Contribution of Peripheral and Central Fatigue in Different Conditions (Gender and Time of Day Differences)

by
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The aim of the present study was to examine the rate of central and peripheral contributions in circadian rhythms and gender differences after exhausted maximal exercise. From 36 volunteers, 12 healthy male and female subjects who had H-reflex (It is elicited by electrical stimulation and it has been suggested to be one of mechanisms that could cause central fatigue) were selected (mean age of 23.5±1.37 years, height of 166.83±5.11 cm, weight of 59.8±9.15 kg, VO2max of 33.14±4.71 ml. kg⁻¹. min⁻¹ in male subjects and mean age of 23±6.32 years, height of 165.17±2.48 cm, weight of 56±2.09 kg and VO2max of 29.05±2.89 ml. kg⁻¹. min⁻¹ in female subjects) and took part in this investigation. Central (MVC, latency of H-reflex, amplitude of H-reflex and ratio of H-reflex to M-wave) and peripheral properties of fatigue (blood lactate and M-wave) were recorded before and after an exhausted maximal exercise at two different times of day: 9 am and 6 pm which is separated by a week. Significant differences were found in HR, blood lactate and latency of H-reflex before and after exercise. However differences between male and female in MVC, amplitude of H-reflex, ratio of H-reflex to M-wave and M-wave was significant. With induced –exercise, central and peripheral fatigue was higher in men in the morning and in women in the afternoon., thus our results recommended designing suitable exercise programs for men and women in the afternoon and morning, respectively. However, further studies are needed to confirm these results, especially in recommending using transcranial magnetic stimulation.

Key words: central and peripheral fatigue, circadian rhythms, gender differences

Introduction

Fatigue is a complex process that is defined as a reduction in force generating capacity, or the inability to maintain performance (Fulco, 1999). The development of this temporary loss of force is a complicated phenomenon which may develop due to dysfunction of any critical cites on the pathways from the central nervous to the peripheral systems (Gandevia, 2001), such as motor unit recruitment, firing rate and chemical transmission across neuromuscular junctions (Bigland-Ritchie, 1984), propagation of action potential along the muscle membrane and T tubules, Ca²⁺ release from sarcoplasmic reticulum, Ca²⁺ binding to troponin C and cross-bridge cycling (Mclester, 1997).

In analyzing the site of force origination, causes of fatigue can be divided into peripheral and central factors, whereby neuromuscular junctions and mus-
cle tissue are regarded as peripheral sites and the higher systems as the central sites (Gandevia, 2001).

Different studies have used EMG as a fatigue index. The strength of the excitation achieved by a same sized compound afferent action potential on its motorneuron pool can be measured by means of Hoffman-reflex (Kent-Braun, 1999). A decrease in the H-reflex can be attributed to decreased motorneuron excitability or increased pre-synaptic inhibition of Ia afferent neuron (Hugon, 1973). Decreased excitation from Ia afferent neuron due to decreased firing frequencies of muscle spindles has been suggested to be one of mechanisms that could cause central fatigue (Nourdland, 2003; Macfield, 1999).

Several processes are involved in converting an axonal action potential in to a sarcomia action potential. Collectively, these processes are referred as neuromuscular propagation. Sustained action can impair some of the processes involved in neuromuscular propagation, and this can contribute to the decline in force associated with fatigue. The most common way to test impairment of neuromuscular propagation is to elicit M-waves before, during and after a fatiguing contraction (Sandiford, 2004). M-wave is measured by applying an electric shock to a nerve to generate action potential in the axons of alpha motorneurons and measuring the EMG response in muscle (Enoka, 1994; Fuglevand, 1993). There are experimental evidences to show that the decrease in intracellular pH, related to the increase in H+ concentration associated with lactate production could well explain the force reduction, observed during fatigue (Stackhouse, 2001). Although fatigue has both central and peripheral components, but whose relative contribution to fatigue may be task dependent. Fatigue ability is various between muscle groups (Behm, 2002), muscle action types (Löscher, 2002) and individuals (Westing, 1999). In addition to variations in the extent and mechanism of fatigue with differences in task performance, some studies have suggested that fatigue may be influenced by gender (Hunter, 2001) and times of day (Castaingts, 2004). The majority of these studies have shown that women exhibit greater resistance to fatigue than men during maximal isometric contractions. Moreover, the results of other studies have suggested that the greater fatigue resistance of women declines as the intensity of the contraction increases (Maughan, 1986; Kent-Braun, 2002). Hunter & Enoka (2001) pointed to a disappearance in related fatigue differences between men and women.

In adult women, circulating estrogen and progesterone levels vary normally throughout the menstrual cycle that can cause variations in cardiovascular, respiratory and metabolic parameters (Constantini, 2005). Nevertheless different times of day may affect women gender steroids with subsequent implications for performance.

Circadian rhythms are internally generated phenomena with periodicity of 24 hours (Aschoff, 1995). Many factors could affect internal body environment, such as core body temperature and hormones result in the body experiences various conditions which can influence on exercise in different times of day (Guette, 2005a). Guette et al (2005b) showed that the quadriceps muscle torque changes in a predictable manner during a 24 h period—changes, which are linked to muscular rather than neural level. Castigaignts et al (2004) reported weaker neuromuscular function in the evening that this weaker performance of triceps sura muscle was related to a higher fatigue in activated motor units (Castigaignts, 2004). Nevertheless it is less known about the effect of times of day on fatigue.

Since no studies have been conducted to investigate fatigability differences between men and women, in different times of day and the importance of these parameters in a train programming. Therefore, our aim was to determine the rate of contribution of central (H-reflex, MVC) and peripheral (M-wave, lactate) fatigue, following exhaustive maximal exercise in various condition (gender and times of day). It was hypothesized that men and women responded differently in the morning and evening to the maximal exhausting exercise.

Methods

Subjects: Thirty-six healthy physical education students from Tehran universities, with at least a 2-year history of physical activity, were volunteered. Six men and six women of whom had H-reflex, were chosen as research subjects (mean age of 23.5±1.37 years, height of 166.83±5.11 cm, weight of 59.8±9.15 kg, VO2max of 33.14±4.71 ml. kg⁻¹ min⁻¹ in male subjects and mean age of 23±6.32 years, height of 165.17±2.48 cm, weight of 56±2.09 kg and VO2max of 29.05±2.89 ml. kg⁻¹. min⁻¹ in female subjects).

Procedure: A couple days before experimental session, subjects familiarized with proper nutrition guidelines, activity and proper exercise clothing during testing. In the experimental session, all subjects completed health questionnaire and general in-
formation form, and written informed consent was obtained. Each subject’s body mass, height and resting heart rate using belt was determined. Following 10 min rest, blood samples were drawn from a finger to measure resting blood lactate concentration using lactometer (CE483, Germany). Subjects then layed prone to obtain an H-reflex. MVC of plantar flexors were measured 3 times, with 30s rest between each trial. The subjects then began the fatiguing protocol (Storer & Davis, 1998) on an ergometer (Monark, Taiwan). The exercise protocol consisted of maximal training until exhaustion. After exhaustion, power and heart rate were registered. Immediately after fatiguing protocol, lactate and MVC were recorded, and H-reflex was performed. All subjects completed tests 2 times: 9 am and 6 pm, within a time interval of 4 to 7 days. Women were tested in the same menstrual cycle phase.

**H-reflex recording**: The H-reflex was elicited by electrical stimulation of the peripheral or tibial nerve in the political fossa of the soleus muscle (Simonsen, 1999). For this reason, subjects lay prone, with the cathode electrode placed on the tibial nerve and proximally, relative to anode electrode. The stimulus consisted of 1 ms, such that no further Ia afferent neuron could be stimulated (Burke, 1989). The optimum site to elicit the H-reflex was marked. The stimulus intensity increased gradually until H-reflex and M-wave were obtained.

**Fatiguing protocol**: After adjusting the seat of cycle ergometer with the heel on the pedal and leg straight (Adams & Beam 2008), the subjects performed an incremental exercise test with a 15 W/min increment until volitional fatigue, after a 4-min warm up with no load. The maximal watts attained during the exercise are used in gender-specific equations that also involve body weight and age for calculating VO2max (Equation 1, 2).

\[
\text{Men: VO2max (ml.min)} = (10.51(W,max)+[6.35(wt,kg)] - [10.49(age,yr)]+519.3) \quad \text{(Eq.1)}
\]

\[
\text{Women: VO2max (ml.min)} = (9.39(W, max)+[7.7(wt,kg)]-[5.88(age,yr)]+136.7) \quad \text{(Eq.2)}
\]

(Robergs & Keteyian 2003).

**Statistical analysis**: The K–S was used to test the normality of data. Descriptive statistics was used to present data as means ± SD. To test for changes in the dependent variables before and after fatigue protocol, repeated measure ANOVAs (gender as between group factor, time and fatigue as within group factors) were preformed for each variable. In the event of a significant main effect, Bonferroni test was used. All statistical significance was established at p≤0.05.

**Results**

Maximal exercise and gender had significant effect on heart rate (respectively F1, 10 = 697.7, p=0.000; F1, 10 = 14.8, p=0.003). Time of day had no significant effect on HR. HR response was greater in the morning in women and in the evening in men.

Lactate concentration increased significantly following maximal exercise (F1, 10 = 0.81, p=0.000) (Figure 1). The data was significantly different in genders after maximal exercise (F1, 10 = 10.85, p=0.008) (Figure 1).

We observed a significant effect of maximal exercise on delaying T (the time between stimulation and response of muscle). In addition, the effect of sex on H-reflex amplitude was detected. Men tended to exhibit a greater decline in H-reflex amplitude than women in the morning, while women showed greater decline in the evening (Figure 2).

No significant changes were found in M-wave after maximal exercise (F1, 10 = 1.62, p=0.023). Meanwhile, time of day and sex had significant effect on M-wave (respectively, F1, 10 =6.06, p=0.034 and F1.
Discussion

Results demonstrated that heart rate increases after maximal exercise in the morning and evening. HR response was significantly different between men and women after exercise. These results are in agreement with some studies (Ryon, 1994; Davis, 2000; O’Toole, 1989) but not with others (Melanson, 2002). It is clear that women and men show different physiologic responses to exercise. Of the different responses, sinus heart rhythm is a function of sex. Ryan et al. (1994) showed that high frequency power spectrum of heart rate and parasympathetic function is greater in women (Hargreaves, 1998). Moreover, differences in utilization of energy supplies (Darleen, 2002; Russ, 2003), energy metabolism pathways (Kent-Braun, 2002), vascular diffusion (Hatzikotoulas, 2004) and rate of recruitment of motor units (Melanson, 2002) can be contributed to these varieties.

Differences in metabolic responses between men and women show more reliance of men to non-oxidative ATP sources (Hicks, 2001). The greater oxidative capacity of women allows them to utilize the available oxygen in a more efficient manner than men; while a greater rate of glycolysis in men would be associated with greater rates of carbohydrate oxidation (Kent-Braun, 2002).

The present findings point to significant changes after maximal exercise, and sex factor differences showed a significant effect on lactate changes, with greater changes occurring in men during the morning and evening. These results were in agreement with Russ & Kent-Braun (2003), which demonstrated...
a correlation between lactate accumulation and fatigue, where \( \text{H}^+ \) is a by-product of glycolysis and thereby involved in fatigue (Russ, 2002). The build-up of \( \text{H}^+ \) within the muscle lowers the blood \( \text{pH} \) and may reduce muscle force by decreasing \( \text{Ca}^{2+} \) from the sarcoplasmic reticulum, thereby decreasing the sensitivity of troponin C to \( \text{Ca}^{2+} \) and interfering with cross-bridge cycling (McLester, 1997).

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No significant differences were observed in strength of plantar flexors after maximal exercise. Changes in strength are dependent on various factors, such as exercise protocol and specific muscle involvement (Garland, 2002). Since the soleus muscle is more oxidative than other muscles during plantar flexion (Gardiner, 2001), despite soleus muscle fatigue, there were no changes observed in MVC (ACSM, 2006). Sometimes fatigue occurs without any change in force (Gardiner, 2001). Moreover, plantar flexors are more resistant to fatigue than dorsiflexors (Gardiner, 2001).

In this experiment, it was shown that time of day had no effect on muscle strength following maximal exercise. Contrary, Castigaignts et al. (2004) observed less neuromuscular efficiency and Gutte et al. (2005a) reported a decline in torque force of the plantar flexors in the evening. Castigaignts et al. (2004) argued that most fatigable motor units could be affected by daily activity. Martin et al. (1989) advanced the hypothesis that Pi concentration and calcium movements in muscle fibers could explain the diurnal variation of strength. In contrast with the diurnal variation observed in concentric contractions, in which some researches relate these variations to the metabolic and ionic factors involved in circadian rhythm, it was found that time of day had no effect upon eccentric tasks (De Ruiter, 2001). However, more investigations are needed to find the effect of time of day on strength.

Gender had significant effect on decline in plantar flexor strength following sub-maximal training. These findings are in agreement with Fulco et al. (1999), Garland & Gossen (1991) and Hicks et al. (2001), who believe that sex differences, may affect fatigability. But in Hatzikotoulas et al. (2004) findings, men and women were matched for strength, and gender differences were not observed. Furthermore, differences between men and women can be attributed to non-matched subjects for strength. H-reflex amplitude showed decrease after maximal exercise but was not statistically significant. This decrease in the H-reflex amplitude can be attributed to decreased motoneuron excitability or increased presynaptic inhibition of Ia afferents (Chen, 2002).

The present study demonstrated no significant change in M-wave amplitude following maximal exercise, but the change was significant between men and women at different times of the day. Women had experienced more declines in the morning and men had shown more declines in the evening. Moreover, peripheral fatigue, which is attributed to neuromuscular fatigue and intracellular metabolism, was observed more in women rather than in men. These results were in agreement with Tanino et al. (2005). A decline in M-wave amplitude is interpreted as an impairment of one or more of the processes involved in converting the axonal action potential in to a muscle action potential (Enoka, 1994).

Hmax/Mmax ratio had no significant change following maximal exercise or between genders at different times of day. To our knowledge no research has investigated the relationships between men and women in the morning and evening after maximal exercise. Guette et al. (2005) observed no change in Hmax/Mmax ratio as an index of efficiency in transition of reflex between Ia afferents and motoneuron pool, at different times of day. Castigaignts et al. (2004) reported similar findings, suggesting a lack of diurnal fluctuation in the proportion of motor units reflexively activated. Thus, it is postulated that since the Hmax/Mmax ratio did not change in the present
study, it seems that the reflex pathway was not affected by daily activity at rest.

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Conclusion

Fatigue has been the subject of many investigations, but the important fact obtained from the results is that measured peripheral fatigue indices (M-wave amplitude and lactate) in women were greater than in men. Men had a greater decrease in the morning and women had bigger decline in the evening in peripheral fatigue factors. These changes were unexpected because reduced vascular occlusion occurred during muscle contraction as a result of lower absolute forces; thus more vascular diffusion and less ischemic condition is expected during contraction. Subsequently, women have more dependency on oxidative capacity. But from the reflective pathways of central fatigue indices (H-reflex, time latency of H-reflex, MVC, Hmax/Mmax ratio), after maximal exercise, the major results of this study revealed a difference in central fatigue between genders. The H-reflex was greater in the morning for men and greater in the evening for women.

With induced–exercise, central and peripheral fatigue was higher in men in the morning and in women during the evening. Based on our results, we recommend designing suitable training programs for men and women in the afternoon and morning, respectively. However, more research is needed to confirm these results, especially when recommending the use of transcranial magnetic stimulation.

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