# UNIVERSIDADE FEDERAL DO RIO GRANDE DO SUL ESCOLA DE EDUCAÇÃO FÍSICA, FISIOTERAPIA E DANÇA PROGRAMA DE PÓS-GRADUAÇÃO EM CIÊNCIAS DO MOVIMENTO HUMANO

PATRICIA FREITAS DOS SANTOS

# EFFECTS OF CONCENTRIC, ECENTRIC AND CONVENTIONAL FORCE TRAINING OF THE LOWER LIMBS ON MUSCLE ADAPTATIONS AND FUNCTIONAL CAPACITY IN ELDERLY ADULTS: A SYSTEMATIC REVIEW WITH META-ANALYSIS OF RANDOMIZED CLINICAL TRIALS

PORTO ALEGRE - RS 2021

### PATRICIA FREITAS DOS SANTOS

# EFFECTS OF CONCENTRIC, ECENTRIC AND CONVENTIONAL FORCE TRAINING OF THE LOWER LIMBS ON MUSCLE ADAPTATIONS AND FUNCTIONAL CAPACITY IN ELDERLY ADULTS: A SYSTEMATIC REVIEW WITH META-ANALYSIS OF RANDOMIZED CLINICAL TRIALS

Master's Dissertation presented to the Graduate Program in Human Movement Sciences of the School of Physical Education, Physiotherapy and Dance of the Federal University of Rio Grande do Sul, as a final requisite for obtaining the title of Master's in Human Movement Sciences.

Supervisor: Prof. Dr. Marco Aurélio Vaz

### **PORTO ALEGRE - RS**

2021

CIP - Catalogação na Publicação

Freitas dos Santos, Patricia EFFECTS OF CONCENTRIC, ECENTRIC AND CONVENTIONAL FORCE TRAINING OF THE LOWER LIMBS ON MUSCLE ADAPTATIONS AND FUNCTIONAL CAPACITY IN ELDERLY ADULTS: A SYSTEMATIC REVIEW WITH META-ANALYSIS OF RANDOMIZED CLINICAL TRIALS / Patricia Freitas dos Santos. --2021. 81 f. Orientador: Marco Aurélio Vaz. Dissertação (Mestrado) -- Universidade Federal do Rio Grande do Sul, Escola de Educação Física, Programa de Pós-Graduação em Ciências do Movimento Humano, Porto Alegre, BR-RS, 2021. 1. Older. 2. Strength training. 3. Functionality. I. Aurélio Vaz, Marco, orient. II. Título.

Elaborada pelo Sistema de Geração Automática de Ficha Catalográfica da UFRGS com os dados fornecidos pelo(a) autor(a).

### PATRICIA FREITAS DOS SANTOS

# EFFECTS OF CONCENTRIC, ECENTRIC AND CONVENTIONAL FORCE TRAINING OF THE LOWER LIMBS ON MUSCLE ADAPTATIONS AND FUNCTIONAL CAPACITY IN ELDERLY ADULTS: A SYSTEMATIC REVIEW WITH META-ANALYSIS OF RANDOMIZED CLINICAL TRIALS

Final Degree.....

Approved in .....

## **EXAMINATION BOARD**

Prof. Dr. Rafael Reimann Baptista Pontifícia Universidade Católica do Rio Grande do Sul (PUCRS)

Prof. Dr. Eduardo Lusa Cadore Universidade Federal do Rio Grande do Sul (UFRGS)

Prof. Bruno Manfredini Baroni Universidade Federal de Ciências da Saúde e Porto Alegre (UFCSPA)

> Supervisor Prof. Dr. Marco Aurélio Vaz Universidade Federal do Rio Grande do Sul (UFRGS)

### AGRADECIMENTOS

- Ao meu orientador Marco Vaz, pela oportunidade de crescimento pessoal e profissional. Obrigada por acreditar e me guiar por essa experiência única que foi o mestrado. Obrigada por me mostrar luz quando as possibilidades pareciam estar esgotadas. Obrigada pela paciência e compreensão com uma pessoa que tinha pouca experiência na área de pesquisa. És uma pessoa incrível.
- Ao grupo GPBIC Klauber, Emmanuel, Isabel, Francesca, Daniele, Rodrigo, Rodrigo, Kelly, Eliane – obrigada por terem me recebido tão calorosamente. Vocês fizeram e fazem a diferença na minha vida, tornaram os dias difíceis mais amigáveis. Obrigada!
- Aos professores que fizeram toda a diferença nesse percurso. Vocês foram responsáveis pelo meu amadurecimento profissional e certamente um pouco de cada um eu levarei comigo.
- A equipe Physique Anna, Anete, Pauline, Alisson, Lucas, Bruno, Claudia, Carolina, Luciane, Juliana, Janaína - obrigada por todo apoio e compreensão durante esse período intenso. Um Obrigada especial a Viviane Frasson, que me incentivou e me apoiou a encarar essa jornada, sempre incentivando meu crescimento pessoal e profissional.
- A Isabel Paz, que me ajudou diretamente com a dissertação, ouvindo meus lamentos e minhas felicidades. Obrigada por se fazer tão presente nesse momento. Um agradecimento especial também ao Klauber Pompeo e Emmanuel Rocha que sempre estiveram disponíveis a me ensinar.
- Aos meus amigos, que sempre estiveram presentes e entenderam muitas vezes as minhas ausências e angustias. Obrigada por sempre se preocuparem comigo e não desistirem de mim. Obrigada em especial a Francine e ao Cleison que sempre se preocuparam e me apoiaram incondicionalmente. A Anna e a Juliana que entendiam exatamente meus sentimentos e estavam ali como um ombro amigo.
- Aos meus familiares pelo apoio imensurável que vocês me deram nesse período. Aos meus pais que sempre incentivaram meu crescimento, que me proporcionaram estudo e acreditaram nas oportunidades as quais eu queria encarar. Ao meu irmão sempre disponível com uma carona, uma ajuda ou outra em momentos de aperto.
- A família "Silva Martins" a qual me deu apoio e suporte inúmeras vezes. Obrigada também por entenderem muitas das minhas ausências. Obrigada por acolherem como família.

• E obrigada ao Rafael Martins, meu companheiro de todas as horas. Obrigada pela paciência gigante, pelo companheirismo, por fazer eu acreditar em mim mesma a todo momento, principalmente nos momentos mais desesperadores. Obrigada por estar ao meu lado, com certeza fizeste a diferença na minha vida.

"There's nothing you can do That can't be done Nothing you can sing that can't be sung Nothing you can say But you can learn how to play the game"

- The Beatles-

## LISTA DE FIGURAS

FIGURA 1.	Flowchart for identification and selection of articles for final inclusion22
FIGURA 2.	
	<i>Figura 2.1.1</i> Isometric torque
	<i>Figura 2.1.2</i> Concentric slow
	<i>Figura 2.1.3</i> Concentric moderate
	<i>Figura 2.1.4</i> Concentric high
	<i>Figura 2.1. 5</i> Eccentric torque31
FIGURA 3.	
	<i>Figura 3.1.1</i> Knee extensors
	<i>Figura 3.1.2</i> Leg press
	<i>Figura 3.1.3</i> Leg press eccentric only
	<i>Figura 3.1.4</i> Leg press eccentric emphasis
FIGURA 4.	
	<i>Figura 4.1.1</i> Vastus lateralis pennation angle35
	<i>Figura 4.1.2</i> Muscle thickness
	<i>Figura 4.1.3</i> Fascicle length35
FIGURA 5.	
	<i>Figura 5.1.1</i> TUG
	Figura 5.1.2 5-repetition Sit-to-Stand
	<i>Figura 5.1.3</i> 5-repetition Sit-to-Stand eccentric emphasis
	<i>Figura 5.1.4</i> 6min-Walk-Test
	<i>Figura 5.1.5</i> 6min-Walk-Test eccentric emphasis

# LISTA DE TABELAS

<b>TABELA 1.</b> Literature search strategy used for the PUBMED database	51
<b>TABELA 2.</b> Characteristics of the included studies	23
<b>TABELA 3.</b> Risk of bias of the included studies	26
<b>TABELA 4.</b> Quality of evidence using the GRADE approach	29
<b>TABELA 1S.</b> Muscle function results	52
<b>TABELA 2S.</b> Muscle function results	53
<b>TABELA 3S.</b> Muscle structure and activation results	58
<b>TABELA 4S.</b> Functional performance results.	60

# LISTA DE ABREVIAÇÕES

CON: Concentric ECC: Eccentric CONV: Conventional RFD: Rate of Force Development GRADE: Grading of Recommendations Assessment SD: Standard Deviation ISOM: Isometric M: Male F: Female ISOK: Isokinetic Dynamometer RM: Maximum Repetition CSA: Cross-Sectional Area RTD: Rate of Torque Development QUAD: Quadriceps Muscle

## APRESENTAÇÃO

Durante o processo de envelhecimento ocorrem diversas mudanças a nível estrutural do corpo humano; dentre elas, estão as alterações musculoesqueléticas que, consequentemente, levam a alterações de funcionalidade dessa população. Essas alterações de funcionalidade são avaliadas por testes, como por exemplo, sentar e levantar de uma cadeira, velocidade de caminhada, testes que avaliam a mobilidade, entre outros. É bem estabelecido, na literatura, que o treinamento de força é eficaz para melhorar parâmetros de funcionalidade, e de estrutura, ativação e função musculares.

Dentre as diferentes modalidades do treinamento de força, existem os treinos com foco nas contrações concêntrica e excêntrica e o treino chamado convencional (contrações concêntricas-excêntricas). Já é destaque na literatura específica da área que diferentes tipos de contração muscular geram diferentes adaptações musculares tanto em populações mais jovens quanto em idosos. Mas ainda há uma carência na literatura de estudos que nos possibilitem chegar a um melhor entendimento se algum tipo treinamento é superior aos demais para melhora dos parâmetros musculoesqueléticos e de desempenho na funcionalidade em idosos.

Essa dissertação possuía como objetivo inicial aplicar os diferentes tipos de treinamento de força (concêntrico, excêntrico, concêntrico-excêntrico), no dinamômetro isocinético, nos membros inferiores de idosos saudáveis, bem como, realizar uma avaliação completa para identificar as adaptações de funcionalidade, função, ativação e estrutura muscular nessa população. Porém, com o avanço da pandemia a nível mundial, com laboratórios e universidades fechadas, e por se tratar de uma população de risco, não foi possível desenvolver o estudo original conforme havíamos planejado.

Em função disso, optamos por realizar uma revisão sistemática com meta-análise sobre o mesmo tema do estudo original, visando identificar na literatura já existente os efeitos dos diferentes tipos ou modalidades de treinamento de força nas adaptações citadas acima. Esse estudo difere da proposta original que foi qualificada apenas por incluir o treinamento de força tanto em aparelhos de musculação quanto em dinamômetro isocinético, pois assim é possível fazer um escaneamento completo dos estudos existentes com esse tema, de modo a tornar o presente estudo mais aplicado à prática clínica.

#### PREFACE

During the aging process, several changes occur at the structural level of the human body, among which are the musculoskeletal changes that, consequently, lead to changes in the elderly's functionality. These changes in functionality are measured by tests, such as sitting on and standing from a chair, walking speed, tests that assess mobility, among others. It is well established in the literature that strength training is effective for improving parameters of functionality, and muscle structure, activation and function.

Among the different strength training modalities, there are the training programs focused on concentric contractions, eccentric contractions and conventional training (concentric-eccentric contractions). It is already highlighted in the literature that different types of muscle contraction generate different muscle adaptations in both younger and elderly populations. However, there is still lack in the literature of a study that thoroughly reviewed the existent studies aimed at gaining a better understanding of whether any type of training is superior to the others to improve musculoskeletal parameters and functionality performance in older people.

This dissertation had, as its initial objective, identifying the adaptations in functionality, and muscle structure, activation and function, while applying the different types of strength training in the isokinetic dynamometer (concentric, eccentric, concentric-eccentric), in the lower limbs of healthy elderly people. However, with the advance of the pandemic worldwide, with laboratories and universities closed, and because the elderly are a population at risk, it was not possible to execute the original study approved during the candidacy exam.

Therefore, we decided to conduct a systematic review with meta-analysis on the same topic of the original study, aiming to identify the adaptations mentioned above in the existing literature. This study differs from the original approved study only by the fact that strength training was included both in weight training machines and in the isokinetic dynamometer, as it is possible to do a complete scan of the existing studies on this topic, thereby making the study more applied to clinical practice.

SUMMARY	
LISTA DE FIGURAS	7
LISTA DE TABELAS	8
LISTA DE ABREVIAÇÕES	9
APRESENTAÇÃO	
CAPÍTULO 1	
EFFECTS OF CONCENTRIC, ECENTRIC AND CON OF THE LOWER LIMBS ON MUSCLE ADAPTATIONS IN ELDERLY ADULTS: A SYSTEMATIC REVIEW RANDOMIZED CLINICAL TRIAL	S AND FUNCTIONAL CAPACITY W WITH META-ANALYSIS OF
ABSTRACT	
1. Introduction	
2. Methods	
2.1 Eligibility criteria	
2.2 Search Strategy	
2.3 Study Selection	
2.4 Data Extraction	
2.5 Risk of Bias Assessment	
2.6 Data Analysis	
2.7 Summary of Evidence	
3. Results	
3.1 Study Selection	
Screening	
Included	
Eligibility	
Identification	
3.2 Studies Methodological Quality	
3.3 Intervention Effects	
3.3.1 Muscle function	
3.3.2 Muscle structure	
3.3.3 Performance in functional tests	
4. Discussion	
5. Conclusion	

6.	Study limitations	.43
7.	References	44
Sup	olmentary material	47
8.	General conclusion	62
An	ехо	63

# EFFECTS OF CONCENTRIC, ECENTRIC AND CONVENTIONAL FORCE TRAINING OF THE LOWER LIMBS ON MUSCLE ADAPTATIONS AND FUNCTIONAL CAPACITY IN ELDERLY ADULTS: A SYSTEMATIC REVIEW WITH META-ANALYSIS OF RANDOMIZED CLINICAL TRIALS

Patrícia Freitas dos Santos, BSc<sup>a,b</sup>, Isabel de Almeida Paz, MSc<sup>a</sup>, Marco Aurélio Vaz, PhD<sup>a,b</sup>

<sup>a</sup> Biomechanics and Kinesiology Research Group, Exercise Research Laboratory, Physical Education, Physiotherapy and Dance School, Federal University of Rio Grande do Sul, Porto Alegre, RS, Brazil
<sup>b</sup> Physique Centro de Fisioterapia, Porto Alegre, RS, Brazil

\*Corresponding author:

Marco Aurélio Vaz

Biomechanics and Kinesiology Research Group, Exercise Research Laboratory, Federal University of Rio Grande do Sul, Felizardo Street, 750, Porto Alegre, RS, Brazil – Postal Code 90690-200.

Email address: <u>marco.vaz@ufrgs.br</u> Phone: +55-51-993851188

### ABSTRACT

Introduction: The current increase in life expectancy of the world population has brought an increase in adaptations due to aging (e.g., neuronal losses, decreased muscle strength, decreased muscle structure and decreased functional capacity). Strength training is an effective measure to improve these lost results due to aging. However, there is still no consensus on which type of training, eccentric (ECC), concentric (CON) or conventional (CONV), is superior in reducing these deleterious aging effects. Methods: Randomized controlled trials (RCTs) that used strength training (CONV, ECC or CON) in healthy elderly were included. Searches were performed in the following databases: Pubmed, PEDro, Cochrane Central, Embase and Web of Science. This systematic review by meta-analysis was based on the recommendations of the Cochrane Collaboration and PRISMA. The main results and parameters of muscle function, muscle adaptations and functionality tests were evaluated by two independent reviewers. For each result, the mean and standard deviation values and the number of participants from both groups were extracted. Qualitative and quantitative analyzes of the included studies were performed. Results: Nine studies were included, presenting a high risk of study bias. Metaanalyses revealed no difference between studies, presenting a very low quality of evidence. While analyzing studies qualitatively and the effect sizes, there was a small advantage in the evaluated parameters for the ECC training group. Conclusion: Similar improvement was observed in all training modalities in healthy older adults, although a qualitative analysis seems to favor ECC strength training. However, a larger number of high-quality randomized controlled trials is needed to confirm this hypothesis.

Keywords: Strength training, older, functionality.

#### 1. Introduction

Remarkably, the world is in a demographic historic mark in which the number of people over 60 years old will practically double until 2050, going from 12% to 22% of the world's population (OPAS, 2018). Increase in life expectancy shows that the population aging will continue. In 2019, there were 703 million people with 65 years of age or more at the global population, and this number should double to 1,5 billion elderly in 2050 (NIA, 2011; UNITED NATIONS, 2019). This increase in life expectancy is often accompanied by structural and functional losses in the neuromuscular system, (JANSSEN et al., 2000; REEVES et al., 2006; PERKISAS et al, 2016), which lead to disability (JANSSEN et al., 2000; REEVES et al., 2006; PERKISAS et al, 2016, HUNTER et al., 2016), decreased functionality (BORZUOLA et al., 2020) and increased mortality (BERGLAND et al., 2017).

With aging, neural losses related neuronal death naturally occur, leading to an atrophy process due to muscle fibers' denervation due to motoneurons death. Consequently, there is a reduction in the number of motor units and an increase in the motor units' size due to muscle fibers re-innervation by the remaining motoneurons (GONZALEZ-FREIRE et al., 2014). As a higher proportion of slow-twitch fibers (type I) compared to fast-twitch fibers (type II) were observed in the vastus lateralis muscle of older people with walking ability (NARO et al., 2019), it appears that there is a selective loss of large motoneurons that innervate the fast-twitch muscle fibers with aging. This preferential loss of fast muscle fibers may be responsible by the muscle mass loss or sarcopenia observed with aging (NARO et al., 2019), as well as the elders' slower movement capacity. These muscular structural losses have been described as parallel muscle hypotrophy (i.e., reduction in the number of myofibrils per muscle fiber) and serial muscle hypotrophy (i.e., reduction in the myofibril length due to sarcomere loss). These muscle fibers' structural losses determine a decrease in the fascicles' pennation angle, in fascicle length and in the muscle's cross-sectional area (CSA), leading to deleterious changes in the force-length and force-velocity relationships (KAWAKAMI; ABE; FUKUNAGA, 1993; KUBO et al., 2003; NARICI et al., 2003).

These neuromuscular changes due to aging reduce the activation capacity of the contractile system, as well as the capacity of force generation (GONZALEZ-FREIRE et al., 2014). Therefore, a decrease in maximum strength, in the rate of force development (RFD), and in maximum power is commonly observed in older people (HUNTER et al., 2016). These

outcomes are directly related to older peoples' functional capacity, featuring slowness in daily life activities (MACALUSO; DE VITO, 2004), slower and/or less responsiveness to environmental stimuli (HUNTER et al., 2016), increasing risk of falls and decreasing quality of life (TROMBETTI et al., 2016; VON HAEHLING; MORLEY; ANKER, 2010).

The structure and function of the knee extensor and flexor muscles play important roles in older peoples' functionality. As previously shown, the vastus lateralis muscle quality and thickness are related to older peoples' walking speed (GUADAGNING et al., 2019). Additionally, the peak torque and power of the knee extensor and flexor muscles have shown a significant relationship with the older peoples' functional performance regarding lower limb mobility, balance and strength tests (DE MOURA et al., 2019). Therefore, older people may benefit from stimuli involving the lower limbs' neuromuscular system, mainly related to the knee extensor and flexor muscles.

It is well known that strength training leads to structural and functional gains, improving neural (i.e., increase in muscle activation and greater motor unit synchronization) and muscle factors (i.e., muscle strength parameters, muscle power, RFD, increase in myofibrils per muscle fiber and in fascicle length) (GRANACHER et al., 2011; LEE et al., 2017). However, the exercise type can influence differently the neuromuscular outcomes, as the specificity training principle indicates that different adaptions occur with different types of training. For example, exercises with an emphasis in concentric contractions (CON) lead to a greater muscle activation and greater parallel hypertrophy compared to eccentric (ECC) contractions (AAGAARD et al., 2000; AMIRIDIS; et al., 1996; WISDOM; DELP; KUHL, 2015). Exercises focused in ECC contractions, in the other hand, lead to muscle fiber length increase, that is, greater serial hypertrophy than CON (FRANCHI et al., 2014, 2015; REEVES et al., 2009; TIMMINS et al., 2016).

However, the literature comparing these two strength training types (CON vs. ECC) and the conventional training (CONV – training involving CON and ECC contractions) is unclear, and it is not possible to establish which of the three modalities the best one for older people is. Evidences have shown similar results between CONV and ECC training for muscle architecture (RAJ et al., 2012; VÁCZI et al., 2014), knee extensors' force (DIAS et al., 2015; VÁCZI et al., 2014; GLUCHOWSKI et al., 2017) and functional tests' performance (RAJ et al., 2012; DIAS et al., 2015; GLUCHOWSKI et al., 2017). However, other studies demonstrated that CONV is capable of generating greater improvement in muscle architecture and knee extensors' force (REEVES et al., 2009) compared to ECC, while ECC training has demonstrated similar effects for knee extensors' force (CHEN et al., 2017) or is more efficient for improving functional performance compared to CON training (CHEN et al., 2017). In order to summarize the data, a recently published review (MOLINARI et al., 2019) compared the effects of the ECC versus CONV training in muscle strength, and the meta-analysis revealed no difference between them. However, there are some limitations regarding the methodological quality of this review. Among these limitations, they included different types of training (power training, high and low intensity training) on the same meta-analysis (MOLINARI et al., 2019), they did not control for training intensity, and they included only studies that measured muscle strength through the maximal repetition (1RM) assessment. Other neuromuscular parameters related to muscle structure and function, as well as performance in functional tests, were not investigated. Additionally, they did not assess the quality of the evidence (GRADE), which reflects the confidence that the effect estimates are correct and are adequate to support a specific recommendation (BALSHEM et al., 2011). Furthermore, we were unable to find a review that encompassed the main neuromuscular and functional parameters that are affected by aging, including the training effects on fascicle length and pennation angle. These limitations compromise clinical decision-making regarding the choice of adequate exercise during the development of a training program for the elderly.

Therefore, for better understanding this subject and filling the literature gaps, we developed a systematic review of high methodological quality, following the Cochrane recommendations, in order to evaluate the effectiveness of different types of muscle training (CON, ECC and CONV) at the neuromuscular and functional parameters in older people, through a systematic review and meta-analysis of randomized clinical trials.

#### 2. Methods

This systematic review and meta-analysis followed the recommendations proposed by the Cochrane Collaboration (HIGGINS; GREEN, 2011) and the PRISMA Declaration (LIBERATI et al., 2009), and was registered in the PROSPERO (CRD42020175489).

#### 2.1 Eligibility criteria

This review included randomized controlled trials that investigated the effects of CON, ECC and CONV resistance training for the knee flexor and/or extensor muscles on neuromuscular parameters in older people. The interventions must had assessed the longitudinal effects on the reported results. The studies exclusion criteria were: inclusion of associated diseases; studies using medications or supplementation associated with the protocol; inclusion of people under 60 years of age; high-speed training; exercises with elastic band; using only body weight as exercise load; assessing exercise intensity by subjective scales; studies that compared CONV, CON or ECC to a control group; and strength training associated with other modalities.

#### 2.2 Search Strategy

The searches were carried out in the following electronic databases: Pubmed, PEDro, Cochrane Central, Embase and Web of Science. The date and language of publication were not limited. Controlled and uncontrolled terms were used for population, intervention and type of study. To establish the study type, previously proposed words were used to identify randomized clinical trials (ROBINSON; DICKERSIN, 2002). The search was performed on January 2020. The full search strategy used in PubMed is shown in Table 1S (Supplementary Material), and the search strategies used in the remaining databases are available upon request.

#### 2.3 Study Selection

Two reviewers (PFS and IAP) independently assessed the titles and abstracts of the studies identified in the search strategy. Duplicate studies were excluded. Titles and abstracts were analyzed to select possible studies to be included in the review, according to the eligibility criteria. Next, two researchers (PFS and IAP) independently evaluated the full-text studies. In this phase, the reviewers also adopted the eligibility criteria. Disagreements were discussed and resolved by consensus or by a third reviewer (MAV).

#### 2.4 Data Extraction

Two reviewers (PFS and IAP) performed data extraction independently to obtain the methodological characteristics and results of the studies. When necessary, the authors discussed disagreements, which were resolved by consensus. When consensus was not reached, disagreements were discussed with and resolved by a third reviewer (MAV). The primary outcome was isometric and dynamic muscle strength, and the secondary outcomes were power; rate of torque or force development; muscle activation; muscle quality; muscle architecture (pennation angle, muscle thickness, fascicle length) and functionality (time-up-and-go test - TUG, 6 min walk test, 5 rep sit-to-stand, 30 sec chair stand, stair climbing, stair descent).

#### 2.5 Risk of Bias Assessment

The studies' quality evaluation was carried out independently by two evaluators (PFS and IAP), using the Cochrane Bias Risk tool (HIGGINS; GREEN, 2011). This tool evaluates the following items: adequate sequence generation, allocation concealment, blinding of patients, blinding of outcome assessors, use of intention-to-treat analysis, and description of losses and exclusions. An analysis of the intention to treat was considered only if studies reported (through tables or text) that the number of randomized participants was equal to the number analyzed. If there was no clear description of these items, they were considered unclear.

#### 2.6 Data Analysis

Qualitative and quantitative data analyzes were performed. For the qualitative analysis, the main characteristics and results of the included studies were presented and discussed. For the quantitative analysis, the pre- to post-training percentage change, the effect sizes and metaanalyzes were determined.

Effect size was calculated by Cohen's Equation (1988) (COHEN, 1988) using the website https://lbecker.uccs.edu/, by determining the mean difference between pre and post values in each group (within-group effect size) and the between-groups mean difference only between post-intervention values (between-group effect size), and then dividing the result by the pooled standard deviation. Calculated effect sizes were categorized as trivial (<0.20), small (0.20-0.49), moderate (0.50-0.79), large (0.80 to 1.29), and very large (>1.30) effect (COHEN,

1988; ROSENTHAL, 1996). In addition, the calculation of the mean relative change between the pre- and post-training was performed for each group.

A meta-analysis was performed for each outcome. In addition to the authors and year of the studies, the mean, standard deviation, and the number of subjects were extracted. The values, extracted from the results, were pre and post mean intervention values, as well as the pre and post standard deviation (SD). As the baseline values were different for some studies, delta values - i.e., the difference between the final value (post) and the initial value (pre) - of each group's result were used. The SD of this value (delta SD) was also calculated according to Cochrane recommendations (HIGGINS; GREEN, 2011) and both were used in the metaanalysis. The calculations in the meta-analysis were performed using a random effects method. A value of p <0.05 was considered statistically significant. The statistical heterogeneity of the effects of the intervention between studies was assessed using the inconsistency test (I2), in which values less than 25% were considered indicative of low heterogeneity, between 25% and 50% were considered indicators of moderate heterogeneity and above 50% were considered to be highly heterogeneous (HIGGINS et al., 2003). All analyzes were performed using the Review Manager software, version 5.3 (HIGGINS; GREEN, 2011). To explore the heterogeneity between the studies, perform subgroup analyzes, considering the types of training performed, if performed only in an eccentric manner or with an emphasis on eccentric contraction.

Due to the different training protocols among the reviewed studies, and to facilitate the results' synthesis, we chose to separate the studies' strength training groups into four groups. Conventional training (CONV) performed exercises with both ECC and CON phases (with similar times in both phases); CON training performed only exercises with CON phase; ECC-only training performed only ECC muscle action during exercises; ECC-emphasis training performed exercises with both ECC and CON phases, but with emphasis in the time or duration of the ECC phase.

## 2.7 Summary of Evidence

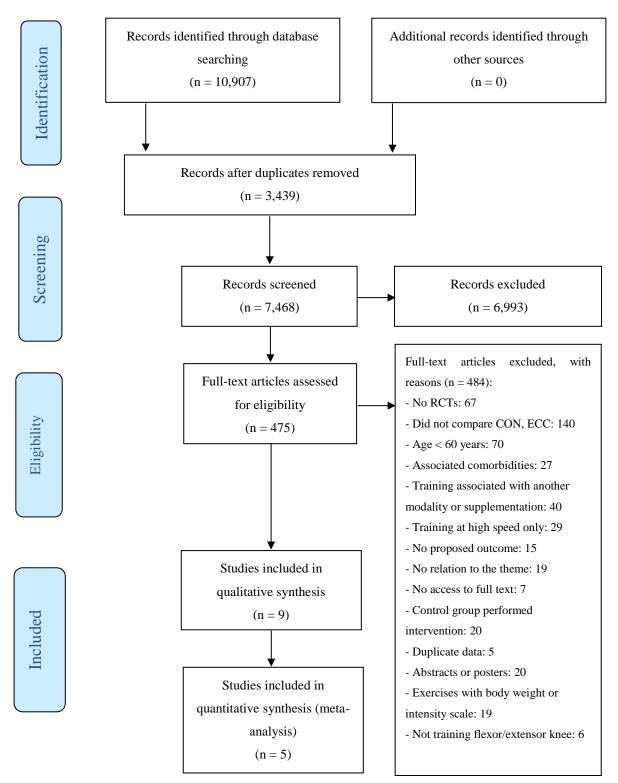
The quality of the evidence was carried out independently by two evaluators (PFS and IAP) using the Grading of Recommendations Assessment, Development and Evaluation (GRADE) (SCHÜNEMANN et al., 2008; HIGGINS; GREEN, 2011). For each presented

result, the quality of the evidence was based on the following factors: (1) risk of bias; (2) inconsistency; (3) indirectness; (4) imprecision; and (5) other considerations (publication bias). The GRADE procedure resulted in four levels of quality of evidence: high, moderate, low, and very low, which are defined according to the abovementioned factors and are applied to a body of evidence (BALSHEM et al., 2011).

#### 3. Results

#### 3.1 Study Selection

Using the search protocol, 10,907 articles (database) were identified. Four hundred and seventy-five of them were identified as eligible (read in full). After applying the inclusion criteria, 9 studies were selected for this review and included in the qualitative analysis, and 5 were included in the meta-analysis (Figure 1, Table 2).



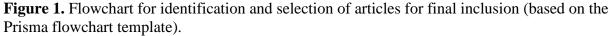


Table 2 – Characteristics	of the included studies.
---------------------------	--------------------------

Study	Sample	Mean age	Comparison	Training	Intensity	Sets/exercise	Time or	Rest	Total	Frequency	Intervention	Main Outcomes
	(gender)	(years)	Training -	device		Reps per set	velocity	between	time per		time in weeks	
			Туре					sets	session		or months	
Chen et al.,	26 (M)	65.9±4.7	ECC-only vs.	Weight	ECC: 10-	3 x 10	3 s	3-min	30-60 min	1x/week	12 weeks	*Knee extensor's maximal
2017			CON	machine	100% of 1RM	6 x 10						ISOM torque
					CON: 50-							*Knee extensor's maximal
					100% of 1RM							CON torque
												*Knee extensor's RM
												*Timed up-and-go test
												*6-m walk test
												*30-s chair stand
Dias et al.,	26 (F)	67±6	ECC-	Weight	45% to 70%	2 x 12	ECC: 1.5s	2-3 min	NI	2x/week	12 weeks	*Knee extensor's RM
2015			emphasis vs.	machine	1RM	2 x 10	and 4.5s					*Leg press' RM
			CONV			3 x 8	CONV:1.5/					*6-m walk test
							1.5s					*Timed up-and-go test
												*Stair-climbing test
												*Chair-rising test
Gluchowski	33 (21F;	67±4.5	ECC-only vs.	Weight	70% of 1RM	4 x 10	CONV and	60 s	NI	2x/week	8 weeks	*Leg press' RM
et al., 2017	12M)		ECC-	machine			ECC-					*6-m walk test
			emphasis vs.				emphasis:					*5 rep sit-to-stand
			CONV				2/1/2s;					*Stair-descent test
							ECC-only:					
							2-1					
Kang et al.,	22 (11F;	ECC:	ECC vs. CON	NK table	70% of the	3 x 10	ECC and	5 min	40 min	3x/week	4 weeks	*Surface electromyography
2016	11M)	67.1±1.8			1RM		CON: 5s					
		CON:68.2										
		±1.4										

Malliou et	52 (26F;	CONV: M:	CON vs.	ISOK &	CONV: 90%	CON: 9 x 12	150°/s and	2 min	45-55 min	3x/week	10 weeks	*Knee extensor's maximal
al., 2003	26F)	70.7±2.5;	CONV	Weight	of the 1RM	CONV: 3 x 12	180/s					CON torque
		F: 66±5.5		machine			CONV: 2/2-					
		CON: M					3s					
		69.7±2.2;										
		F: 68±5.1										
Raj et al.,	28 (11F;	68±5	ECC-	Weight	CONV: 75%	CONV: 2 x 10	NI	3 min	NI	2x/week	16 weeks	*Leg press' maximum
2012	17M)		emphasis vs.	machine	of 1RM;	ECC: 3 x 10						repetition
			CONV		ECC: 50% of	and 3 x 5						*Knee extensor's maximal
					the 1RM							ISOM torque
												*Knee extensor's maximal
												CON torque
												*Muscle architecture (VL
												fascicle pennation angle,
												fascicle length, and muscle
												thickness)
												*Timed up-and-go test
												*6-m walk test

Reeves et	19 (10F;	CONV:	ECC-only vs.	Weight	$\sim 80\%$ of the	2 x 10	CONV: ~2-	NI	NI	3x/week	14 weeks	*Leg press' RM
al., 2009	9M)	74±3	CONV	machine	5RM		3s					*Knee extensor's RM
		ECC: 67±2					ECC: ~3s					*Knee extensor's maximal
												ISOM torque
												*Knee extensor's maximal
												CON torque
												*Knee extensor's maximal
												ECC torque
												*Muscle architecture (VL
												fascicle pennation angle,
												fascicle length, and muscle
												thickness)
Symons et	37 (19F;	CON:	CON vs. ECC	ISOK	Maximal	3 x 10	NI	2 min	NI	3x/week	12 weeks	*Maximal CON torque
al., 2005	18M)	71.8±3.1;			ISOM &							*Maximal ECC torque
		ECC:			ISOK							
		$70.5 \pm 5.2$			resistance							
					training							
Váczi et al.,	16 (M)	CONV:	ECC vs.	ISOK	Stretch-load	4 x 8	NI	2 min	NI	2-3x/week	10 weeks	*Knee extensor's maximal
2014		64.4±4.1;	CONV		ranged	4 x 10						CON torque
		ECC:			between 86 J	4 x 12						*Knee extensor's maximal
		65.7±5.3			and 120 J	4 x 12						ECC torque
						4 x 13						*RTD
						4 x 14						*QUAD anatomical CSA

M: male; F: female; CONV: conventional; ECC: eccentric; CON: concentric; ISOK: isokinetic dynamometer; ISOM: isometric; RM: maximum repetition; NI: not informed; CSA: cross-sectional area; RTD: rate of torque development; VL: vastus lateralis; QUAD: quadriceps muscle; NK table: specific brand of exercise chair.

#### 3.2 Studies Methodological Quality

Regarding the studies' methodological quality, 22% described a sequence of random generation; no study described the allocation concealment and blinding of patients, and only one study blinded the evaluator to the outcomes (11%). In addition, only 44% of the studies reported the description of losses and exclusions, and only 22% described the analysis by intention to treat. Overall, a high risk of bias was observed for all analyzed items (Table 3).

Study	Adequate sequence generation	Allocation concealment	Blinding of patients	Blinding of outcome assessors	Description of losses and exclusions	Intention-to- treat analysis	Overall Risk of Bias for each study
Chen et al. 2017	LOW	NI	NI	NI	LOW	NI	HIGH
Dias et al. 2015	LOW	NI	NI	NI	LOW	LOW	HIGH
Gluchowski et al.	UNCLEAR	NI	NI	NI	LOW	LOW	HIGH
2017							
Kang et al. 2016	UNCLEAR	NI	NI	NI	NI	NI	HIGH
Malliou et al. 2003	UNCLEAR	NI	NI	NI	NI	HIGH	HIGH
Raj et al. 2012	UNCLEAR	NI	NI	LOW	NI	NI	HIGH
Reeves et al. 2009	UNCLEAR	NI	NI	NI	NI	NI	HIGH
Symons et al. 2005	UNCLEAR	HIGH	NI	HIGH	LOW	NI	HIGH
Váczi et al. 2014	UNCLEAR	NI	NI	NI	NI	NI	HIGH
Overall Risk of Bias for each criterion	HIGH	HIGH	HIGH	HIGH	HIGH	HIGH	

Table 3 – Risk of bias of the included studies.

NI = not informed.

#### 3.3 Intervention Effects

#### 3.3.1 Muscle function

#### 3.3.1.1 Isometric Torque – ECC-only or ECC-emphasis versus CONV

The knee extensors' isometric torque was compared in three studies (RAJ et al., 2012; REEVES et al., 2009; VÁCZI et al., 2014) that performed ECC versus CONV training, with subjects positioned on the isokinetic dynamometer at 60° and 70° of knee extension. Qualitative analysis showed no differences between groups, and trivial or small between-groups effect size (Table 2S, Supplementary Material). Quantitative analysis (Figure 2.1.1) showed no statistical

difference between the groups, with low heterogeneity. Based on the GRADE approach, the quality of the evidence for this outcome was considered very low (Table 4).

#### 3.3.1.2 Isometric Torque – ECC-only versus CON

Only one study (CHEN et al., 2017) evaluated the effects of ECC-only versus CON on isometric torque. Qualitative analysis showed that ECC-only induced greater increase at the knee extensors' isometric torque compared to CON, and large between-groups effect size (Table 2S, Supplementary Material).

#### 3.3.1.3 Concentric Torque - ECC versus CONV

Two articles (RAJ et al., 2012; REEVES et al., 2009) carried out the knee extensors' concentric torque analysis, comparing ECC versus CONV training at three different angular velocities: slow  $(50^{\circ}.s^{-1} - 60^{\circ}.s^{-1})$ , moderate  $(100^{\circ}.s^{-1} - 120^{\circ}.s^{-1})$  and fast  $(200^{\circ}.s^{-1} - 240^{\circ}.s^{-1})$ . At the qualitative analysis, Reeves et al. (2009) did not report between-groups comparisons, while Raj et al. (2012) showed no difference between training modalities for slow, moderate and fast angular velocities. Both studies showed a trivial or small between-group effect size (Table 2S, Supplementary Material). The meta-analyzes of these comparisons (Figures 2.1.2, 2.1.3, 2.1.4) showed no between-groups difference at any of the angular velocities, and low heterogeneity. Based on the GRADE approach, the quality of the evidence for this outcome was considered low and very low (Table 4).

#### 3.3.1.4 Concentric Torque – CON versus CONV

One study (MALLIOU et al., 2003) compared the CON versus CONV group at the angular velocities of  $60^{\circ}$ .s<sup>-1</sup> and  $180^{\circ}$ .s<sup>-1</sup>. There was a larger increase in CON torque at  $60^{\circ}$ .s<sup>-1</sup> for the CONV compared to CON group (large effect size), while in  $180^{\circ}$ .s<sup>-1</sup> there was no between-groups difference (small effect size) (Table 2S, Supplementary Material).

#### 3.3.1.5 Concentric Torque – ECC-only versus CON

Two articles (CHEN et al., 2017; SYMONS et al., 2005) analyzed the CON torque, but it was not possible to perform a meta-analysis because of the studies' methodological differences. However, in the qualitative analysis (Table 2S, Supplementary Material), Chen et al. (2017) demonstrated greater increase in CON torque for the ECC-only group compared to CON, and a moderate between-groups effect size. Symons et al. (2005) did not find differences between the two groups and demonstrated a trivial effect size when the groups were compared (Table 2S, Supplementary Material).

#### 3.3.1.6 Eccentric Torque - ECC-only versus CONV

Two studies (REEVES et al., 2009; VÁCZI et al., 2014) evaluated ECC torque at the angular velocities of 30°.s<sup>-1</sup> and 50°.s<sup>-1</sup> comparing ECC-only to CONV. In the qualitative analysis, the between-groups difference was not reported by Reeves et al. (2009), while Váczi et al. (2014) demonstrated that the ECC-only induced a greater ECC torque increase compared to CONV. However, the between-groups effect size was trivial (Table 2S, Supplementary Material). In the quantitative analysis (Figure 2.1.5), there was no difference between the groups and the heterogeneity was low.

#### 3.3.1.8 Eccentric Torque – ECC-only versus CON

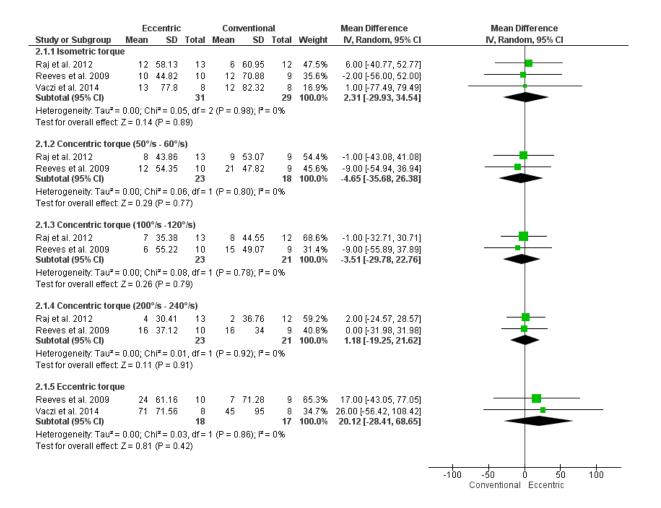
Only one study (SYMONS et al., 2005) evaluated the ECC torque between CON versus ECC-only training. Quantitative analysis (Table 2S, Supplementary Material) showed no difference between the groups, with a small effect size (Table 2S, Supplementary Material).

		(	Certainty assessr	nent		l	N	Absolute	Certainty
Outcome	N (RCTs)	<b>Risk of Bias</b>	Inconsistency	Indirectness	Imprecision	Interv	Comp	(95% CI)	
ISOMETRIC TORQUE									
ECC vs. CONV	3	very serious <sup>a</sup>	not serious	not serious	serious <sup>c</sup>	31	29	2.31 [-29.93 - 34.54]	VERY LOW
CONCENTRIC TORQUE									
ECC vs. CONV at low intensity	2	very serious <sup>a</sup>	not serious	not serious	very serious <sup>d</sup>	23	18	4.65 [-35.68 - 26.38]	VERY LOW
ECC vs. CONV at moderate intensity	2	very serious <sup>a</sup>	not serious	not serious	very serious <sup>d</sup>	23	21	3.51 [-29.78 - 22.76]	VERY LOW
ECC vs. CONV at high intensity	2	very serious <sup>a</sup>	not serious	not serious	serious <sup>c</sup>	23	21	1.18 [-19.25 - 21.62]	VERY LOW
ECCENTRIC TORQUE									
ECC vs. CONV at low intensity	2	very serious <sup>a</sup>	not serious	not serious	very serious <sup>d</sup>	18	17	20.12 [-28.41 - 68.65]	VERY LOW
KNEE EXTENSOR (RM)									
ECC vs. CONV	2	very serious <sup>a</sup>	very serious <sup>b</sup>	not serious	very serious <sup>d</sup>	20	19	12.55 [-16.24 - 41.35]	VERY LOW
LEG PRESS (RM)									
ECC vs. CONV	5	very serious <sup>a</sup>	not serious	not serious	very serious <sup>d</sup>	55	42	7.47 [-14.82 - 29.77]	VERY LOW
SA: ECC-only vs. CONV	2	very serious <sup>a</sup>	very serious <sup>b</sup>	not serious	very serious <sup>d</sup>	21	20	29.33 [-51.20 - 109.86]	VERY LOW
SA: ECC-emphasis vs. CONV	3	very serious <sup>a</sup>	not serious	not serious	not serious	34	33	0.05 [-25.05 - 24.15]	LOW
PENNATION ANGLE									
ECC vs. CONV	2	very serious <sup>a</sup>	very serious <sup>b</sup>	not serious	serious <sup>c</sup>	23	21	1.65 [-7.33 - 4.02]	VERY LOW
THICKNESS									
ECC vs. CONV	2	very serious <sup>a</sup>	not serious	not serious	not serious	23	21	0.05 [-0.30 - 0.41]	LOW
FASCICLE LENGTH									
ECC vs. CONV	2	very serious <sup>a</sup>	not serious	not serious	very serious <sup>d</sup>	23	21	0.93 [-0.30 - 2.17]	VERY LOW
TUG									
ECC vs. CONV	2	very serious <sup>a</sup>	not serious	not serious	not serious	22	23	0.05 [-0.51 - 0.60]	LOW
5 REP SIT-TO-STAND									
ECC vs. CONV	3	very serious <sup>a</sup>	not serious	not serious	not serious	32	21	0.13 [-1.10 - 1.35]	LOW
SA:ECC-emphasis vs. CONV	2	very serious <sup>a</sup>	not serious	not serious	not serious	21	21	0.38 [-0.80 - 1.56]	LOW
6M WALK TEST									

# **Table 4** – Quality of evidence using the GRADE approach.

ECC vs. CONV	4	very serious <sup>a</sup>	not serious	not serious	not serious	45	33	0.03 [-0.26 - 0.21]	LOW
SA:ECC-emphasis vs. CONV	3	very serious <sup>a</sup>	not serious	not serious	not serious	34	33	0.00 [-0.28 - 0.28]	LOW

SA: Subgroup analysis; a: High risk bias (3 or more items); b: High heterogeneity (over 50%); c: Moderate confidence interval (CI); d: Large confidence interval (CI); Comp: Comparison; Interv: intervention



**Figure 2.** Isometric (2.1.1), concentric (2.1.2, 2.1.3, 2.1.4) and eccentric (2.1.4) torques: comparison between ECC and CONV training at isometric (2.1.1), concentric slow (2.1.2), moderate (2.1.3) and high (2.1.4) speeds, and eccentric (2.1.5) contractions.

#### 3.3.1.9 Knee extensor's maximum repetition – ECC-only or ECC-emphasis versus CONV

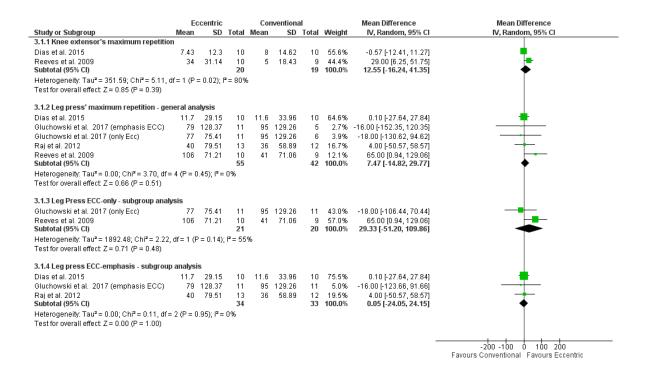
Two studies (DIAS et al., 2015; REEVES et al., 2009) evaluated a maximum repetition in the knee extensor machine comparing the ECC-only (REEVES et al., 2009) or ECCemphasis (DIAS et al., 2015) to CONV training. Qualitative analysis showed divergent results. While Dias et al. (2015) showed no between-groups differences, Reeves et al. (2009) demonstrated that ECC-only induced a greater increase in the knee extensors' maximum strength compared to CONV, with a very large effect size (Table 2S, Supplementary Material). Regarding the quantitative analysis (Figure 3.1.1), there was no between-groups difference with a high heterogeneity (80%). Based on the GRADE approach, the quality of the evidence for this outcome was considered very low (Table 4).

#### 3.3.1.10 Knee extensor's maximum repetition – ECC-only versus CON

Only one study (CHEN et al., 2017) compared the effects of ECC-only versus CON at the knee extensors maximum repetition test. Qualitative analyzes demonstrated a greater increase at the knee extensors maximum repetition for the ECC-only compared to CON, with a large between-groups effect size (Table 2S, Supplementary Material).

#### 3.3.1.11 Leg press maximum repetition - ECC-only or ECC-emphasis versus CONV

Four studies (DIAS et al., 2015; GLUCHOWSKI et al., 2017; RAJ et al., 2012; REEVES et al., 2009) carried out the analysis of the leg press maximum repetition test for the knee extensors, comparing the group that trained ECC-only versus CONV. Qualitative analysis showed contradictory results. While three studies (DIAS et al., 2015; GLUCHOWSKI et al., 2017; RAJ et al., 2012) found no between-groups differences with a small effect size, one study (REEVES et al., 2009) showed a larger increase in maximum repetition for the ECC-only group compared to CONV, with a very large between-groups effect size (Table 2S, Supplementary Material). Considering that only one study (GLUCHOWSKI et al., 2017) performed both ECC-emphasis and ECC-only training compared to CONV, we performed a meta-analysis with the results of all studies, followed by a subgroup analysis comparing CONV to ECC-only or ECC-emphasis (Figure 3.1.2, 3.1.3, 3.1.4). There was no between-groups difference and the heterogeneity was low. Only in the subgroup analysis with ECC-only contraction there was a heterogeneity of 55%. Based on the GRADE approach, the quality of the evidence for this outcome was considered low for ECC-emphasis versus CONV and very low for CONV versus ECC-only (Table 4).



**Figure 3.** Knee extensors (3.1.1) and leg press (3.1.2, 3.1.3, 3.1.4) maximum repetition: comparison between ECC and CONV

#### 3.3.1.10 Power – ECC vs. CON

Only one article (SYMONS et al., 2005) investigated the effects of the ECC versus the CON group on Power. Qualitative analysis showed a greater increase for the group trained CON, with a medium effect size (Table 2S, Supplementary Material) compared to the ECC group, with an insignificant effect size.

#### 3.3.1.11 RTD

Only one article (VÁCZI et al., 2014) evaluated the knee extensors' rate of torque development for times of 30ms, 50ms and 100ms. It was not possible to perform a metaanalysis, but it was possible to calculate the effect size for the three variables. The rate of torque development showed a small effect size for all comparisons (Table 2S, Supplementary Material).

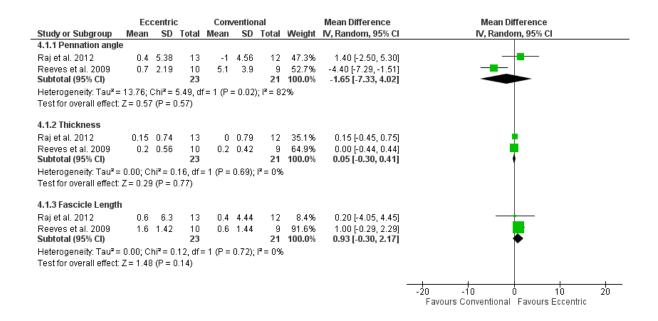
#### 3.3.1.12 Surface electromyography – ECC vs. CON

Only one article (KANG et al., 2016) evaluated muscle activation of the rectus femoris, vastus medialis and vastus lateralis between the ECC and CON training groups. It was not possible to perform a meta-analysis, but it was possible to calculate the effect size for the three variables. Muscle activation for the vastus medialis and vastus lateralis muscles showed a small effect size (0.27 and 0.25, respectively) while the rectus femoris showed a large effect size (0.92) when comparing the groups (Table 3S, Supplementary Material).

#### 3.3.2 Muscle structure

#### 3.3.2.1 Pennation Angle, Muscle Thickness and Fascicle Length - ECC versus CONV

Three studies evaluated muscle architecture parameters (RAJ et al., 2012; REEVES et al., 2009; VÁCZI et al., 2014) and compared ECC versus CONV. Qualitative analysis showed divergent results. While one study (RAJ et al., 2012) found no between-training differences for pennation angle, fascicle length and muscle thickness (trivial and small between-group effect size), results in favor of CONV were demonstrated for the pennation angle (very large betweengroup effect size), and results in favor of ECC were demonstrated for fascicle length (very large between-group effect size) (REEVES et al., 2009). When summarizing the data, only two of these studies (RAJ et al., 2012; REEVES et al., 2009) were included in the meta-analysis for pennation angle, muscle thickness and fascicle length (Figures 4.1.1, 4.1.2, 4.1.3). The results of all the meta-analyzes demonstrated no difference between comparisons and low heterogeneity (Figures 4.1.1, 4.1.2, 4.1.3), except for an 82% heterogeneity in the pennation angle assessment (Figure 4.1.1). Based on the GRADE approach, the quality of the evidence for these outcomes was considered low for muscle thickness and very low for pennation angle and fascicle length (Table 4). Only one study (VÁCZI et al., 2014) evaluated the quadriceps muscle CSA, and the qualitative analysis demonstrated that both ECC-only and CONV training are able to increase the CSA, although the results showed no difference between the two training modalities, and the between-groups effect size was moderate (Table 3S, Supplementary Material).



**Figure 4.** Vastus lateralis pennation angle (4.1.1), muscle thickness (4.1.2) and fascicle length (4.1.3): comparison between ECC and CONV.

# 3.3.3 Performance in functional tests

# 3.3.3.1 Timed Up and Go (TUG) - ECC-emphasis vs. CONV

Two studies (DIAS et al., 2015; RAJ et al., 2012) compared the effects between ECCemphasis versus CONV training at the TUG's test performance. In both qualitative (Table 4S, Supplementary Material) and quantitative analyzes (Figure 5.1.1), there was no between-groups difference, with trivial and small effect sizes, respectively. Based on the GRADE approach, the quality of the evidence for this outcome was considered low (Table 4).

# 3.3.3.2 Timed Up and Go (TUG) - ECC-only vs. CON

A single study (CHEN et al., 2017) compared the effects of ECC-only versus CON training at TUG's performance. Qualitative analysis demonstrated greater reduction in TUG

values for the ECC-only compared to CON, and a large between-groups effect size (Table 4S, Supplementary Material).

# 3.3.3.3 Sit and Stand Test (SST) - ECC-only or ECC-emphasis vs. CONV

Two studies (DIAS et al., 2015; GLUCHOWSKI et al., 2017) evaluated SST through the time to perform five repetitions. As one of the studies (GLUCHOWSKI et al., 2017) compared CONV to ECC-only or ECC-emphasis training, it was possible to perform metaanalysis and subgroup analysis (Figure 5.1.2, 5.1.3). The results for the SST five repetitions test demonstrated no between-groups difference in both qualitative (Table 4S, Supplementary Material) and quantitative analyzes (Figure 5.1.2, 5.1.3). The effect size between the groups was small and moderate, respectively (Table 4S, Supplementary Material). Based on the GRADE approach, the quality of the evidence for this outcome was considered low (Table 4).

# 3.3.3.4 Sit and Stand Test (SST) - ECC-only vs. CON

Only one study (CHEN et al., 2017) compared the effects of ECC-only and CON at the number of times the individual was able to sit and stand during 30s. The results showed an increase in the number of SST repetitions during 30s for both groups (pre- vs. post-training). However, ECC-only showed a greater increase compared to CON, with a large between-groups effect size (Table 4S, Supplementary Material).

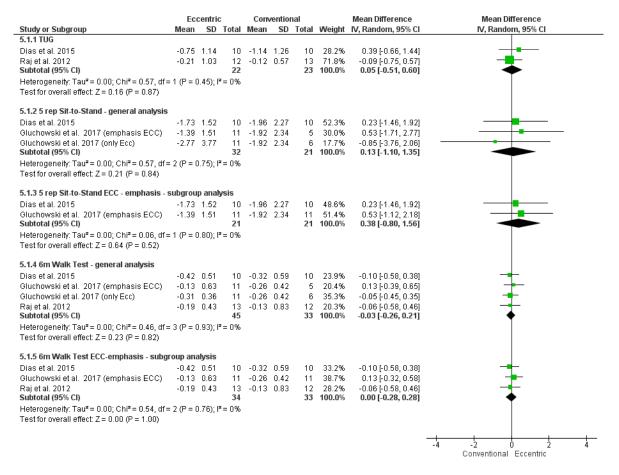
# 3.3.3.5 6m Walk Test (6MWT) - ECC-emphasis or ECC-only versus CONV

Three studies (DIAS et al., 2015; GLUCHOWSKI et al., 2017; RAJ et al., 2012) investigated the 6MWT, comparing the ECC versus the CONV training. One of the studies (GLUCHOWSKI et al., 2017) also compared the CONV training to ECC-emphasis or ECC-only, which allowed us to perform meta-analysis and subgroup analysis (Figure 5.1.4, 5.1.5). The results demonstrated no between-groups difference in both qualitative (Table 4S, Supplementary Material) and quantitative (Figure 5.1.4, 5.1.5) analyzes, and the between-groups effect size was small and moderate, respectively (Table 4S, Supplementary Material).

Based on the GRADE approach, the quality of the evidence for this outcome was considered low (Table 4).

### 3.3.3.6 6m Walk Test (6MWT) - ECC-only vs. CON

Only one study (CHEN et al., 2017) compared the effects of ECC-only versus CON training at 6MWT. Qualitative analyzes demonstrated no between-groups difference for the 6MWT values, with a small between-groups effect size (Table 4S, Supplementary Material).



**Figure 5.** Functional tests. TUG (5.1.1), 5-repetition Sit-to-Stand (5.1.2, 5.1.3) and 6m Walk-Test (5.1.4, 5.1.5) comparison between ECC and CONV.

### 3.3.3.7 Stair Climbing - ECC-emphasis versus CONV

Only one study (DIAS et al., 2015) compared the effects of ECC-emphasis versus CONV training at the stair climbing test performance. Qualitative analysis demonstrated that both groups reduced the total time for stair climbing (pre vs. post). Although the study found no significant differences between the groups, the percentage time reduction was greater for ECC-emphasis and there was a very large between-groups effect size (Table 4S, Supplementary Material).

# 3.3.3.8 Stair Descent - ECC-only or ECC-emphasis versus CONV

Only one study (GLUCHOWSKI et al., 2017) compared the effects of ECC-only or ECC-emphasis versus CONV training at the stair descent test. Qualitative analysis demonstrated a reduction in the time to stair descent for all groups (pre vs. post values). The percentage reduction (delta between pre and post values) was greater post ECC-only, followed by CONV and finally ECC-emphasis training. Although the study found no significant differences between the groups, it was possible to observe a moderate effect size (ECC-only vs. CONV) and a large effect size (ECC-emphasis vs. CONV) (Table 4S, Supplementary Material).

# 4. Discussion

The objective of this study was to identify which type of training (CON, ECC or CONV) is most effective to improve parameters of strength, muscle architecture and functionality in older people. Although the evidences might give us an idea of some of the effects of the different training modalities in the above parameters, the existent studies showed a low methodological quality (Table 3), suggesting that additional high quality studies should be performed. In addition, despite no between-groups difference was observed for most of the surveyed parameters, the qualitative analysis of the studies was able to identify some differences among the training modalities (Tables 2S, 3S and 4S, Supplementary Material).

Regarding the torque of CON, ECC and isometric contractions, there was no difference between the studies that compared ECC versus CONV training (RAJ et al., 2012; REEVES et al., 2009; VÁCZI et al., 2014). The qualitative analysis of the CON, ECC and isometric torque results, and of concentric power (SYMONS et al., 2005), are in line with the specificity principles of sports training (RELLY et al., 2009), as long-term adaptations demonstrated a slight increase in the ECC peak torque when subjects were trained in an ECC way. As for the groups that trained CON (CHEN et al., 2017; SYMONS et al., 2005; MALLIOU et al., 2003), there was an improvement in peak torque in all contraction types in disagreement with the specificity principle, but in agreement with a previous study (BAPTISTA et al., 2016); therefore, there is no difference between strength training modalities.

Purely CON or ECC contractions, as well as CON-ECC contractions used during CONV strength training, generate different neural and active and passive mechanical demands of the musculoskeletal system. Therefore, one would expect that specific adaptations should occur in muscle structure, in muscle function and in functionality after each training modality. Muscle strength gains usually occur according to the muscle contraction type used during training (AAGAARD et al., 2000; AMIRIDIS et al., 1996), although some studies found a greater increase in muscle strength for the ECC training group compared to CON (HORTOBAGYI et al., 1996; FARTHING, CHILIBECK, 2003; LASTAYO et al., 2014). This appears to occur mainly in exercises performed with faster contractions (CRESS et al., 1992; FARTHING, CHILIBECK, 2003). However, our qualitative analysis revealed that, in addition to maximal strength, the ECC training also determined a functionality improvement, as the greatest increase in maximum repetitions in the leg press and knee extension tests occurred in the ECC training group.

The rate of torque development is directly related to the functionality of the elderly, especially when it comes to preventing falls (AAGAARD et al., 2002; AAGAARD et al., 2007; (CASEROTTI et al., 2008, OSAWA et al., 2018) and it is also related to muscle activation (AAGAARD et al., 2002; MAFFIULETTI et al, 2016), because for a greater increase in RFD, greater recruitment of motor units is necessary. In the study by Váczi et al. (2014) there was a greater increase in the group trained in a CONV way, which is in line with the literature, as possibly the CONV training was superior to the ECC because it has CON contractions, which have a characteristic of greater muscular activation (AAGAARD et al., 2000; AMIRIDIS et al., 1996). However, what was identified by this systematic review is that ECC training determined greater increases in muscle activation (KANG et al., 2016), contradicting the existing literature and leaving a gap for a better understanding of the mechanisms of ECC contractions.

Muscle structure losses occur with aging. More specifically, aging determines a decrease in fascicle length (NARICI et al., 2003), in fascicle pennation angle and in muscle thickness (KUBO et al., 2003), and these losses can be decreased by strength training (e.g., KAWAKAMI; ABE; FUKUNAGA, 1993). Depending on the training specificity, one would

expect that ECC training should produce the best mechanical stimulus to revert these losses, as the mechanical demands are generated with the muscle being actively stretched. This should produce an increase in both serial and parallel sarcomere numbers, thereby increasing fascicle length, pennation angle and muscle thickness. Therefore, one could expect that both ECC and CONV training modalities should produce the abovementioned structural adaptations.

Only two studies compared the effects of different strength training modalities on fascicle length (RAJ et al., 2012; REEVES et al., 2009). Our qualitative analysis revealed an increase in fascicle length only for the group that trained ECC, which is in line with the findings of an increase in serial sarcomeres post ECC training (FRANCHI et al., 2014; TIMMINS et al., 2016). Regarding the pennation angle, there was a divergence in the quantitative results, where Raj et al. (2012) showed a decrease for the CONV training group and Reeves et al. (2009) showed an increase in the angle of pennation for the same group. The increase in the angle of pennation post CONV training is well described in the literature (EMA et al., 2016; BLAZEVICH et al., 2003), and the results' divergence might be explained by the low reproducibility for the assessment of the angle of pennation, both for young and old people in the vastus lateralis muscle (STRASSER et al., 2013). As for the muscular thickness, evaluated by two studies (RAJ et al., 2012; REEVES et al., 2009), there was a slight superiority for the ECC training group (RAJ et al., 2012). This may be explained by the largest training volume performed in the ECC group (RAJ et al., 2012; FRANCHI et al., 2017). However, when performed at the same intensity and training volume, there was no difference in muscle thickness post training between the groups (REEVES et al., 2009). In the case of aging, the cross-sectional area is an important parameter to control because it is proportional to the maximum muscle strength (LIEBER and FRIDÉN, 2001). In the study by Váczi et al. (2014), both groups showed improvement in the anatomical cross-sectional area, with slight superiority for the ECC group, as already demonstrated in the literature (BARONI et al., 2015, MAEO et al., 2018). The reason for that is because the ECC training produces greater muscle forces (LASTAYO et al., 2014) or a greater mechanical stimulus, being therefore capable of generating greater muscle hypertrophy (HIGBIE et al., 1996).

Changes in functionality are evident with aging (CIOLAC; RODRIGUES-DA-SILVA, 2016), but only four studies (DIAS et al., 2015; GLUCHOWSKI et al., 2017; RAJ et al., 2012; CHEN et al., 2017) investigated the strength training effects in functionality. The gait speed test has been shown to be effective in screening functionality (CRUZ-JENTOFT et al., 2019). In the qualitative analysis, the ECC training was slightly more effective than the CONV for improving performance on the 6m Walk Test (DIAS et al., 2015; GLUCHOWSKI et al., 2017; RAJ et al., 2012). This is in line with the literature, which reported a correlation of the gait speed test with maximum strength (KO et al., 2012). Our qualitative analysis support this idea, as there were improvements both in the gait speed test and in the maximum strength test in the ECC training group (Table 2S, Supplementary Material). In the 6-minute walk test, there was also an improvement in performance for the ECC group (CHEN et al., 2017), as well as an improvement in the strength parameters (CHEN et al., 2017; Table 2S, Supplementary Material). Similar results were observed in pathological patients, where ECC exercise was effective in improving performance in this test (ISNER-HOROBETI et al., 2013). However, they disagree with another study in which there was no improvement in the 6-minute walk test after training (REIS et al., 2012).

Tsubaki et al. (2016) also found a correlation between the functional TUG test performance and the knee extensors peak torque. In our qualitative analysis, one study (CHEN et al., 2017) showed better TUG performance in the ECC training group, with large effect size, and also presented a greater increase in the isometric and CON peak torques for the same training group. However, two other studies (DIAS et al., 2015; RAJ et al., 2012) found divergent results when comparing ECC versus CONV training, which is in line with the findings of a literature review (GAULT, WILLEMS, 2013). This divergent result may have occurred due to the fact that these two studies used different training intensities.

ECC training has been shown to be superior to CON or CONV training to improve muscle strength, hypertrophy and, consequently, functionality in the elderly due to various adaptive factors of this training type (HODY et al., 2019; KOWALCHUK, BUTCHER, 2019; VOGT, HOPPELER, 2014). This may explain the slightly better functional performance of the ECC versus the CONV training group for the 30s chair stand test (CHEN et al., 2017), stair climbing (DIAS et al., 2015) and stair descent (GLUCHOWSKI et al., 2017). In the 5-rep sit-to-stand test, there was little qualitative difference between the groups with divergence in the results (DIAS et al., 2015; GLUCHOWSKI et al., 2017), presenting a slightly better clinical performance for the ECC group. What explains this improvement is the hypothesis that this test requires faster contractions and greater muscle strength production, which has already been described in the literature as ECC training adaptations (CRESS et al., 1992; FARTHING,

CHILIBECK, 2003; HODY et al., 2019; KOWALCHUK, BUTCHER, 2019; VOGT, HOPPELER, 2014).

# 5. Conclusion

This study compared different types of lower limb strength training in healthy older people and showed little difference in the structural and functional parameters between training modalities. Although the quantitative analysis showed no difference among the types of training, and the GRADE varied from low to very low, for the analyzed outcomes, qualitatively the ECC training proved to be slightly superior to improve the parameters of muscle function, muscle architecture and functional performance in this population. However, there are several methodological limitations in the reviewed literature, which do not allow us to conclude which training type is the most beneficial for the elderly. Therefore, more studies with better methodological quality are needed before we can reach a clear definition of which modality between CON, ECC and CONV training produces the most effective and beneficial effects in older people.

# 6. Study limitations

This review evaluated only studies that applied different types of training to lower limbs in healthy elderly people. Therefore, the findings may not be applied to elderly people with associated pathologies and young individuals. Additionally, we cannot generalize the effects of training to other muscle groups.

# 7. References

Aagaard, P.; Simonsen, E.B; Andersen, J.L.; Magnusson, S.P.; Halkjær-Kristensen, J.; Dyhre-Poulsen, P. Neural inhibition during maximal eccentric and concentric quadriceps contraction: effects of resistance training. J Appl Physiol, v. 89, n. 6, p. 2249-2257, 2000. doi: 10.1152/jappl.2000.89.6.2249

- Aagaard, P.; Simonsen, E.B; Andersen, J. L.; Magnusson, S. P.; dyhre-poulsen, P Increased rate of force development and neural drive of human skeletal muscle following resistance training. J Appl Physiol, v. 93, n. 4, p. 1318–1326, 2002. doi: 10.1152/japplphysiol.00283.2002.
- Amiridis, I.G.; Martin A.; Morlon, B.; Martin, L.; Cometti, G.; Pousson, M.; Hoecke, J.V. Coactivation and tension-regulating phenomena during isokinetic knee extension in sedentary and highly skilled humans. Eur J Appl Physiol, v. 73, n 1-2, p. 149-156, 1996. doi: 10.1007/BF00262824.
- Balshem, H.; Helfand, M.; Schünemann, H.J.; Oxman, A.D.; Kunz, R.; Brozek, J.; Vist, G.E.; Falck-Ytter, Y.; Meerpohl, J.; Norris, S. GRADE guidelines: 3. Rating the quality of evidence. J Clinical Epid, v. 64, n. 4, p. 401-406, 2011. doi: 10.1016/j.jclinepi.2010.07.015.
- Baptista, R.; Onzi, E.; Goulart, N.; Santos, L.; Makarewicz, G.; Vaz, M. Effects of concentric versus eccentric strength training on the elderly's knee extensor structure and function. J Exerc Physiol, v. 19, n. 3, p. 120-131. 2016.
- Baroni, B.M.; Pinto, R.S.; Herzog, W.; Vaz, M.A. Eccentric resistance training of the knee extensor muscle: Training programs and neuromuscular adaptations. Isok Exerc Sci, v. 23, n. 3, p. 183-198, 2015. doi: 10.3233/IES-150580
- Blazevich, A.J.; Gill, N.D.; Bronks, R.; Newton, R.U. Training-specific muscle architecture adaptation after 5-wk training in athletes. Sports Exerc, v. 35, n. 12, p. 2013-2022, 2003. doi: 10.1249/01.MSS.0000099092.83611.20
- Bergland, A.; Jørgensen, L.; Emaus, N.; Strand, B. H. Mobility as a predictor of all-cause mortality in older men and women: 11.8 year follow-up in the Tromsø study. BMC Health Serv Res, v. 17, n. 1, p. 1-7, 2017. doi: 10.1186/s12913-016-1950-0
- Borzuola, R.; Giombini, A.; Torre, G.; Campi, S.; Albo, E.; Bravi, M.; Borrione P., Fossati, C.;
  Macaluso, A. Central and peripheral neuromuscular adaptations to ageing. J Clin Med, v.
  9, n. 741, p. 2-16, 2020. doi: 10.3390/jcm9030741
- Caserotti, P. et al. Explosive heavy-resistance training in old and very old adults: Changes in rapid muscle force, strength and power. Scand J Med Sci Sports, v. 18, n. 6, p. 773–782, 2008. doi: 10.1111/j.1600-0838.2007.00732.x.
- Chen, T.C.; Tseng, W.; Huang, G.; Chen, H.; Tseng, K.; Nosaka, K. Superior effects of eccentric to concentric knee extensor resistance training on physical fitness, insulin

sensitivity and lipid profiles of elderly men. Front Physiol, v. 10, n. 8, p. 209, 2017. doi: 10.3389/fphys.2017.00209.

- Ciolac, E.G.; Rodrigues-Da-Silva, J.M. Resistance Training as a Tool for Preventing and Treating Musculoskeletal Disorders. Sports Med, v. 46, n. 9, p. 1230-1248, 2016. doi:
- Cress, N.M.; Peters, K.S.; Chandler, J.M. Eccentric and concentric force-velocity relationships of the quadriceps femoris muscle. J Orthop Sports Phys Ther, v. 16, n. 2, p. 82-86, 1992. doi: 10.1007/s40279-016-0507-z
- Cruz-Jentoft, A.J.; Bahat, G.; Bauer, J.; Boirie, Y.; Bruyere, O.; Landi, F.; Rolland, Y.; Sayer, A.A.; Sieber, C.C.; Visser, M.; Zamobni, M. Sarcopenia: Revised European consensus on definition and diagnosis. Age Ageing, v. 48, n. 1, p. 16-31, 2019. doi: 10.1093/ageing/afy169.
- De Moura, T. G.; De Almeida Nagata, C.; Garcia, P. A. The influence of isokinetic peak torque and muscular power on the functional performance of active and inactive communitydwelling elderly: a cross-sectional study. Braz J Phys Ther, v. 24, n. 3, p. 256-263, 2020. doi: 10.1016/j.bjpt.2019.03.003.
- Dias, C.P.; Toscan, R.; Camargo, M.; Pereira, E.P.; Griebeler, N.; Baroni, B.M.; Tiggemann, G.L. Effects of eccentric-focused and conventional resistance training on strength and functional capacity of older adults. Age (Dordr), v. 37, n. 5, p. 99, 2015. doi: 10.1007/s11357-015-9838-1.
- Ema, R.; Akagi, R.; Wakahara, T.; Kawakami, Y. Training-induced changes in architecture of human skeletal muscles: Current evidence and unresolved issues. J Phys Fit Sports Med, v. 5, n. 1, p. 37-46, 2016. doi: 10.7600/jpfsm.5.37
- Farthing, J.; Chilibeck, P.D. The effects of eccentric and concentric training at different velocities on muscle hypertrophy. J Appl Physiol, v. 89, p. 578-586, 2003. doi: 10.1007/s00421-003-0842-2
- Franchi, M.V.; Wilkinson, D.J.; Quinlan, J.I.; Mitchell, W.L.; Lund, J.N.; Willians, J.P.; Reeves, N.D.; Smith, K.; Atherton, P.J.; Narici, M.V. Early structural remodeling and deuterium oxide-derived protein metabolic responses to eccentric and concentric loading in human skeletal muscle. Physiol Rep, v. 3, n. 11, 2014. doi: 10.14814/phy2.12593.
- Gault, M.L.; Willems, M.E.T. Aging, Functional Capacity and Eccentric Exercise Training. Aging Dis, v. 4, n. 6, p. 351-363, 2013. doi: 10.14336/AD.2013.0400351.

- Gluchowski, A.; Dulson D.; Merien, F.; Plank, L.; Harris, N. Comparing the effects of two distinct eccentric modalities to traditional resistance training in resistance trained, higher functioning older adults. Exp Geront, v. 98, p. 224-229, 2017. doi: 10.1016/j.exger.2017.08.034.
- Gonzalez-Freire, M.; Cabo, R.; Studenski, S.A.; Ferrucci, L. The neuromuscular junction: Aging at the crossroad between nerves and muscle. Front Aging Neurosci, v. 6, p. 1-11, 2014. doi: 10.3389/fnagi.2014.00208
- Granacher, U.; Muehlbauer, T.; Zahner, L.; Gollhofer, A.; Kressig, R.W. Comparison of Traditional and Recent Approaches in the Promotion of Balance and Strength in Older Adults. Sports Med, v. 41, n. 5, p. 377-400, 2011. doi: 10.2165/11539920-00000000-00000
- Guadagnin, E. C.; Priario, L. A.; Carpes, F. P.; Vaz, M. A. Correlation between lower limb isometric strength and muscle structure with normal and challenged gait performance in older adults. Gait & Posture, v. 73, p. 101-107, 2019. doi: 10.1016/j.gaitpost.2019.07.131
- Higbie, E.J.; Cureton, K.J.; Warren Iii, G.L.; Prior, B.M. Effects of concentric and eccentric training on muscle strength, cross-sectional area, and neural activation. J Appl Physiol, v. 81, n. 5, p. 2173-2181, 1996. doi: 10.1152/jappl.1996.81.5.2173.
- Higgins, J.P.; Altman, D.G.; Gøtzsche, P.C.; Jüni, P.; Moher, D.; Oxman, A.D.; Savović, J.; Schulz, K.F.; Weeks, L.; Sterne, J.A. The Cochrane Collaboration's tool for assessing risk of bias in randomised trials. British Med J, v.343, p. d5928, 2011. doi: 10.1136/bmj.d5928
- Hody, S.; Croisier, J.; Bury, T.; Rogister, B.; Leprince, P. Eccentric muscle contractions: Risks and Benefits. Front Physiol, v. 10, n. 536, 2019. doi: 10.3389/fphys.2019.00536
- Hortobágyi, T.; Hill, J.P.; Houmard, J.A.; Fraser, D.D.; Lambert, N.J.; Israel, R.G. Adaptive responses to muscle lengthening and shortening in humans. J Appl Physiol, v. 80, n. 3, p. 765-772, 1996. doi: 10.1152/jappl.1996.80.3.765.
- Hunter, S.K.; Pereira, H.M; Keenan, K.G. The aging neuromuscular system and motor performance. J Appl Physol, v. 121, n. 4, p. 982-995, 2016. doi: 10.1152/japplphysiol.00475.2016.
- Isner-Horobeti, M.; Dufour, S.P.; Vautravers, P.; Geny, B.; Coudeyre, E.; Richard, R. Eccentric exercise training: modalities, applications and perspectives. Sports Med, v. 43, n. 6, p. 483-512, 2013. doi: 10.1007/s40279-013-0052-y

- Janssen, I.; Heymsfield, S.B.; Ross, R. Low relative skeletal muscle mass (sarcopenia) in older persons is associated with functional impairment and physical disability. Amer Geriat Soci, v. 50, n. 5, p. 889-896, 2000. doi: 10.1046/j.1532-5415.2002.50216.x.
- Kang, J.; Jeong, D.; Chol, H. The effect of intervention according to muscle contraction type on the cerebral cortex of the elderly. J Phys Ther Sci, v. 28, p. 2560-2564, 2016. doi: 10.1589/jpts.28.2560
- Kawakami, Y.; Fukunaga, T. Muscle-Fiber Pennation Angles Are Greater In Hypertrophied Than In Normal Muscles. J Appl Physiol, v. 74, n. 6, p. 2740-2744, 1993.
- Kowalchuk, K.; Butcher, S. Eccentric overload flywheel training in older adults. J Funct Morphol Kinesiol, v. 4, n. 3, p. 61, 2019. doi: 10.1152/jappl.1993.74.6.2740.
- Kubo, K.; Kanehisa, H.; Azuma, K.; Ishizu, M.; Kuno, S.Y.; Okada, M; Fukunanaga, T. Muscle Architectural Characteristics in Young and Elderly Men and Women. Int J Sports Med, v. 24, n. 2, p. 125-130, 2003. doi: 10.1055/s-2003-38204.
- Lastayo, P.; Marcus, R.; Dibble, L.; Wong, B.; Pepper, G. Eccentric versus traditional resistance exercise for older adult fallers in the community: a randomized trial within a multicomponent fall reduction program. BMC Geriatrics, v. 17, p. 149, 2017. doi: 10.1186/s12877-017-0539-8
- Lastayo, P.; Mercus, R.; Dibble, L.; Frajacomo, F.; Lindstedt, S. Eccentric exercise in rehabilitation: safety, feasibility, and application. J Appl Physiol (1985), v. 116, n. 11, p. 1426-34, 2014. doi: 10.1152/japplphysiol.00008.2013.
- Lee, P.G.; Jackson, E.A; Richardson, C.R. Exercise Prescriptions in Older Adults. Am Fam Phys, v. 95, n. 7, p. 425-232, 2017.
- Liberati, A.; Altman, D.G.; Tetzlaff, J.; Mulrow, C.; Gøtzsche, P.C.; Ioannidis, J.P.; Clarke, M.; Devereaux, P.J.; Kleijnen, J.; Moher, D. The PRISMA statement for reporting systematic reviews and meta-analyses of studies that evaluate health care interventions: explanation and elaboration. PLoS Med, v.6, n.7, 2009. doi: 10.1136/bmj.b2700
- Lieber, R.L.; Fridén, J. Clinical Significance of Skeletal Muscle Architecture. Clin Orthop Relat Res, n. 383 p. 140-151, 2001. doi: 10.1097/00003086-200102000-00016
- Macaluso, A.; De Vito, G. Muscle strength, power and adaptations to resistance training in older people. Eur J Appl Physiol, v. 91, n. 4, p. 450-72, 2004. doi: 10.1007/s00421-003-0991-3

- Maeo, S.; Shan, X.; Otsuka, X.; Kanehisa, H.; Kawakami, Y. Neuromuscular adaptations to work-matched maximal eccentric vs concentric training. Med Sci Sports Exerc, v. 50, n. 8, p. 1629-1640, 2018. doi: 10.1249/MSS.000000000001611
- Maffiuletii, N. A.; Aagaard, P.; Blazevich, A. J.; Folland, J., Tillin, N.; Duchateau, J. Rate of force development: physiological and methodological considerations. Eur J Appl Physiol, v. 116, n. 6, p. 1091-1116, Mar 2016. doi: 10.1007/s00421-016-3346-6.
- Malliou, P.; Fatouros, I.; Beneka, A.; Gioftsidou, A.; Zissi, V.; Godolias, G.; Fotinakis, P. Different training programs for improving muscular performance in healthy inactive elderly. Isok Exerc Sci, v. 11, n. 4, p. 189-195, 2003. doi: 10.3233/IES-2003-0146
- Molinari, T.; Steffens, T.; Roncada, C.; Rodrigues, R.; Dias, C.P. Effects of eccentric-focused versus conventional training on lower limb muscular strength in older people: a systematic review with meta-analysis. J Aging Phys Act, v. 27, n. 4, p. 823-830, 2019. doi: 10.1123/japa.2018-0294.
- Narici, M.V.; Maganaris, C.N.; Reeves, N.D.; Capodaglio, P. Effect of Aging on Human Muscle Architecture. J Appl Physiol, v. 95, n. 6, p. 2229-2234, 2003. doi: 10.1152/japplphysiol.00433.2003.
- Naro, F.; Venturelli, M.; Monaco, L.; Toniolo, L.; Muti, E.; Milanese, C.; Zhao, J.; Richardson, R. S.; Schena, F.; Reggiani, C. Skeletal muscle fiber size and gene expression in the oldest-old with differing degrees of mobility. Front Physiol, v.10, p. 313, 2019. doi: 10.3389/fphys.2019.00313
- National Institute of Aging. US. 2011. Global health and ageing. https://www.who.int/ageing/publications/global\_health/en/
- Osawa Y.; Studensk S.A.; Ferruci L. Knee extension rate of torque development and peak torque: associations with lower extremity function. J Cach Sarcop Muscl, v. 9, p. 530-539, 2018. doi: 10.1002/jcsm.12285.
- OPAS (Organização Pan-Americana de Saúde). Folha informativa Envelhecimento e saúde, 2018.

https://www.paho.org/bra/index.php?option=com\_content&view=article&id=5661:folhainformativa-envelhecimento-e-saude&Itemid=820

Perkisas, S.; Cock, A.; Verhoeven, V.; Vandewoude M. Physiological and architectural changes in the ageing muscle and their relation to strength and function in sarcopenia. Eur Geriat, v. 7, n. 3, p. 201-206, 2016. doi:10.1016/j.eurger.2015.12.016

- Raj, I.S.; Bird, S., Westfold, B.; Shield, A. Effects of eccentrically biased versus conventional weight training in older adults. Med Sci Sports Exerc, v. 44, n. 6, p. 1167-1176, 2012. doi: 10.1249/MSS.0b013e3182442ecd.
- Reeves, N.D.; Maganaris, C.N.; Longo, S.; Narici, M.V. Differential adaptations to eccentric versus conventional resistance training in older humans. Exp Physiol, v. 94, n. 7, p. 825-833, 2009. doi: 10.1113/expphysiol.2009.046599.
- Reeves, N.D.; Maganaris, C.N.; Narici, M.V. Plasticity of dynamic muscle performance with strength training in elderly humans. Muscle Nerve, v. 31, p. 355-364, 2005. doi: 10.1002/mus.20275.
- Reis, J.G.; Costa, G.C.; Schimidt, A.; Ferreira, C.H.; Abreu, D.C. Do muscle strengthening exercises improve performance in the 6-minute walk test in postmenopausal women? Braz J Phys Ther, v. 16, n. 3, p. 236-40, 2012. doi: 10.1590/S1413-35552012005000022.
- Robinson, K.A.; Dickersin, K. Development of a highly sensitive search strategy for the retrieval of reports of controlled trials using PubMed. Inter J Epid, v.31, n.1, p. 150-153, 2002. doi: 10.1093/ije/31.1.150.
- Schünemann, H.J.; Oxman, A.D.; Brozek, J.; Glasziou, P.; Jaeschke, R.; Vist, G.E.; Williams, J.W.; Kunz, R.; Craig, J.; Montori, V.M. Grading quality of evidence and strength of recommendations for diagnostic tests and strategies. British Med J, v. 336, n. 7653, p. 1106-1110, 2008. doi: 10.1136/bmj.39500.677199.AE.
- Strasser, E.M.; Draskovits, T.; Praschak, M.; Quittan, M.; Graf, A. Association between ultrasound measurements of muscle thickness, pennation angle, echogenicity and skeletal muscle strength in the elderly. Age, v, 35, n. 6, p. 2377-2388, 2013. doi: 10.1007/s11357-013-9517-z.
- Symons, T.B.; Vandervoort, A.A.; Rice, C.L.; Overend, T.J.; Marsh, G.D. Effects of maximal isometric and isokinetic resistance training on strength and functional mobility in older adults. J Geront, v. 60A, n. 6, p. 777-781, 2005. doi: 10.1093/gerona/60.6.777.
- Timmins, R.G.; Ruddy, J.D.; Presland, J.; Maniar, N.V.; Shield, A.J.; Williams, M.D.; Opar, D.A. Architectural changes of the biceps femoris long head after concentric or eccentric training. Med Sci Sports Exerc, v. 48, n. 3, p. 499-508, 2016. doi: 10.1249/MSS.000000000000795.
- Trombetti, A.; Reid, K.F.; Hars, M.; Herrmann, F.R.; Pasha, E.; Phillips, E.M.; Fielding, R.A. Age-associated declines in muscle mass, strength, power, and physical performance:

impact on fear of falling and quality of life. Osteoporos Int., v. 27, n. 2, p. 463-471, 2016. doi: 10.1007/s00198-015-3236-5

- Tsubaki, A.; Kubo, M.; Kobayashi, R.; Jogami, H.; Sugawara, K.; Takarashi, H.E. Maximum Power during Vertical Jump and Isometric Knee Extension Torque Alter Mobility Performance: A Cross-Sectional Study of Healthy Individuals. PM & R, v. 8, n. 1, p. 19-27, 2016. doi: 10.1016/j.pmrj.2015.04.012.
- United Nations. US. 2019. World Population Ageing 2019. https://www.un.org/en/development/desa/population/publications/pdf/ageing/WorldPopul ationAgeing2019-Highlights.pdf
- Váczi, M.; Nagy, S.A.; Koszegi, T.; Amburs, M.; Bogner, P.; Perlaki, G; Orsi, G.; Toh, G.; Hostobagyi, T. Mechanical, hormonal, and hypertrophic adaptations to 10 weeks of eccentric and stretch-shortening cycle exercise training in old males. Exp Geront, v. 58, p. 69-77, 2014. doi: 10.1016/j.exger.2014.07.013
- Vogt, M.; Hoppeler, H.H. Eccentric exercise: mechanisms and effects when used as training regime or training adjunct. J Appl Physiol, v. 116, p. 1446-1454, 2014. doi: 10.1152/japplphysiol.00146.2013.
- Von Haehling, S.; Morley, J.E.; Anker, S.D. An overview of sarcopenia: facts and numbers on prevalence and clinical impact. J Cachex Sarcop Muscle, v. 1, n. 2, p. 129-133, 2010. doi: 10.1007/s13539-010-0014-2.
- Wisdom, K.M.; Delp, S.L.; Kuhl, E. Use it or lose it: multiscale skeletal muscle adaptation to mechanical stimuli. Biomech Model Mechanobiol, v. 14, p. 195-215, 2015. doi: 10.1007/s10237-014-0607-3.

# **Credit Authorship Contribution Statement**

The authors have made substantial contributions to conception, drafting and revising the manuscript critically for relevant intellectual content, and approved the final version and agreed to be accountable for all aspects of the study.

# **Declaration of Competing Interest**

The authors declare that they have no conflict of interest concerning the content of the manuscript.

# Funding

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

Highlights (85 characters max/bullet point)

- Eccentric and conventional training produce similar muscle adaptations in older people.
- Eccentric and conventional training improve functionality similarly in older people.
- Qualitatively, eccentric training was slightly superior to conventional training.
- Low-quality of reviewed literature suggests improvement in methodological quality.

# **Supplementary Material**

Table 1S. Studies' search strategy used in PubMed.

Table 1S – Literature search strategy used for the PUBMED database

- #1 "Aged"[Mesh] OR "Aged" OR "Elderly" OR "Aging"[Mesh] OR "Aging" OR "Senescence" OR "Biological Aging" OR "Aging, Biological" OR "Ageing Elderly" OR "older" OR "old" OR "Ageing" OR "Aged, 80 and over" [Mesh] OR "Aged, 80 and over" OR "Oldest Old" OR "Nonagenarians Older" OR "Old" OR "Nonagenarians Older" OR "Old" OR "Centenarian" OR "Octogenarians" OR "Centenarian"
- #2 "Resistance Training" [Mesh] OR "Resistance Training" OR "Training, Resistance" OR "Strength Training" OR "Training, Strength" OR "Weight-Lifting Strengthening Program" OR "Strengthening Program, Weight-Lifting" OR "Strengthening Programs, Weight-Lifting" OR "Weight-Lifting Strengthening Programs" OR "Weight-Lifting Exercise Program" OR "Weight-Lifting Strengthening Programs" OR "Weight-Lifting" OR "Exercise Program, Weight-Lifting OR "Weight-Lifting" OR "Weight-Lifting Exercise Programs, Weight-Lifting OR "Weight-Bearing" OR "Weight-Bearing" OR "Weight-Bearing OR "Weight-Bearing" OR "Weight-Bearing" OR "Weight-Bearing" OR "Weight-Bearing OR "Weight-Bearing" OR "Weight-Bearing" OR "Weight-Bearing" OR "Weight-Bearing OR "Weight-Bearing Strengthening Programs" OR "Weight-Bearing Programs Weight-Bearing" OR "Weight-Bearing Contraction" OR "Weight-Bearing OR "Weight-Bearing Strengthening Programs" OR "Weight-Bearing" OR "Weight-Bearing OR "Weight-Bearing Strengthening Programs" OR "Weight-Bearing Contraction" OR "Concentric Training" OR "Eccentric Exercise" OR "Eccentric Contraction" OR "Concentric Training" OR "Concentric Exercise" OR "Concentric Contraction" OR "Eccentric" OR "Lengthening Contraction" OR "Negative Work" OR "Positive Work" OR "Shortening Contraction" OR "concentric eccentric" OR "concentric eccentric" OR "eccentric" OR "eccentric" OR "Concentric" OR "Negative Work" OR "Positive Work" OR "Shortening Contraction" OR "concentric eccentric"
- "Muscle Strength" [Mesh] OR "Muscle Strength" OR "Strength, Muscle" OR "Arthrogenic Muscle #3 Inhibition" OR "Arthrogenic Muscle Inhibitions" OR "Inhibition, Arthrogenic Muscle" OR "Muscle Inhibition, Arthrogenic" OR "Strength, Muscle" OR "Torque" [Mesh] OR "Torque" OR "Torques" OR "Walking Speed" [Mesh] OR "Walking Speed" OR "Speed, Walking" OR "Speeds, Walking" OR" Walking Speeds" OR "Gait Speed" OR "Gait Speeds" OR "Speed, Gait" OR "Speeds, Gait" OR "Walking Pace" OR "Pace, Walking" OR "Paces, Walking" OR "Walking Paces" OR "force" OR "power" OR "rate of torque development" OR "rate of torque production" OR "rate of force production" OR "rate Supplemental Digital Content 1 of force development" OR "muscle activation" OR "muscular activation" OR "pennation angle" OR "fascicle length" OR "muscle thickness" OR "echo intensity" OR "echo-intensity" OR "muscle quality" OR "muscular quality" OR "functionality" OR "functional" OR "mobility" OR "gait velocity" OR "timed up and go test" OR "timed up and go" OR "TUG OR TUG test" OR "sit to stand" OR "sit to stand test" OR "30 second sit to stand test" OR "30 second sit to stand" OR "sit-to-stand" OR "sit-to-stand test" OR "jump" OR "vertical jump" OR "Stair Climbing" OR "Climbing, Stair" OR "Stair Navigation" OR "Navigation, Stair" OR "6 min walk test" OR "6 min walking test" OR "6-min-walk-test" OR "6-min-walkingtest" OR "6 minutes walk test" OR "6 minutes walking test" OR "6- minutes-walk-test" OR "6minutes-walking-test" OR "six min walk test" OR "six min walking test" OR "six six-min-walktest" OR "six-min-walking-test" OR "six minutes walk test" OR "six minutes walking test" OR "sixminutes-walk-test" OR "six-minutes-walking-test" OR "Functional testing"
- #4 randomized controlled trial[pt] OR controlled clinical trial[pt] OR randomized controlled trials[mh] OR random allocation[mh] OR double-blind method[mh] OR single-blind method[mh] OR clinical trial[pt] OR clinical trials[mh] OR ("clinical trial"[tw]) OR ((singl\*[tw] OR doubl\*[tw] OR trebl\*[tw] OR tripl\*[tw]) AND (mask\*[tw] OR blind\*[tw])) OR ("latin square"[tw]) OR placebos[mh] OR placebo\*[tw] OR random\*[tw] OR research design[mh:noexp] OR follow-up studies[mh] OR prospective studies[mh] OR cross-over studies[mh] OR control\*[tw] OR prospectiv\*[tw] OR volunteer\*[tw]) NOT (animal[mh] NOT human[mh]).
  #5

Table 2S. Muscle function results.

Study	Outcome	Muscle Testing	Mean±SD Pre	Mean±SD Post	% Change Pre-to-Post	Effect Size Within- Groups Pre vs Post	Effect Size Between- Groups Post	P-value
Dias et al. 2015	1RM (Kg)	Knee extension	ECC-emphasis: 29.5±5.7 CONV: 29.7±6.3	ECC-emphasis: 37.0±10.9 CONV: 37.8±13.2	ECC-emphasis: +25.4% CONV: +27.3%	ECC: 0.86 CONV: 0.78	ECC x CONV: 0.06	Within-groups ECC: 0.007 CONV: 0.007 Between- groups: 0.82
Dias et al. 2015	1RM (Kg)	Leg press	ECC-emphasis: 90.5±21.9 CONV: 85.1±12.9	ECC-emphasis: 102.2±19.2 CONV: 96.7±31.4*	ECC-emphasis: +12.9% CONV: +13.6%	ECC: 0.56 CONV: 0.48	ECC x CONV: 0.21	Within-groups ECC: NS CONV: NS Between- groups: 0.89
Gluchowski et al. 2017	1RM (Kg)	Leg press	ECC-emphasis: 251.0±94.3 ECC-only: 219.0±58.0 CONV: 222.0±87.0	ECC-emphasis: 330.0±87.1 ECC-only: 296.0±48.2 CONV: 317.0±95.6	ECC-emphasis: +31.5% ECC-only: +35.1% CONV: +42.8%	ECC- emphasis: 0.87 ECC-only: 1.44 CONV: 1.03	ECC-emphasis x CONV: 0.14 ECC-only x CONV: 0.27	Within-groups ECC- emphasis: p<0.01 ECC-only: p<0.01 CONV: p<0.01 Between- groups: NS
Raj et al. 2012	1RM (Kg)	Leg press	ECC-emphasis: 171.0±51.0 CONV: 159.0±38.0	ECC-emphasis: 211.0±61.0 CONV: 195.0±45.0	ECC-emphasis: +23.4% CONV: +22.6%	ECC: 0.71 CONV: 0.86	ECC x CONV: 0.29	Within-groups ECC: p<0.01 CONV: p<0.01 Between- groups: NS
Raj et al. 2012	Isometric knee extension (Nm)	Isokinetic dynamometer	ECC-emphasis: 175.0±38.0 CONV: 160.0±40.0	ECC-emphasis: 187.0±44.0 CONV: 166.0±46.0	ECC-emphasis: +6.8% CONV: +3.7%	ECC: 0.29 CONV: 0.13	ECC x CONV: 0.46	Within-groups ECC: p<0.05 CONV: NS Between- groups: NS
Raj et al. 2012	Concentric torque 60°/s (Nm)	Isokinetic dynamometer	ECC-emphasis: 129.0±30.0	ECC-emphasis: 137.0±32.0	ECC-emphasis: +6.2% CONV: +7.1%	ECC: 0.25 CONV: 0.23	ECC x CONV: 0.05	Within-groups ECC: p<0.05 CONV: p<0.05

			CONV: 126.0±36.0	CONV: 135.0±39.0				Between- groups: NS
Raj et al. 2012	Concentric torque 120°/s (Nm)	Isokinetic dynamometer	ECC-emphasis: 101.0±24.0 CONV: 101.0±31.0	ECC-emphasis: 108.2±26.0 CONV: 109.0±32.0	ECC-emphasis: +6.9% CONV: +7.9%	ECC: 0.28 CONV: 0.25	ECC x CONV: 0.02	Within-groups ECC: p<0.01 CONV: p<0.01 Between- groups: NS
Raj et al. 2012	Concentric torque 240°/s (Nm)	Isokinetic dynamometer	ECC-emphasis: 74.0±21.0 CONV: 75.0±26.0	ECC-emphasis: 78.0±22.0 CONV: 77.0±26.0	ECC-emphasis: +5.4% CONV: +2.6%	ECC: 0.18 CONV: 0.07	ECC x CONV: 0.04	Within-groups ECC: p<0.05 CONV: NS Between- groups: NS
Raj et al. 2012	Concentric torque 360°/s (Nm)	Isokinetic dynamometer	ECC-emphasis: 56.0±16.0 CONV: 59.0±21.0	ECC-emphasis: 62.0±19.0 CONV: 60.0±24.0	ECC-emphasis: +10.7% CONV: +1.7%	ECC: 0.34 CONV: 0.04	ECC x CONV: 0.09	Within-groups ECC: p<0.01 CONV: NS Between- groups: p<0.05
Reeves et al. 2009	5RM (Kg)	Knee extension	ECC-only: 69.0±23.0 CONV: 44.0±12.0	ECC-only: 103.0±21.0 CONV: 49.0±14.0	ECC-only: +49.3% CONV: +11.4%	ECC: 1.54 CONV: 0.38	ECC x CONV: 3.02	Within-groups ECC: NI CONV: NI Between- groups: p<0.01
Reeves et al. 2009	5RM (Kg)	Leg press	ECC-only: 252.0±56.0 CONV: 178.0±45.0	ECC-only: 358.0±44.0 CONV: 219.0±55.0	ECC-only: +42.06% CONV: +23%	ECC: 2.10 CONV: 0.81	ECC x CONV: 2.79	Within-groups ECC: NI CONV: NI Between- groups: p<0.01
Reeves et al. 2009	Isometric extension (Nm)	Isokinetic dynamometer	ECC-only: 119.0±35.0 CONV: 115.1±51.5	ECC-only: 129.0±28.0 CONV: 126.6±48.7	ECC-only: +8.4% CONV:+10.0%	ECC: 0.31 CONV: 0.22	ECC x CONV: 0.06	Within-groups ECC: NS CONV: NS Between- groups: NS
Reeves et al. 2009	Concentric torque 50°/s (Nm)	Isokinetic dynamometer	ECC-only: 95.0±42.0 CONV: 80.0±34.0	ECC-only: 107.0±34.0 CONV: 101.4±33.0	ECC-only: +12.6% CONV: +26.8%	ECC: 0.31 CONV: 0.63	ECC x CONV: 0.16	Within-groups ECC: NS CONV: p<0.05 Between- groups: NI

Reeves et al. 2009	Concentric torque 100°/s (Nm)	Isokinetic dynamometer	ECC-only: 86.0±41.0 CONV: 69.0±37.0	ECC-only: 92.0±37.0 CONV: 84.0±33.0	ECC-only: +7.0% CONV: +21.7%	ECC: 0.15 CONV: 0.42	ECC x CONV: 0.22	Within-groups ECC: NS CONV: p<0.05 Between- groups: NI
Reeves et al. 2009	Concentric torque 150°/s (Nm)	Isokinetic dynamometer	ECC-only: 69.0±37.5 CONV: 52.3±34.2	ECC-only: 73.5±32.0 CONV: 71.7±28.0	ECC-only: +6.5% CONV: +37.0%	ECC: 0.12 CONV: 0.62	ECC x CONV: 0.05	Within-groups ECC: NS CONV: p<0.05 Between- groups: NS
Reeves et al. 2009	Concentric torque 200°/s (Nm)	Isokinetic dynamometer	ECC-only: 46.0±27.0 CONV: 40.0±26.0	ECC-only: 61.0±26.0 CONV: 56.0±21.0	ECC-only: +34.0% CONV: +40.0%	ECC: 0.56 CONV: 0.67	ECC x CONV: 0.21	Within-groups ECC: p<0.05 CONV: p<0.01 Between- groups: NI
Reeves et al. 2009	Eccentric torque -50°/s (Nm)	Isokinetic dynamometer	ECC-only: 146.0±44.0 CONV: 162.0±58.0	ECC-only: 169.0±43.0 CONV: 169.0±41.0	ECC-only: +16.1% CONV: +4.3%	ECC: 0.52 CONV: 0.13	ECC x CONV: 0.00	Within-groups ECC: p<0.01 CONV: NS Between- groups: NI
Reeves et al. 2009	Eccentric torque -100°/s (Nm)	Isokinetic dynamometer	ECC-only: 156.0±44.0 CONV: 156±47.0	ECC-only: 172.0±37.0 CONV: 168.0±45.0	ECC-only: +10.2% CONV: +7.6%	ECC: 0.39 CONV: 0.26	ECC x CONV: 0.09	Within-groups ECC: p<0.05 CONV: NS Between- groups: NI
Reeves et al. 2009	Eccentric torque -150°/s (Nm)	Isokinetic dynamometer	ECC-only: 160.0±41.5 CONV: 166.2±57.7	ECC-only: 177.0±49.0 CONV: 167.8±43.7	ECC-only: +10.6% CONV: +0.96%	ECC: 0.37 CONV: 0.03	ECC x CONV: 0.19	Within-groups ECC: p<0.05 CONV: NS Between- groups: NI
Reeves et al. 2009	Eccentric torque -200°/s (Nm)	Isokinetic dynamometer	ECC-only: 158.0±46.5 CONV: 162.0±58.1	ECC-only: 170.5±44.5 CONV: 171.9±38.7	ECC-only: +7.9% CONV: +6.1%	ECC: 0.27 CONV: 0.20	ECC x CONV: 0.03	Within groups ECC: p<0.05 CONV: NS Between- groups: NI
Váczi et al. 2014	Isometric extension (Nm)	Isokinetic dynamometer	ECC-only: 216.0±47.0 CONV: 218.0±53.0	ECC-only: 229.0±62.0 CONV: 230.0±63.0	ECC-only: +6.0% CONV: +5.5%	ECC: 0.23 CONV: 0.20	ECC x CONV: 0.01	Within-groups ECC: p<0.05 CONV: p<0.05

								Between- groups: NS
Váczi et al. 2014	Eccentric torque 90°/s (Nm)	Isokinetic dynamometer	ECC-only: 235.0±39.0 CONV: 264.0±62.0	ECC-only: 306.0±60.0 CONV: 309.0±72.0	ECC-only: +30.2% CONV: +17.0%	ECC: 1.40 CONV: 0.66	ECC x CONV: 0.04	Within-groups ECC: p<0.01 CONV: p<0.01 Between- groups: NS
Váczi et al. 2014	RTD 30ms (Nm*ms <sup>-1</sup> )	Isokinetic dynamometer	ECC-only: 1.22±0.52 CONV: 1.28±0.38	ECC-only: 1.10±0.87 CONV: 1.67±1.05	ECC-only: -9.83% CONV: +30.4%	ECC: 0.16 CONV: 0.49	ECC x CONV: 0.59	Within-groups ECC: NS CONV: p<0.05 Between- groups: NS
Váczi et al. 2014	RTD 50ms (Nm*ms <sup>-1</sup> )	Isokinetic dynamometer	ECC-only: 1.36±0.45 CONV: 1.44±0.39	ECC-only: 1.34±0.94 CONV: 1.86±1.00	ECC-only: -1.47% CONV: +29.16%	ECC: 0.02 CONV: 0.55	ECC x CONV: 0.53	Within-groups ECC: NS CONV: p<0.05 Between- groups: NS
Váczi et al. 2014	RTD 100ms (Nm*ms <sup>-1</sup> )	Isokinetic dynamometer	ECC-only: 1.03±0.22 CONV: 1.14±0.32	ECC-only: 1.04±0.43 CONV: 1.13±0.40	ECC-only: +0.97% CONV: -0.87%	ECC: 0.02 CONV: 0.02	ECC x CONV: 0.21	Within-groups ECC: NS CONV: NS Between- groups: NS
Chen et al. 2017	1 RM (Kg)	Knee extension	ECC-only: 29.8 ±2.9 CON: 30.5±3.1	ECC-only: 45.2±3.2 CON: 41.7±3.3	ECC-only: +51.7% CON: +36.7%	ECC: 5.04 CON: 3.5	ECC x CON: 1.07	Within-groups ECC: NI CON: NI Between- groups: p<0.05
Chen et al. 2017	Isometric extension (Nm)	Isokinetic dynamometer	ECC-only: 149.2 ±11.7 CON: 152.7±12.6	ECC-only: 195.7 ±13.2 CON: 180.0±13.5	ECC-only: +31.2% CON: +17.9%	ECC: 3.72 CON: 2.09	ECC x CON: 1.17	Within-groups ECC: NI CON: NI Between- groups: p<0.05
Chen et al. 2017	Concentric torque 30°/s (Nm)	Isokinetic dynamometer	ECC-only: 119.8±10.3 CON: 121.1±10.7	ECC-only: 141.2±11.0 CON: 133.7±11.	ECC-only: +17.9% CON: +10.4%	ECC: 2.00 CON: 1.13	ECC x CON: 0.66	Within-groups ECC: NI CON: NI Between- groups: p<0.05
Symons et al. 2005	Isometric extension (Nm)	Isokinetic dynamometer	ECC-only: 142.0±39.8	ECC-only: 176.0±44.7	ECC-only: +23.9%	ECC: 0.80 CON: 0.41	ECC x CON: 0.52	Within-groups ECC: p<0.01

			CON: 130.6±54.0	CON: 151.8±48.3	CON: +16.2%			CON: p<0.01 Between- groups: NS
Symons et al. 2005	Concentric torque 90°/s (Nm)	Isokinetic dynamometer	ECC-only: 107.5±30.7 CON: 93.6±40.4	ECC-only: 116.3±26.1 CON: 113.9±48.3	ECC-only: +8.2% CON: +21.7%	ECC: 0.30 CON: 0.45	ECC x CON: 0.06	Within-groups ECC: p<0.01 CON: p<0.01 Between- groups: NS
Symons et al. 2005	Eccentric torque 90°/s (Nm)	Isokinetic dynamometer	ECC-only: 168.5±40.0 CON: 161.9±62.6	ECC-only: 207.1±35.6 CON: 191.4±76.8	ECC-only: +22.9% CON: +18.2%	ECC: 1.01 CON: 0.42	ECC x CON: 0.26	Within-groups ECC: p<0.01 CON: p<0.01 Between- groups: NS
Symons et al. 2005	Peak Concentric Power (W)	Isokinetic dynamometer	ECC-only: 83.5±32.8 CON: 75.5±37.2	ECC-only: 98.1±28.7 CON: 104.4±41.0	ECC-only: +17.4% CON: +38.2%	ECC: 0.47 CON: 0.73	ECC x CON: 0.17	Within-groups ECC: p<0.01 CON: p<0.01 Between- groups: NS
Malliou et al. 2003	Concentric torque 60°/s (Nm)	Isokinetic dynamometer	CON: 106.3±12.2 CONV: 109.0±8.9	CON: 118.3±12.8 CONV: 128.1±10.8	CON: +11.3% CONV: +17.5%	CON: 0.95 CONV: 1.93	CON x CONV: 0.82	Within-groups CON: p<0.05 CONV: p<0.05 Between- groups: p<0.05
Malliou et al. 2003	Concentric torque 180°/s (Nm)	Isokinetic dynamometer	CON: 68.4±9.2 CONV: 70.1±8.2	CON: 81.3±5.8 CONV: 79.2±6.8	CON: +18.8% CONV: +13.0%	CON: 1.67 CONV: 1.20	CON x CONV: 0.33	Within-groups CON: p<0.05 CONV: p<0.05 Between- groups: NS

ECC: eccentric training; CONV: conventional training; CON: concentric training; RTD: rate of torque development; NS: not significant; NI: not informed.

Table 3S. Muscle structure and activation results.

Study	Outcome	Assessment equipment Muscle	Mean±SD Pre	Mean±SD Post	% Change Pre-to-Post	Effect Size Within- Groups Pre vs Post	Effect Size Between- Groups Post	P-value
Raj et al. 2012	Pennation angle (°)	Ultrasonography VL	ECC-emphasis: 11.5±4.4 CONV: 12.9±2.8	ECC-emphasis: 11.9±3.1 CONV: 11.9±3.6	ECC: +3.5% CONV: - 14.7%	ECC: 0.10 CONV: 0.31	ECC x CONV: 0.00	Within-group: ECC: NS CONV: NS Between- groups: NS
Raj et al. 2012	Fascicle length (cm)	Ultrasonography VL	ECC-emphasis: 12.3±3.8 CONV: 11.1±2.6	ECC-emphasis: 12.9±5.1 CONV: 11.5±3.6	ECC: +4.9% CONV: +3.6%	ECC: 0.13 CONV: 0.12	ECC x CONV: 0.31	Within-group: ECC: NS CONV: NS Between- groups: NS
Raj et al. 2012	Thickness (cm) VL1*	Ultrasonography VL	ECC-emphasis: 4.03±0.62 CONV: 3.88±0.46	ECC-emphasis: 4.22±0.68 CONV: 3.96±0.47	ECC: +4.7% CONV: +2.1%	ECC: 0.29 CONV: 0.17	ECC x CONV: 0.44	Within-group: ECC: p<0.05 CONV: NS Between- groups: NS
Raj et al. 2012	Thickness (cm) VL2**	Ultrasonography VL	ECC-emphasis: 4.49±0.53 CONV: 4.71±0.5	ECC-emphasis: 4.64±0.52 CONV: 4.71±0.62	ECC: +3.3% CONV: 0.0%	ECC: 0.28 CONV: 0.00	ECC x CONV: 0.12	Within-group: ECC: NS CONV: NS Between- groups: NS
Reeves et al. 2009	Pennation angle (°)	Ultrasonography VL	ECC-only: 13.7±1.6 CONV: 14.7±2.5	ECC-only: 14.4±1.5 CONV: 19.8±3.0	ECC: +5.1% CONV: +34.7%	ECC: 0.45 CONV: 1.86	ECC x CONV: 2.27	Within-group: ECC: NS CONV: p<0.01 Between- groups: p<0.01
Reeves et al. 2009	Fascicle length (cm)	Ultrasonography VL	ECC-only: 7.9±0.9 CONV: 7.2±0.8	ECC-only: 9.5±1.1 CONV: 7.8±1.2	ECC: +20.3% CONV: +8.3%	ECC: 1.59 CONV: 0.58	ECC x CONV: 1.47	Within-group: ECC: p<0.01 CONV: p<0.05 Between- groups: p<0.05

Reeves et al. 2009	Thickness (cm) 50% VL	Ultrasonography VL	ECC-only: 1.8±0.4 CONV: 1.8±0.3	ECC-only: 2.0±0.4 CONV: 2.0±0.3	ECC: +11.1% CONV: +11.1%	ECC: 0.50 CONV: 0.66	ECC x CONV: 0.00	Within-group: ECC: p<0.05 CONV: p<0.05 Between- groups: NS
Váczi et al. 2014	ACSA (mm <sup>2</sup> ) Quadriceps	Magnetic resonance	ECC-only: 5,752±977 CONV: 5,139±879	ECC-only: 5,938±985 CONV: 5,245±866	ECC: +3.2% CONV: +2.1%	ECC: 0.19 CONV: 0.12	ECC x CONV: 0.75	Within-group: ECC: p<0.05 CONV: p<0.05 Between- groups: NS
Kang et al. 2016	Surface EMG (%) Rectus femoris	EMG Biopac System	ECC-only: 104.8±13.2 CON: 106.8±18.2	ECC-only: 127.7±13.6 CON: 114.1±15.7	ECC: +21.8% CON: +6.8%	ECC: 1.70 CON: 0.42	ECC x CON: 0.92	Within-group: ECC: p<0.05 CON: NS Between- groups: NI
Kang et al. 2016	Surface EMG (%) Vastus medialis	EMG Biopac System	ECC-only: 98.8±14.8 CON: 95.7±15.3	ECC-only: 107.3±16.3 CON: 103.2±13.8	ECC: +8.6% CON: +7.8%	ECC: 0.54 CON: 0.51	ECC x CON: 0.27	Within-group: ECC: NS CON: NS Between- groups: NI
Kang et al. 2016	Surface EMG (%) VL	EMG Biopac System	ECC-only: 100.6±13.3 CON: 99.9±14.8	ECC-only: 109.2±14.2 CON: 105.4±15.2	ECC: +8.5% CON: +5.5%	ECC: 0.62 CON: 0.36	ECC x CON: 0.25	Within-group: ECC: NS CON: NS Between- groups: NI

VL: vastus lateralis muscle; ECC: eccentric training; CONV: conventional training; CON: concentric training; EMG: electromyography; NS: not significant; NI: not informed; \* 62.5% between the anterior superior iliac spine and the superior patella in the midsagittal (VL1); \*\* 62.5% between the anterior superior iliac spine and the superior of the patella in the mid-coronal (VL2).

Study	Outcome	Mean±SD Pre	Mean±SD Post	% Change Pre-to-Post	Effect Size Within-Groups Pre vs Post	Effect Size Between- Groups Post	P-value
Dias et al. 2015	6m Walk Test (s)	ECC-emphasis: 3.41±0.44 CONV: 3.32±0.49	ECC-empahsys: 2.99±0.27 CONV: 3.0±0.33	ECC-emphasis: -12.3% CONV: -9.6%	ECC: 1.15 CONV: 0.76	ECC x CONV: 0.03	Within-groups ECC: p<0.002 CONV: p<0.002 Between-groups: NS
Dias et al. 2015	TUG (s)	ECC-emphasis: 6.54±1.09 CONV: 6.45±0.92	ECC-emphasis: 5.40±0.63 CONV: 5.71±0.69	ECC-emphasis: -17.4% CONV: -11.5%	ECC: 1.28 CONV: 0.91	ECC x CONV: 0.46	Within-groups ECC: p<0.001 CONV: p<0.001 Between-groups: NS
Dias et al. 2015	Stair Climbing test (s)	ECC-emphasis: 3.35±0.59 CONV: 3.42±0.44	ECC-emphasis: 2.91±0.14 CONV: 3.12±0.18	ECC-emphasis: -13.1% CONV: -8.8 %	ECC: 1.02 CONV: 0.89	ECC x CONV: 1.30	Within-groups ECC: p<0.002 CONV: p<0.002 Between-groups: NS
Dias et al. 2015	5 REP Sit-to- stand (s)	ECC-emphasis: 11.82±1.24 CONV: 12.27±1.83	ECC-emphasis: 10.09±0.88 CONV: 10.32±1.36	ECC-emphasis: -14.6% CONV: -15.9%	ECC: 1.60 CONV: 1.20	ECC x CONV: 0.20	Within-groups ECC: p<0.001 CONV: p<0.001 Between-groups: NS
Gluchowski et al. 2017	6m Walk Test (s)	ECC-emphasis: 2.60±0.57 ECC-only: 2.81±0.28 CONV: 2.63±0.37	ECC-emphasis: 2.47±0.28 ECC-only: 2.50±0.24 CONV: 2.37±0.2	ECC-emphasis: -5% ECC-only: -11% CONV: -9.8%	ECC-emphasis: 0.28 ECC-only: 1.18 CONV: 0.87	ECC-emphasis x CONV: 0.41 ECC-only x CONV: 0.59	Within-groups ECC-emphasis: NS ECC-only: p<0.05 CONV: p<0.05 Between-groups: NS
Gluchowski et al. 2017	Stair descent (s)	ECC-emphasis: 4.60±0.76 ECC-only: 4.93±0.84 CONV: 4.30±0.48	ECC-emphasis: 4.11±0.56 ECC-only: 4.19±0.81 CONV: 3.68±0.49	ECC-emphasis: -10.7% ECC-only: -15% CONV: -14.4%	ECC-emphasis: 0.73 ECC-only: 0.89 CONV: 1.27	ECC-emphasis x CONV: 0.81 ECC-only x CONV: 0.76	Within-groups ECC-emphasis: p<0.01 ECC-only: p<0.01 CONV: p<0.01 Between-groups: NS
Gluchowski et al. 2017	5 REP Sit-to- stand (s)	ECC-emphasis: 7.01±1.20 ECC-only: 10.29±2.67	ECC-emphasis: 5.62±0.92 ECC-only: 7.52±2.67	ECC-emphasis: -19.8% ECC-only: -26.9%	ECC-emphasis: 1.30 ECC-only: 1.03 CONV: 1.15	ECC-emphasis x CONV: 0.37 ECC-only x CONV: 0.56	Within-groups ECC-emphasis: p<0.01 ECC-only: p<0.05

Table 4S. Functional performance results.

		CONV: 8.11±1.32	CONV: 6.19±1.94	CONV: -23.7%			CONV: p<0.01 Between groups: NS
Raj et al. 2012	6m Walk Test (s)	ECC-emphasis: 2.79±0.32 CONV: 2.79±0.57	ECC-emphasis: 2.60±0.29 CONV: 2.66±0.61	ECC-emphasis: -6.8% CONV: -4.7%	ECC: 0.62 CONV: 0.22	ECC x CONV: 0.12	Within-groups ECC: p<0.01 CONV: p<0.01 Between-groups: NS
Raj et al. 2012	TUG (s)	ECC-emphasis: 4.51±0.43 CONV: 4.55±0.81	ECC-emphasis: 4.39±0.38 CONV: 4.34±0.64	ECC-emphasis: -2.7% CONV: -4.6%	ECC: 0.29 CONV: 0.28	ECC x CONV: 0.09	Within-groups ECC: NS CONV: p<0.01 Between-groups: NS
Chen et al. 2017	6m Walk Test (s)	ECC-only: 515.8±35.5 CON: 510.4±38.3	ECC-only: 541.6±38.3 CON: 529.3±40.0	ECC-only: +5.0% CON: +3.7%	ECC: 0.69 CON: 0.48	ECC x CON: 0.32	Within-groups ECC: p<0.05 CON: p<0.05 Between-groups: NS
Chen et al. 2017	TUG (s)	ECC-only: 7.2±0.4 CON: 7.3±0.4	ECC-only: 5.3±0.37 CON: 5.7±0.37	ECC-only: -26.4% CON:-21.9%	ECC: 4.93 CON: 4.15	ECC x CON: 1.08	Within-groups ECC: p<0.05 CONC: p<0.05 Between-groups: p<0.05
Chen et al. 2017	30s Chair Stand (REP)	ECC-only: 17.1±2.7 CON: 16.0±2.0	ECC-only: 23.5±3.0 CON: 19.9±2.2	ECC-only: +34.7% CON: +24.4%	ECC: 2.24 CON: 1.85	ECC x CON: 1.36	Within-groups ECC: p<0.05 CONC: p<0.05 Between-groups: p<0.05

TUG: time-up-and-go test; REP: repetitions; ECC: eccentric training; CONV: conventional training; CON: concentric training; NS: not significant; NI: not informed.

# 8. General conclusion

Although the studies included in this dissertation presented low methodological quality and the quantitative analysis did not show any difference between the types of training, in a qualitative way the ECC strength training was slightly superior to improve the parameters of muscle function, muscle architecture and functional performance in healthy elderly people than the other training modalities.

# Anexo

A seguir segue as normas de submissão de artigos para a revista:



# EXPERIMENTAL GERONTOLOGY

**p.4 ISSN:** 0531-5565

# AUTHOR INFORMATION PACK

# TABLE OF CONTENTS

•	Description	
•	Audience	p.1
•	Impact Factor Abstracting and Indexing	p.1
•	Editorial Board Guide for Authors	p.1
•	Guide for Authors	p.1
		p.2



# 9. DESCRIPTION

*Experimental Gerontology* is a multidisciplinary journal for the publication of work from all areas of **biogerontology**, with an emphasis on studies focused at the systems level of investigation, such as whole organisms (e.g. invertebrate genetic models), immune, endocrine and cellular systems, as well as whole population studies (e.g. epidemiology).

The journal also publishes studies into the **behavioural** and **cognitive** consequences of **aging**, where a clear biological causal link is implicated. Studies aimed at bridging the gap between basic and clinical aspects of **gerontology**, such as papers on the basic aspects of **age-related diseases**, are welcomed, as is research orientated toward the modulation of the **aging process**. Original research manuscripts, special issues, short reports, reviews, mini-reviews, and correspondence are published. Manuscripts on social aspects of aging and reports on clinical studies do not fall within the scope of the journal.

# **10. AUDIENCE**

Gerontologists, Biological Scientists, Immunologists, Endocrinologists, Neuroscientists, Pathologists, Nutritionists

# **11.IMPACT FACTOR**

2019: 3.376 © Clarivate Analytics Journal Citation Reports 2020

# **12. ABSTRACTING AND INDEXING**

BIOSIS Citation Index PubMed/Medline Nutrition Abstracts Embase Elsevier BIOBASE Current Contents EMBiology Abstracts in Social Gerontology Scopus

# **13. EDITORIAL BOARD**

Editor-in-Chief

Christiaan Leeuwenburgh, University of Florida College of Medicine, Gainesville, Florida, United States

Section Editors Cell

Biology

Werner Zwerschke, University of Innsbruck, Innsbruck, Austria

Demography, Epidemiology and Population Genetics

Eva Grill, Ludwig Maximilians University Munich, Munich, Germany

#### Immunology and Inflammation

Daniela Frasca, University of Miami School of Medicine, Miami, Florida, United States

#### Invertebrate Models and Genetics

Sean P. Curran, University of Southern California Davis School of Gerontology, Los Angeles, California, United States

#### Musculoskeletal System and Exercise

Christiaan Leeuwenburgh, University of Florida College of Medicine, Gainesville, Florida, United States

Emanuele Marzetti, Catholic University of the Sacred Heart Institute of Public Health, Rome, Italy

Anna-Karin Welmer, Karolinska Institute Department of Neurobiology Care Sciences and Society, Huddinge, Sweden

#### Neuroscience

Stéphane Baudry, University Libre of Brussels, Laboratory of Applied Biology and Neurophysiology, Faculty for Motor Sciences, Brussels, Belgium

Tibor Hortobagyi, University of Groningen Center for Human Movement Sciences, Groningen, Netherlands

#### Nutrition, Metabolism and Endocrinology

Cheryl A. Conover, Mayo Foundation for Medical Education and Research, Rochester, Minnesota, United States

Michał Masternak, University of Central Florida College of Medicine, Orlando, Florida, United States

#### **Previous Editors-in-Chief**

Thomas.E Johnson, University of Colorado Boulder, Boulder, Colorado, United States

#### Editorial Board

David Allison, Indiana University Bloomington, Bloomington, Indiana, United States

Stephen Always, The University of Tennessee Health Science Center College of Health Professions Department of Physical Therapy, Memphis, Tennessee, United States

Cédric Annweiler, University of Angers, Angers, France

Stephen D. Anton, University of Florida Department of Aging and Geriatric Research, Gainesville, Florida, United States

Mylène Aubertin-Leheudre, University of Quebec in Montreal Department of Human Kinetics, Montréal, Quebec, Canada

Ivan Bautmans, VUB University, Brussel, Belgium

Olivier Beauchet, McGill University Department of Medicine, Montréal, Quebec, Canada

Vilhelm Bohr, Biomedical Research Center, Lab. of Molecular Gerontology, National Institute of Aging (NIA), Baltimore, Maryland, United States

Holly Brown-Borg, University of North Dakota, Grand Forks, North Dakota, United States Thomas Buford, The University of Alabama at Birmingham, Birmingham, Alabama, United States Gillian Butler-Browne, Sorbonne University Pierre and Marie Curie Campus, Paris, France

Eduardo Cadore, Federal University of Rio Grande do Sul School of Physical Education Physiotherapy and Dance, PORTO ALEGRE, Brazil

Shi-qing Cai, Institute of Neuroscience Chinese Academy of Sciences, Shanghai, China

Riccardo Calvani, University Hospital Agostino Gemelli Department of Geriatrics Neurosciences and Orthopedics, Roma, Italy

Judith Campisi, Buck Institute for Research on Aging, Novato, California, United States

James Carey, University of California Davis, Davis, California, United States

Siriporn Chattipakorn, Chiang Mai University Faculty of Medicine Cardiac Electrophysiology Research and Training Center, Chiang Mai, Thailand

Liang-Kung Chen, National Yang-Ming University School of Medicine, Taipei, Taiwan

Rita Effros, University of California Los Angeles Health System, Los Angeles, California, United States

Luigi Ferrucci, National Institutes of Health (NIH), National Institute on Aging, Baltimore, Maryland, United States

Thomas C. Foster, University of Florida, Gainesville, Florida, United States

Claudio Franceschi, University of Bologna, Bologna, Italy

William M. Freeman, Oklahoma Medical Research Foundation, Oklahoma City, Oklahoma, United States

Tamas Fülöp, Sherbrooke University of Applied Sciences Institute, Quebec, Canada

Efstathios Gonos, National Hellenic Research Foundation, Athens, Greece

Vassilis Gorgoulis, National and Kapodistrian University of Athens School of Medicine, Athens, Greece

Jorg Goronzy, Stanford University Division of Immunology and Rheumatology, Stanford, California, United States

Kristin Gribble, Marine Biological Laboratory, Woods Hole, Massachusetts, United States

Beatrix Grubeck-Loebenstein, University of Innsbruck Research Institute for Biomedical Aging Research, Innsbruck, Austria

Mark W. Hamrick, Augusta University Medical College of Georgia, Augusta, Georgia, United States

Josep Maria Haro, University of Barcelona, Barcelona, Spain

Lorna Harries, University of Exeter, Exeter, United Kingdom

Christian Humpel, Medical University of Innsbruck, Innsbruck, Austria

S. Michal Jazwinski, Tulane University, New Orleans, Louisiana, United States

Tom Kirkwood, Newcastle University, Newcastle Upon Tyne, United Kingdom

Dimitris Kletsas, Ethniko Kentro Ereunas Physikon Epistemon 'Demokritos', Athens, Greece

Carel Meskers, VU Amsterdam, Amsterdam, Netherlands

Richard Miller, University of Michigan Medical School, Ann Arbor, Michigan, United States

Heinz Osiewacz, Goethe University Frankfurt, Frankfurt am Main, Germany

Giuseppe Passarino, Institute for Biomedical Research and Innovation National Research Council Cosenza Branch, Mangone, Italy

Giuseppe Passarino, Institute for Biomedical Research and Innovation National Research Council Cosenza Branch, Mangone, Italy

Graham Pawelec, Eberhard Karls University Tübingen, Munich, Germany

Anna Picca, Catholic University of the Sacred Heart Institute of Internal and Geriatric Medicine, Roma, Italy

Kaisu Pitkälä, University of Helsinki, Helsinki, Finland

M. Cristina Polidori, Medical Faculty of the University of Cologne, Köln, Germany

Arlan Richardson, The University of Oklahoma Health Sciences Center, Oklahoma City, Oklahoma, United States Erik Rosendahl, Umea University Department of Community Medicine and Rehabilitation, Umeå, Sweden Jaime Ross, University of Rhode Island, Kingston, Rhode Island, United States

Markus Schosserer, University of Natural Resources and Life Sciences Vienna, Wien, Austria

Colin Selman, University of Glasgow, Glasgow, Scotland, United Kingdom

Joseph Signorile, University of Miami School of Education and Human Development Department of Kinesiology and Sport Sciences, Coral Gables, Florida, United States

Andreas Simm, University Hospital Halle, Department of Cardiac Surgery, Halle (Saale), Germany

Rafael Solana, National University of Cordoba, Cordoba, Argentina Ioakim Spyridopoulos, Newcastle University, Newcastle Upon Tyne, United Kingdom Matt Stock, University of Central Florida, Orlando, Florida, United States Josef Syka, Czech Academy of Sciences, Praha, Czech Republic Tamar Tchkonia, Mayo Clinic Robert and Arlene Kogod Center on Aging, Rochester, Minnesota, United States LaDora Thompson, Boston University, Boston, Massachusetts, United States John Tower, University of Southern California, Los Angeles, California, United States Stella Trompet, Leiden University Medical Center, Leiden, Netherlands Diana Van Heemst, Leiden University Medical Center Gerontology and Geriatrics, Leiden, Netherlands Nicola Veronese, University of Palermo School of Medicine and Surgery, Palermo, Italy Thomas Von Zglinicki, Newcastle Upon Tyne Hospitals NHS Foundation Trust, Newcastle Upon Tyne, United Kingdom Ashley Webb, Brown University, Providence, Rhode Island, United States

Jean Woo, The Chinese University of Hong Kong Department of Medicine and Therapeutics, Hong Kong, Hong Kong

# Your Paper Your Way

We now differentiate between the requirements for new and revised submissions. You may choose to submit your manuscript as a single Word or PDF file to be used in the refereeing process. Only when your paper is at the revision stage, will you be requested to put your paper in to a 'correct format' for acceptance and provide the items required for the publication of your article. *To find out more, please visit the Preparation section below.* 

# Submission checklist

You can use this list to carry out a final check of your submission before you send it to the journal for review. 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 the Internet)
- A competing interests statement is provided, even if the authors have no competing interests to declare
- 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.

# **BEFORE YOU BEGIN**

# Ethics in publishing

Please see our information pages on Ethics in publishing and Ethical guidelines for journal publication.

# Declaration of interest

All authors must disclose any financial and personal relationships with other people or organizations that could inappropriately influence (bias) their work. Examples of potential competing interests include employment, consultancies, stock ownership, honoraria, paid expert testimony, patent applications/registrations, and grants or other funding. Authors must disclose any interests in two places: 1. A summary declaration of interest statement in the title page file (if double-blind) or the manuscript file (if single-blind). If there are no interests to declare then please state this: 'Declarations of interest: none'. This summary statement will be ultimately published if the article is accepted. 2. Detailed disclosures as part of a separate Declaration of Interest form, which forms part of the

journal's official records. It is important for potential interests to be declared in both places and that the information matches. More information. *Submission declaration and verification* 

Submission of an article implies that the work described has not been published previously (except in the form of an abstract, a published lecture or academic thesis, see 'Multiple, redundant or concurrent publication' for more information), that it is not under consideration for publication elsewhere, that

its publication is approved by all authors and tacitly or explicitly by the responsible authorities where the work was carried out, and that, if accepted, it will not be published elsewhere in the same form, in English or in any other language, including electronically without the written consent of the copyright- holder. To verify originality, your article may be checked by the originality detection service Crossref Similarity Check.

#### Preprints

Please note that preprints can be shared anywhere at any time, in line with Elsevier's sharing policy. Sharing your preprints e.g. on a preprint server will not count as prior publication (see 'Multiple, redundant or concurrent publication' for more information). Use of inclusive language

Inclusive language acknowledges diversity, conveys respect to all people, is sensitive to differences, and promotes equal opportunities. Content should make no assumptions about the beliefs or commitments of any reader; contain nothing which might imply that one individual is superior to another on the grounds of age, gender, race, ethnicity, culture, sexual orientation, disability or health condition; and use inclusive language throughout. Authors should ensure that writing is free from bias, stereotypes, slang, reference to dominant culture and/or cultural assumptions. We advise to seek gender neutrality by using plural nouns ("clinicians, patients/clients") as default/wherever possible to avoid using "he, she," or "he/she." We recommend avoiding the use of descriptors that refer to personal attributes such as age, gender, race, ethnicity, culture, sexual orientation, disability or health condition unless they are relevant and valid. These guidelines are meant as a point of reference to help identify appropriate language but are by no means exhaustive or definitive.

# Author contributions

For transparency, we encourage authors to submit an author statement file outlining their individual contributions to the paper using the relevant CRediT roles: Conceptualization; Data curation; Formal analysis; Funding acquisition; Investigation; Methodology; Project administration; Resources; Software; Supervision; Validation; Visualization; Roles/Writing - original draft; Writing - review & editing. Authorship statements should be formatted with the names of authors first and CRediT role(s) following. More details and an example

### Changes to authorship

Authors are expected to consider carefully the list and order of authors **before** submitting their manuscript and provide the definitive list of authors at the time of the original submission. Any addition, deletion or rearrangement of author names in the authorship list should be made only **before** the manuscript has been accepted and only if approved by the journal Editor. To request such a change, the Editor must receive the following from the **corresponding author**: (a) the reason for the change in author list and (b) written confirmation (e-mail, letter) from all authors that they agree with the addition, removal or rearrangement. In the case of addition or removal of authors, this includes confirmation from the author being added or removed.

Only in exceptional circumstances will the Editor consider the addition, deletion or rearrangement of authors **after** the manuscript has been accepted. While the Editor considers the request, publication of the manuscript will be suspended. If the manuscript has already been published in an online issue, any requests approved by the Editor will result in a corrigendum.

Article transfer service

This journal is part of our Article Transfer Service. This means that if the Editor feels your article is more suitable in one of our other participating journals, then you may be asked to consider transferring the article to one of those. If you agree, your article will be transferred automatically on your behalf with no need to reformat. Please note that your article will be reviewed again by the new journal. More information. *Copyright* 

Upon acceptance of an article, authors will be asked to complete a 'Journal Publishing Agreement' (see more information on this). An e-mail will be sent to the corresponding author confirming receipt of the manuscript together with a 'Journal Publishing Agreement' form or a link to the online version of this agreement.

Subscribers may reproduce tables of contents or prepare lists of articles including abstracts for internal circulation within their institutions. Permission of the Publisher is required for resale or distribution outside the institution and for all other derivative works, including compilations and translations. If

excerpts from other copyrighted works are included, the author(s) must obtain written permission from the copyright owners and credit the source(s) in the article. Elsevier has preprinted forms for use by authors in these cases.

For gold open access articles: Upon acceptance of an article, authors will be asked to complete an 'Exclusive License Agreement' (more information). Permitted third party reuse of gold open access articles is determined by the author's choice of user license.

### Author rights

As an author you (or your employer or institution) have certain rights to reuse your work. More information.

#### Elsevier supports responsible sharing

Find out how you can share your research published in Elsevier journals.

# Role of the funding source

You are requested to identify who provided financial support for the conduct of the research and/or preparation of the article and to briefly describe the role of the sponsor(s), if any, in study design; in the collection, analysis and interpretation of data; in the writing of the report; and in the decision to submit the article for publication. If the funding source(s) had no such involvement then this should be stated. *Open access* 

Please visit our Open Access page for more information.

## Elsevier Researcher Academy

Researcher Academy is a free e-learning platform designed to support early and midcareer researchers throughout their research journey. The "Learn" environment at Researcher Academy offers several interactive modules, webinars, downloadable guides and resources to guide you through the process of writing for research and going through peer review. Feel free to use these free resources to improve your submission and navigate the publication process with ease.

### Language (usage and editing services)

Please write your text in good English (American or British usage is accepted, but not a

mixture of these). Authors who feel their English language manuscript may require editing to eliminate possible grammatical or spelling errors and to conform to correct scientific English may wish to use the English Language Editing service available from Elsevier's Author Services.

### Submission

Our online submission system guides you stepwise through the process of entering your article details and uploading your files. The system converts your article files to a single PDF file used in the peer-review process. Editable files (e.g., Word, LaTeX) are required to typeset your article for final publication. All correspondence, including notification of the Editor's decision and requests for revision, is sent by e-mail.

#### Referees

Please submit the names and institutional e-mail addresses of several potential referees. For more details, visit our Support site. Note that the editor retains the sole right to decide whether or not the suggested reviewers are used. PREPARATION

# NEW SUBMISSIONS

Submission to this journal proceeds totally online and you will be guided stepwise through the creation and uploading of your files. The system automatically converts your files to a single PDF file, which is used in the peer-review process.

As part of the Your Paper Your Way service, you may choose to submit your manuscript as a single file to be used in the refereeing process. This can be a PDF file or a Word document, in any format or lay- out that can be used by referees to evaluate your manuscript. It should contain high enough quality figures for refereeing. If you prefer to do so, you may still provide all or some of the source files at the initial submission. Please note that individual figure files larger than 10 MB must be uploaded separately.

### References

There are no strict requirements on reference formatting at submission. References can be in any style or format as long as the style is consistent. Where applicable, author(s) name(s), journal title/ book title, chapter title/article title, year of publication, volume number/book chapter and the article number or pagination must be present. Use of DOI is highly encouraged. The reference style used by the journal will be applied to the accepted article by Elsevier at the proof stage. Note that missing data will be highlighted at proof stage for the author to correct.

#### Formatting requirements

There are no strict formatting requirements but all manuscripts must contain the essential elements needed to convey your manuscript, for example Abstract, Keywords, Introduction, Materials and Methods, Results, Conclusions, Artwork and Tables with Captions.

If your article includes any Videos and/or other Supplementary material, this should be included in your initial submission for peer review purposes.

Divide the article into clearly defined sections.

### Figures and tables embedded in text

Please ensure the figures and the tables included in the single file are placed next to the relevant text in the manuscript, rather than at the bottom or the top of the file. The corresponding caption should be placed directly below the figure or table.

# Peer review

This journal operates a single anonymized review process. All contributions will be initially assessed by the editor for suitability for the journal. Papers deemed suitable are then typically sent to a minimum of two independent expert reviewers to assess the scientific quality of the paper. The Editor is responsible for the final decision regarding acceptance or rejection of articles. The Editor's decision is final. Editors are not involved in decisions about papers which they have written themselves or have been written by family members or colleagues or which relate to products or services in which the editor has an interest. Any such submission is subject to all of the journal's usual procedures, with peer review handled independently of the relevant editor and their research groups. More information on types of peer review.

REVISED SUBMISSIONS

### Use of word processing software

Regardless of the file format of the original submission, at revision you must provide us with an editable file of the entire article. Keep the layout of the text as simple as possible. Most formatting codes will be removed and replaced on processing the article. The electronic text should be prepared in a way very similar to that of conventional manuscripts (see also the Guide to Publishing with Elsevier). See also the section on Electronic artwork.

To avoid unnecessary errors you are strongly advised to use the 'spell-check' and 'grammar-check' functions of your word processor.

### Article structure

#### Subdivision - numbered sections

Divide your article into clearly defined and numbered sections. Subsections should be numbered

1.1 (then 1.1.1, 1.1.2, ...), 1.2, etc. (the abstract is not included in section numbering). Use this numbering also for internal cross-referencing: do not just refer to 'the text'. Any subsection may be given a brief heading. Each heading should appear on its own separate line.

#### Introduction

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

#### Material and methods

Provide sufficient details to allow the work to be reproduced by an independent researcher. Methods that are already published should be summarized, and indicated by a reference. If quoting directly from a previously published method, use quotation marks and also cite the source. Any modifications to existing methods should also be described.

# Theory/calculation

A Theory section should extend, not repeat, the background to the article already dealt with in the Introduction and lay the foundation for further work. In contrast, a Calculation section represents a practical development from a theoretical basis.

### Results

Results should be clear and concise.

### Discussion

This should explore the significance of the results of the work, not repeat them. A combined Results and 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 stand alone 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 in appendices 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.

# Essential title page information

• *Title*. Concise and informative. Titles are often used in information-retrieval systems. Avoid abbreviations and formulae where possible.

• *Author names and affiliations.* Please clearly indicate the given name(s) and family name(s) of each author and check that all names are accurately spelled. You can add your name between parentheses in your own script behind the English transliteration. Present the authors' affiliation addresses (where the actual work was done) below the names. Indicate all affiliations with a lower- case superscript letter immediately after the author's name and in front of the appropriate address. Provide the full postal address of each affiliation, including the country name and, if available, the e-mail address of each author.

• *Corresponding author*. Clearly indicate who will handle correspondence at all stages of refereeing and publication, also post-publication. This responsibility includes answering any future queries about Methodology and Materials. Ensure that the e-mail address is given and that contact details are kept up to date by the corresponding author.

• *Present/permanent address*. If an author has moved since the work described in the article was done, or was visiting at the time, a 'Present address' (or 'Permanent address') may be indicated as a footnote to that author's name. The address at which the author actually did the work must be retained as the main, affiliation address. Superscript Arabic numerals are used for such footnotes.

# **Highlights**

Highlights are mandatory for this journal as they help increase the discoverability of your article via search 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 abstract is required. The abstract should state briefly 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). Also, non-standard or uncommon abbreviations should be avoided, but if essential they must be defined at their first mention in the abstract itself.

Graphical abstract

Although a graphical abstract is optional, its use is encouraged as it draws more attention to the online article. The graphical abstract should summarize the contents of the article in a concise, pictorial form designed to capture the attention of a wide readership. Graphical abstracts should be submitted as a separate file in the online submission system. Image size: Please provide an image with a minimum of  $531 \times 1328$  pixels (h  $\times$  w) or proportionally more. The image should be readable at a size of  $5 \times 13$  cm using a regular screen resolution of 96 dpi. Preferred file types: TIFF, EPS, PDF or MS Office files. You can view Example Graphical Abstracts on our information site.

Authors can make use of Elsevier's Illustration Services to ensure the best presentation of their images and in accordance with all technical requirements.

## 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 sparing with abbreviations: only abbreviations firmly established in the field may be eligible. These keywords will be used for indexing purposes.

### Abbreviations

Define abbreviations that are not standard in this field in a footnote to be placed on the first page of the article. Such abbreviations that are unavoidable in the abstract must be defined at their first mention there, as well as in the footnote. Ensure consistency of abbreviations throughout the article.

#### Acknowledgements

Collate acknowledgements in a separate section at the end of the article before the references and do not, therefore, include them on the title page, as a footnote to the title or otherwise. List here those individuals who provided help during the research (e.g., providing language help, writing assistance or 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. When funding is from a block grant or other resources available to a university, college, or other research institution, 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, or not-for-profit sectors.

Units

Follow internationally accepted rules and conventions: use the international system of units (SI). If other units are mentioned, please give their equivalent in SI.

### Math formulae

Please submit math equations as editable text and not as images. Present simple formulae in line with normal text where possible and use the solidus (/) instead of a

horizontal line for small fractional terms, e.g., X/Y. In principle, variables are to be presented in italics. Powers of e are often more conveniently denoted by exp. Number consecutively any equations that have to be displayed separately from the text (if referred to explicitly in the text).

Footnotes

Footnotes should be used sparingly. Number them consecutively throughout the article. Many word processors build footnotes into the text, and this feature may be used. Should this not be the case, indicate the position of footnotes in the text and present the footnotes themselves separately at the end of the article.

Artw ork Electro nic artwor k Gener al points

- Make sure you use uniform lettering and sizing of your original artwork.
- Preferred fonts: Arial (or Helvetica), Times New Roman (or Times), Symbol, Courier.
- Number the illustrations according to their sequence in the text.
- Use a logical naming convention for your artwork files.
- Indicate per figure if it is a single, 1.5 or 2-column fitting image.
- For Word submissions only, you may still provide figures and their captions, and tables within a single file at the revision stage.
- Please note that individual figure files larger than 10 MB must be provided in separate source files.

### A detailed guide on electronic artwork is available.

# You are urged to visit this site; some excerpts from the detailed information are given here.

#### Formats

Regardless of the application used, when your electronic artwork is finalized, please 'save as' or convert the images to one of the following formats (note the resolution requirements for line drawings, halftones, and line/halftone combinations given below): EPS (or PDF): Vector drawings. Embed the font or save the text as 'graphics'. TIFF (or JPG): Color or grayscale photographs (halftones): always use a minimum of 300 dpi. TIFF (or JPG): Bitmapped line drawings: use a minimum of 1000 dpi. TIFF (or JPG): Combinations bitmapped line/half-tone (color or grayscale): a minimum of 500 dpi is required.

Please do not:

- Supply files that are optimized for screen use (e.g., GIF, BMP, PICT, WPG); the resolution is too low.
- Supply files that are too low in resolution.
- Submit graphics that are disproportionately large for the content.

#### Color artwork

Please make sure that artwork files are in an acceptable format (TIFF (or JPEG), EPS (or PDF) or MS Office files) and with the correct resolution. If, together with your accepted article, you submit usable color figures then Elsevier will ensure, at no additional charge, that these figures will appear in color online (e.g., ScienceDirect and other sites) in addition to color reproduction in print. Further information on the preparation of

# electronic artwork.

### Figure captions

Ensure that each illustration has a caption. A caption 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 these references 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. Please note 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 the DOI 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. Any further 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 a different heading if desired, or can be included in the reference list.

#### Data references

**For reference style 2 Harvard:** [dataset] Oguro, M., Imahiro, S., Saito, S., Nakashizuka, T., 2015. Mortality data for Japanese oak wilt disease and surrounding forest compositions. Mendeley Data, v1. http://dx.doi.org/10.17632/xwj98nb39r.1.

### 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 in the text) to other articles in the same Special Issue.

#### Reference management software

Most Elsevier journals have their reference template available in many of the most popular reference management software products. These include all products that support Citation Style Language styles, such as Mendeley. Using citation plug-ins from these products, authors only need to select the appropriate journal template when preparing their article, after which citations and bibliographies will be automatically formatted in the journal's style. If no template is yet available for this journal, please follow the format of the sample references and citations as shown in this Guide. If you use reference management software, please ensure that you remove all field codes before submitting the electronic manuscript. More information on how to remove field codes from different reference management software.

Users of Mendeley Desktop can easily install the reference style for this journal by clicking the following link:

http://open.mendeley.com/use-citation-style/experimental-gerontology

When preparing your manuscript, you will then be able to select this style using the Mendeley plug- ins for Microsoft Word or LibreOffice.

### Reference formatting

There are no strict requirements on reference formatting at submission. References can be in any style or format as long as the style is consistent. Where applicable, author(s) name(s), journal title/ book title, chapter title/article title, year of publication, volume number/book chapter and the article number or pagination must be present. Use of DOI is highly encouraged. The reference style used by the journal will be applied to the accepted article by Elsevier at the proof stage. Note that missing data will be highlighted at proof stage for the author to correct. If you do wish to format the references yourself they should be arranged according to the following examples:

Ref ere nce Ref ere nce styl e

Text: Indicate references by first author last name and date of publication in parentheses in line with the text. The actual authors can be referred to, however the complete reference must always be given. Example: 'Barnaby (2001) obtained a different result...' Reference to a journal publication:

J. van der Geer, J.A.J. Hanraads, R.A. Lupton, 2010. The art of writing a scientific article, J. Sci. Commun. 163,51 59.

Reference to a book: W. Strunk Jr., E.B. White, 2000. The Elements of Style, fourth ed., Longman, New York. Reference to a chapter in an edited book: G.R. Mettam, L.B. Adams, 2009. How to prepare an electronic version of your article, in: B.S. Jones, R.Z. Smith (Eds.), Introduction to the Electronic Age, E-Publishing Inc., New York, 281304.

## Journal abbreviations source

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

# Data visualization

Include interactive data visualizations in your publication and let your readers interact and engage more closely with your research. Follow the instructions here to find out about available data visualization options and how to include them with your article.

# Supplementary material

Supplementary material such as applications, images and sound clips, can be published with your article to enhance it. Submitted supplementary items are published exactly as they are received (Excel or PowerPoint files will appear as such online). Please submit your material together with the article and supply a concise, descriptive caption for each supplementary file. If you wish to make changes to supplementary material during any stage of the process, please make sure to provide an updated file. Do not annotate any corrections on a previous version. Please switch off the 'Track Changes' option in Microsoft Office files as these will appear in the published version.

### Research data

This journal encourages and enables you to share data that supports your research publication where appropriate, and enables you to interlink the data with your published articles. Research data refers to the results of observations or experimentation that validate research findings. To facilitate reproducibility and data reuse, this journal also encourages you to share your software, code, models, algorithms, protocols, methods and other useful materials related to the project.

Below are a number of ways in which you can associate data with your article or make a statement about the availability of your data when submitting your manuscript. If you are sharing data in one of these ways, you are encouraged to cite the data in your manuscript and reference list. Please refer to the "References" section for more information about data citation. For more information on depositing, sharing and using research data and other relevant research materials, visit the research data page.

# Data linking

If you have made your research data available in a data repository, you can link your article directly to the dataset. Elsevier collaborates with a number of repositories to link articles on ScienceDirect with relevant repositories, giving readers access to underlying data that gives them a better understanding of the research described.

There are different ways to link your datasets to your article. When available, you can directly link your dataset to your article by providing the relevant information in the submission system. For more information, visit the database linking page.

For supported data repositories a repository banner will automatically appear next to

your published article on ScienceDirect.

In addition, you can link to relevant data or entities through identifiers within the text of your manuscript, using the following format: Database: xxxx (e.g., TAIR: AT1G01020; CCDC: 734053; PDB: 1XFN).

### Mendeley Data

This journal supports Mendeley Data, enabling you to deposit any research data (including raw and processed data, video, code, software, algorithms, protocols, and methods) associated with your manuscript in a free-to-use, open access repository. During the submission process, after uploading your manuscript, you will have the opportunity to upload your relevant datasets directly to *Mendeley Data*. The datasets will be listed and directly accessible to readers next to your published article online.

For more information, visit the Mendeley Data for journals page.

### Data in Brief

You have the option of converting any or all parts of your supplementary or additional raw data into a data article published in *Data in Brief*. A data article is a new kind of article that ensures that your data are actively reviewed, curated, formatted, indexed, given a DOI and made publicly available to all upon publication (watch this video describing the benefits of publishing your data in *Data in Brief*). You are encouraged to submit your data article for *Data in Brief* as an additional item directly alongside the revised version of your manuscript. If your research article is accepted, your data article will automatically be transferred over to *Data in Brief* where it will be editorially reviewed, published open access and linked to your research article on ScienceDirect. Please note an open access fee is payable for publication in *Data in Brief*. Full details can be found on the Data in Brief website. Please use this template to write your *Data in Brief* data article.

#### Data statement

To foster transparency, we encourage you to state the availability of your data in your submission. This may be a requirement of your funding body or institution. If your data is unavailable to access or unsuitable to post, you will have the opportunity to indicate why during the submission process, for example by stating that the research data is confidential. The statement will appear with your published article on ScienceDirect. For more information, visit the Data Statement page.

# AFTER ACCEPTANCE

# Online proof correction

To ensure a fast publication process of the article, we kindly ask authors to provide us with their proof corrections within two days. Corresponding authors will receive an e-mail with a link to our online proofing system, allowing annotation and correction of proofs online. The environment is similar to MS Word: in addition to editing text, you can also comment on figures/tables and answer questions from the Copy Editor. Web-based proofing provides a faster and less error-prone process by allowing you to directly type your corrections, eliminating the potential introduction of errors.

If preferred, you can still choose to annotate and upload your edits on the PDF version. All instructions for proofing will be given in the e-mail we send to authors, including alternative methods to the online version and PDF.

We will do everything possible to get your article published quickly and accurately. Please use this proof only for checking the typesetting, editing, completeness and correctness of the text, tables and figures. Significant changes to the article as accepted for publication will only be considered at this stage with permission from the Editor. It is important to ensure that all corrections are sent back to us in one communication. Please check carefully before replying, as inclusion of any subsequent corrections cannot be guaranteed. Proofreading is solely your responsibility. *Offprints* 

The corresponding author will, at no cost, receive a customized Share Link providing 50 days free access to the final published version of the article on ScienceDirect. The Share Link can be used for sharing the article via any communication channel, including email and social media. For an extra charge, paper offprints can be ordered via the offprint order form which is sent once the article is accepted for publication. Both corresponding and co-authors may order offprints at any time via Elsevier's Author Services. Corresponding authors who have published their article gold open access do not receive a Share Link as their final published version of the article is available open access on ScienceDirect and can be shared through the article DOI link. AUTHOR INQUIRIES

AUTHON INQUINES

Visit the Elsevier Support Center to find the answers you need. Here you will find everything from Frequently Asked Questions to ways to get in touch.

You can also check the status of your submitted article or find out when your accepted article will be published.

© Copyright 2018 Elsevier | https://www