: 321–332.

\_\_\_\_\_, ATKINSON G, WATERHOUSE J. Chronobiology and physical performance. In: GARRETT W JR, KIRKENDALL D (eds) *Exercise and sport science*. Philadelphia: Lippincott, 2000.

SATINOFF E, LI H, TCHENG TK, LIU C, MCARTHUR AJ, MEDANIC M, GILLETTE MU. Do the suprachiasmatic nuclei oscillate in old rats as they do in young ones? *Am J Physiol* 1993; 265(5 Pt 2): R1216-22.

\_\_\_\_\_, PROSSER RA. Suprachiasmatic nuclear lesions eliminate circadian rhythms of drinking and activity, but not of body temperature, in male rats. *J Biol Rhythms* 1988; 3(1): 1-22.

\_\_\_\_\_, LIRAN J, CLAPMAN R. Aberrations of circadian body temperature rhythms in rats with medial preoptic lesions. *Am J Physiol* 1982; 242(3): R352-7.

SCHEER FAJL, VAN DOORMEN LJP, BUIJS RM. Light and diurnal cycle affect human heart rate: possible role for the circadian pacemaker. *J Biol Rhythms* 1999; 14: 202-212.

\_\_\_\_\_, TER HORST GJ, VAN DER VILET J, BUIJS RM. Physiological and anatomic evidence for regulation of the heart by suprachiasmatic nucleus in rats. *Am J Physiol* 2001; 280: H1391-H1399.

\_\_\_\_\_, VAN DOORMEN LJP, BUIJS RM. Light and diurnal cycle affect autonomic cardiac balance in human: possible role for the biological clock. *Auton Neurosci* 2004; 110: 44-48.

\_\_\_\_\_, PIROVANO C, VAN SOMEREN EJW, BUIJS RM. Environmental light and suprachiasmatic nucleus interact in the regulation of body temperature. *Neuroscience* 2005; 132: 465–477.

TANAKA H, YANASE M, KANOSUE K, NAKAYAMA T. Circadian variation of thermoregulatory responses during exercise in rats. *Am J Physiol* 1990; 258 (Regulatory Integrative Comp. Physiol. 27): R836-R841.

WACHULEC M, LI H, TANAKA H, PELOSO E, SATINOFF E. Suprachiasmatic nuclei lesions do not eliminate homeostatic thermoregulatory responses in rats. *J Biol Rhythms* 1997; 12(3): 226-34.

WANNER SP, GUIMARAES JB, PIRES W, LA GUARDIA RB, HAIBARA AS, MARUBAYASHI U, COIMBRA CC, LIMA NR. Physical exercise-induced cardiovascular adjustments are modulated by muscarinic cholinergic receptors within the ventromedial hypothalamic nucleus. *Physiol Res* 2010; 59(2): 165-75.

YAMAOKA I, NAKAYAMA M, MIKI T, YOKOYAMA T, TAKEUCHI Y. Dietary protein modulates circadian changes in core body temperature and metabolic rate in rats. *J Physiol Sci* 2008; 58, n. 1: 75-81.