Aerobic Program in Persons with Stroke: A Systematic Review



Treino Aeróbico em Indivíduos após Acidente Vascular Cerebral: Uma Revisão Sistemática

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ABSTRACT

Introduction: The aim of this study was to perform a systematic review on the exercise trials post stroke.

Material and Methods: Therefore, we conducted a systematic review of randomized controlled clinical trials published in PubMed and PEDro. The inclusion criteria were: Studies – randomized or controlled clinical trials; Participants – Adults of any age with a clinical diagnosis of stroke; Interventions – any aerobic physical training aimed at improving cardiovascular capacity and/or function. Two independent reviewers categorized the selected trials, assessed methodological quality and extracted the relevant data. Various protocols were analyzed and used for both the assessment and physical training of post-stroke subjects.

Results: According to clinical trials analyze, a wide range of instruments were used to measure functional capacity. Ergospirometry and ergometry were found to be main devices in the assessment of cardiovascular capacity. Cycle ergometer training was the most commonly used training strategy, and conventional physiotherapy (based on stretching, strengthening, balance, coordination and gait training) was the most frequent control therapy.

Discussion: Training duration ranged from 4 weeks to 6 months; the mean weekly frequency was 3 sessions per week while training intensity ranged from 40 to 80% of maximal heart rate or VO2 peak. Duration of training sessions ranged from 25 minutes to 1 hour. **Conclusion:** Exercise training is a promising tool for the management of post-stroke patients, both in terms of improved cardiovascular and functional capacity. Further studies are needed to widen the scope of therapeutic management of this population. **Keywords:** Exercise: Stroke; Walking; Exercise Therapy.

RESUMO

Introdução: O objetivo deste estudo foi realizar uma revisão sistemática sobre o exercício após acidente vascular cerebral.

Material e Métodos: Para isso, foi realizada uma revisão sistemática de ensaios clínicos randomizados e controlados publicados no PubMed e PEDro. Os critérios para inclusão dos artigos foram: Estudos - ensaios clínicos randomizados ou controlados; participantes - adultos de qualquer idade com diagnóstico clínico de acidente vascular cerebral; intervenções - qualquer treino físico aeróbio que vise melhorar a capacidade e/ou função cardiovascular. Dois revisores independentes categorizaram os ensaios selecionados, avaliaram a qualidade metodológica e extraíram os dados relevantes. A análise realizada foi dos vários protocolos usados tanto para a avaliação e treino de indivíduos pós-acidente vascular cerebral.

Resultados: De acordo com estudos clínicos analisados, uma vasta gama de instrumentos foi utilizada para medir a capacidade funcional dos pacientes. Ergometria e ergoespirometria foram os principais dispositivos da avaliação da capacidade cardiovascular. Treino\exercício na bicicleta ergométrica foi a estratégia de treino mais comumente usada, e fisioterapia convencional (baseada em alongamentos, fortalecimentos, treino de equilíbrio, coordenação e marcha) foi a terapia de controle mais frequente.

Discussão: A duração do programa variou de 4 semanas a 6 meses, a frequência semanal média foi de três sessões por semana, enquanto que a intensidade do treinamento variou de 40 a 80% da frequência cardíaca máxima ou VO2 máximo. A duração de sessões de treino variou entre 25 minutos a 1 hora.

Conclusão: O treino físico é uma ferramenta promissora para o tratamento de pacientes pós-AVC, tanto em termos de melhoria da capacidade funcional e da capacidade cardiovascular. No entanto, mais estudos são necessários para ampliar a área de atuação terapêutica nesta população.

Palavras-chave: Exercício Físico; Acidente Vascular Cerebral; Marcha.

INTRODUCTION

Stroke is the third most common cause of mortality in developed countries¹ and a leading cause of a permanent disability.² The long-term disability and institutionalization of stroke sufferers pose a substantial economic burden in many countries.³ For this reason, it is necessary further effective rehabilitation programs to improve the recovery of functional status and quality of life of individuals who have suffered a stroke.⁴

Several disabilities potentially follow stroke, including loss of motor, sensory and cognitive functions.¹ Hemiparesis is the main neurologic deficit, affecting an estimated

85% of acute stroke patients.⁵ In most of these individuals, hemiparesis persists after 6 months of the onset, often resulting in impaired mobility and sedentary lifestyle.⁵ Considering the motor sequelae, most studies have focused primarily on restoration of motor control and independence of the affected subject. However, the majority of post-stroke individuals also suffer from cardiac patology,⁶ and have low exercise capacity as a consequence of immobility.⁷

The physical and functional benefits of stroke survivors' engagement in various types of physical activity are now well established. These benefits may include improved

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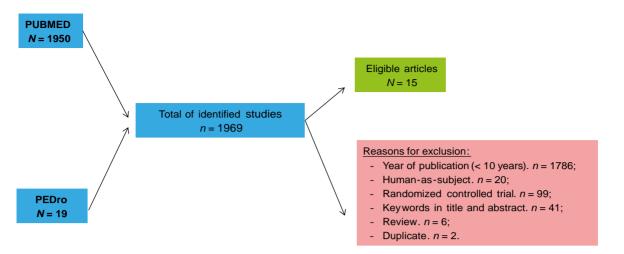


Figure 1 - Flow chart demonstrating search strategy and exclusion criteria

recovery, function and various aspects of fitness. As a result, group exercise classes are emerging as a valuable strategy to promote engagement in regular exercise.⁸

Hill et al⁹ have argued that one factor limiting the incorporation of aerobic exercise into rehabilitation programs is the difficulty in obtaining an exercise stress test from a stroke patient. Neurologic deficits in gait and balance after stroke often preclude exercise testing and training using traditional equipment such as treadmills and cycle ergometers. For this reason, adapted testing protocols and exercise training programs have been developed to evaluate the exercise tolerance e to promote physical benefits in patients after stroke.

Taking into account the range and effectiveness of different training and physical capacity evaluation protocols in the literature, we decided to analyze the literature in this area to survey the techniques and procedures applied in stroke rehabilitation.

MATERIAL AND METHODS

A systematic literature review was conducted from August through January 2013, using the PubMed (a service of the U.S. National Library of Medicine and the National Institutes of Health) and PEDro (Physiotherapy Evidence Database) electronic databases. The PEDro has free access, facilitating the use by professionals, students and researchers in the area. It is available in five languages: English, Mandarin, French, German and Portuguese, and offers a simple version without technical terms to consumers of physiotherapy services. The findings included the articles that contained the key words: 'Stroke' [Mesh] AND 'Physical Training' AND 'Aerobic'.

During the search, the 1950 studies observed presented quite different and sometimes conflicting results, so we decided to narrow our scope to the articles that a) dealt with the evaluation protocols for post-stroke aerobic training; b) used randomized controlled trials; c) were published in English; d) were published in the last 10 years (01/2002 to 10/2012); e) used only humans as research subjects; f) and had keywords in the title and/or abstract. Additionally, studies which failed to adequately describe the characteristics of the evaluation or physical training were excluded. For the selection of articles, the compatibility between titles and abstracts of articles was independently analyzed by two researchers. Each researcher created a single list with a written justification for either inclusion or exclusion of articles, and the two lists were then compared and analyzed by a senior researcher who determined the potential relevance of the articles

This was followed by the reading of the articles in full, and two researchers independently performed data extraction of information about authorship, country of origin, journal title, country of journal, year of publication, impact factor of journal, and methodology used for evaluation and physical training.

When we performed the first key word search in PubMed, we obtained a total of 1950 articles. This number fell to 165 when the 10 years of publication criteria was applied and to 145 when human-as-subjects requirement was addressed. After the criteria of randomized controlled trials and availability in English were applied the number of articles dropped to 46. Of these, 32 failed to meet the criteria of key words in the titles and abstracts. The remaining 14 articles fully met our inclusion criteria.

When keyword search was performed using the PEDro, 19 articles were selected, of which 6 were excluded because they were reviews, 9 because they lacked keywords in the title or abstract (mismatch with the research topic), 2 because they had appeared previously on our PubMed search, and 1 was excluded because it was published in 1999. Only one study could be used for analysis (Fig. 1).

RESULTS

When analyzing the authorship of articles, we found that all studies had multiple authors. 33.3% were written by 4 authors, 20% by 5 authors, 20% by 7 authors, and 13% by 8 authors, with an average 5.1 ± 1.8 authors per article. Country of origin was quite heterogeneous but UK and American journals largely prevailed as sites of publication. These data are shown in Table 1.

The most commonly found protocols for assessment of cardiovascular performance used three instruments which are well known in clinical practice: the six minutes walking test (6MWT - 5 studies) ergometry (7 studies) and ergospirometry (8 studies). The articles showed a concern for quantifying not only the cardiorespiratory but also the functional ability of patients and for this purpose they used a series of tests designed to measure different functions, such as the Force Platform (one study), the Berg Balance Scale (4 studies), the Rivermead Mobility Index (3 studies), the French Activities Index (FAI - 2 studies), the Fugl-Meyer Sensorimotor Test (1 study), the Get up and Go (GUG -1 study), the Functional Independence Measure (FIM - 3 studies), motor assessment scale (1 study), and the Functional Ambulation Category (FAC - 2 studies). These data are shown graphically in Fig. 2.

For a better understanding of the assessment protocols used in the studies analyzed, we chose to divide the results into: 1) Population characteristics (time of stroke, sample, age and gender); 2) Protocols of aerobic training (training duration, weekly frequency, intensity, duration of sessions, type of training); and 3) Main results. Looking at the evaluation protocol, we found that all studies were randomized controlled trials, with one group being selected for aerobic training, and the other as a control group. As to the instruments used for aerobic training, we observed that 60% of studies used cycle ergometer, 33.3% used a treadmill and 6.7% a combination of the two. With regard to control group, 73.3% of studies used the conventional physiotherapy as a therapeutic option, 6.7% used functional training, and 20% had chosen alternative protocols,¹⁰⁻¹² which are shown in Table 2.

We also observed that the total duration of the training ranged between 4 weeks and 6 months (6.7% - 4 week / 20.1% - 6 weeks / 20.1% - 8 weeks / 6.7% - 10 to 12 weeks / 6.7% - 3 months / 6.7% - 14 weeks / 20.1% - 6 months). Regarding weekly frequency, in 9 studies the subjects underwent 3 x/week training, while in 3 studies subjects underwent 5x/week training. As to exercise intensity, 6 studies used fixed values, 40% (1 study), 60% (4 studies) and 70% (1 study) of maximal HR or VO2 peak. Other studies have chosen progressive load ranging from 50-60% (1 study),

Table 1 - Descri	iption of author	and journal cha	aracteristics of	choice for publishing
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Study	N⁰ Authors	Country of Author	Journal	Country of Journal	Impact Factor
Globas et al ¹⁵	8	Switzerland	Neurorehabilitation and Neural Repair	USA	4.495
Shin et al ¹⁴	5	Korea	J Phys Ther Sci	Japan	0.292
Toledano-Zarhi et al ¹⁶	4	Israel	NeuroRehabilitation	International	1.635
Ivey et al ¹⁷	4	USA	Stroke	USA	5.729
Quaney et al ¹⁸	7	USA	Neurorehabilitation and Neural Repair	USA	4.495
Rimmer et al ¹³	5	USA	Arch Phys Med Rehabil	USA	2.282
Janssen et al ¹¹	7	Netherlands	Arch Phys Med Rehabil	USA	2.282
Lee et al ¹²	7	Australia	JAGS	USA	3.737
Lennon et al ¹⁹	4	Ireland	Clinical Rehabilitation	England	2.123
Ivey et al ²⁰	5	USA	Stroke	USA	5.729
Katz-Leurer & Shochina ²¹	2	Israel	NeuroRehabilitation	International	1.635
Macko et al ²²	8	USA	Stroke	USA	5.729
Eich et al ¹⁰	4	Germany	Clinical Rehabilitation	England	2.123
Katz-Leurer et al ²³	3	Israel	Clinical Rehabilitation	England	2.123
Katz-Leurer et al ²⁴	4	Israel	Arch Phys Med Rehabil	USA	2.282

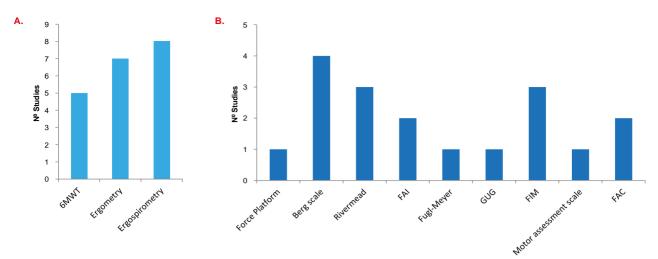


Figure 2 - A) Characteristics of evaluation protocol to determination of cardiorespiratory capacity; B) Characteristics of evaluation protocol to determination of functional capacity. Abbreviations: 6MWT = Six Minutes Walking Test; Berg scale = Berg Balance Scale; Rivermead = Rivermead Mobility Index; FAI = French Activities Index; Fugl-Meyer = Fugl-Meyer Sensoriomotor Test; GUG = Get up and Go; FIM = Functional Independence Measure; FAC = Functional Ambulation Category.

50-70% (2 studies), 60-70% (3 studies) and 60-80% (1 study). However, when we observed session length, 8 studies chose 30 minutes fixed time (4 studies), 40 minutes (3 studies) and 45 minutes (1 study). Some researchers have chosen to create protocols with progression of time, such as 25-30 minutes (1 study), 30-50 minutes (1 study), 35-55 minutes (1 study) and 30-60 minutes (1 study). Three of the 15 studies had different time combinations, ^{10,14,15} which can be seen in Table 2.

Concerning the main results presented, we observed: improvement in cardiorespiratory capacity (11 studies), walking speed (3 studies), as well as functional improvement (3 studies), increase in balance (2 studies) and in muscle strength (1 study). Only one study did not show any improvement after training proposed. Other benefits demonstrated were related to improved blood flow in the paretic muscles, improvement in hemodynamic and metabolic variables (Table 2).

DISCUSSION

Authorship analysis revealed a tendency towards multiple authorship. This can be explained by the complexity of the studies and their clinical nature, composed of samples involving a large numbers of subjects in both aerobic and conventional treatment. Of note was the heterogeneity of country of origin (at least 1 study by geographic continent), possibly due to the magnitude of chronic sequelae of stroke-associated morbidities, which has crossed borders and generated increasing interest in research involving new therapeutic approaches for the affected population. We also observed a tendency for authors to publish their experiments in American or UK journals, a fact that can be explained by the high impact factor of these journals, since these journals follow a tradition of strict scientific selection criteria, promoting greater visibility to the studies.

In the assessment of cardiorespiratory capacity, we identified three methods most used, the 6MWT, ergome-

try and ergospirometry. However, we observed that the 5 studies using the 6MWT, occasionally used the results of this evaluation not to prescribe aerobic exercise, but as a parameter to measure the ability to walk, since the maximal HR data obtained from this evaluation was not taken into consideration for training prescription. However, different functional evaluations were used, and related to the specific objectives for each experiment. For example, the Berg Scale was most frequently used to balance evaluation, Rivermead Mobility Index for mobility, GUG for agility, FIM and FAC for functional independence, and Fugl-Meyer to evaluate sensorimotor characteristics.

The main tool used for aerobic exercise training was cycle ergometer, probably because it is generally regarded as safe for a population with critical balance deficits. Moreover, the choice of conventional physical therapy as control therapy is easy to understand, since in many situations it means a continuation of a treatment patients have been undergoing.

We were surprised to observe a mismatch between the periods, as 40.2% of articles reported short workouts (between 6 to 8 weeks) while 20.1% reported much longer ones (6 months). We detected a somewhat regular cycle, whereby once short-term studies have been proposed, then immediately there was an attempt to replicate, or even to overcome, the same effects by using a long-term protocol. Likewise, studies using long-term protocols were followed by short-term protocol studies, and so forth.

According to our data analysis, we observed that 1 month of aerobic training promoted significant improvement in balance. However, aerobic training between 1 and 3 months of duration promoted increase of cardiovascular capacity, walking speed, balance, strength and improved quality of life. Six months of training induced additional benefits, such as improvement in hemodynamic and metabolic parameters.

In most studies, subjects underwent a 3x/week frequency of training, probably because this population presented important physical deconditioning, needing a rest period between one session and another for full physical recovery. Regarding the duration of the session, it seems clear that, although many studies have chosen constant intensities (range 30-40 minutes), the majority opted for a progressive increase (range 25-60 minutes).

Table 2 - Description of type of study, population and aerobic training protocol characteristics, as well as summary of results

STUDY	DESIGN	POPULATION	DURATION	INTERVENTION	RESULTS
Globas et al ¹⁵	Randomized controlled trial - Exercise group - Control group	Ischemic Stroke > 6 months Middle cerebral artery N = 38 Age > 60 years Male/ Female	3 months	Exercise group 3x/ week 30-50min/ session 60-80% HRR Treadmill Control group Conventional Physiotherapy	↑ Peak VO ₂ ↑ walking velocity ↑ balance Improve in SF-12
Shin et al ¹⁴	Randomized controlled trial - Combined exercise group - Conventional exercise group	Chronic stroke (6 months – 5 years) N = 21 Mean age = 57 years Male/ Female	4 weeks	60 min/ session 5x/ week Combined exercise group 30 min: functional strength training 30 min aerobic exercise: (40%HRR) 15 min: treadmill 15 min: treadmill 15 min: cycle ergometer Conventional exercise group Functional training	Improvement in static and dynamic balance in both groups; Greater improvement in dynamic balance in combined exercise group.
Toledano-Zarhi et al ¹⁶	Randomized controlled trial - Exercise group - Control group	Ischemic Stroke 1-3 weeks Modified Rankin Scale ≤ 2 N = 28 Mean age = 65 years Male	6 weeks	Exercise group 3 hours/week 35-55 min/session 50-70% HRmax Upper cycle ergometer Control group Conventional Physiotherapy	None significant difference between groups, and previous and post intervention.
Ivey et al ¹⁷	Randomized controlled trial - Exercise group - Control group	Ischemic stroke > 6 months <i>N</i> = 53 Mean age: 62 years Male/ Female	6 months	Exercise group 3 x/week 40 min/session 60-70% HRR Treadmill Control group Conventional Physiotherapy	↑ Blood flow – paretic side ↑ Cardiorespiratory function (18%)
Quaney et al ¹⁸	Randomized controlled trial - Exercise group - Control group	Ischemic Stroke < 6 months N = 38 Mean age = 58 years Male/ Female	8 weeks	Exercise group 3 x/week 45 min/session 70% HRR Cycle ergometer Control group Conventional Physiotherapy	↑ VO2 peak, processing velocity and reaction, strength and accuracy during grip test, gait velocity and motor learn.

STUDY	DESIGN	POPULATION	DURATION	INTERVENTION	RESULTS
Rimmer et al ¹³	Randomized controlled trial - Intensity group - Duration group - Control group	Ischemic Stroke < 6 months <i>N</i> = 55 Mean age 59.6 years Male/ Female	14 weeks	3x/week (all groups) Intensity group Moderate intensity 30 min/session 60-70% VO _{2 peak} Cycle ergometer Duration group Progressive intensity 60 min/ session 40-70% VO _{2 peak} Cycle ergometer Cycle ergometer Control group Conventional Physiotherapy	Intensity group ↓ SAP, DAP, triglycerides and total cholesterol Duration group ↓ triglycerides
Janssen et al ¹¹	Randomized controlled trial - Exercise group - Control group	Stroke > 5 months <i>N</i> = 12 Age 54 ± 10 years Male/ Female	6 weeks	2 x/ week 25-30 min/ session Exercise group (Cycling + maximally tolerable to electric stimulation) Control group (Cycling + sensible electric stimulation)	↑ aerobic capacity and functional performance after short aerobic training.
Lee et al ¹²	Randomized controlled trial - Aerobic: cycle training - Progressive resistance training (PRT)	Stroke > 3 months <i>N</i> = 52 Mean age = 63 years Male/ Female	10-12 weeks	30 sessions 50-70% VO _{2 peak} 30-60 min/ session 1) aerobic cycling plus sham PRT 2) sham cycling plus PRT 3) aerobic cycling plus PRT 4) sham cycling plus sham PRT	↑ aerobic capacity Improve functional performance, but strength was not different between groups, as well as, previous and post intervention.
Lennon et al ¹⁹	Randomized controlled trial - Exercise group - Control group	Ischemic Stroke > 1 year N = 48 Mean age = 60 years Male/ Female	10 weeks	Exercise group 16 sessions 30 min/ session 50-60% HRR Cycle ergometer Control group Conventional Physiotherapy	↑ VO _{2 peak} ↓ cardiac risk score, perception exertion, major depression
lvey et al ²⁰	Randomized controlled trial - Exercise group - Control group	Ischemic Stroke < 6 months N = 46 Age > 45 years Male/ Female	6 months	Exercise group 3 x/ week 40 min/ session 60-70% HRR Treadmill Control group Conventional Physiotherapy	↑ 15% VO _{2 peak} ↓ insulin Improvement in glucose response

Table 2 - Description of type of study, population and aerobic training protoco	ol characteristics, as well as summary of results (cont.)
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Table 2 - Description of type of study, population and aerobic training protocol characteristics, as well as summary of results (end)

STUDY	DESIGN	POPULATION	DURATION	INTERVENTION	RESULTS
Katz-Leurer et al ²¹	Randomized controlled trial - Exercise group - Control group	Acute ischiemic stroke N = 64 Mean age = 62 years Male/ Female	8 weeks	Exercise group 3 x/ week 30 min/ session 60% HRR Cycle ergometer Control group Conventional Physiotherapy	↑ endurance and walking performance. No significant interaction effects between autonomic impairment and training.
Macko et al ²²	Randomized controlled trial - Exercise group - Control group	Stroke > 6 months <i>N</i> = 45 Age > 45 years Male/Female	6 months	Exercise group 3 x/ week 40 min/ session 60-70% HRmax Treadmill Control group Conventional Physiotherapy	↑ 17% cardiovascular capacity
Eich et al ¹⁰	Randomized controlled trial - Exercise group - Control group	Acute stroke Supratentorial N = 40 Age: 50-75 years Male/ Female	6 weeks	Exercise group 6 x/ week 30 min/ session treadmill 30 min/ session physioteherapy 60% HRR Control group 60 min/ session Gait Bobath	 ↑ velocity and capacity to walking ↑ 30% performance 6MWT ↑ VO_{2 peak} ↑ functional mobility
Katz-Leurer et al ²³	Randomized controlled trial - Exercise group - Control group	Acute Stroke (30 days) N = 92 Mean age = 63 years Male/ Female	8 weeks	Exercise group 5 x/ week 30 min/ session 60% HRmax Cycle ergometer Control group Conventional Physiotherapy	↓ rest HR ↑ load and working time
Katz-Leurer et al ²⁴	Randomized controlled trial - Exercise group - Control group	Acute stroke Follow-up 6 months N = 90 Mean age = 63 years Male/ female	8 weeks	Exercise group 5 x/ week 30 min/ session 60% HRmax Cycle ergometer Control group Conventional Physiotherapy	None significant difference between groups in daily activities, as well as previous and post exercise intervention.

HRmax (Maximal heart rate); HRR (Heart Rate Reserve); HRmax (Maximal heart rate).

Similarly, we detected a clear disparity among researchers with respect to the intensity of exercises. Six studies used fixed intensity, while in other 7 studies progressive intensities were chosen. One study did not set any intensity for training¹¹ and another study compared three types of protocol, with 2 of them involving aerobic training, one with high intensity (60-70%) and one with low intensity (40-70%).¹³ Both duration and intensity may have been preferentially

chosen because they may facilitate the adaptation of the subjects to training, minimizing discomfort and the possibility of withdrawal.

CONCLUSION

In our overall evaluation, aerobic exercise training emerged as a valuable strategy to improve cardiorespiratory capacity, often associated with significant changes in metabolism in post-stroke patients. The outlook seems to identify very favorable outcomes for both cardiovascular risk reduction and reduced risk of a new ischemic event. Also, the data offered a clearer understanding of the impact on functional variables provided by this training, which may be safely associated with conventional therapy or even as an alternative to it.

Exercise prescription has been hindered by the lack of standardized assessment and aerobic training protocols, which would otherwise help professionals to a fuller informed decision-making in the safe management of poststroke patients. However, the data here presented seems

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promising, both in terms of improved functional capacity and improved cardiovascular capacity. Further studies are needed to help standardize and expand the therapeutic tools for the treatment this population.

CONFLICTS OF INTEREST

The authors declare that there are no conflicts of interest.

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