ABSTRACT

Loss of sensory motor function is one of the main causes of physical and activity limitations among individual with spinal cord injury (SCI). SCI may lead to muscle paralysis, weakness and disused muscle atrophy. Evidences have shown electrical stimulation and strengthening exercise might improve lower limb muscle strength and size among individual with SCI. Functional electrical stimulation (FES) evoked cycling is one of the methods that can elicit leg muscle contractions in order to produce a cycling motion and promote the integrity of the involved muscles. Therefore, this review is to synthesize the scientific literature regarding the effects of multiple dosages of FES-evoked lower limb cycling on muscle properties. A systematic literature search from 1946 to 2016 was performed. From over 1,139 articles retrieved from the database, about 31 potentially relevant articles were retained for possible inclusion. However, only 10 articles out of 31 articles fulfilled the inclusion criteria. Although the available evidence is compelling, there is insufficient quantity and quality evidence to draw conclusions regarding the specific parameter of FES-CE that may optimally increase muscle strength, mass, and circumference. However, it can be safely concluded that an effective training session would spend for 45-60 min, 3 times a week for at least 4 weeks to see changes in muscle size and strength.

Keywords: Cycling exercise; functional electrical stimulation; muscle strength; spinal cord injury

INTRODUCTION

Spinal cord injury (SCI) is one of the leading causes of physical disability worldwide. In the USA, approximately 17,000 new cases of SCI were reported each year due to road traffic accident, falls, and sports (National Spinal Cord Injury Database 2015). It can be pronounced that the consequences after SCI is usually a profound reduction in the muscle mass, strength, and muscle volume (Dionyssiotis et al. 2015). Thus, as a result of this muscle integrity degeneration, it could lead to lower limb muscle weakness and muscle atrophy. These can affect their quality of life in which long-term recovery and rehabilitation process is required (LiVecchi 2011; Westcott & Rosa 2010).

It has been widely established that exercise and training programs for individual with SCI could enhance their muscle power function and prevent muscle atrophy (American College of Sports Medicine 2010). Thus, a suitable exercise regime can help in achieving these goals and reduce the risk of secondary health problems accompanying with physical inactivity. One of the technique is by applying the functional electrical stimulation (FES). This device will assist exercise through the application of a low-level
electrical stimulation current on muscle motor control to evoke muscle contraction (Lynch et al. 2008; Peckham & Knutson 2005). There have been so many method with FES, however, FES evoked cycling exercise (FES-CE) has been widely used in clinical settings for aerobic and strength training applications (Estigoni et al. 2011; Fornusek et al. 2013; Kuhn et al. 2014; Thrasher et al. 2013). Technically, FES-CE is a computer-controlled electrical stimulation that elicits leg muscle contractions in order to produce a cycling motion and promote the integrity of the involved muscles (Ragnarsson 2008). Based on a study by Hamzaid and Davis (2009), the FES-CE could increase muscle size and volume, and prevent the alteration of muscle morphology in individuals with lower motor neuron lesions SCI (Hamzaid & Davis 2009). Therefore, the ultimate goal of this intervention is to provide therapeutic gains for persons with SCI (Estigoni et al. 2011; Fornusek & Davis 2004; Hasan et al. 2013).

Despite numerous studies of FES-CE effect on muscle size and muscle strength among individual with SCI, no evidence concerning about its appropriate dosage of training. In this narrative review, inclusive review of obtainable literature was provided to verify the effects of FES-CE on muscle size and muscle strengths among individual with SCI. By providing the quality of evidence and dosage for this kind of training, it may help the medical practitioners to decide on its proper application. Thus, the aim of this review was to synthesize the scientific literature to explore the multiple dosages of FES-CE that is adequate to improve muscle properties and characteristics including muscle mass, volume, and strength for individuals with SCI.

**METHODS: STUDY SELECTION**

A systematic literature search was conducted by using electronic databases: ProQuest, Science Direct, MEDLINE between 1946 and October 2016; Ovid MEDLINE Daily Update; and Web of Science between 1980 and 2016. For a wider coverage, the literature searches also included manual searches of clinical rehabilitation specific journals including Clinical Rehabilitation, Spinal Cord, Physiotherapy, Physical therapy and multidisciplinary journals for examples, BMI, Lancet, and JAMA. The search strategy for reviews on our topic was also carried out using the Cochrane Library and Cochrane Database for Systematic Reviews. The literature search was, however, limited to human research studies, based on any language if the translation of English abstract provides a clear description of the study and the outcome measures. All systematic literature findings were explored by the author based on certain inclusion criteria until it reaches ‘saturation point.’

The inclusion criteria for this review was that the studies must involve the following: functional electrical stimulation (FES); or functional neuromuscular stimulation (FNS) evoked cycling; lower limbs cycling; outcome should be measured on the effects of the cycling exercise on improvement or otherwise of muscle strength, mass, and volume; and in persons with spinal cord injury. In addition, studies that involved a combination of FES-evoked upper limb and lower limb exercise were included, if they provide lower limbs muscle strength as one of the outcome measures. Keywords used in this review were: FES, FNS, functional electrical stimulation, functional neuromuscular stimulation, electrical stimulation, functional stimulation, neuromuscular stimulation, neurostimulation; training, exercise, cycling; SCI, spinal cord survivors, paraplegic, paraplegia, quadriplegic, tetraplegia, tetraplegic, paralyzed, paralysis; subjects, participants, patients, human, persons; muscle strength, muscle atrophy, muscle mass, muscle volume, muscle circumference and muscle properties. The title of all the potentially eligible studies were scanned manually using Google Scholar search engine. On the first round of search, 1,139 experimental studies were discovered. Thus, the second round of search included randomized controlled trial (RCT), quasi-experimental research (randomized only or controlled only), case-controlled study, observation studies and crossover designs. As highlighted earlier only studies that recruited persons with SCI for the experiment and control group were retained. Studies that recruited able-bodied participants as the control were excluded. This was due to the knowledge that the inference or conclusion from such studies may not be sufficient to verify the proposed hypothesis (Hamzaid & Davis 2009). In the present review, the RCT and quasi-RCT retained were meant to provide a strong evidence whereas the other studies were used as supporting evidence for the hypothesis of the study.

**RESULTS**

The articles from RCT, quasi-RCT, nonrandomized or controlled studies and cross-sectional or crossover designs were retained for this review as they provide the scientific evidence regarding the FES-CE. The article search was conducted from February 2016 until October 2016. From over 1,139 articles retrieved from the database, about 31 potentially relevant articles were retained for possible inclusion. However, only 10 articles out of 31 articles fulfilled the inclusion criteria. All the 10 studies reported muscle size and strength benefits effect following FES-CE in individuals with SCI (Table 1). All studies had samples sizes ranging from 8-45 participants.

Muscle size has generally been measured as body lean mass by using bioelectrical impedance analysis and Lunar DPX x-ray to determine lesser fat or bigger muscle after FES-CE training (Baldi et al. 1998; Liu et al. 2007). Muscle circumference usually measured in the supine position using a tape measure (Fornusek et al. 2013; Kuhn et al. 2014; Liu et al. 2007). The measurements were usually reported before and after the FES-CE training based on the anatomical landmarks chosen by researchers. Significant improvement of muscle peak torque has been reported in individuals with SCI after FES-CE training.
<table>
<thead>
<tr>
<th>References</th>
<th>Sample</th>
<th>Duration</th>
<th>Intervention</th>
<th>Main result</th>
</tr>
</thead>
<tbody>
<tr>
<td>Johnston et al. (2015)</td>
<td>N= 17 Complete SCI</td>
<td>6 months</td>
<td>FES cycling 1-hour 3 times per week Low cadence vs. high cadence</td>
<td>Muscle volume Low cadence group increased by 19%, high cadence group increase by 10%</td>
</tr>
<tr>
<td>Sköld et al. (2002)</td>
<td>N= 15 Complete SCI</td>
<td>6 months</td>
<td>FES cycling 30 min × 3 times per week Freq= 30 Hz PW= 350 μsec Stim= up to 130 mA Pedal cadence= 35 rpm</td>
<td>Muscle volume Group trained with FES increased 1300 cm$^3$ ($p &lt; 0.001$) than those in the control group</td>
</tr>
<tr>
<td>Kuhn et al. (2014)</td>
<td>N=30 Complete SCI vs. Incomplete SCI</td>
<td>4 weeks</td>
<td>FES-CE 20 min × 3 times per week Freq= 30 Hz PW= 250 μsec Stim= 10-130 mA Pedal cadence= 15-55 rpm</td>
<td>Circumferences Greater improvement in incomplete group ($p&lt;0.05$) Manual muscle testing Those in incomplete group reported 43.5% increase ($p&lt;0.001$) in quadriceps femoris muscle and 25% ($p&lt;0.001$) in gluteus maximus muscle</td>
</tr>
<tr>
<td>Thrasher et al. (2013)</td>
<td>N= 11 Complete SCI vs. incomplete SCI</td>
<td>13 weeks</td>
<td>FES-LCE training 1 h × 3 times per week, 40 sessions Freq= 30 Hz PW= 250 μsec Stim= up to 140 mA Pedal cadence= 45 rpm</td>
<td>Peak torque Both groups had improved knee extension torque 3.8 to 16.9 Nm ($p = 0.006$). However, those in incomplete group shown greater improvement</td>
</tr>
<tr>
<td>Formussek et al. (2013)</td>
<td>N=8 Complete and Incomplete SCI</td>
<td>6 weeks</td>
<td>FES-CE 30 min × 3 times per week 1 leg on low cadence 10 rpm and the other leg on high cadence 50 rpm Freq= 35 Hz PW= 250 μsec Stim= 40-140 mA</td>
<td>Thigh girth Both groups reported greater improvement on low cadence Quadriceps peak torque Quadriceps peak torque has increased 87% in low cadence</td>
</tr>
<tr>
<td>Sadowsky et al. (2013)</td>
<td>N= 45 Complete and Incomplete SCI</td>
<td>29.1 month</td>
<td>FES-CE 50-60 min × 3 times per week Freq= 100 Hz PW= 500 μsec Stim= up to 140 mA Pedal cadence= 50 rpm</td>
<td>Muscle volume Those in FES group had significantly higher anterior (36%; 96.3 cc; $p \leq 0.001$) and posterior (30%; 63.9 cc; $p = 0.005$) thigh compartment muscles than those in the control group Muscle strength trained muscles of the FES group was significantly greater (quadriceps, $p = 0.006$; hamstrings, $p = 0.011$) than in controls</td>
</tr>
<tr>
<td>Duffell et al. (2008)</td>
<td>N= 11 Complete SCI</td>
<td>1 year</td>
<td>FES cycling 1 h × 3 times/week Freq= 50 Hz if rapid fatigue 10 Pedal cadence= 45-55 rpm</td>
<td>Quadriceps Maximal Torque Increased progressively and significantly throughout training ($p = 0.012$). It had increased by 399% and 673% after 3 and 12 months</td>
</tr>
<tr>
<td>Liu et al. (2007)</td>
<td>N=18 Complete and Incomplete SCI</td>
<td>8 weeks</td>
<td>FES-CE 30 min × 3 times per week Freq= 30 Hz PW= 300 μsec Stim= 10-130 mA Pedal cadence= 45 rpm</td>
<td>Body lean mass Mild increase after 8 weeks training ($p=0.03$) Limb girth Significant increase in both upper and lower limb girth after 4 and 8 weeks of training ($p&lt;0.05$) Peak torque Shown improvement only after 8 weeks training ($p&lt;0.05$)</td>
</tr>
<tr>
<td>Demchak et al. (2005)</td>
<td>N= 10 Complete SCI</td>
<td>13 weeks</td>
<td>FES-CE 30 min 3x per week Freq= 35 Hz Pedal cadence= 35 rpm</td>
<td>Cross-sectional area (muscle biopsy) Group trained with FES-CE increased CSAf 171% greater than the CSAf in the control group ($p &gt; 0.05$)</td>
</tr>
<tr>
<td>Baldi et al. (1998)</td>
<td>N=26 Complete SCI</td>
<td>3 month + 6 month</td>
<td>FES-CE 30 min × 3 times per week, FES-IC 1 h × 5 times per week Freq= 60 Hz PW= 375 μsec Stim= up to 140 mA Pedal cadence= 35-50 rpm</td>
<td>Lean body mass FES-CE showed a significant hypertrophy after 6 months on both lower limb and gluteal lean body mass in comparison to FES alone</td>
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</table>

FES-CE = Functional Electrical Stimulation-Cycling Ergometer; FES-LCE= Functional electrical stimulation Leg Cycling Ergometer; FES-IC= Functional Electrical Stimulation Isometric Contraction, N= Number of participants, Freq= Frequency, PW= Pulse Width, Stim= Stimulation
Based on the literature reviewed, we can safely infer that FES-CE has the potential to prevent the loss of muscle mass (Fornusek et al. 2013; Kuhn et al. 2014). Therefore, available evidence showed that the strength gains during FES-CE training were positively related to the spared muscle function after SCI trauma. This may suggest that SCI individuals with higher residual muscle strength might achieve better results from the cycling training.

**DISCUSSION**

This review yielded only 10 articles that emphasize the effect of FES-evoked cycling exercise on muscle size and muscle strength among individual with SCI. It has been proven that this type of exercise is highly relevant for individuals with chronic SCI as it has a significant impact on their capability to perform activities of daily living. Additionally, it has been identified that the FES-CE does not only improve muscle size, it also promotes the quality of life of incomplete SCI individuals in particular.

**EFFECTS OF FES-CE ON MUSCLE SIZE**

Individuals who were trained with FES-CE had shown significant improvement in muscle mass and muscle girth after 4 weeks (Kuhn et al. 2014). However, it has been identified that the training should start within the first 3 months after a SCI to ensure the efficacy of FES-CE training (Baldi et al. 1998). Baldi et al. (1998) have shown that FES-CE could prevent the progression of muscle mass reduction after 6 months of training. Paralysis and episode of sedentary lifestyle after chronic SCI could lead to elevation of fat mass and reduction of lean body mass percentage (Giangregorio et al. 2005). Notably, individuals with acute SCI only have 60-65% lean mass than their able-bodied counterparts and their fat gain can be up to 100-113% of their able-bodied counterparts (Giangregorio et al. 2005). Therefore, this may lead to a serious long-term health complication that may compromise with their quality of life (Hicks et al. 2011).

Nevertheless, the significant changes in leg lean mass following FES-CE training for incomplete SCI has been reported to be greater than in complete SCI individuals even in a short-term (8 weeks) cycling training (Liu et al. 2007). Hypertrophy of stimulated muscle has also been observed after both short and long-duration FES-CE protocol (Fornusek et al. 2013; Kuhn et al. 2014). Additionally, although a non-significant general increase of body lean mass in completely paralyzed individuals has been reported, such increase has a positive effect in preventing muscle atrophy of stimulated muscles (Baldi et al. 1998). Based on the literature reviewed, we can safely infer that FES cycling with 3 days per week training for more than 4 weeks could prevent the loss of muscle mass (Fornusek et al. 2013; Kuhn et al. 2014).

**EFFECTS OF FES-CE ON MUSCLE STRENGTH**

Evidence suggested that in order to achieve greater dramatic effects of muscle strength, the stimulation parameters used in the FES-CE training must be able to increase fatigue-resistance of muscles (Fornusek et al. 2013; Johnston et al. 2015). Low-cadence (10 rpm) but not high-cadence (i.e., 50 rpm) cycling has been shown to promote increased fatigue-resistance of the quadriceps muscle (Fornusek et al. 2013). This result was supported by earlier recommendation that the health effects from FES-CE should be maximized for SCI survivors (Duffell et al. 2008; Liu et al. 2007). As evident from the reviewed literature, most FES-CE cycle ergometers are designed to operate at a cadence of 50 rpm. However, fatigue develops more slowly during FES-CE at lower pedal cadences, allowing greater muscle forces to be maintained (Johnston et al. 2015). Therefore, continuous stimulation in muscle’s normal physiological frequency range (30-35 Hz) (Demchak et al. 2005; Duffell et al. 2008; Sadowsky et al. 2013) and pulse width (250-300 µsec) (Baldi et al. 1998; Johnston et al. 2015; Liu et al. 2007) may offer a better electrical stimulus effect that could promote adaptation of a muscle to be more fatigue resistant. In a total of 10 studies, not all studies stated the resistance used while cycling with FES. However, it seemed that by increasing the resistance every ~3W~6W if participants able to maintain rotation per minutes of the cadence by three consecutive 30 min training sessions it could give more impact to the strength of the muscle (Demchak et al. 2005; Liu et al. 2007; Thrasher et al. 2013).

**CONCLUSION**

Although the available evidence is compelling, there is insufficient quantity and quality evidence to draw conclusions regarding the specific parameter of FES-CE that may optimally increase muscle strength, mass, and circumference. Nevertheless, based on Table 1, it can be safely concluded that an effective training session would spend between 45 and 60 min, 3 times a week for at least 4 weeks to see changes in muscle size and strength. The FES stimulation parameters of 250-300 microseconds pulse width, current amplitude 40mA-140mA, 30-35Hz frequency are optimally administered for each participant. Studies on leg FES-cycling have only been carried out in persons with complete SCI or those with a combination of complete and incomplete SCI. However, a lesser attention has been paid to the effects of FES-cycling on persons with incomplete SCI. Although FES-cycling in persons with incomplete SCI may give a greater improvement in muscle strength, the affected population may not be able to tolerate electrical stimulation of a similar intensity as administered to complete SCI survivors. Thus, further quality research is needed to examine the most suitable parameters that can be used to maximize its effectiveness. For clinical rehabilitation applications, a standardized FES-evoked cycling protocol will be guiding especially for clinicians...
and other allied health professional administering FES-cycling for strength training.

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