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The influence of stimulus phase duration on discomfort and electrically induced torque of quadriceps femoris

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ABSTRACT | Background: Although a number of studies have compared the influence of different electrical pulse parameters on maximum electrically induced torque (MEIT) and discomfort, the role of phase duration has been poorly investigated. **Objective:** To examine the variation in muscle torque and discomfort produced when electrically stimulating quadriceps femoris using pulsed current with three different phase durations in order to establish whether there are any advantages or disadvantages in varying the phase duration over the range examined. **Method:** This is a two repeated-measures, within-subject study conducted in a research laboratory. The study was divided into 2 parts with 19 healthy young adults in each part. In part 1, MEIT was determined for each phase duration (400, 700, and 1000 μ s), using a biphasic pulsed current at a frequency of 50 Hz. In part 2, stimulus amplitude was increased until the contractions reached 40% of maximum voluntary isometric contraction (MVIC) and the associated discomfort produced by each phase duration was measured. **Results:** In part 1 of the study, we found that the average MEITs generated with each phase duration (400, 700, and 1000 μ s) were 55.0, 56.3, and 58.0% of MVIC respectively, but the differences were not statistically significant ($p=.45$). In part 2, we found a statistically significant increase in discomfort over the same range of phase durations. The results indicate that, for a given level of torque production, discomfort increases with increasing phase duration ($p=.008$). **Conclusions:** Greater muscle torque cannot be produced by increasing the stimulus phase duration over the range examined. Greater discomfort is produced by increasing the stimulus phase duration.

Keywords: electric stimulation; torque; quadriceps muscle; physical therapy.

Trial Registration: NCT01862393 (ClinicalTrials.org).

HOW TO CITE THIS ARTICLE

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• Introduction

Neuromuscular Electrical Stimulation (NMES) is a well-substantiated strategy to augment muscle performance in healthy and non-healthy subjects¹⁻⁹. It is also claimed to be useful for delaying or preventing muscle atrophy, increasing muscle thickness, fascicle length, and knee extensor torque in patients with knee osteoarthritis, hastening recovery from knee ligament surgery and from chondromalacia patellae, reducing spasticity and contractures, preventing deep vein thrombosis, and treating a range of clinical conditions¹⁰⁻¹⁹. The claims are extensive but the published evidence for some of these claims is either lacking or unconvincing²⁰⁻²². Nonetheless, if a condition is known to benefit from induced muscle contraction, it is reasonable to conclude that electrical stimulation intervention, at levels high enough to produce a forceful contraction, is likely to be beneficial²³.

When attempting to increase muscular strength with electrical stimulation, as with voluntary exercise, there is a strong correlation between the intensity of the muscle contraction and the strength improvement obtained²³⁻²⁵. It is also known that the main limiting factor to electrical stimulation for muscle strengthening is the discomfort caused by the stimulating current. Studies have shown successful muscle strengthening after NMES training regimens using 33 to 91% of maximal voluntary contraction^{20,25,26}. To electrically elicit muscular forces within the upper limits of this range requires high current intensities, with accompanying pain that may approach an intolerable level²⁵. As a consequence, this modality of muscle training is more efficient when subjects are both highly motivated and have learned by experience to tolerate progressively higher levels of electrical stimulation during training⁵.

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The issue of discomfort is not restricted to strengthening. Patient acceptance and tolerance is an important factor in determining whether electrical stimulation intervention in rehabilitation more generally will be successful. Thus it is important to know whether changing stimulus parameters such as the phase duration will make a difference to either the maximum electrically induced torque (MEIT) or the perceived level of discomfort at lower torque levels. Since achieving high electrically-induced torque is limited by pain, investigators have compared different stimulus waveforms and different types of stimulators in terms of both force production and relative discomfort based on the assumption that a more comfortable stimulus might allow higher stimulation intensities, consequently optimizing the results obtained with NMES^{24,25,27-30}.

Two variables that can normally be adjusted by the clinician using a commercially available pulsed current (PC) stimulator are the phase duration of the stimulus and the phase amplitude. At stimulation intensities above the motor threshold, an increase in either the stimulus amplitude or the phase duration will result in an increase in the muscle torque elicited. There will also be an accompanying increase in discomfort. This raises the question of whether it is preferable to increase the phase duration or increase the amplitude in order to increase the torque while minimizing the increase in discomfort. An increase in stimulus phase duration (at fixed amplitude) will certainly produce an increase in the force of muscle contraction. This is because activation of nerve fibers depends on the charge moved across the fiber membrane, and with a longer phase duration, more α -motoneurons will experience sufficient charge movement to produce an action potential. But pain will also be increased as greater charge movement means that more A- δ and C fibers will also be activated. An increase in stimulus amplitude (at a fixed phase duration) will also produce an increase in the force of muscle contraction and an associated increase in pain fiber activity due to the greater charge movement. What is not known is whether the effects of increasing the phase charge by increasing the phase duration or phase amplitude are different. The present study was designed to address this issue.

Although a number of studies have compared different kinds of stimulus waveform, there are only a few studies which have directly investigated the effect of phase duration on MEIT³¹⁻³³. Scott et al.³¹ compared monophasic squared-wave pulses with duration of 50 μ s and 200 μ s, showing that 200 μ s produces a greater phase charge and a greater MEIT. Similar results were presented by Gorgey

and Dudley³² when comparing two NMES protocols (100 Hz and 450 μ s versus 60 Hz and 250 μ s) on the evoked torque. These authors concluded that the longer pulse duration (450 μ s) resulted in a greater torque when compared to the shorter pulse duration (250 μ s)³². Gregory et al.³³ investigated the effects of several combinations of frequencies (10, 20, 30, 40, 50, 60, 70, and 100 Hz) and pulse durations (100, 200, 300, 400, 500, 600, and 700 μ s) on muscle torque production and fatigue. At constant current amplitude, torque increased as pulse duration was increased, and total charge was a predictor of torque production. Higher frequency trains caused greater fatigue. Nevertheless, none of these studies assessed the discomfort produced by different pulse durations, and no studies comparing longer pulse durations were found in the literature. Accordingly, we chose to compare both the MEIT of quadriceps femoris and the discomfort associated with PC of different phase durations using a commercially available stimulator. The purpose of this study was to establish whether, over the range of phase durations investigated, there are any advantages or disadvantages in varying the phase duration in terms of (i) muscle torque production and (ii) associated discomfort.

• Method

Subjects

Ethics approval for the study was given by the Institutional Review Board (IRB) of Universidade Cidade de São Paulo (UNICID), São Paulo, SP, Brazil (PP 13323922). A convenience sample of undergraduate physical therapy students was recruited from UNICID. All participants were aware of the experimental procedures and gave informed consent. For each of the two parts of the study, separate groups of twenty sedentary healthy university students were recruited as participants (19.5 ± 3.2 years). The participants were all female, reflecting the gender imbalance in the undergraduate physical therapy student population. Each participant was informed about the aim of the study and given details, including what would be required of them and each gave written informed consent before the experimental measures were made. All participants had previously used and experienced neuromuscular electrical stimulation during electrotherapy classes and were therefore familiar with electrical stimulation.

None of the participants in this study had a history of knee surgery or injury, musculoskeletal, cardiovascular or neurological disease including

impaired sensation. An exclusion criterion for part 1 of the study was if the participant did not reach MEIT with the stimulator used. Two participants were excluded from the analysis. One participant tolerated the maximum output of the stimulator and was therefore excluded. Another could not tolerate the discomfort of stimulation to 40% of MVIC and was therefore excluded. Thus, parts 1 and 2 each had 19 participants.

Apparatus

The stimulator used for the study was a mains-powered, commercially available, constant current unit called Dualpex 961 (Quark Produtos Médicos, Piracicaba, São Paulo, Brazil). The Dualpex was set to symmetric biphasic square pulsed current at a frequency of 50 Hz ramped up in amplitude for 2 seconds then constant for 4 seconds of 'on' time (part 1) or 6 seconds of 'on' time (part 2). The shorter 'on' time in part 1 of the study was because MEIT was measured, so it was important to minimize the duration of the discomfort, which was maximum tolerable. A longer 'on' time was used in part 2 to allow the subject to better assess the discomfort associated with stimulation at 40% of MVIC. Stimulus phase durations were set to 400, 700, and 1000 μ s. At the frequency used (50 Hz), these phase durations are not likely to pose a risk of skin irritation or damage²². The current was applied to the right quadriceps femoris muscle^{29,34} using two conductive (carbon-impregnated) rubber electrodes (size 10x4.5 cm) and electrode gel (Sonic-Plus Gel, Hal Indústria e Comércio Ltda, São Paulo, Brazil). The proximal electrode was placed over the femoral triangle of the quadriceps and the distal electrode was over the motor point of the vastus medialis²⁸. Electrodes were secured in place with adhesive tape. A 3-minute interval between each repetition was standardized among the MEITs performed to minimize the effects of muscle fatigue^{8,35}.

An isokinetic dynamometer (Cybex Norm 6000, Cybex Division of Lumex, Inc., Ronkonkoma, NY, USA) was used for torque measurement. It was set to isometric mode at an angle of 60° degrees of knee flexion^{18,29,36,37}.

Procedure

Each participant undertook a 5-minute warm-up on an exercise bicycle before testing³⁶, cycling at approximately 75 revolutions per minute with no extra resistance. All participants then performed three maximum voluntary isometric contractions (MVIC) of the right quadriceps femoris muscle^{29,34,38}.

Each contraction was sustained for 4 seconds (part 1) and 6 seconds (part 2) and there was a 3 minute rest period between each contraction. The highest of the three torque measurements recorded by the Cybex dynamometer was considered to be MVIC. Subsequently this value was used to calculate maximum electrically induced torque (MEIT) as a percentage of MVIC.

In part 1 of the study, three minutes after the determination of MVIC, maximum electrically induced torque was determined for each phase duration (400, 700, and 1000 μ s). Participants were instructed to relax and neither oppose or assist the contraction. First the amplitude was increased gradually (using continuous output) until the participant indicated that the stimulus intensity was the maximum they were willing or able to tolerate. After a three-minute rest period, the same stimulus amplitude was applied using a two-second ramp-up and four seconds of 'on' time. During the 'on' time, electrically induced torque was recorded. If the participant indicated that they could tolerate a higher stimulus intensity, the amplitude was increased. Once the maximum amplitude was established, three measures were made for each phase duration and the highest value was taken as the MEIT. The order of presentation of the phase durations was randomized so as to avoid order-induced bias. Each electrically elicited contraction was sustained for four seconds, after which there was a rest period of three minutes.

In part 2 of the study, electrical stimulation using phase durations of 400, 700 or 1000 μ s was used. The order of presentation of the three phase durations was randomized to avoid order-induced bias. To avoid muscle fatigue, stimulations were again spaced three minutes apart. First, MEIT was measured (as in part 1). Next the stimulus amplitude was increased until the contractions produced were 40% of MVIC. The peak amplitude (in mA) was recorded and each volunteer rated the associated discomfort using a 10 cm visual analogue scale (VAS) ranging from zero (no discomfort) to 10 (worst discomfort imaginable).

Data analysis

Data were analyzed using the SPSS version 14 software package (SPSS Inc., Chicago, IL, USA) and retrospective power analysis as described by Portney and Watkins³⁹ using Cohen's⁴⁰ sample size tables. Preliminary analysis found no extreme outliers and Q-Q plots identified that the data were normally distributed. Accordingly, parametric tests were conducted. In part 1, to better isolate and identify any variation in MEIT with phase duration,

a one-way, repeated-measures analysis of variance (ANOVA) was conducted. In order to separately examine the effect of MVIC on MEIT, data were further analyzed by regression analysis using raw MEIT values (uncorrected for MVIC) with stimulus phase duration and MVIC as the covariates.

In part 2, linear regression analysis was performed as the three conditions compared (three phase durations) are values chosen from a continuum. Stimulus phase charge was calculated by multiplying the stimulus amplitude (in mA) by the phase duration (in μs) to give the phase charge in microcoulombs (μC). A linear regression analysis of discomfort on phase charge was then performed. To establish whether phase charge or phase duration was the important factor or whether both made some independent contribution, the relationship between phase charge and phase duration was examined, using separate linear regression analyses. Separate analyses were needed as multiple regression cannot be performed unless the factors are independent, and in this case, phase duration and phase charge are not independent.

Results

Part 1

Figure 1 shows the variation in MEIT, expressed as a percentage of MVIC, over the three phase durations examined. Again the smallness of the variation is evident. An interesting feature is that although an increase in mean MEIT with increasing phase duration is evident, the variation is small (53 to 58% of MVIC).

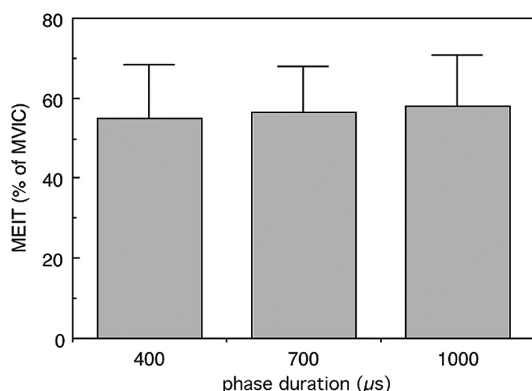


Figure 1. Average MEIT of quadriceps femoris muscle with stimulus phase durations of 400, 700, and 1000 μs . Torque is expressed as a percentage of maximum voluntary isometric contraction (MVIC). Values shown are means. Error bars are standard deviations.

A repeated-measures ANOVA showed that the between subject variance in MEIT was, as might be expected, significant ($F(18,2)=7.07$, $F_c=1.90$, $p<.001$), but the variation with phase duration was not statistically significant ($F(2,18)=0.82$, $F_c=3.26$, $p=.45$). As a consequence, no post-hoc comparisons of phase duration effects were justified.

Compared to the 400 μs pulse duration, the MEIT was 2.4% higher at 700 μs and 5.5% higher at 1000 μs . While this suggests an increasing trend, the variation is too small to demonstrate statistical significance with the subject number used. A power analysis was used to estimate the number of subjects that would be needed to demonstrate statistical significance with the effect size found ($d=0.06$)^{39,40}. Cohen's⁴⁰ tables indicate that to achieve a p value of .05 or less, over 600 participants would have been needed. This indicates that, over the range examined, the variation in MEIT with phase duration is not only statistically insignificant but also clinically insignificant.

Part 2

Figure 2 shows the variation in discomfort (mean VAS and standard deviation), over the three phase durations examined. Linear regression analysis showed that there is a clear and statistically significant increase in group-averaged discomfort rating (VAS) with phase duration (Pearson's $r=0.34$, $p=.008$). The relatively low value of r indicates that phase duration can account for only a fraction of the variation actually observed. The large error bars reflect a large between-subject variation whilst the very low value of p indicates that, despite this, the increase in discomfort with phase duration is significant.

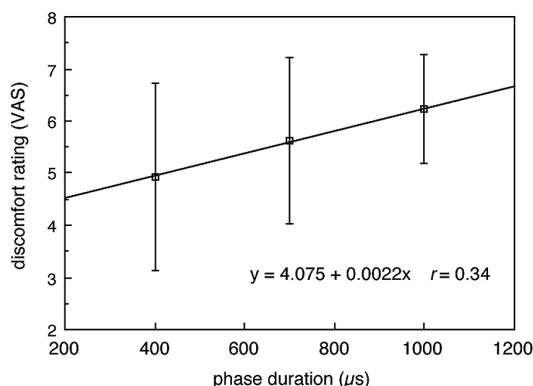


Figure 2. The variation in discomfort (VAS) with phase duration at stimulation intensities sufficient to produce a torque equal to 40% of MVIC. Values shown are means ($n=19$). Error bars are standard deviations.

Figure 3 shows the variation in discomfort (mean VAS and standard deviation), over the three phase durations examined, this time expressed as group-average phase charge. Regression analysis again demonstrates significance i.e. that there is a clear and statistically significant increase in discomfort rating (VAS) with phase charge (Pearson's $r=0.28$, $p=.032$) but the correlation is not as high as discomfort rating with phase duration (Pearson's $r=0.34$, $p=.008$).

The significant increase in group-averaged discomfort rating (VAS) with both phase duration (Figure 2) and phase charge (Figure 3) raises the question whether phase duration is the important factor, or phase charge, or whether both contribute independently in determining the discomfort of stimulation. This question could only be answered by conventional statistical analysis (either a 2-way ANOVA or regression analysis) if phase charge and phase duration were independent parameters, which they are not. Phase charge is equal to phase duration \times amplitude (mA). The high correlation between phase charge and phase duration was confirmed by the positive results of a regression analysis (Pearson's $r=0.72$, $p=.000$), meaning that conventional factor analysis would be unable to separate their contributions to the VAS measures.

Data for each phase duration (400, 700, and 1000 μ s) were therefore examined separately. Scatter graphs of discomfort (VAS) versus phase charge were plotted for each phase duration. These graphs showed no correlation between discomfort and phase charge. By way of illustration, Figure 4 shows a plot of discomfort (VAS) versus phase charge for the 700 μ s phase duration stimulus. The apparent lack of correlation is evidenced by the calculated correlation coefficient (Pearson's $r=0.032$, $p=.895$).

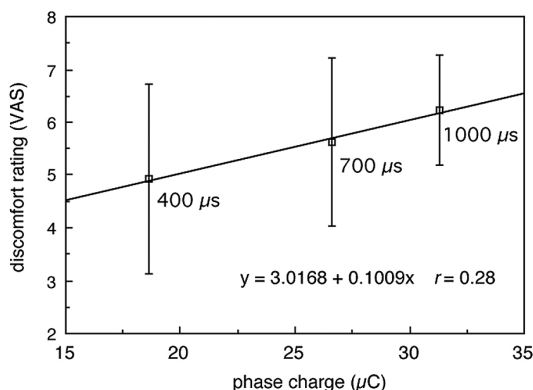


Figure 3. The group averaged variation in discomfort (VAS) with phase charge at stimulation intensities sufficient to produce a torque equal to 40% of MVIC. Values shown are means ($n=19$). Error bars are standard deviations.

Similar results were obtained at phase durations of 400 μ s ($r=0.264$, $p=.260$) and 1000 μ s ($r=-0.110$, $p=.644$). The r values seem haphazard, with an average of 0.061 and associated p values which do not even approach significance. It thus seems that there is no significant correlation between phase charge and discomfort if the phase duration is constant. If the phase duration varies, there is a significant variation in discomfort (Figure 2). Hence it is reasonable to conclude that phase duration is the decisive factor and that the variation in discomfort with phase charge (Figure 3) is only a secondary consequence.

• Discussion

MEIT and stimulus phase duration

In part 1 of the study, data analysis showed no significant variation with stimulus phase duration. This contradicts the idea that greater MEIT can be achieved by increasing the phase duration of the stimulus when using longer pulse durations. It appears that increasing pulse duration will only increase the MEIT in a shorter (50 μ s) to medium (450 μ s) pulse duration range^{31,32}, or when the current amplitude is kept constant during the increase in pulse duration³³. Over the range of phase durations examined in the present study, the increase in MEIT was both statistically and (based upon the small effect size) clinically insignificant.

Discomfort and stimulus phase duration

The relative discomfort of different forms of electrical stimulation is clearly a clinically relevant issue. In part 2 of this study, discomfort was

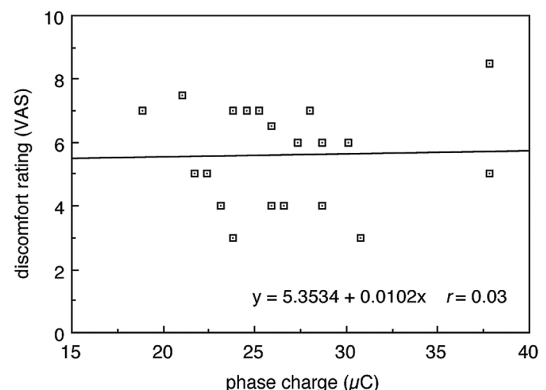


Figure 4. The variation in discomfort (VAS) with phase charge at stimulation intensities sufficient to produce a torque equal to 40% of MVIC. Values shown were obtained using a stimulus phase duration of 700 μ s.

measured at a fixed level of electrically induced torque (40% of MVIC). We found that discomfort, subjectively assessed and reported using a VAS, increases significantly with phase duration over the range examined ($p=.008$). Strength-duration curves for sensory, motor, and pain tolerance thresholds converge at longer phase durations, i.e. there is less separation between pain and motor thresholds, so the increase in discomfort with phase duration is perhaps not unexpected⁴¹⁻⁴³.

MEIT values in the present study

A number of studies have shown that electrical stimulation is useful for improving muscle performance but the findings indicate that electrical stimulation will only be effective if sufficient torque (expressed as % of MVIC) is induced by the electrical stimulation²². Several authors reported positive results for strengthening with electrically induced torques ranging from 33% to 91% of MVIC^{5,10,24,26,30,44,45}. The values obtained in part 1 of this study (mean 56% of MVIC, range 30-86%) are well within the range of these positive findings. The discomfort measurements (VAS) in part 2 of the present study were taken at electrically induced torques equal to 40% of MVIC: again within the effective range.

A perhaps surprising finding of the present study is that little increase in MEIT can be achieved by increasing the phase duration of the stimulus. Certainly if the stimulus amplitude (current or voltage) is kept constant, an increase in phase duration will result in a proportional increase in the charge per pulse and hence greater recruitment of motor nerve fibers and a more forceful contraction^{21,22,33,43}. However, there will also be greater recruitment of pain fibers, so in practice, because MEIT is dictated by discomfort associated with the stimulation, the maximum tolerable stimulation amplitude may decrease significantly with increasing phase duration. It appears that, over the range of phase durations examined in the present study, the increase in discomfort with phase duration almost completely negates the potential increase in MEIT.

Implications

Our finding that discomfort increases significantly with phase duration but that the corresponding increase in MEIT is not significant raises the question whether there is any advantage in having a choice of phase durations on clinical stimulators to be used for NMES. On the basis of our findings, the answer to this question would be 'no'. There is, however, an important practical issue with commercially available stimulators and that is the maximum output available.

Stimulators have an upper limit to the current they can deliver. At shorter phase durations, a greater stimulus amplitude (current) is needed in order to provide the necessary phase charge (which is the amplitude multiplied by the duration). So if the phase duration is too short, the maximum output of the machine (in mA) may not be sufficient to produce a maximal contraction. As noted above, one participant had to be excluded from part 1 of the present study as they had not reached MEIT when maximum output of the stimulator was reached using the shortest (400 μ s) phase duration. Having the option of increasing the phase duration means that the clinician can always elicit a maximal contraction though the trade-off is increased discomfort with a longer phase duration.

Conclusions

The results suggest a trend of increasing MEIT with stimulus phase duration but the measured variation was not statistically significant over the range of phase durations examined. Power analysis demonstrated that any variation which might exist is clinically insignificant.

For a fixed level of muscle torque, discomfort increases with increasing phase duration over the range examined. For this reason we advocate use of the shortest phase duration (400 μ s) to minimize discomfort without compromising torque production.

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