Is the use of computerized electrical stimulation associated with cycloergometrics in individuals with medullary lesion beneficial for the muscular parameters?

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ABSTRACT
The aim of this study was to determine whether the use of FES produces muscular benefits for individuals with spinal cord injuries. Method: A review of the literature was done in the electronic database MEDLINE, PubMed, LILACS and SciELO, without limitation of time or language. The PICO strategy has been used for this research. Results: 554 articles were found. From these, 432 were excluded by title, resulting in 122 articles left. Out of these articles the duplicates were excluded resulting in 73 articles; 36 were then excluded after reading the abstract and 33 more after reading the full text. Four studies were selected. Two articles included males and females in their studies and two only included males. Three studies included quadriplegics and paraplegics in the same study; one included only quadriplegics. One of the studies used a more frequent training routine, seven times a week; three trained only three times a week. The duration of the studies was varied considerably, from six weeks to one year. The resulting measurements for strength and resistance evaluation were performed in different manners, by muscle transverse section area measurement, limb circumference, and muscle biopsy; however, all studies presented at least one of the measurements provided by the equipment, power output or work output. Improvement of the power output and work output has been shown in all studies. Despite the heterogeneity encountered in these studies, the outcomes evaluated by them indicate a significant increase in the power output and work output after training periods, with gains starting from six weeks of training at least three times a week. Conclusion: Future studies are needed to assess different responses in different groups of subjects, paraplegics or quadriplegics, under different frequencies and periods of training, and thereby provide the elaboration of more directed training protocols.

Keywords: electric stimulation, exercise, paraplegia, quadriplegia, spinal cord injuries
INTRODUCTION

Spinal cord injuries affect approximately 32 people per million and about 7,800 people a year in the United States. Worldwide, the variation is from 12.1 to 57.8 cases per million people. It occurs predominantly with males, in about 82% of the cases and with the average age of 33.4 years.

In general, 85% of the patients who survived after the first 24 hours of injury are still alive 10 years later. Projections suggest the need to expand the capacity of attendance/treatment of patients with spinal cord injuries and the need to prepare for the aging of this population. Advances in technology and in medicine have provided greater longevity for patients with spinal cord injuries. However, these patients become susceptible to diseases related to aging. Studies indicate that the occurrence of asymptomatic cardiovascular diseases in people with spinal cord injuries is between 25 and 50%, while the occurrence of symptomatic cardiovascular diseases varies from 30 to 50%. Nevertheless, among people without spinal cord injuries the rate varies between 5 and 10%. This makes cardiovascular diseases one of the main causes of death in older patients with spinal cord injuries. About a third of individuals with spinal cord injury older than 65 and approximately half of those injured for more than 30 years have died of cardiovascular diseases. Pulmonary and cardiac complications are among the main causes of death 12 years after this type of injury, being the first and the second cause of death, respectively. Mortality in people with spinal cord injury is three times greater than in the general population, when compared by age.

From the cardiovascular point of view, when compared to a healthy person, the individual with spinal cord injury presents alterations that interfere with his or her capacity to take in large quantities of oxygen (VO2). In the healthy person, blood is redistributed from inactive tissues to supply muscular activity. This vasoregulation is attributed to the action of the sympathetic nervous system, which in the patient with spinal cord injury is totally or partially absent according to the level of injury. This results in the reduction of cardiac frequency and of the myocardial contractibility, limiting the cardiac output and maximum volume ejected, which reduces the potential to improve the cardiovascular performance.

Nevertheless, muscular atrophy and increase in body fat occur soon after the injury and continue to increase with aging. These changes in the body composition and inactivity predispose people with spinal cord injuries to metabolic abnormalities and accelerate the development of cardiovascular diseases.

Including these cardiovascular and vaso-regulatory mechanisms in patients with spinal cord injuries, the publication of studies that use computerized functional electrical stimulation (CFES) has increased in order to evaluate physiological responses and to develop efficient therapies for the cardiorespiratory conditioning of these patients.

The CFES is a method for therapeutic purposes that stimulates peripheral nerves modulated by a computer through electrodes attached to the skin. It is a form of coordinated electrical stimulation that provokes sequential contraction of the quadriceps femoris, ischiobial, and gluteus muscles, reproducing the pedaling of a bicycle.

Its greatest advantage lies in the possibility of producing the pedal movement of a cycle ergometer through the active contraction of the lower limbs muscles mediated by the electrical stimulation effect. This makes such activity possible even for the individual without any control of these muscles. In the course of training, there is an adaptive response by the individual and the resistance to the movement in the pedals can be modified, provoking more aerobic overload and better cardiovascular conditioning. The improvement of cardiorespiratory conditioning with the CFES is related to the improvement of training load parameters (resistance imposed by the cycle ergometer). In a study with 20 subjects with spinal cord injuries, Glaser compared the physiological responses of healthy people submitted to the same treatment as the injured subjects, and observed the gradual increase of VO2, concomitant to the load increase. In a study with five quadriplegic patients, Hjeltnes evaluated the alterations in body composition after training in the CFES and he observed an increase of VO2 with the load increase, which is in agreement with the study mentioned before. Fignoni evaluated the responses to the exercise in quadriplegic patients, observing the increase of VO2 and of the cardiac output with load increase from 9 W.

CFES is an interesting alternative to aerobic training for people with spinal cord injuries, with the muscular component having an important role in the performance of this training.

OBJECTIVE

This study seeks to determine whether the use of CFES produces beneficial muscular outcomes for individuals with spinal cord injuries.

METHOD

Inclusion criteria

The inclusion and exclusion criteria are defined based on the question that guides the review:

• Without limiting the period of publication;
• Without limiting the language

Types of studies

We reviewed published studies of controlled and random clinical tests, semi-random and controlled clinical tests, controlled clinical tests, series of cases, and case studies.

Participant characteristics

Adult individuals with traumatic spinal cord injury, defined as a penetrating or non-penetrating spinal cord injury resulting from external forces.

Types of intervention

The use of computerized electrical stimulation (CFES) in patients with spinal cord injuries, defined as the use of computerized electrical stimulators. This is equipment that generates low intensity pulses transmitted by surface electrodes, causing coordinated contractions in the large muscles of the legs. It has sensors that provide continuous feedback for the proper control of the muscular contractions and for the resistance of the pedal.

Outcomes

Muscular trophic alterations

Search method to identify the studies

The research had no limitations of data or language

Electronic data bases

• Medline
• Pubmed
• Lilacs
• Portal SciELO

The search strategy was based on questions structured in the P.I.C.O. form (“Paciente” “Patient”, “Intervenção” “Intervention”, “Controle” “Control”, “Outcome/Desfecho” “Outcome”). The following descriptors were
utilized: (Spinal Cord Injuries OR Quadriplegia OR Paraplegia) AND (Electric Stimulation) AND (Ergometry); (Spinal Cord Injuries OR Quadriplegia OR Paraplegia) AND (Electric Stimulation) AND (Bicycling); (Spinal Cord Injuries OR Quadriplegia OR Paraplegia) AND (Electric Stimulation) AND (Exercise); (Spinal Cord Injuries OR Quadriplegia OR Paraplegia) AND (Electric Stimulation) AND (Exercise test); Tetraplegia AND Estimulação elétrica; Paraplegia AND Estimulação elétrica; Traumatismos da medula espinal AND Estimulação elétrica.

Definitions of descriptors
Spinal Cord Injuries/ Traumatismos da medula espinal; Traumas to the spinal cord: penetrating and non-penetrating injuries to the spinal cord resulting from external traumatic forces (e.g. wounds from firearms, whiplash traumas, etc.)
Electric Stimulation/ Estimulação elétrica: Use of electrical currents or potentials to obtain biological responses.
Quadriplegia/Tetraplegia: Severe or complete loss of motor function in all the four limbs that may result from cerebral diseases, spinal cord diseases, peripheral nervous system diseases, neuromuscular diseases, or, rarely, from muscular diseases.
Paraplegia/Paraplegia: Severe or complete loss of motor function in the lower extremities and portions of the trunk. This affliction is more frequently associated with spinal cord diseases, cerebral diseases, peripheral nervous system diseases, neuromuscular diseases, or, rarely, from diseases that can also cause bilateral weakness of the legs.
Exercise Test/Teste de esforço: Controlled physical activity, more active than resting, performed to allow the evaluation of physiological functions, especially cardiovascular and pulmonary, but also of aerobic capacity. Generally, the maximum exercise (more intense) is demanded, but the sub-maximum exercise is also used. The intensity of the exercise is frequently graduated using criteria such as frequency of the work done, consumption of oxygen, and cardiac frequency.
Exercise/ Exercício: Physical activity generally regular and done with the intention of improving or maintaining physical fitness or health. It is different from physical effort, which is directed mainly to physiological and metabolic responses to the use of energy.
Bicycling/ Ciclismo: The use of a bicycle for transport or recreation. This does not include the use of the bicycle in the study of body responses to physical exercises.

Ergometry/Ergometria: Any method to measure the quantity of work done by an organism, generally during physical effort. Ergometry also includes measurements of strength. Some instruments used in these determinations include the manual crank and ergometric bicycle.

The selection of the studies was made through the evaluation of titles and abstracts. When the title and abstract were not clarifying, the entire article was sought out, in order not to leave important studies out of the systematic review. The disagreements that occurred were resolved by consensus.

The data was extracted from each study through a standardized form, with information on the age of the subjects, number of subjects included in the study, type of intervention, the duration, number, and frequency of sessions, types of results measured, and authors’ conclusion on the outcomes of the interventions.

RESULTS

The initial search identified 554 articles (Table 1), from which any duplicates were excluded. The reading of the title (432) and of the abstract (36) excluded 468 articles. From the 37 remaining, after reading the complete text, only 4 satisfied the inclusion criteria in this review.

Bibliographic review articles, studies where the subjects were not adults, studies with associated therapies (strengthening protocol with functional electrical stimulation and computerized electrical stimulation, for example), studies on subjects with incomplete spinal cord injuries and motor activity (ASIA C, D, E), and studies about metabolism and muscle oxygen consumption were excluded. Articles that used non-stationary cycle ergometers were included.

Despite the concern with selecting more homogeneous studies in relation to the characteristics of the injury (complete or incomplete), type of intervention, and type of measurements, some studies presented certain heterogeneity, as presented on Table 2.

Two articles included males and females in their studies,16,19 two other articles, only males. Three studies included quadriplegic and paraplegic subjects in the same study,16-20 one other study included only quadriplegic subjects.16 Two studies included at least one subject with complete injury in the study,15,19 two other studies included only one subject with complete injury.15,19 One of the studies used a greater training frequency, seven times a week,16 three others used a three times a week frequency.18-20 The duration of the studies varied greatly, from six weeks21 to one year.19

The studies’ population samples presented a reduced number of subjects15,18-20 and no studies of random and controlled clinical tests were found.

The result measurements for evaluating strength and resistance were taken by various forms, by measuring the cross-section of muscles16,19 (both studies observed significant increases in the cross-sections), by circumference of the limbs15,20 (one of the studies reported an increase of thigh circumference,15 while another reported maintenance of circumference),20 and by muscular biopsy,18,19 (one of the studies reported an increase in the muscle fibers’ area,18 while another study did not observe any alteration),19 however, all the studies presented at least one of the measurements supplied by the equipment, the evaluation of potency (power output) or of the work performed (work output).16,18-20

DISCUSSION

Despite the wide research done in this study, the number of articles selected was reduced. The use of keywords and broad terms like these in the search for articles is justified, for the studies on computerized electrical stimulation could be associated with various descriptors. Even though the technology associated with CFES is not recent, the availability of equipment is not easy and the demand for regularity in the training requires supplementary efforts by the users to obtain the cardiovascular and musculoskeletal responses desired, which may have contributed to reduce the final number of articles that satisfied the criteria defined.

Thus, the number of articles that really were about the CFES muscular effects was not great. Davis et al.16,17 made a literature review on various outcomes with the use of CFES, muscle morphology and biochemistry, cardiac and hemodynamic responses, bone density, functional changes in the capacity to perform the exercise, psychosocial aspects, and generalized metabolic responses, finding a total of 865 articles, and selecting 177 articles. Peng et al.19 made another literature review evaluating cardiovascular, bone, and muscular aspects, among others. Eight articles were selected regarding the muscular aspect, including evaluations...
on muscular tropism and spasticity, not excluding those studies with subjects who had incomplete injury. These authors observed that the use of CFES promoted increases in the muscle mass and resistance, reduced the frequency and the duration of muscle spasms, increased sensitivity to insulin, increased ventilatory parameters, and increased oxygen consumption.

Coincidently, in this review, the studies selected used the same brand of equipment, Ergys (Therapeutic Alliances Inc, Ohio, USA), which allows us to affirm that the results have a greater consistency. However, other CFES equipment, or cycle ergometer equipment associated with another electrical stimulation apparatus was also already used. Kakebeeke et al. used StimMaster (ELA, Dayton, OH), while Berry and col used Mobile Recumbent Tricycle (Inspired Cycle Engineering Ltd., Falmouth, Cornwall, UK) adapted for coupling with electrical stimulation. Dufel et al.25 made their study using the tricycle Trice (Inspired Cycle Engineering, Ltd., UK) and the Stanmore stimulator as an electrical stimulator, Scremin et al.25 used REGYS (Therapeutic Alliances Inc., Ohio, USA).

Table 1. Number of articles found/selected, by association of descriptors

<table>
<thead>
<tr>
<th>Descriptors</th>
<th>Spinal cord trauma</th>
<th>Paraplegia</th>
<th>Quaeriplegia</th>
<th>Spinal cord injuries OR quadriplegia OR paraplegia</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td>Electric stimulation AND exercise test</td>
<td>*</td>
<td>*</td>
<td>*</td>
<td>93/24</td>
<td>93/24</td>
</tr>
<tr>
<td>Electric stimulation AND exercise</td>
<td>*</td>
<td>*</td>
<td>301/50</td>
<td>301/50</td>
<td></td>
</tr>
<tr>
<td>Electric stimulation AND bicycling</td>
<td>*</td>
<td>*</td>
<td>57/17</td>
<td>57/17</td>
<td></td>
</tr>
<tr>
<td>Electric stimulation AND ergometry</td>
<td>*</td>
<td>*</td>
<td>90/31</td>
<td>90/31</td>
<td></td>
</tr>
<tr>
<td>Electric stimulation</td>
<td>08/00</td>
<td>02/00</td>
<td>03/00</td>
<td>*</td>
<td>13/00</td>
</tr>
<tr>
<td>Total</td>
<td>08/00</td>
<td>02/00</td>
<td>03/00</td>
<td>*</td>
<td>554/122</td>
</tr>
</tbody>
</table>

Table 2. Description of articles used in this review

<table>
<thead>
<tr>
<th>Author</th>
<th>Number of participants (males/ females)</th>
<th>Average age in years ± standard deviation (variation in years)</th>
<th>Level of injury</th>
<th>Average time of Injury in years ± standard deviation (variation in years)</th>
<th>Complete injury (C) Incomplete injury (I)*</th>
<th>Equipment</th>
</tr>
</thead>
<tbody>
<tr>
<td>Chilibeck 1999aa</td>
<td>6 (5 males, 1 female)</td>
<td>(31-50)</td>
<td>CS-T8</td>
<td>(3-25)</td>
<td>A1 C</td>
<td>ERGYS 2 (Therapeutic Alliances Inc., Ohio, USA)</td>
</tr>
<tr>
<td>Gerths 2000b</td>
<td>7 (males)</td>
<td>40.4 ± 11.0 (28-61)</td>
<td>CS-T8</td>
<td>9.8 ± 10.0 (1-27)</td>
<td>5C and 2 I</td>
<td>ERGYS 2 (Therapeutic Alliances Inc., Ohio, USA)</td>
</tr>
<tr>
<td>Hettnes 1997c</td>
<td>5 (males)</td>
<td>35 ± 6</td>
<td>CS-C7</td>
<td>10.2 ± 7.6 (4-23)</td>
<td>A1 C</td>
<td>ERGYS1 (Therapeutic Alliances Inc., Ohio, USA)</td>
</tr>
<tr>
<td>Mohr 1997d</td>
<td>10 (8 males, 2 females)</td>
<td>35.3 ± 7.2 (27-45)</td>
<td>C6-T4</td>
<td>12.5 ± 6.2 (3-23)</td>
<td>9 C and 1 I</td>
<td>REGYS 1 (Therapeutic Alliances Inc., Ohio, USA)</td>
</tr>
</tbody>
</table>

Results

<table>
<thead>
<tr>
<th>Description of randomization</th>
<th>Training frequency per week</th>
<th>Duration of each session</th>
<th>Duration of training</th>
<th>Losses reported*</th>
<th>Type of measurement</th>
</tr>
</thead>
<tbody>
<tr>
<td>Non-randomized</td>
<td>3</td>
<td>30 min</td>
<td>8 weeks</td>
<td>No</td>
<td>Muscle biopsies of the lateral vastus muscle, obtained before and after training. Work output.</td>
</tr>
<tr>
<td>Non-randomized</td>
<td>3</td>
<td>30 min</td>
<td>6 weeks</td>
<td>No</td>
<td>Circumference of thigh, measured at 5 and 20 cm from the patella. Work output.</td>
</tr>
<tr>
<td>Non-randomized</td>
<td>7</td>
<td>30 min</td>
<td>8 weeks</td>
<td>Yes, and justifies losses</td>
<td>Transversal section through computed tomography of gluteus maximus and medius, ischiofibrosis muscles, and quadriceps. Power output.</td>
</tr>
<tr>
<td>Non-randomized</td>
<td>3</td>
<td>30 min</td>
<td>12 months</td>
<td>No</td>
<td>Circumference of thigh, measured at 10 and 20 cm from the patella. Transversal section of thigh muscle, medium portion, by functional magnetic resonance; and muscle biopsy of the lateral vastus muscle. Power output.</td>
</tr>
</tbody>
</table>

The present study shows clearly that 3 weeks of training in the FES-LCE can improve the physiological properties of the quadriceps in individuals with SCI in relation to resistance to fatigue.

Results

| Average increase of 23% in the muscle fibers area and of 39% in the number of capillaries | Eight weeks of training 3x/week of CFES promotes the proportional increase of the fibers' area and of the number of capillaries. These changes can increase functional parameters, such as muscle resistance in individuals with spinal cord injuries. |
| Significant increase of muscles’ cross-section at 22% or 58 cm increase in the load (6W) to (22W). Four subjects were able to perform the training session in the CFES for 30 min with loads varying from 6W (subject 3 and 4) to 12.2W (subject 2) and 18.3W (subject 1), but subject 4 felt fatigue after 20 min at 6W. | The training done 7w/week for 8 weeks promoted significant increase in muscular resistance in quadriplegic patients. |
| The thigh circumference, at 20 cm from the patella, increased 0.9 ± 0.2 cm (p < 0.05) after 6 months and 0.9 ± 0.2 cm (p < 0.05) after 12 months, respectively 3% and 5% increases. Increase of 5% in the cross-section of the thigh muscles. There was no significant increase of the muscle fiber areas. | The musculoskeletal changes that occurred in the subject with spinal cord injury after the injury can be reverted. The training of legs with electrostimulation is an effective rehabilitation tool. |

* Complete injury, sensory and motor complete; incomplete injury, sensory incomplete, but motor complete.
The CFES devices were introduced into the market in the 1980s with the first commercialized equipment - REGYS (Therapeutic Alliances Inc.). Since then other companies have introduced new CFES or cycloergometer models and protocols into the market associated with other electrical stimulation devices. Nevertheless, the high cost and maintenance of the equipment, and the clinical complications inherent to the population studied, may have contributed to the reduced number of studies made.

The evaluation measurements used in the studies of this review were the muscular microstructure evaluation by biopsy, the measurement of limb circumference (thigh), the evaluation of the muscle cross-section (thigh), work output, and power output.

The evaluation of muscle cross section used in two studies\textsuperscript{16,17} showed an increase in area of the muscle fibers examined. However, this increase in the muscle fiber areas was not observed through the muscular biopsy.\textsuperscript{19} According to the authors of this study, muscle fiber areas smaller than 200\textmu m\textsuperscript{2} were not counted, for they could be confused with artifacts, this could justify this controversial finding. Conversely, Gillibeck et al.\textsuperscript{18} evaluated the muscle fiber areas by biopsy and observed an increase in the fibers' areas after the period of training.

The thigh circumference measuring was used by Mohr et al.\textsuperscript{19} and Gerrits et al.\textsuperscript{20} the former observed an increase of this measurement, while the latter did not observe any such change. Gerrits et al.\textsuperscript{20} report that this result may be associated with the fact of this measurement not being sensitive and subject to measuring errors, despite its easy application.

The work output and power output parameters used in these studies indicate the productivity of the exercise, that is, the capacity of the individual to perform the exercise. The work output parameter refers to the quantity of energy spent during the exercise, measured in Joules (J). The power output parameter refers to the velocity at which the quantity of energy is spent per unit of time during the exercise, measured in Watts (W).

FIGONI\textsuperscript{27} correlated cardiorespiratory improvement in quadriplegic individuals submitted to exercise based on their performance at the potency of 9 W (210 Kcal/hour), compared to a walk at the velocity of 3.2 km/h for people without spinal cord injuries. In another study\textsuperscript{28} the authors identified that individuals with spinal cord injuries trained in the CFES presented maximum peak of O\textsubscript{2} consumption equivalent to an individual without spinal cord injury walking at 4.8 km/h or pedaling at 50 W.

Spinal cord injuries create important alterations in the muscle tissue over time. There is an important muscular atrophy in the paralyzed muscles in the first year after the injury, maintaining a "stationary state" of atrophy after that.\textsuperscript{26,27} Nevertheless, in the paralyzed muscles of these individuals with SCI, there is a transformation of fibers type I (slow twitch and resistant to fatigue) into muscle fibers type IIB (fast twitch and quick to fatigue), with the consequent predominance of muscle fibers type IIB (fast twitch and quick to fatigue) in relation to muscle fibers type I and type IIA (fast twitch and resistant to fatigue).\textsuperscript{28}

Training with CFES is capable of partially reversing these alterations in the muscle tissue. Mohr et al.\textsuperscript{19} observed a reversal of muscular atrophy and a reduction of adipose tissue, with a significant increase in muscle fibers type IIA in relation to fibers type IIB.

The findings described above may be criticized due to none of the selected studies being randomized, controlled, or masked clinical tests. However, clinical knowledge allows us to declare with confidence that individuals with complete spinal cord injury do not improve their muscular tropism without some kind of direct stimulation to the musculature, such as electrical current, and that the mere passive cyclic movement only results in maintenance of articular amplitude and muscle tone regulation. Since there were no control groups, it would make no sense to randomize the subjects. Finally, masking in clinical studies aims to reduce the impression that there may be bias in the reporting of results by interference from the research subject or from the evaluator. Since the muscle outcomes selected were obtained objectively by equipment or biopsy, interferences of this nature are not expected. Thus, we can affirm that this synthesis of results from literature offers clear answers as to the effect of exercise associated to CFES on muscular parameters, except for it being applied only to a restricted group of patients with complete spinal cord injury.

CONCLUSION

Despite the heterogeneity found in these studies, the outcomes evaluated indicate that training using CFES in individuals with spinal cord injuries promotes improvement in the muscle parameters, with a significant increase of power and work output after the periods of training, with gains reported after six weeks.\textsuperscript{29} Frequency of training starting from three times a week\textsuperscript{10,15,20} with 30-minute sessions, which means that these patients better support the overload of imposed physical activity. This study still leaves some questions open: what would be the effect of this type of training on individuals with incomplete injury and what is the dose-response relationship, that is, whether more density in training could be more efficient or if there is a ceiling effect.

REFERENCES

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