



UNIVERSIDADE DE BRASÍLIA-UnB FACULDADE DE CEILÂNDIA-FCE CURSO DE FISIOTERAPIA

DANIELLY NERY TOMAZ DA SILVA

INFLUENCE OF ELECTRICAL STIMULATION IN THE QUALITY OF LIFE OF INDIVIDUALS WITH KNEE OSTEOARTHRITIS: A SYSTEMATIC REVIEW

DANIELLY NERY TOMAZ DA SILVA

INFLUENCE OF ELECTRICAL STIMULATION IN THE QUALITY OF LIFE OF INDIVIDUALS WITH KNEE OSTEOARTHRITIS: A SYSTEMATIC REVIEW

Trabalho de Conclusão de Curso apresentado à Universidade de Brasília – UnB – Faculdade de Ceilândia como requisito parcial para obtenção do título de bacharel em Fisioterapia.

Orientador (a): Prof. Dr. João Luiz Quagliotti Durigan

DANIELLY NERY TOMAZ DA SILVA

INFLUENCE OF ELECTRICAL STIMULATION IN THE QUALITY OF LIFE OF INDIVIDUALS WITH KNEE OSTEOARTHRITIS: A SYSTEMATIC REVIEW

Brasília,02/05/2022

COMISSÃO EXAMINADORA

Prof. Dr. João Luiz Quagliotti Durigan
Faculdade de Ceilândia - Universidade de Brasília-UnB
Orientador

Dra. Camila Cadena de Almeida Secretaria de Saúde – Prefeitura de Macapá

Dr. Ivo Vieira de Souza Neto

E-lastic equipamentos e serviços de medição Ltda. – Brasília

Dedicatória

Este trabalho é dedicado à minha mãe, minha família e meus amigos que tanto me ajudaram nessa caminhada.

AGRADECIMENTOS

Gostaria de primeiramente agradecer a Deus, meu Senhor e Salvador, aquele que me amou primeiro, pois sem Ele nada disso seria possível, a importância d'Ele na minha vida é imensurável, nada posso senão por meio do Senhor.

Neste documento, vejo a necessidade de agradecer as pessoas que foram importantes para que eu pudesse chegar neste momento, é um item opcional neste tipo de documento, no entanto este é o único registro formal durante toda minha graduação da minha gratidão para cada uma das pessoas que passou pela minha vida, me fortaleceu, me deu apoio e me fez mais feliz até mesmo nos dias mais difíceis.

Quero começar agradecendo minha mãe, Edgenia, que apesar de todas as dificuldades da vida, sempre me deu o melhor que ela pode, a melhor educação, os melhores ensinamentos, me abriu portas no mundo e me deu a oportunidade de focar na minha graduação, muito obrigada mãe por toda a dedicação e cuidado, eu sei que a senhora sempre fez o melhor que pode. Quero agradecer minha irmã, Géssica, que é quase como uma segunda mãe e minha melhor amiga, me deu o aporte necessário, me acompanhou nessa jornada e em breve também será uma fisioterapeuta excelente, ao meu irmão, Igor, pelo apoio nos momentos de estresse com os problemas na universidade, a minha avó, que me deu um teto e companhia durante todo esse tempo, que brigou comigo e cuidou de mim sempre que preciso, agradecer aos meus tios William e Rosângela, por todos os conselhos, por serem como pais e por estarem sempre presentes, meus tios Wellington e Martina pelos momentos divertidos, minha tia Vanessa por ser minha parceira, meu tio Worlens (in memorian), aos meus primos por me tirarem dos momentos tristes, me fazer companhia e me apoiar e ao Marcio que deu todo o apoio nesse final de graduação. Quero agradecer também ao meu namorado, Mateus, que me apoiou em cada passo dessa graduação, que fez tudo o que estava a seu alcance para me ajudar e me fazer chegar aqui, meus sogros e minha cunhada por todo o apoio, minha madrinha e meu padrinho que me ensinaram muito da vida e que me deram um empurrão para a área da saúde.

Quero agradecer aos meus amigos que a graduação me trouxe, Arielle, Beatriz, Breno, Elane e Lucas, sem vocês a faculdade não seria a mesma, como já conversamos, chegaríamos ao ponto final sem a ajuda um do outro? Sim, mas o fardo foi mais leve, os dias foram mais felizes e sei que tenho em vocês amigos para levar para a vida, quero agradecer minha parceira de estágio obrigatório Lorrane que viveu comigo dias tensos, dias de muito aprendizado, dias de luto e foi um grande suporte para mim, minhas parceiras de estágio da rede Sarah Eleonaia, Ludimila e Tatiana que fizeram esses momentos de grande aprendizado mais e feliz.

Gostaria de agradecer também aos meus preceptores de estágio que moldaram muito do meu perfil profissional, aprendi muito com cada um de vocês, Taíssa, Cristina, Major Dayvison, Major Brandão, e toda a equipe da rede Sarah.

Quero agradecer aos meus professores, que me ensinaram muito dentro e fora de sala, em especial aos professores Aline Teixeira, Aline do Carmo, Aline Toledo, Fernanda, Josevan, João Paulo, Liana, Osmair, Patrícia, Sergio Thomaz, e Vera que marcaram minha vida de uma forma extremamente positiva, nunca esquecerei os seus nomes, seus ensinamentos, aprendi muito com vocês como professores e como pessoas.

Não posso me esquecer do meu orientador João Durigan e sua esposa Rita, que abriram portas para mim durante essa graduação, me abriram os olhos para a área de pesquisa e me incluíram em um grupo cheio de pessoas extremamente inteligentes com muito conhecimento para compartilhar. Quero agradecer também a

todo o grupo Gplast por estarem sempre disponíveis para discutir casos, estudos e compartilhar conhecimento, e ainda dentro desse grupo gostaria de agradecer especialmente a Camila Cadena que foi a doutoranda, hoje doutora que me auxiliou nas orientações de PIBIC, ao Klaus, que se tornou um amigo, que me traduziu as coisas que eu não entendi que o João dizia, que me ajudou na escolha do tema do meu TCC e me mostrou muito dessa área de pesquisa e provou que tem como seguir na área de pesquisa sem abandonar a prática da fisioterapia.

Finalmente, gostaria de agradecer também ao CNPq e a Fundação Universidade de Brasília - FUB pelo apoio financeiro como bolsista de iniciação científica e como tutora, respectivamente.

Epígrafe

Portanto, quer comais quer bebais, ou façais outra qualquer coisa, fazei tudo para glória de Deus.

1 Coríntios 10:31

RESUMO

Introdução: Osteoartrite (OA) é uma doença articular degenerativa que atinge principalmente a população idosa, ela pode atingir diversas articulações, mais comumente em membros inferiores, e levam a diversas limitações funcionais e redução da qualidade de vida. Dessa forma, são necessárias estratégias para atenuar essas respostas prejudiciais. A eletroestimulação é um recurso comumente utilizada com fins terapêuticos, onde uma corrente elétrica altera padrões fisiológicos da área levando melhora ao tecido afetado. Qualidade de vida é um desfecho muito importante que busca avaliar a percepção do indivíduo quanto a diversos aspectos. Objetivo: O objetivo deste estudo foi identificar, a partir de uma revisão sistemática, se o uso da estimulação elétrica melhora efetivamente a qualidade de vida em indivíduos com OA de joelho em comparação com um grupo controle com tratamento padrão ou placebo, Métodos: estudos foram identificados em diversas bases de dados e pesquisados a partir de termos controlados buscados no MeSH. A qualidade metodológica dos estudos foi avaliada pela escala PEDro. Estudos que aplicaram eletroestimulação em participantes com OA de joelho e avaliaram qualidade de vida por meio de questionários verificados foram incluídos, estudos foram excluídos quando trataram outras doenças, ou quando usaram uma outra terapêutica. Resultados: Dentre 917 estudos, apenas 4 atendiam os critérios após a seleção. Os estudos incluídos tiveram uma média de 7,75 na PEDro. Os estudos utilizaram TENS com modulações de parâmetros semelhantes, aplicadas por aparelho portátil em casa e como comparação havia TENS placebo e terapia convencional. Amostra de 199 participantes. A partir do SF-36, três estudos publicaram resultados de melhora na qualidade de vida sem diferença estatisticamente significante, e uma publicação demonstrou efeito. três dos estudos demonstraram diferenças significativas na melhora da dor, enquanto um apresentou melhora, mas não estatisticamente significativa. Conclusão: o estudo não apresentou melhora estatisticamente significante na qualidade de vida desses participantes, no entanto é importante ressaltar que os estudos não utilizaram parâmetros baseados em evidência cientifica para aplicação nessa população, o que poderia levar a diferentes resultados.

Palavras chave: Osteoartrite; eletroestimulação; qualidade de vida.

ABSTRACT

Introduction: Osteoarthritis is a degenerative articular condition that primarily affects the elderly population, it can affect various joints, most commonly lower member, leading to functional limitations and lowering quality of life. Thus, strategies are needed to mitigate these harmful responses. Electrical stimulation is a widely used therapeutic resource in which an electrical current alters the physiology of the treated area leading to the tissue improvement. Quality of life is a new topic that evaluates one's perception of different aspects of life. Objective: The objective of this study is to identify, through a systematic review, if the use of electrical stimulation effectively improves quality of life in participants with knee OA in comparison with a control group. Method: Studies were identified from various databases, searched using keywords of MeSH. The methodological quality of the included studies was assessed using PEDro scale. Studies that used electrical stimulation in participants with knee OA and evaluated quality of life using verified questionnaires were included and studies were excluded when treating other diseases or using another therapeutic modality. Results: In 917 studies only 4 fit the criteria after selection. The studies had a 7,75 mean in PEDro Scale. Studies used TENS with similar parameter modulation applied with a portable device at home and as a control group they used TENS placebo or standard care. The sample was of 199 participants. Through SF-36 three studies showed improvement in quality of life but not statistically significant while one did. three of the studies showed significant improvement in pain relief while one wasn't significant. Conclusion: this study did not show statically significant improvement in quality of life of these participants, however it is important to highlight that the studies did not follow evidencebased parameters to these population which could have changed the results.

Key words: Osteoarthritis; Electric Stimulation; electrical Stimulation; Transcutaneous Electric Nerve Stimulation; Electric Stimulation Therapy; Quality of Life.

LISTA DE TABELAS E FIGURAS

Figura 1. PRISMA flow diagram	19
Tabela 1. Study characteristics	21
Tabela 2. PEDro scale	22
Tabela 3. Characterization of the sample in age gender and BMI	23
Tabela 4. Results of SF-36 in each study	24
Tabela 5. Results of VAS in each study	25

LISTA DE ABREVIATURAS

APTA - American physical therapy association

BMI - Body Mass Index

DeCS - Descritores em Ciências da Saúde

LEFS - Lower Extremity Functional Scale

MCID - minimal clinically important difference

MCS - Mental Component Summary

MeSH - Medical Subject Headings

OA - Osteoarthritis

OAKHQOL - Osteoarthritis Knee and Hip Health-related Quality of Life

PCS - Physical Component Summary

PRISMA - Preferred Reporting Items for Systematic Reviews and Meta-Analyses

QOL - Quality of Life

SF-36 - Short Form Health Survey 36

TUGT - Timed up and go test

VAS - Visual Analogue Scale

WHO - World Health Organization

WHOQOL - World Health Organization Quality of Life

WOMAC - Western Ontario and McMaster Universities Osteoarthritis Index

SUMÁRIO

1. INTRODUÇÃO	14
2. OBJETIVOS	16
3. METODOLOGIA	16
4. RESULTADOS	18
5. DISCUSSÃO	25
6. CONCLUSÃO	28
REFERÊNCIAS	30
ANEXOS	35
ANEXO A – Normas da Revista Científica	35

1. INTRODUCTION

Osteoarthritis (OA) is an inflammatory, biomechanical, and degenerative articular condition characterized by the articular cartilage deterioration, resulting in pain, stiffness, edema, and functional impairments¹. It is most common in knee and hip¹. The three most affected tissues are cartilage, bone, and the synovial fluid².

OA is a condition most commonly found in the elderly population, and its risk factors are age, gender – since women tend to develop the most severe form of the disease and be most affected by it³ – trauma, obesity, metabolic illnesses in which the cause is unknown¹ the hypothesis is that within aging our metabolic rate decreases and when there is a release of normal enzymes in the tissue due to the metabolic imbalance it occurs the degeneration of the joint and all tissues affected by OA. According to Callahan et al⁴ 15% of the population worldwide is affected by OA, being 32.5 million in the US alone.

The population affected with this condition has various functional limitations in daily activities such as walking, climbing stairs, sitting down, and others. These limitations are caused by symptoms of the condition namely swelling, loss of strength and range of movement and pain⁵, culminating in a worsening of the physical health and quality of life⁶.

Electrotherapy or electrical therapy uses energy to achieve a therapeutic goal⁷. It has and it has had different terms along the years. The American Physical Therapy Association⁸ (APTA) has already published an Electrotherapeutic Terminology in Physical Therapy however, according to Cameron⁹ it is not consistently respected. Electrotherapy consists in the use of the electrical current in which the charged particle flow when applied to a body induces physiological changes to achieve a therapeutic effect in the involved tissue.

The electrical stimulation has various goals, such as reducing pain, and increase strength. The electroanalgesia has the goal of reducing pain, and it is based on the gate control theory, and endogenous opioid release mechanisms, activation of delta opioid receptors and the strengthening electrical stimulation is obtained through the stimulation of a muscle contraction to gain strength of the muscle involved in the affected joint to reduce pain¹⁰.

Quality of life (QOL) is also a term with many definitions, and none of them are consistently accepted¹¹. In general, it is a synonym for health or general satisfaction in life, and it varies concept according to different areas of study. The World Health Organization (WHO) defines QOL as an individual's perception of their position in life in the context of the culture and value systems in which they live and concerning their goals, expectations, standards and concerns¹².

The QOL is evaluated through a questionnaire they are practical tools for physical therapists and clinicians because they are low-cost time effective and easy to interpret. There are different questionnaires to evaluate QOL like World Health Organization Quality of Life (WHOQOL) which is a WHO questionnaire with about 100 questions divided into 6 domains: physical, psychological, level of independence, social relationships, environment, and spirituality/ religion/ personal beliefs. The questionnaire Short Form Health Survey 36 (SF-36) is another method to evaluate QOL. It has 36 questions divided into 8 domains: physical functioning, role-physical, bodily pain, mental health, role-emotional, social functioning, vitality and general health. Taft et al divides them into two big domains¹³ the Physical Component Summary (PCS) and the Mental Component Summary (MCS). Besides the general QOL questionnaires, some of them are specific to a condition like the Osteoarthritis

Knee and Hip Health-related Quality of Life (OAKHQOL)¹⁴ that consist of 43 questions divided into 5 domains 3 independent items.

2. OBJECTIVE

The aim of this study is to verify if the use of electrical stimulation effectively improves quality of life in participants with knee OA compared to a control group with standard care or placebo.

3. METHOD

Protocol and Registration

This study is a systematic review. This study selection process included database search, checking for duplicates, screening of titles and abstract, evaluation of inclusion and exclusion criteria, and full-text reading were conducted based on the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA). PROSPERO registration number: CRD42021260207.

Eligibility criteria

We included studies that applied electrical stimulation in participants with knee OA and evaluated QOL through a valid questionnaire. A secondary outcome was assessed: pain through valid scales, since this outcome is an important variant for this condition. There was no restriction on the language of the studies or year of publication or methodological quality also there was no restrictions as to age or gender due to the lack of studies in previous and the actual search.

Information Sources

The Search was conducted in January 2022 in the following databases: Pubmed, Pubmed Central, BVS, Web of Science, Scopus, EBSCO, Embase, Cochrane Central Register of Controlled Trials (CENTRAL).

Search strategies

The search strategy was selected initially through Descritores em Ciências da Saúde (DeCS) to obtain all the controlled vocabulary in English and later applied in the Medical Subject Headings (MeSH) to verify the controlled vocabulary and also to obtain synonyms to create a search strategy for the databases that do not recognize MESH. The search strategy used in the databases BVS, Pubmed, Pubmed Central, Scopus, EBSCO – Medline, Embase e Cochrane Central Register of Controlled Trials (CENTRAL) was Osteoarthritis AND ("Electric Stimulation" OR "Electric Stimulation Therapy" OR "Transcutaneous Electric Nerve Stimulation") AND "Quality of Life" while in Web of Science for not recognizing the controlled vocabulary of MESH the search strategy used will be Osteoarthritis AND ("Electric Stimulation" OR "Electric Stimulation" Therapy" OR "Electrical Stimulation" OR "Therapeutic Electrical Stimulation" OR "Transcutaneous Electric Stimulation" OR "Percutaneous Electric Nerve Stimulation" OR TENS OR Electroanalgesia OR "Analgesic Cutaneous Electrostimulation" OR "Transcutaneous Electrical Nerve Stimulation" OR "Transcutaneous Stimulation") AND ("Quality of Life" OR "Life Quality" OR "Health-Related Quality Of Life" OR "Health Related Quality Of Life").

Study selection

We exclude studies that treated or compared OA with other conditions or use other therapies, except when in the control group (conventional therapy). Within the end of the search process, one independent author selected the studies. In case of dispute, a second reviewer was consulted. If needed, a third one was also consulted. Data collection process

Two authors independently assessed trials for inclusion, and evaluated the risk of bias with PEDro scale the authors will resolve any disagreement regarding the evaluation through discussion.

The following data were obtained in each study: age, gender, body mass index (BMI), type of current and parameters, the primary outcome (QOL), and a secondary outcome, pain, for being an essential factor in this condition.

Risk of bias in individual studies

To evaluate the methodological quality of the included studies two authors independently analyzed PEDro scale. This scale has 11 items to evaluate methodological quality of Clinical Trials. Each item must be given a point if present in the study.

Statistical analysis

The statistical analysis was conducted through mean differences and standard deviations. The Microsoft Excel software was used to fill out the spreadsheet and to do the data analysis.

4. RESULTS

Study selection

Nine hundred and seventeen studies were identified through the search in the previously mentioned databases. After removing all duplicates, there was eight hundred and eighty-four articles out of these. It was identified twenty-one through the screening of titles and abstract which seventeen were excluded for not fitting in the inclusion and exclusion criteria. Four studies^{15,16,17,18} were selected. Most of the articles were excluded due to combined therapies or other treating other pathologies. The flow diagram is available in the picture 1. The study characteristics of all the included are available in Table 1.

Risk of bias

The PEDro scale as mentioned previously evaluates the methodological quality of the selected studies. The selected articles obtained a total score of 6 to 10 with mean of 7,75. Most of the studies did the allocation concealed (75%)^{15,17,18}.

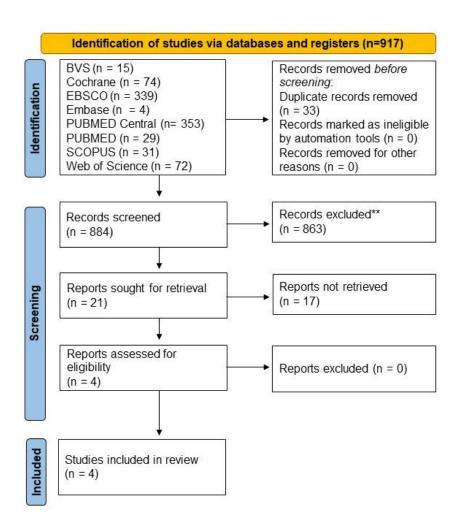


Figure 1: PRISMA Flow Diagram.

All the studies obtained points in random allocation, baseline similarities, less than 15% drop outs, intention to treat analysis, between group comparison reported and post intervention & variability measures. This scale is available in table 2.

Intervention

In all studies, the only type of electrical stimulation used was the eletroanalgesia

Title	Authors	Objectives	Ν	Intervention	Control Group	Results
Use of transcutaneous Eletrical Nerve Stimulation Device in Early Osteoarthritis of the Knee	Cherian , J et al.	Evaluate the effects of the application of TENS in patients in the initial stage of knee osteoarthritis (Kellgren-Lawrence grade 1 and 2) over the following factors: pain reduction, subjective functional improvements, objective functional improvements, QOL measure improvements and isokinetic strength.	23	TENS	Non operative therapy with self-directed exercise therapy and corticosteroid injections	TENS had significant improvements in Timed up and go test (TUGT) and objective Knee society score (KSS) when compared with the control group and had significant improvement in the physical component of their Lower Extremity Functional Scale (LEFS), SF-36, quadriceps strength and the timed stair climb.
Knee Osteoarthritis: Does Transcutaneous Electrical Nerve Stimulation Work?	Cherian , J et al.	Evaluate TENS device efficacy comparing to the standard treatment for Kellgren-Lawrence grade 3 and 4 of knee OA.	36	TENS	9 received nonoperative treatment involving corticosteroid injections and 9 received 6 weeks of physical therapy.	TENS group improved in Visual Analogue Scale (VAS) pain. TUGT, Timed stair climb, 2 minutes walk test, 5-repetition chair rise test, 6-inch step test, and passive flexion motion. However, the control group improved in the active range of motion.

	Do the Effects of Transcutaneous Electrical Nerve Stimulation on Knee Osteoarthritis Pain and Function Last?	Cherian , J et al.	compare TENS treatment with standard care. the study objective is to determinate the differences in: pain perception, subjective medication use, subjective functional abilities, QOL, dispositive use and TKA conversion between groups.	70	TENS	Standard care including corticosteroid injections and physical therapy exercises.	TENS lower scores in VAS and showed significant improvement in subjective and functional outcomes also patients in the TENS group reduced the amount of pain medication.
_	The Effectiveness of Pulse Electrical Stimulation in the Management of Osteoarthritis of the Knee	Fary, R et al.	Determine efficacy of subsensorial pulsed electrical stimulation (PES) in the management of symptoms of knee OA	70	TENS	TENS placebo: an identical device was used however the electrical pulse was turned off after 3 minutes.	There was no difference between groups on pain, function or stiffness scores in Western Ontario and McMaster Universities Osteoarthritis Index (WOMAC); 56% achieved a clinically relevant 20-mm improvement in VAS pain score at 26 weeks compared with 44% of controls;

Table 1: study characteristics. N= sample.

obtained from Transcutaneous Electrical Nerve Stimulation (TENS).

In all studies the participants received a portable stimulator with pre modulated currents, in the three of the studies^{15,16,17} they were meant to use the stimulator throughout the whole day except when in water activities with the following parameters: a pulsed

	Cherian et al (2015 v.28)	Cherian et al (2016 v.29)	Cherian et al (2016 v.39)	Fary et al (2011 v.63)
Eligibility criteria				
specified	\checkmark	\checkmark	\checkmark	\checkmark
Random allocation Allocation	1	1	1	1
concealed	1	0	1	1
Baseline similarity	1	1	1	1
Patient blinding	1	0	0	1
Physical therapist blinding	0	0	0	1
Assessor blinding	0	0	0	1
< 15% drop outs	1	1	1	1
Intention to treat analysis Between group comparison	1	1	1	1
reported	ı	ı	ı	ı
Post intervention point & variability measures	1	1	1	1
Total score	8/10	6/10	7/10	10/10

Notes: $\sqrt{\ }$ = Eligibility criteria satisfied (Not used in total score calculation); 1 = yes; 0 = no Table 2: PEDro scale.

current with a simple asymmetrical wave, a pulse duration of 48 to 100 µs with intervals of 12 seconds, an alternating pulse frequency and the intensity was not mentioned, and the fourth study¹⁸ they were meant to use the stimulator for 7 hours a day hours preferably during sleep hours, with the following parameters: a pulsed current with an asymmetrically biphasic wave, a pulse frequency of 100 Hz, a pulse duration of 4 ms

and the intensity was turned up until they felt pins and needles and after achieving it they were supposed to turn it down the intensity until they could no longer feel it.

As a control group, the studies choose either the placebo¹⁸ characterized by the Transcutaneous electrical nerve stimulation (TENS) application but in a short duration of time or corticosteroid injections or a six weeks protocol of physical therapy^{15,16,17} for strengthening and range of motion.

Sample

The sample analyzed was a total of 199, 50,75% being the control group and 49,25% electrical stimulation. All studies claim there was no significant difference in the sample when comparing age, gender, and body mass index (BMI) with one exception one of the studies did not have the BMI data. All the characteristics of the sample can be found in table 3.

			erian v. 28	Cherian 2016 v. 29		Cherian 2016 v. 39		Fary v. 63		Mean	
		ES	CG	ES	CG	ES	CG	ES	CG	ES	CG
Age		55	54	62	58	63	61	70,7	68,9	62,67	60,47
Condor	F	11	7	19	27	7	9	17	20	13,5	15,75
Gender	M	2	3	14	10	11	9	17	16	11	9,5
BMI		32,5	34,8	N/A	N/A	33,1	34,3	29,4	26,8	23,75	23,97

Table 3: Characterization of the sample in age, gender and BMI. ES: electrical stimulation; CG: Control Group; F: female; M: male; N/A: not available.

The mean age of all participants in electrical stimulation as approximately 63 years old in a sample of 54 women and 44 men. In the control group was approximately 60 years old in a sample of 63 women and 38 men.

The mean BMI of those who explored this variant was 31,67 in electrical stimulation with a standard deviation of approximately 1,6. The mean in the control group was 31,97 with a standard deviation of approximately 3,7.

	Electrical Stimulation														
	Cherian 15				Cherian 16 v.29 Cherian 16 v.39						Fary				
_	-36		-36		SF-36 SF-36			SF-36		SF-36		SF-36		SF-36	
Phy:	sical	Me	ntal	Phy	sical	Mental		Mental Physical		Mental		Phy	sical	Mental	
Pre	Post	Pre	Post	Pre	Post	Pre	Post	Pre	Post	Pre	Post	Pre	Post	Pre	Post
28,13	39,53	50,46	54,52	29,2	41	48	48 52,6		Δ		37	36	52,7	51,5	
						Con	trol Gr	oup							
	Cherian 15				Cherian 16 v.29			Cherian 16 v.39			Fary				
SF.	-36	SF	-36	SF	-36	SF	-36	SF	-36	SF	-36	SF	-36	SF	-36
Phy:	sical	Me	ntal	Phy	sical	Mental		Phy	/sical	Me	ental	Phy	sical	Ме	ntal
Pre	Post	Pre	Post	Pre	Post	Pre	Post	Pre	Post	Pre	Post	Pre	Post	Pre	Post
٨				32.8	36.4	51.3	48 2		/	١		36.5	33 9	53.7	51.3

Table 4: Results of SF-36 in each study. Δ : Variation.

Quality of life

As mentioned previously quality of life is of each individual, and to evaluate it some questionnaires were developed for example, SF-36, WHOQOL, and others. All studies in this research used SF-36.

All of the included studies analyzed the SF-36 data through two domains, physical and mental. Three of the studies noticed an improvement in the physical and mental domain however the improvement was not statistically significant ^{16,17,18}, while one of the studies affirm to have seen a statistically significant improvement in both domains ¹⁵. Commonly in all studies the highest improvement was seen in the physical domain. The results of each study are available in table 4.

No other scale or questionnaire was used to evaluate the quality of life in none of the studies.

Pain

Pain is an important factor for patients with OA due to its incapacitating power.

All studies included in this review address pain somehow. Although it is a sensorial and subjective experience, it is possible to quantify and or qualify through scales.

All studies used the visual analogue scale (VAS) as an evaluation method. In All studies used the visual analogue scale (VAS) as an evaluation method. In three of the studies there was an improvement statistically significant in the individual's pain, however, one of them did not show a significant difference. The analysis of pain can be found in table 5.

Electrical Stimulation										
Che	rian 15	Cheria	n 16 v.29	Cheri	an 16 v.39	Fary				
	/AS	V	'AS		VAS	VAS				
Pre	Post	Pre	Post	Pre	Post	Pre	Post			
5	2,38	4,95	4,55		Δ	51	20			
			Control	Group						
Che	rian 15	Cheria	n 16 v.29	Cheri	an 16 v.39	Fary				
VAS VA		'AS		VAS		VAS				
Pre	Post	Pre	Post	Pre	Post	Pre	Post			
Δ		4,5	5,12		Δ	52	19			

Table 5: Results of VAS in each study. Δ: Variation.

5. DISCUSSION

The primary outcome study in this review was found in all articles included, and so was the secondary outcome. The SF-36 questionnaire is valid in Australia¹⁹ and in the United States of America²⁰ where the studies were developed. It is vast the use of electrical stimulation for the target population of this study. Several studies were conducted using electric stimulation for this population¹⁰. One of the most used currents was TENS²¹ due to the pain relief capacities of this current, achieved based on the gate control theory and the mechanisms of endogenous opioid release. All the studies included in this review used TENS, but they did not follow modulation oriented through systematic review to decide the best parameters^{22,23} this way, the results could have changed if applied the proper modulations based on evidence which are pulse frequency of 80Hz, pulse width of 140us, current intensity the highest tolerable by the participant in a sensorial level and the duration of 30 minutes²⁴.

Pain is one of the main incapacitating factors of OA. According to Cohen²⁵, pain is an unpleasant emotional and sensory experience associated with real or potential damage to the tissue. According to Capela²⁶, the quality of life of an individual has a strong correlation to pain especially in individuals with chronic pain²⁷ since they usually have the most intense symptoms of pain and have the worst quality of life thus these two variables have a negative correlation.

It is known that individuals with OA tend to have the functional capacity, functional limitation, and pain affected in the SF-36²⁸, and both of the first ones are strongly correlated to pain which usually increases with degeneration and age²⁹. As mentioned before, the quality of life of an individual is highly affected by pain, studies^{27,30} conducted on individuals with chronic and back pain affirm that those individuals have a reduced quality of life when compared to individuals who did not experience chronic pain, and this variable was independent of the gender of the sample.

Electrical stimulation is recommended for pain relief in other situations such as labor³¹, post episiotomy³², OA²⁴ it is related to the release of endogenous opioids and activation of descendant inhibitory pathways to reduce hyperalgesia. Individuals with intervertebral disc degeneration in low back³³ reported significant improvement on pain when using TENS, so did the individuals that went through colonoscopy³⁴ and did not use any sedation. This information corroborates that pain relief using electrical stimulation does not keep itself to few studies it has lots of possibilities of usage when we think about the intensity or the location or the condition appliable.

The usage of TENS to improve quality of life is also used in other pathologies such as low back pain³⁵, diabetic peripheral neuropathy and others³⁶. A study conducted by one of the greatest references and electrical stimulation Kathleen Sluka³⁶

demonstrated that reducing pain improves the quality of life of an individual in various pathologies.

Limitations

These studies conducted by Cherian et al¹⁵ and Cherian et al¹⁷ presented its results in a way that could interfere with the possible evaluation of the results. The study Cherian et al¹⁵ show the results of the pre and post treatment only with the TENS group however it did not bring the same the results in the control group only the variation of the pre- and post-treatment results. The study Cherian et al¹⁷ showed only the variation from pre- and post-intervention in both groups.

It can be said that the usage of electrical stimulation can be recommended due to the QOL correlation to pain, but there is not strong evidence to refute the application of the current. Without significant statical relevance we could indicate clinical improvement of the participants through the minimal clinically important difference (MCID). The MCIDs are obtained from a specific study to determine the scores of clinically significant improvement using the mean pre and post intervention scores but it is not applicable to this study because the Cherian et al¹⁵ and Cherian et al¹⁷ studies lack the data already mentioned. Searching the studies of MCID it was also noticed that there were very few studies on this approach on OA³⁷ one of the studies showed the clinically improvement scores on SF-36, the questionnaire outcome in this present study, individuals with osteoarthritis however, this study contemplates seven of the eight domains in SF-36 and also did not use the two domains (mental and physical) analysis making it unfeasible for this evaluation of the results in the present study. Other studies were found correlating to different situations like total knee arthroplasty³⁸ but it also analyzes the SF-36 in the format of eight domains not in two big ones.

The analysis of PEDro scale shows that one¹⁸ of the studies executed a triple blind study one¹⁵ of the other three did blinding of the participants. Studies that do not follow the blinding method tend to have better results and according to John P A lodannidis³⁹ blind studies should be done to avoid biased results.

Also, another limitation is the lack of studies and this approach seen as we have only four studies included.

6. CONCLUSION

This study showed that there was no significant statistical improvement in QOL, evaluated through SF-36, in participants with osteoarthritis after the treatment with electrical stimulation. But it also showed that electrical stimulation promotes improvements in the physical domain which correlates to the pain relief provided by the electrical stimulation. It can be said that there is a significant statistically improvement in pain of these participants which can lead to improvement of the quality of life seeing as they have a strong correlation. The studies included did not follow the recommended parameters²⁴ which could have changed the results. Thus, electrical stimulation can be considered as a recommendation of a good resource in the rehabilitation of people with knee OA.

Therefore as the National Institute for Health and Clinical Excellence⁴⁰ recommends the usage of TENS in the treatment of people with knee OA for pain relief and improvement of the stiffness especially in the short-term treatment it is indicated the use of electrical stimulation with the following parameters: pulse frequency of 80Hz, pulse width of 140us, current intensity the highest tolerable by the participant in a sensorial level and the duration of 30 minutes²⁴ furthermore it can also be applied with other interventions as pain education for the participant and the family, physical

exercises for the reduction of mechanical factors on the joint with the goal of pain relief and improvement of quality of life.

Lastly, it is recommended the development of new studies on the subject so that we can have a higher power of evidence.

This research did not receive any specific grant from funding agencies in the public, commercial, or not-for-profit sectors.

REFERENCES

- Marieb EN, Wilhelm PB, Mallatt J. Anatomia Humana. 7th ed. São Paulo: Pearson Education do Brasil; 2014.
- Rezende MUD, Campos GCD, Pailo AF. Conceitos Atuais em Osteoartrite.
 2013 Janeiro 5; 21: p. 120-122.
- Cançado FAX, Doll J, Gorzoni ML. Tratado de Geriatria e Gerontologia.
 3rd ed. Rio de Janeiro: Guanabara Koogan; 2013.
- Callahan LF, Cleveland RJ, Allen KD, Golightly Y. Racial/Ethnic, Socioeconomic, and Geographic Disparities in the Epidemiology of Knee and Hip Osteoarthritis. Rheumatic Disease Clinics of North America. 2021 Fevereiro; 47(1).
- McDonough CM, Jette AM. The contribution of osteoarthritis to functional limitations and disability. Clinics in Geriatric Medicine. 2010 Agosto; 26: p. 387-399.
- Meng Z, Huang R. Topical Treatment of Degenerative Knee Osteoarthritis.
 The American Journal Of The Medical Sciences. 2018 Janeiro; 355: p. 6-12.
- 7. Pereira MdFL, Paganin C. Eletroterapia: Difusão; 2016.
- American Physical Therapy Association. Electrotherapeutic Terminology in Physical Therapy. Journal of Orthopaedic & Sports Physical Therapy. 1996; 23: p. 233.
- 9. Cameron MH. Agentes Físicos na reabilitação. 3rd ed.; 1999.
- 10. Giggins OM, Fullen BM, Coughlan GF. Neuromuscular electrical stimulation in the treatment of knee osteoarthritis: a systematic review and meta-analysis. clinical rehabilitation. 2012; 26: p. 867-881.
- 11. Pereira ÉF, Teixeira CS, Santos Ad. Qualidade de vida: abordagens, conceitos e avaliação. Revista Brasileira de Educação Física e Esporte. 2012 Abril/ Junho; 26: p. 241-250.
- 12. Fleck MPdA. O instrumento de avaliação de qualidade de vida da Organização Mundial da Saúde (WHOQOL-100): características e perspectivas. Revista Ciência & Saúde Coletiva. 2000; 5: p. 33-38.

- 13. Taft C, Karlsson J, Sullivan M. Do SF-36 summary component scores accurately summarize subscale scores? Quality of Life Research. 2001; 10: p. 395-404.
- 14. Choojaturo S, Sindhu S, Utriyaprasit K, Viwatwongkasem C. Factors associated with access to health services and quality of life in knee osteoarthritis patients: a multilevel crosssectional study. BMC Health Services Research. 2019; 19.
- 15. Cherian JJ, Kapadia BH, Bhave A, McElroy MJ, Cherian C, Harwin SF, et al. Use of Transcutaneous Electrical Nerve Stimulation Device in Early Osteoarthritis of the Knee. Journal of knee surgery. 2015; 28: p. 321-328.
- 16. Cherian JJ, Harrison PE, Benjamin SA, Bhave A, Harwin SF, Michael A. Mont. Do the Effects of Transcutaneous Electrical Nerve Stimulation on Knee Osteoarthritis Pain and Function Last? Journal of knee surgery. 2016; 29: p. 497-501.
- 17. Cherian JJ, Kapadia BH, McElroy MJ, Johnson AJ, Bhave A, Harwin SF, et al. Knee Osteoarthritis: Does Transcutaneous Electrical Nerve Stimulation Work? Orthopedics. 2016; 39: p. e180-e186.
- 18. Fary RE, Carroll GJ, Briffa TG, Briffa NK. The Effectiveness of Pulsed Electrical Stimulation in the Management of Osteoarthritis of the Knee. Arthritis & Rheumatism. 2011 Maio; 63: p. 1333-1342.
- 19. Butterworth P, Crosier T. The validity of the SF-36 in an Australian National Household Survey: demonstrating the applicability of the Household Income and Labour Dynamics in Australia (HILDA) Survey to examination of health inequalities. BMC Public Health. 2004 Outubro; 4(44).
- 20.RAND Corporation. Rand Corporation. [Online]. [cited 2022. Available from: https://www.rand.org/health-care/surveys_tools/mos/36-item-short-form.html.
- 21. Adam CT, Ferrarini EG, Avelar NCPd. Effects of neuromuscular electrical stimulation on pain, function performance and muscle strength in patients with knee osteoarthritis. Fisioterapia Brasil. 2018; 19: p. 561-567.
- 22. Bjordala JM, Johnsonb MI, Ljunggreen AE. 1. Transcutaneous electrical nerve stimulation (TENS) can reduce postoperative analgesic

- consumption. A meta-analysis with assessment of optimal treatment parameters for postoperative pain. European Journal Pain. 2003; 7: p. 181-188.
- 23. Almeida CCd, Silva VZMd, Júnior GC, Liebano RE, Durigan JLQ. Transcutaneous electrical nerve stimulation and interferential current demonstrate similar effects in relieving acute and chronic pain: a systematic review with meta-analysis. Brazilian Journal of Physical Therapy. 2018; 22: p. 347-354.
- 24. Morgan CR, Santos FS. Estudo da estimulação elétrica nervosa transcutânea (TENS) nível sensório para efeito de analgesia em pacientes com osteoartrose de joelho. Fisioterapia em movimento. 2011 Dezembro; 24: p. 637-646.
- 25. Cohen M, Quintner J, Rysewyk Sv. Reconsidering the International Association for the Study of Pain definition of pain. Pain Reports. 2018 Março/ Abril; 3: p. 1-7.
- 26. Capela C, Marques AP, Assumpção A, Sauer JF, Cavalcante AB, Chalot SD. Associação da qualidade de vida com dor, ansiedade e depressão. Fisioterapia e Pesquisa. 2009 Julho/Setembro; 16: p. 263-268.
- 27. Ferretti F, Castanha AC, Padoan ER, Lutinski J, Silva MRd. Quality of life in the elderly with and without chronic pain. Brazilian journal of pain. 2018 Abril/Junho; 1: p. 111-115.
- 28. Kawano MM, Araújo ILA, Castro MC, Matos MA. Assessment of quality of life in patients with knee osteoarthritis. Acta Ortopédica Brasileira. 2015 Novembro/Dezembro; 23: p. 307-310.
- 29. Rosemann T, Laux G, Kuehlein T. Osteoarthritis and functional disability: results of a cross sectional study among primary care patients in Germany. BMC Musculoskeletal Disorders. 2007; 8.
- 30. Nasution IK, Lubis NDA, Amelia S, Hocin K. The correlation of pain intensity and quality of life in chronic LBP patients in Adam Malik general hospital.. IOP Conf. Series: Earth and Environmental Science.; 125.

- 31. Czech Iea. Pharmacological and Non-Pharmacological Methods of Labour Pain Relief—Establishment of Effectiveness and Comparison. International journal ofenvironment Research and public health. 2018 Dezembro; 15.
- 32. Pitangui AC, Araújo RC, Bezerra MJ, Ribeiro CO, Nakano AM. Low and high-frequency TENS in post-episiotomy pain relief: a randomized, double-blind clinical trial.. Brazilian Journal of Physical Therapy. 2014 Janeiro/fevereiro; 18: p. 72-78.
- 33. Pop T, Austrup H, Preuss R, Niedziałek M, Zaniewska A, Sobolewski M, et al. Effect of TENS on pain relief in patients with degenerative disc disease in lumbosacral spine. Ortopedia Traumatologia Rehabilitacja. 2010 Julho/Agosto; 12: p. 289-300.
- 34. Amer-Cuenca JJ, Goicoechea C, Girona-López A, Andreu-Plaza JL, Palao-Román R, Martínez-Santa G, et al. Pain relief by applying transcutaneous electrical nerve stimulation (TENS) during unsedated colonoscopy: a randomized double-blind placebo-controlled trial.. European Journal of Pain. 2011 Janeiro; 15: p. 29-35.
- 35. Zaniewska R, Okurowska-Zawada B, Kułak W, Domian K. Analysis of quality of life in patiens with low back pain after receiving transcutaneous electrical nerve stimulation (TENS).. Medycyna pracy. 2012; 63: p. 295-302.
- 36. Vance CG, Dailey DL, Rakel BA, Sluka. KA. Using TENS for pain control: the state of the evidence.. Pain Management. 2014 Maio; 4: p. 197-209.
- 37. Angst F, Benz T, Lehmann S, Aeschlimann A, Angst J. 1. Multidimensional minimal clinically important differences in knee osteoarthritis after comprehensive rehabilitation: a prospective evaluation from the Bad Zurzach Osteoarthritis Study. Rheumatic & musculoeskeletal diseases Open. 2018; 4: p. 1-9.
- 38. Keurentjes JC, Tol FRV, Fiocco M, Schoones JW, Nelissen RG. Minimal clinically important differences in health-related quality of life after total hip or knee replacement. Bone joint research. 2012 Maio; 1: p. 71-77.
- 39. Ioannidis. JPA. Why Most Published Research Findings Are False. PLOS Medicine. 2005 Agosto; 2: p. e124.

40. National Institute for Health and Clinical Excellence (NICE). steoarthritis. The care and management of osteoarthritis in adults. NICE Clinical Guideline. 2008;: p. 59.

ANEXOS

NORMAS DA REVISTA CIENTÍFICA

GUIDE FOR AUTHORS

INTRODUCTION

Types of article

a) Systematic reviews: studies that analyze and/or synthesize the literature on a topic related to the scope of the BJPT. Systematic reviews that include meta-analysis will have priority over other systematic reviews. Those that have an insufficient number of articles or articles with low quality in the Methods section and do not include an assertive and valid conclusion about the topic will not beconsidered for peer-review analysis.

The authors must follow the Preferred Reporting Items for Systematic Reviews and Meta- Analyses (PRISMA) checklist to format their systematic reviews. The checklist is available at http://www.prisma-statement.org/PRISMAStatement/Default.aspx and must be filled in and submitted with the manuscript.

Potential authors are encouraged to read the following tutorial, which contains the minimum requirements for publication of systematic reviews in the BJPT: Mancini MC, Cardoso JR, Sampaio RF, Costa LCM, Cabral CMN, Costa LOP. Tutorial for writing systematic reviews for the Brazilian Journal of Physical Therapy (BJPT). Braz J Phys Ther. 2014 Nov-Dec; 18(6):471-480.

Submission checklist

You can use this list to carry out a final check of your submission before you send it to the journal forreview. Please check the relevant section in this Guide for Authors for more details.

Ensure that the following items are present:

One author has been designated as the corresponding author with contact details:

- E-mail address
- Full postal address

All necessary files have been uploaded:

Manuscript:

- Include keywords
- All figures (include relevant captions)
- All tables (including titles, description, footnotes)
- Ensure all figure and table citations in the text match the files provided
- Indicate clearly if color should be used for any

figures in print Graphical Abstracts / Highlights files

(where applicable) Supplemental files (where applicable)

Further considerations

- Manuscript has been 'spell checked' and 'grammar checked'
- All references mentioned in the Reference List are cited in the text, and vice versa
- Permission has been obtained for use of copyrighted material from other sources (including theInternet)
- A competing interests statement is provided, even if the authors have no competing interests todeclare
- Journal policies detailed in this guide have been reviewed

Referee suggestions and contact details provided, based on journal requirements

For further information, visit our Support Center.

PREPARATION

Article structure

Subdivision - unnumbered sections

Divide your article into clearly defined sections. Each subsection is given a brief heading. Each heading should appear on its own separate line. Subsections should be used as much as possible when cross-referencing text: refer to the subsection by heading as opposed to simply 'the text'.

Introduction

State the objectives of the work and provide an adequate background, avoiding a detailed literaturesurvey or a summary of the results.

Material and methods

Provide sufficient detail to allow the work to be reproduced.

Results

Results should be clear and concise.

Discussion

This should explore the significance of the results of the work, not repeat them. A combined Resultsand Discussion section is often appropriate. Avoid extensive citations and discussion of published literature.

Conclusions

The main conclusions of the study may be presented in a short Conclusions section, which may standalone or form a subsection of a Discussion or Results and Discussion section.

Appendices

If there is more than one appendix, they should be identified as A, B, etc. Formulae and equations inappendices should be given separate numbering: Eq. (A.1), Eq. (A.2), etc.; in a subsequent appendix, Eq. (B.1) and so on. Similarly for tables and figures: Table A.1; Fig. A.1, etc.

Hiahliahts

Highlights are mandatory for this journal as they help increase the discoverability of your article viasearch engines. They consist of a short collection of bullet points that capture the novel results of your research as well as new methods that were used during the study (if any). Please have a look at the examples here: example Highlights.

Highlights should be submitted in a separate editable file in the online submission system. Please use 'Highlights' in the file name and include 3 to 5 bullet points (maximum 85 characters, including spaces, per bullet point).

Abstract

A concise and factual structured abstract is required. The abstract should briefly state the purpose of the research, the principal results and major conclusions. An abstract is often presented separately from the article, so it must be able to stand alone. For this reason, References should be avoided, but if essential, then cite the author(s) and year(s).

Keywords

Immediately after the abstract, provide a maximum of 6 keywords, using American spelling and avoiding general and plural terms and multiple concepts (avoid, for example, 'and', 'of'). Be sparingwith abbreviations: only abbreviations firmly established in the field may be eligible. These keywordswill be used for indexing purposes.

Acknowledgements

Collate acknowledgements in a separate section at the end of the article before the

references and donot, therefore, include them on the title page, as a footnote to the title or otherwise. List here thoseindividuals who provided help during the research (e.g., providing language help, writing assistanceor proof reading the article, etc.).

Formatting of funding sources

List funding sources in this standard way to facilitate compliance to funder's requirements:

Funding: This work was supported by the National Institutes of Health [grant numbers xxxx, yyyy]; the Bill & Melinda Gates Foundation, Seattle, WA [grant number zzzz]; and the United States Institutes of Peace [grant number aaaa].

It is not necessary to include detailed descriptions on the program or type of grants and awards. Whenfunding is from a block grant or other resources available to a university, college, or other researchinstitution, submit the name of the institute or organization that provided the funding.

If no funding has been provided for the research, please include the following sentence:

This research did not receive any specific grant from funding agencies in the public, commercial, ornot-for-profit sectors.

Figure captions

Ensure that each illustration has a caption. Supply captions separately, not attached to the figure. Acaption should comprise a brief title (**not** on the figure itself) and a description of the illustration. Keep text in the illustrations themselves to a minimum but explain all symbols and abbreviations used.

Tables

Please submit tables as editable text and not as images. Tables can be placed either next to the relevant text in the article, or on separate page(s) at the end. Number tables consecutively in accordance with their appearance in the text and place any table notes below the table body. Be sparing in the use of tables and ensure that the data presented in them do not duplicate results described elsewhere in the article. Please avoid using vertical rules and shading in table cells.

References

Citation in text

Please ensure that every reference cited in the text is also present in the reference list (and vice versa). Any references cited in the abstract must be given in full. Unpublished results and personal communications are not recommended in the reference list, but may be mentioned in the text. If thesereferences are included in the reference list they should follow the standard reference style of the journal and should include a substitution of the publication date with either 'Unpublished results' or 'Personal communication'. Citation of a reference as 'in press' implies that the item has been accepted for publication.

Reference links

Increased discoverability of research and high quality peer review are ensured by online links to the sources cited. In order to allow us to create links to abstracting and indexing services, such as Scopus, CrossRef and PubMed, please ensure that data provided in the references are correct. Pleasenote that incorrect surnames, journal/book titles, publication year and pagination may prevent link creation. When copying references, please be careful as they may already contain errors. Use of theDOI is highly encouraged.

A DOI is guaranteed never to change, so you can use it as a permanent link to any electronic article. An example of a citation using DOI for an article not yet in an issue is: VanDecar J.C., Russo R.M., James D.E., Ambeh W.B., Franke M. (2003). Aseismic continuation of the Lesser Antilles slab beneath northeastern Venezuela. Journal of Geophysical Research, https://doi.org/10.1029/2001JB000884. Please note the format of such citations should be in the same style as all other references in the paper.

Web references

As a minimum, the full URL should be given and the date when the reference was last accessed. Anyfurther information, if known (DOI, author names, dates, reference to a source publication, etc.), should also be given. Web references can be listed separately (e.g., after the reference list) under adifferent heading if desired, or can be included in the reference list.

Data references

This journal encourages you to cite underlying or relevant datasets in your manuscript by citing them in your text and including a data reference in your Reference List. Data references should include the following elements: author name(s), dataset title, data repository, version (where available), year, and global persistent identifier. Add [dataset] immediately before the reference so we can properly identify it as a data reference. The [dataset] identifier will not appear in your published article.

References in a special issue

Please ensure that the words 'this issue' are added to any references in the list (and any citations inthe text) to other articles in the same Special Issue.

Reference style

Text: Indicate references by (consecutive) superscript arabic numerals in the order in which they appear in the text. The numerals are to be used *outside* periods and commas, *inside* colons and semicolons. For further detail and examples you are referred to the AMA Manual of Style, A Guide for Authors and Editors, Tenth Edition, ISBN 0-978-0-19-517633-9.

List: Number the references in the list in the order in which they appear in the text. *Examples:*

Reference to a journal publication:

1. Van der Geer J, Hanraads JAJ, Lupton RA. The art of writing a scientific article. *J Sci Commun.*

2010;163:51-59.

https://doi.org/10.1016/j.Sc.2010.00372.Reference to a journal publication with an article number:

- 2. Van der Geer J, Hanraads JAJ, Lupton RA. The art of writing a scientific article. *Heliyon*.2018;19:e00205. https://doi.org/10.1016/j.heliyon.2018.e00205. Reference to a book:
- 3. Strunk W Jr, White EB. *The Elements of Style.* 4th ed. New York, NY: Longman; 2000.Reference to a chapter in an edited book:
- 4. Mettam GR, Adams LB. How to prepare an electronic version of your article. In: Jones BS, SmithRZ, eds. *Introduction to the Electronic Age.* New York, NY: E-Publishing Inc; 2009:281–304. Reference to a website:
- 5. Cancer Research UK. Cancer statistics reports for the UK. http://www.cancerresearchuk.org/aboutcancer/statistics/cancerstatsreport/; 2003 Accessed 13 March 2003.

Reference to a dataset:

[dataset] 6. Oguro, M, Imahiro, S, Saito, S, Nakashizuka, T. Mortality data for Japanese oak wilt disease and surrounding forest compositions, Mendeley Data, v1; 2015. https://doi.org/10.17632/xwj98nb39r.1.

Journal abbreviations source

Journal names should be abbreviated according to the List of Title Word Abbreviations.