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**Efeito da fotobiomodulação
associada ao treinamento de força
em homens idosos: um ensaio
clínico randomizado duplo-cego
placebo-controlado**

**Universidade Federal de Ciências da Saúde
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Efeito da fotobiomodulação associada ao treinamento de força em homens idosos: um ensaio clínico randomizado duplo-cego placebo-controlado

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placebo-controlado**

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RESUMO

Introdução: A terapia de fotobiomodulação tem demonstrado capacidade de aumentar o desempenho físico e otimizar a recuperação pós exercício. Além disto, quando associada ao treinamento de força, tem demonstrado melhoras nos ganhos induzidos pelo exercício na população jovem. Entretanto, este efeito na população idosa permanece incerto. **Objetivo:** O objetivo do presente estudo foi de investigar o efeito da associação ao treinamento de força em idosos saudáveis na força e na massa muscular dos extensores de joelho e na capacidade funcional de homens idosos. **Metodologia:** Foi realizado um ensaio clínico randomizado duplo-cego placebo-controlado. Homens idosos saudáveis (idade 60-80 anos) realizaram 12 semanas de treinamento de força (2x/semana) com aplicação de fotobiomodulação placebo (n = 13) ou ativa (n = 11) no quadríceps (850nm, 30J por ponto, 8 pontos por membro) antes de cada sessão. Os seguintes testes foram realizados antes e depois da intervenção: teste de uma repetição máxima (1 RM) dos exercícios “leg press” e extensão de joelhos, pico de torque concêntrico e isométrico, espessura muscular dos músculos reto femoral (RF) e vasto lateral (VL), teste “timed up-and-go” (TUG) e teste de sentar e levantar da cadeira (TSL). **Resultados:** Houve melhoras significativas em todas as variáveis para ambos os grupos após a intervenção ($p < 0,05$), com exceção da espessura muscular do RF para o grupo placebo ($p = 0,09$). Tamanhos de efeito grandes ($ES > 0,8$) foram encontrados para 1 RM de “leg press” e extensão de joelhos e TSL para ambos os grupos, assim como para o teste TUG para o grupo fotobiomodulação ativa. Pico de torque isocinético para ambos os grupos e o teste TUG para o grupo placebo apresentaram tamanhos de efeito moderados ($ES > 0,5$). Pico de torque isométrico e espessuras musculares obtiveram pequenos tamanhos de efeito ($ES > 0,2$). A análise de inferência baseada em magnitude não demonstrou nenhuma vantagem da terapia ativa em relação à placebo. **Conclusão:** A aplicação de fotobiomodulação com os parâmetros utilizados no presente estudo não proporciona efeitos adicionais aos ganhos de força, massa muscular e capacidade funcional gerados pelo treinamento de força em idosos saudáveis.

Palavras-chave: fototerapia; laser de baixa potência; treinamento de resistência; envelhecimento.

ABSTRACT

Introduction: Photobiomodulation therapy (PBMT) has demonstrated capacity to enhance exercise performance and post-exercise recovery. Moreover, when associated to resistance training, it has shown a capacity to optimize training induced gains in the young population. However, its effect in the elderly remains unclear. **Objective:** The purpose of this study was to investigate the effects of PBMT combined to resistance training on knee extensor muscle mass, strength and physical function in elderly men. **Methodology:** A randomized double-blinded placebo-controlled trial was developed. Healthy elderly men (age 60-80 years) completed 12 weeks of resistance training (2x/week) with application of placebo (n = 13) or active PBMT (n = 11) on quadriceps muscles (850 nm, 30 J per point, 8 points per limb) before each training session. Leg press and leg extension one-repetition maximum (1 RM) tests, isometric and concentric peak torques, rectus femoris (RF) and vastus lateralis (VL) muscle thickness, timed-up-and-go (TUG) and chair rise to standing (CRS) tests were performed before and after the intervention period. **Results:** There were significant improvements in all outcomes for both groups ($p < 0.05$), except for RF muscle thickness for the placebo group ($p = 0.09$). Large effect sizes ($ES > 0.8$) were observed for leg press and leg extension 1 RM and CRS tests for both groups, as well as for TUG test for PBMT group. Isokinetic peak torque for both groups and TUG for placebo group had moderate increases ($ES > 0.5$). Muscle thicknesses and isometric peak torque had small increases ($ES > 0.2$) in both groups. Magnitude-based inference analysis does not support any advantages for using PBMT compared to placebo. **Conclusion:** In conclusion, PBMT with the parameters used in this study did not provide any additional benefits in comparison to placebo application on muscle mass, strength and physical function of healthy elderly men engaged in a resistance-training program.

Keywords: phototherapy; low-level laser therapy; strength training; older.

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LISTA DE ABREVIATURAS E SIGLAS

Revisão da Literatura

ATP	Adenosina Trifosfato
CK	Creatina Quinase

Artigo

1-RM	One-repetition maximum
ATP	Adenosine triphosphate
CRS	Chair rise to standing
ICC	Intraclass correlation coefficient
J	Joules
PBMT	Photobiomodulation therapy
RF	Rectus femoris
TUG	Timed up-and-go
VL	Vastus lateralis

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1 INTRODUÇÃO

Já está bem estabelecido na literatura que a população idosa apresenta declínios nas funções fisiológicas de todos os sistemas corporais, incluindo o sistema musculoesquelético (ALI; GARCIA, 2014). O processo de envelhecimento diminui a massa e a qualidade muscular assim como altera a ativação neuromuscular (BARONI et al., 2013; BROCCA et al., 2017; STEVENS et al., 2003). Esta redução de massa muscular foi classificada inicialmente como “sarcopenia” (MITCHELL et al., 2012). Entretanto, o termo também passou a ser utilizado para nomear a perda de força muscular associada à perda de massa (MITCHELL et al., 2012). Atualmente, existem diferentes definições para o termo e não há um consenso mundial acerca de sua utilização ideal (MITCHELL et al., 2012). Segundo o Grupo de Estudos Europeu sobre Sarcopenia, esta é definida como uma síndrome caracterizada por perdas progressivas e generalizadas na massa e na força muscular que podem levar a efeitos adversos como incapacidade física, baixa qualidade de vida e morte (CRUZ-JENTOFT et al., 2010). A sarcopenia é considerada um componente-chave para o desenvolvimento da síndrome da fragilidade (CLEGG et al., 2013) e possui uma prevalência mundial de até 33% em adultos com mais de 50 anos (CRUZ-JENTOFT et al., 2014). Devido à relação entre massa e força muscular com a longevidade (EVANS; CAMPBELL, 1993; SRIKANTHAN; KARLAMANGLA, 2014), a manutenção e/ou o aumento da massa e da força muscular de idosos se tornou uma grande preocupação da área da saúde mundialmente (CRUZ-JENTOFT et al., 2014; EVANS; CAMPBELL, 1993).

O treinamento de força tem a capacidade de retardar parte do efeito do envelhecimento sobre o organismo humano (CSAPO; ALEGRE, 2016), levando a melhoras importantes na força muscular e na funcionalidade tanto em idosos saudáveis quanto em idosos sarcopênicos e frágeis (CSAPO; ALEGRE, 2016; LOPEZ et al., 2017; VAN ROIE et al., 2013; YOSHIMURA et al., 2017). O treinamento de força também gera aumentos na espessura e na área de secção transversa muscular (CORREA et al., 2013; CSAPO; ALEGRE, 2016), sendo que estes ganhos variam de 1,5% a 15,6% (CSAPO; ALEGRE, 2016; STEWART; SAUNDERS; GREIG, 2014). Além disto, diferentes estratégias ergogênicas vem sendo estudadas com o objetivo de maximizar os ganhos de força muscular e, principalmente, massa muscular

induzidas pelo treinamento, como suplementação nutricional (MALAFARINA et al., 2013), reposição hormonal (HILDRETH et al., 2013), entre outros.

A terapia de fotobiomodulação vem sendo estudada como um recurso ergogênico inovador, não-invasivo e não-farmacológico (VANIN et al., 2018). O seu efeito se dá através da absorção da luz por cromóforos específicos da mitocôndria, o que gera uma série de reações intracelulares culminando em aumento da produção de ATP e do metabolismo celular, dentre outros (FERRARESI et al., 2015a; KARU, 2010). Consequentemente, as evidências apontam a fotobiomodulação como uma terapia eficaz para a melhora do desempenho físico e da recuperação pós-exercício (FERRARESI; HUANG; HAMBLIN, 2016; VANIN et al., 2018). Além disto, estudos que associaram a fotobiomodulação ao treinamento de força encontraram um maior ganho de força e massa muscular em comparação ao treinamento associado ao placebo (BARONI et al., 2015; FERRARESI et al., 2011, 2016; VANIN et al., 2016). Entretanto, grande parte da literatura acerca dos efeitos da fotobiomodulação no desempenho foram encontrados em jovens e/ou atletas (BARONI et al., 2015; FERRARESI et al., 2011, 2016; FERRARESI; HUANG; HAMBLIN, 2016; VANIN et al., 2016, 2018) e há poucos estudos com a população idosa, sendo que eles apresentam resultados contraditórios (TOMA et al., 2013, 2016; VASSÃO et al., 2016). Ao meu conhecimento, há apenas um estudo avaliando o efeito crônico da aplicação da fotobiomodulação associada ao treinamento de força em idosos e este não encontrou efeitos adicionais da terapia (TOMA et al., 2016). Contudo, o estudo não utiliza parâmetros apropriados, segundo a revisão sistemática mais recente (VANIN et al., 2018). Sendo assim, o real efeito da fotobiomodulação sobre as adaptações ao treinamento de força em idosos permanece incerto.

Devido à grande importância da força e da massa muscular para idosos (EVANS; CAMPBELL, 1993; SRIKANTHAN; KARLAMANGLA, 2014), tratamentos com potencial de aumentá-las são extremamente relevantes e devem ser investigados. Já que a fotobiomodulação se mostrou eficaz na maximização das adaptações induzidas pelo treinamento de força em jovens (BARONI et al., 2015; FERRARESI et al., 2011; VANIN et al., 2016), ela pode ser muito útil para idosos. Portanto, o objetivo do presente estudo foi verificar o efeito da associação da fotobiomodulação ao treinamento de força na força e na massa muscular dos extensores de joelho e na funcionalidade de idosos saudáveis.

2 REVISÃO DE LITERATURA

2.1 ENVELHECIMENTO

A proporção de pessoas com 60 anos ou mais está crescendo mundialmente (WHO, 2015). Estima-se que países em desenvolvimento, como Brasil, China e Índia, vão apresentar um aumento de 100% na sua população idosa em um pouco mais de 20 anos e calcula-se que os idosos passem a representar mais de 30% da população de países desenvolvidos no ano de 2050 (WHO, 2015). Segundo o Censo Brasileiro de 2015, houve um aumento de aproximadamente 46% no número de pessoas com 65 anos ou mais entre os anos 2005 e 2015 no Brasil (IBGE, 2016), o que reflete a importância de intervenções de saúde para esta população tanto a nível nacional quanto internacional.

O envelhecimento está associado a diversas modificações nos sistemas corporais, incluindo o sistema musculoesquelético (ALI; GARCIA, 2014; BROCCA et al., 2017; CARTER; CHEN; HOOD, 2015). Evidências sugerem uma diminuição significativa em massa, força e qualidade muscular (força por área muscular) a partir da quinta década de vida, principalmente em membros inferiores (JANSSEN et al., 2000; LYNCH et al., 1999). Estudos mostram que, a partir dos 70 anos, há diminuição em torno de 1% ao ano na massa muscular, enquanto a diminuição na força muscular pode chegar a até 5% ao ano (ALI; GARCIA, 2014; DELMONICO et al., 2009). Estas alterações ocorrem independentemente do nível de atividade física e da presença de doenças (BROCCA et al., 2017) e estão associadas a diversas alterações locais (BARONI et al., 2013; BROCCA et al., 2017; CRUZ-JENTOFT et al., 2010; MARZETTI et al., 2013) e sistêmicas (CRUZ-JENTOFT et al., 2010; DALLE; ROSSMEISLOVA; KOPPO, 2017; MARZETTI et al., 2013). Dentre as alterações locais estão a diminuição da área de secção transversa tanto da fibra muscular quanto de todo o grupo muscular, a redução no comprimento do fascículo e o aumento de tecidos gordurosos e não contráteis intramusculares, além de alterações mitocondriais (BARONI et al., 2013; BROCCA et al., 2017; CRUZ-JENTOFT et al., 2010; JANSSEN et al., 2000; LYNCH et al., 1999; NARICI et al., 2003; YOSHIMURA et al., 2017). Já as alterações sistêmicas incluem mudanças nos níveis hormonais e no mecanismo de ativação muscular, presença de estado

pró-inflamatório e de nutrição inadequada/má absorção (CRUZ-JENTOFT et al., 2010; DALLE; ROSSMEISLOVA; KOPPO, 2017; STEVENS et al., 2003).

A redução de massa muscular foi classificada inicialmente como “sarcopenia” (MITCHELL et al., 2012). Entretanto, o termo também passou a ser utilizado para nomear a perda de força muscular associada à perda de massa (MITCHELL et al., 2012). Atualmente, existem diferentes definições para o termo e não há um consenso mundial acerca de sua utilização ideal. Há classificações que também consideram a redução de funcionalidade consequente às diminuições de massa e força muscular, que são reconhecidas pelo Instituto de Saúde Nacional Americano (MITCHELL et al., 2012).

Neste documento, a classificação do Grupo de Estudos Europeu sobre Sarcopenia será utilizada, o qual define sarcopenia como uma síndrome caracterizada por perdas progressivas e generalizadas na massa e na força muscular que podem levar a efeitos adversos como incapacidade física, baixa qualidade de vida e morte (CRUZ-JENTOFT et al., 2010). Revisões sistemáticas recentes encontraram prevalências de até 17% e 33% de sarcopenia na população idosa do Brasil e de países desenvolvidos, respectivamente (CRUZ-JENTOFT et al., 2014; DIZ et al., 2017), dependendo da amostra estudada. Estas prevalências expressivas de sarcopenia associadas à relação direta entre perda de massa e força muscular com diminuição da funcionalidade e da longevidade (CRUZ-JENTOFT et al., 2010; EVANS; CAMPBELL, 1993; SRIKANTHAN; KARLAMANGLA, 2014), tornam intervenções que objetivem o aumento/a manutenção do sistema muscular de idosos ainda mais importantes (CRUZ-JENTOFT et al., 2014; EVANS; CAMPBELL, 1993).

2.2 TREINAMENTO DE FORÇA

O treinamento de força é uma modalidade de exercício que se popularizou nas últimas décadas devido à sua capacidade de aumento no desempenho físico (KRAEMER; RATAMESS, 2004). Esta melhora ocorre através do aumento de força, potência e massa muscular, assim como do desempenho muscular local, do controle motor, do equilíbrio e da coordenação (KRAEMER; RATAMESS, 2004). Inicialmente, o treinamento de força era realizado somente por pessoas que buscavam ganho de força e massa muscular, como fisiculturistas (KRAEMER; RATAMESS, 2004). Entretanto, a

partir de década de 1990, após uma melhor compreensão dos seus benefícios para a saúde, houve sua popularização e recomendação para a maioria das populações, incluindo a idosa (FIATARONE et al., 1994; KRAEMER; RATAMESS, 2004).

Apesar de não impedir o envelhecimento, o treinamento de força possui um grande potencial de minimizar os seus efeitos sobre o organismo humano (BORDE; HORTOBAGYI; GRANACHER, 2015). Diferentes revisões sistemáticas já foram realizadas e evidenciaram aumentos de força muscular em idosos saudáveis (BORDE; HORTOBAGYI; GRANACHER, 2015; CSAPO; ALEGRE, 2016), sarcopênicos (YOSHIMURA et al., 2017) e frágeis (LOPEZ et al., 2017). Os resultados reportados na literatura diferem muito em relação aos ganhos de força máxima em idosos saudáveis, variando entre 20-60% em estudos com diferentes amostras e metodologias (CADOORE et al., 2010, 2012, 2013; CORREA et al., 2013; CSAPO; ALEGRE, 2016; FERRARI et al., 2013; LIXANDRÃO et al., 2016; TIGGEMANN et al., 2016; VAN ROIE et al., 2013). O fortalecimento se deve, em parte, às adaptações de ordem neuromuscular, em especial à capacidade de ativação muscular (HÄKKINEN et al., 2000). Já do ponto de vista das adaptações estruturais, o treinamento de força gera aumento na área de secção transversa e na espessura muscular, os quais variam em torno de 1,5% a 15% (BORDE; HORTOBAGYI; GRANACHER, 2015; CSAPO; ALEGRE, 2016; STEWART; SAUNDERS; GREIG, 2014).

O treinamento de força também incrementa a funcionalidade da população idosa (TIGGEMANN et al., 2016; VAN ROIE et al., 2013). Apesar de o treinamento de força não envolver atividades funcionais, ele tem gerado melhoras em torno de 5-20% em testes funcionais como o “Timed up-and-go” e o “Teste de Sentar e Levantar” (TIGGEMANN et al., 2016; VAN ROIE et al., 2013). Estas melhoras em tarefas motoras consideradas mais próximas das demandas diárias do indivíduo representam a transferência dos ganhos obtidos com o treinamento (ISSURIN, 2013). Ou seja, o treinamento de força consegue melhorar a capacidade funcional de idosos sem exercitar estas atividades na rotina de treinos. Entretanto, é importante salientar que a periodização do treinamento (frequência, duração, volume, etc.) e as características individuais (idade, sexo, nível de atividade física) interferem diretamente na magnitude de

todas as adaptações ao treinamento de força (BORDE; HORTOBAGYI; GRANACHER, 2015; BROCCA et al., 2017).

Devido à relação da força e da massa muscular com a longevidade de idosos (EVANS; CAMPBELL, 1993; SRIKANTHAN; KARLAMANGLA, 2014), diferentes estratégias vem sendo investigadas de modo associado ao treinamento a fim de maximizar estas melhoras (MALAFARINA et al., 2013; TOMA et al., 2016). Contudo, estudos investigando os efeitos de suplementos alimentares (CHALÉ et al., 2013; MALAFARINA et al., 2013; TIELAND et al., 2012) e reposições hormonais (HILDRETH et al., 2013; KENNY et al., 2010) associados a diferentes modalidades de exercícios em idosos tem encontrado efeitos contraditórios. Desta forma, faz-se necessária a investigação de diferentes modalidades terapêuticas para a otimização das adaptações neuromusculares associadas ao treinamento em idosos.

2.3 TERAPIA DE FOTOBIMODULAÇÃO

“Terapia de fotobiomodulação” é o termo mais atual e abrangente para descrever a terapia até então conhecida como “fototerapia”, a qual envolve as intervenções com energia proveniente de fontes luminosas com objetivos terapêuticos (ANDERS; LANZAFAME; ARANY, 2015). As formas mais comuns de fotobiomodulação são as que utilizam LASER (Light Amplification by Stimulated Emission of Radiation) e LED (Light Emitting Diode) como fontes luminosas (ANDERS; LANZAFAME; ARANY, 2015). A ação da fotobiomodulação ocorre através da absorção da energia luminosa por cromóforos específicos na mitocôndria gerando uma série de reações que levam a aumentos da produção do ATP, do consumo de oxigênio, de síntese proteica e do metabolismo celular (FERRARESI et al., 2015a; HUANG et al., 2011; KARU, 2010). Ao longo dos anos, inúmeros trabalhos têm relatado o efeito da fotobiomodulação na cicatrização de feridas cutâneas (AL-WATBAN; ZHANG; ANDRES, 2007; MINATEL et al., 2009), na regeneração de tecidos tendinosos (ENWEMEKA; REDDY, [s.d.]; OLIVEIRA et al., 2009), musculares (CRESSONI et al., 2008) e nervosos (ROCHKIND, 2009); além de sua ação analgésica (BJORDAL et al., 2007; CHOW et al., 2009) e anti-inflamatória (LOPES-MARTINS et al., 2005; YAMAURA et al., 2009). Mais recentemente, uma inovadora área de atuação desta terapia vem recebendo atenção da comunidade

científica: a relação da fotobiomodulação com o processo de contração muscular.

Lopes-Martins et al. (2006) foram os primeiros a demonstrar que a aplicação da fotobiomodulação em músculos isolados de ratos previamente a um protocolo de repetidas contrações musculares estimuladas eletricamente era capaz de amenizar o processo de fadiga e incrementar o desempenho do músculo ao longo do protocolo (LOPES-MARTINS et al., 2006). Estes achados foram sustentados por estudos posteriores em modelo animal com diferentes protocolos (FERRARESI et al., 2015b, 2015c; LEAL JUNIOR et al., 2010a). Já em seres humanos, Leal Junior et al. (2008) foram os primeiros a demonstrar a melhora do desempenho físico por meio da fotobiomodulação. Atletas de voleibol conseguiram realizar um maior número de repetições quando tiveram seus músculos flexores de cotovelo irradiados com terapia ativa em comparação à placebo previamente a um protocolo de exercício até a exaustão (LEAL JUNIOR et al., 2008). Resultados positivos também foram verificados em estudos subsequentes com este mesmo modelo experimental (LEAL JUNIOR et al., 2009a, 2009b, 2010b), assim como em estudos envolvendo exercício isométrico (DE ALMEIDA et al., 2012), exercício isocinético (BARONI et al., 2010a), exercício em esteira (DE MARCHI et al., 2012) e em testes de ciclismo (LANFERDINI et al., 2017, 2018).

Somados aos efeitos na performance e na fadiga muscular, estudos demonstraram a atenuação de marcadores de dano muscular induzido por exercício, como dor muscular tardia, queda da capacidade de produção de força e níveis de CK, redução do processo inflamatório e do processo de fadiga com a aplicação da fotobiomodulação (ANTONIALLI et al., 2014; BARONI et al., 2010b; FRITSCH et al., 2016). Este corpo de evidências levou ao desenvolvimento de revisões sobre o tema (BORSA; LARKIN; TRUE, 2013; FERRARESI; HUANG; HAMBLIN, 2016; LEAL-JUNIOR et al., 2015; VANIN et al., 2018), incluindo duas meta-análises (LEAL-JUNIOR et al., 2015; VANIN et al., 2018), as quais concluíram que a fotobiomodulação apresenta evidências promissoras como um agente não-farmacológico capaz de potencializar o

desempenho de seres humanos na execução de exercício físico e otimizar o processo de regeneração muscular pós-exercício.

Além dos efeitos agudos da terapia sobre o desempenho físico, redução do processo da fadiga e otimização do processo de recuperação muscular (FERRARESI; HUANG; HAMBLIN, 2016; VANIN et al., 2018), a aplicação crônica da fotobiomodulação associada ao treinamento de força também tem se mostrado eficaz para maximização dos ganhos induzidos pelo treinamento (BARONI et al., 2015; FERRARESI et al., 2011; VANIN et al., 2016). Foi evidenciado que quando aplicada antes (BARONI et al., 2015; VANIN et al., 2016) ou depois (FERRARESI et al., 2011) do treinamento, a fotobiomodulação é capaz de aumentar os ganhos de força (BARONI et al., 2015; FERRARESI et al., 2011; VANIN et al., 2016) e de espessura muscular proporcionados pelo treinamento de força (BARONI et al., 2015; VANIN et al., 2016), conforme Tabela 1. Tal resultado é explicado pelos efeitos terapêuticos já mencionados, como aumento do desempenho muscular, redução da fadiga e aceleração da recuperação pós-exercício (BARONI et al., 2015; FERRARESI et al., 2011; MIRANDA et al., 2016). Entretanto, estes estudos foram realizados com adultos jovens, sendo que o seu efeito em outras populações ainda permanece incerto.

Em vista dos efeitos positivos da fotobiomodulação em adultos jovens (FERRARESI; HUANG; HAMBLIN, 2016; VANIN et al., 2018) e dos efeitos do envelhecimento no sistema musculoesquelético (MITCHELL et al., 2012; NARICI et al., 2003), seria de enorme importância encontrar parâmetros ideais para a aplicação desta terapia em idosos. Contudo, poucos estudos foram realizados com esta população e os efeitos encontrados foram contraditórios (TOMA et al., 2013, 2016; VASSÃO et al., 2016). A aplicação de fotobiomodulação antes de protocolos de fadiga de extensão de joelhos com pesos livres gerou aumento no número de repetições (TOMA et al., 2013), enquanto a sua aplicação prévia a um protocolo de extensão de joelho no dinamômetro isocinético não aumentou o desempenho durante o exercício (VASSÃO et al., 2016). Além disto, o único estudo investigando o efeito da aplicação crônica da terapia em associação ao treinamento de força em idosos não encontrou diferenças em relação ao grupo placebo após o período de treinamento (TOMA et al., 2016). Entretanto, este estudo utilizou um probe simples (“probe caneta”) para a aplicação da terapia em um músculo grande

como o quadríceps totalizando uma dose de energia de 56 J (TOMA et al., 2016). Estes parâmetros não estão de acordo com as recomendações da revisão sistemática mais recente para melhores efeitos no desempenho de grandes grupos musculares (VANIN et al., 2018), o que pode ter limitado os resultados do estudo de Toma *et al.* (2016) e ressalta a necessidade de maiores investigações acerca do efeito de parâmetros ideais de fotobiomodulação em idosos.

TABELAS

Tabela 1. Descrição dos estudos avaliando a associação da fotobiomodulação ao treinamento de força

Autores (ano)	Amostra	Protocolo de treinamento	Protocolo de fotobiomodulação	Variáveis analisadas	Resultados
Baroni et al. (2015)	Homens jovens saudáveis (idade 20-35 anos) GC (n = 10) GPT (n = 10) GFT (n = 10)	Treinamento excêntrico isocinético de extensores de joelho, 2x/sem, 8 semanas 1ª-5ª sem: 3 x 10 repetições contrações excêntricas máximas 5ª-8ª sem: 4 x 10 repetições contrações excêntricas máximas	Antes do treinamento Cluster com 5 diodos de laser; Comprimento de onda 810nm; Potência 200mW; Dose por diodo 6 J; Dose por ponto 30 J; 8 pontos de aplicação no quadríceps; Dose total por membro 240 J; Tempo total de aplicação por membro 4 min	Espessura muscular do VL, RF, VM e do quadríceps total (ecografia), PTiso, PTcon e PTexc de extensores de joelhos (dinamometria isocinética)	GFT obteve melhores resultados na hipertrofia total do quadríceps, no PTisom e no PTexc (p < 0,05)
Ferraresi et al. (2011)	Homens jovens saudáveis fisicamente ativos (idade 18-28 anos) GC (n = 10) GT (n = 10) GFT (n = 10)	Exercícios de “leg press” e extensão de joelhos, 2x/sem, 12 semanas 5 x 10 repetições ou até a falha concêntrica a 80% 1 RM	Após o treinamento Feixe com 6 diodos de laser; Comprimento de onda 808nm; Potência 60mW; Dose por diodo 0,6 J; 7 pontos de aplicação no quadríceps; Dose total por membro 25,2 J Tempo total de aplicação por membro 70 s	Perimetria da coxa, 1 RM “leg press”, 1 RM extensão de joelhos, CVM e PT médio de extensores de joelho do membro dominante (dinamometria isocinética)	GTF obteve maiores ganhos percentuais apenas no 1 RM “leg press” (p < 0,05)
Toma et al. (2016)	Mulheres idosas ativas/insuficientemente ativas (idade 60-70 anos) GC (n = 11) GTP (n = 13) GTF (n = 14)	Exercício de extensão de joelhos 2x/sem, 8 semanas 3 x 12 repetições a 60-80% 1RM	Depois do treinamento Sonda com 1 diodo de laser; Comprimento de onda 808nm; Potência 100mW; Dose por diodo 7 J; Dose por ponto 7 J; 8 pontos de aplicação no quadríceps; Dose total por membro 56 J; Tempo total de aplicação por membro ~9,5 min	TC6, trabalho total, PTcon, potência e índice de fadiga de extensores de joelho do membro dominante (dinamometria isocinética), nível de lactato 0, 3, 6 e 30 min depois da avaliação isocinética, IFE (eletromiografia de superfície), 1RM extensão de joelhos	GC, GT e GTF não apresentaram diferenças para o TC6, índice de fadiga e IFE. GTF apresentou melhores resultados que GC mas não que GT para trabalho total, PTcon, potência, nível de lactato em 3 min e 1RM extensão de joelhos.
Vanin et al. (2016)	Homens jovens saudáveis (idade 18-35 anos)	Exercícios de “leg press” e extensão de joelhos, 2x/sem, 12	Antes e/ou depois do treinamento; Cluster com 12 diodos (4 de laser	1RM “leg press”, 1RM extensão de joelhos, CVM	GFTP aprendeu melhores

GFTF (n=12)	semanas	super-pulsado, 4 de LED vermelho, 4 de LED infravermelho);	(dinamometria isocinética) e	incrementos que os
GFTP (n = 12)	5 x 10 repetições ou até a falha	Comprimentos de ondas de 905nm	perimetria de extensores de	demaís grupos para
GPTF (n = 12)	concêntrica a 80% 1RM	(laser super-pulsado), 640nm (4	joelho bilateralmente	1RM "leg press", 1RM
GPTP (n = 12)	unilateralmente, para ambos os	LEDs vermelhos) e 875nm (4 LEDs		extensão de joelhos e
	membros	infravermelhos);		CVM para ambos os
		Potência 1,25mW, 60mW, 70mW,		membros ($p < 0,05$).
		respectivamente;		O GPTP apresentou
		Dose por diodos 0,07 J, 3,42 J e 3,99		pior incremento de
		J, respectivamente;		1RM "leg press" do
		Dose por ponto 30 J;		membro esquerdo
		6 pontos de aplicação no quadríceps;		que os demais
		Dose total por membro 180 J;		grupos ($p < 0,05$).
		Tempo total de aplicação 228 s		Não houveram
		(todos pontos tratados		diferenças entre
		simultaneamente)		grupos para
				perimetria ($p > 0,05$).

Legenda: 1RM: 1 Repetição Máxima; CVM: Contração Voluntária Máxima; GC: Grupo Controle; GFT: Grupo Fotobiomodulação + Treinamento; GPT: Grupo Placebo + Fotobiomodulação; GT: Grupo Treinamento; GTF: Grupo Treinamento + Fotobiomodulação; GTP: Grupo Treinamento + Placebo; GFTF: Grupo Fotobiomodulação + Treinamento + Fotobiomodulação; GFTP: Grupo Fotobiomodulação + Treinamento + Placebo; GPTF: Grupo Placebo + Treinamento + Fotobiomodulação; GPTP: Grupo Placebo + Treinamento + Placebo; J: Joules; IFE: Índice de Fadiga por Eletromiografia; PT: Pico de Torque; PTcon: Pico de Torque concêntrico; PTexc: Pico de Torque excêntrico; PTiso: Pico de Torque isométrico

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4 ARTIGO

EFFECTS OF PHOTOBIMODULATION THERAPY ASSOCIATED TO RESISTANCE TRAINING IN ELDERLY MEN: A RANDOMIZED DOUBLE- BLINDED PLACEBO-CONTROLLED TRIAL

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ABSTRACT

Purpose: The purpose of this study was to investigate the effects of photobiomodulation therapy (PBMT) combined to resistance training on knee extensor muscle mass, strength and functional capacity in elderly men. **Methods:** In this randomized double-blinded placebo-controlled trial, healthy elderly men (age 60-80 years) completed 12 weeks of resistance training (2x/week) with application of placebo (n = 13) or active PBMT (n = 11) on quadriceps muscles (850 nm, 30 J per point, 8 points per limb) before each training session. Leg press and knee extension one-repetition maximum (1RM) tests, isometric and concentric peak torques, rectus femoris (RF) and vastus lateralis (VL) muscle thickness, timed-up-and-go (TUG) and chair rise to standing (CRS) tests were performed before and after the intervention period. **Results:** There were significant improvements in all outcomes for both groups ($p < 0.05$), except for RF muscle thickness for the placebo group ($p = 0.09$). Large effect sizes ($ES > 0.8$) were observed for leg press and leg extension 1RM and CRS tests for both groups, as well as for TUG test for PBMT group. Isokinetic peak torque for both groups and TUG for placebo group had moderate increases ($ES > 0.5$). Muscle thicknesses and isometric peak torque had small increases ($ES > 0.2$) in both groups. Magnitude-based inference analysis did not support any advantages for using PBMT compared to placebo. **Conclusion:** Differently than previously evidenced in young subjects, PBMT with the parameters used in this study did not provide any additional benefits in comparison to placebo application on muscle mass, strength and functional capacity of healthy elderly men engaged in a resistance-training program.

Keywords: phototherapy, low-level laser therapy, strength training, older.

List of abbreviations:

1-RM	One-repetition maximum
ATP	Adenosine triphosphate
CRS	Chair rise to standing
ICC	Intraclass correlation coefficient
J	Joules
PBMT	Photobiomodulation therapy
RF	Rectus femoris
TUG	Timed up-and-go
VL	Vastus lateralis

INTRODUCTION

It is well known that elderly individuals present multifactorial declines in physiological functions of all body systems, including the musculoskeletal system (ALI; GARCIA, 2014). The ageing process is associated with decreases in the muscle quantity and quality, as well as in the neural factors related to muscle strength and power output; consequently, older people present reduced muscle strength and limited functional capacity (MITCHELL et al., 2012). Sarcopenia (i.e., the age-related phenomenon characterised by progressive loss of muscle mass and strength) has been found in up to 33% of adults with 50 years old or older living in the community (CRUZ-JENTOFT et al., 2014), and is considered a key component of frailty syndrome (i.e., a state of dramatic response to a stimulus, leading to change in health status) (CLEGG et al., 2013). Because of the relationship between muscle mass and strength with elderly longevity (MITCHELL et al., 2012), maintaining and/or improving muscle mass and strength of older people has become a major health concern in the last decades (CRUZ-JENTOFT et al., 2014; MITCHELL et al., 2012).

Resistance training has been proved to counteract at least part of the ageing process (CSAPO; ALEGRE, 2016). Conditioning status of knee extensors is considered a factor for physical function of older subjects (MITCHELL et al., 2012), and resistance training programs in that population have been proven effective for increasing thickness and maximal strength by around 40% and 10%, respectively (CSAPO; ALEGRE, 2016). That localized changes on muscle structure and strength lead to significant improvements on the elderly's systemic functional capacity, such those measured through tests involving functional motor tasks (i.e., stair climb test, chair rise to standing test, 6 Minute Walk Test) (RAYMOND et al., 2013). Therefore, increasing attention has been devoted to strategies that are potentially capable of maximizing the adaptations promoted by resistance training in the elderly (MALAFARINA et al., 2013; MITCHELL et al., 2012), such as nutritional supplementation (MALAFARINA et al., 2013) and hormonal reposition (HILDRETH et al., 2013). However, there is no consensus about safety and effectiveness of these approaches to enhance the muscular response to resistance training in elderly (MITCHELL et al., 2012).

Photobiomodulation therapy (PBMT) is an innovative, non-invasive and non-pharmacological ergogenic approach (for a review, see Vanin et al. 2018). The absorption of light energy by specific chromophores at mitochondria lead to a series on intracellular reactions that increase the ATP production and cellular metabolism, among other responses (KARU, 2010). Evidences have supported that PBMT application leads to greater energy availability in high intensity exercise, which may also be related to the integration between aerobic and anaerobic

ATP production (GOREHAM et al., 1999). Consequently, these effects are believed to be related to the increase in exercise performance and the improve post-exercise recovery (FERRARESI et al., 2011; VANIN et al., 2018). In addition, studies that associated PBMT to resistance training programs have found greater muscle hypertrophy and strengthening compared to resistance training alone or combined to placebo treatment (BARONI et al., 2015; FERRARESI et al., 2011; MIRANDA et al., 2016). However, the positive effects of PBMT have been tested almost only in young and athletic populations. When applied in the older population, contradictory results have been found regarding its acute effects (TOMA et al., 2013; VASSÃO et al., 2016), and the single study that investigated the chronic effect of PBMT associated to resistance training in elderly found no advantages compared to resistance training alone (TOMA et al., 2016). Nevertheless, TOMA *et al.*,(2016) used a single probe instead a cluster probe to treat the whole quadriceps muscle, and their total energy dose of 56 J is below the suggestions of the most recent systematic review for treating large muscular groups (i.e., 60-300 J) (VANIN et al., 2018). PBMT has a typical biphasic dose-response effect, and optimal treatment parameters are specific to the therapy's aim, the target tissue, and the subject/tissue status (HUANG et al., 2011). Thus, further investigations are needed to elucidate whether PBMT may maximize resistance training adaptations in elderly.

The objective of the present study was to investigate the effect of PBMT combined to resistance training on knee extensor muscle mass, strength and functional capacity of healthy elderly men. Our hypothesis was that applying PBMT before each resistance training session with proven effective parameters (BARONI et al., 2015) volunteers would have a greater exercise performance during the training session and an optimized post-exercise recovery (Vanin et al. 2018), leading to faster increments in training load and to greater mechanical work in comparison to volunteers treated with placebo, what could maximize training-induced adaptations (CSAPO; ALEGRE, 2016; RAYMOND et al., 2013).

METHODS

Study design

This study was a randomized double-blinded placebo-controlled trial. Volunteers were engaged in a 12-week resistance training program. Placebo or PBMT treatments were applied to their quadriceps muscles before each training session, according to the groups' allocation. Knee extensors ultrasonography, isokinetic dynamometry, one-repetition maximum (1RM) tests, and functional capacity tests were performed before and after the resistance training program. This study was approved by the Ethics Committee of the Federal University of

Health Sciences of Porto Alegre (Porto Alegre, Brazil), prospectively registered at ClinicalTrials.gov (NCT03287284) and conducted according with the 1964 Helsinki declaration and its later amendments. All subjects were informed on the procedures, benefits and risks prior to signing an informed consent document to participate in the study. There were no changes in the methods after trial commencement.

Volunteers

Individuals were recruited to participate in the study through newspaper and social media advertisement. During the first visit, all volunteers were informed about the study procedures and performed a health screening. To be included in the study participants had to be 60 to 80 years old, have no cardiovascular, neurological or musculoskeletal restrictions to resistance training; and, be at least three months without performing any systematic, regular resistance training. Participants had also to present a recent exercise electrocardiogram test reviewed by a cardiologist evidencing their capacity to perform physical exercises. If they did not present one, they were indicated to consult with the cardiologist of the university. Participants were excluded if they presented difficulties in understanding or performing tests and training exercises.

Sample size calculation

Sample size was calculated using the G*Power software (version 3.1.9.2) based on previous trials (CADORE et al., 2013; VAN ROIE et al., 2013) that determined that a sample of $n = 9$ participants would provide a statistical power over 0.90. The outcomes used to calculate the sample size were leg press and leg extension 1RM, as well as rectus femoris muscle thickness.

Randomization and blinding

As shown in Figure 1, 40 men volunteered for the study. Thirty-one volunteers met the inclusion/exclusion criteria and were allocated into the two experimental groups through an on-line computer randomization program: Placebo Group ($n=16$) and PBMT Group ($n=15$). There were 7 drop outs during the training period, as shown in Figure 1. Therefore, 24 volunteers completed the study schedule and were included in the final analyses: Placebo group ($n = 13$), and PBMT group ($n = 11$).

Concealment was guaranteed as the researcher who applied the therapy was the only one aware of the randomization, while the researchers who were involved in the training sessions and assessments remained blinded. Because PBMT provoked no thermic or other perceptive effects and PBMT and placebo treatments followed the same application procedures, volunteers were also blinded to their allocation into PBMT or placebo groups.

<<Insert Figure 1 around here>>

Assessment procedures

All assessments were performed twice within three weeks before the start of the training program. Volunteers responded to the International Physical Activity Questionnaire (BENEDETTI et al., 2007) and were familiarized for each test before evaluation sessions, and the test-retest served to assess the reliability of our measures. In addition, the evaluation protocol was divided in two days with a minimum 48-hour interval period: day 1 – ultrasound and isokinetic dynamometry; day 2 – 1RM and functional tests.

Ultrasonography

Muscle thickness of rectus femoris (RF) and vastus lateralis (VL) muscles were measured with the SSD 4000 ultrasound equipment (Aloka Inc., Japan) with a linear-array probe (60 mm; 7.5 MHz). Before each assessment, subjects were required to rest for 10 minutes in the supine position with knees and hips in neutral position. The measurements were performed at midway between the femur lateral condyle and the articular line of the knee at the anterior side of the thigh for RF and at the lateral side of the thigh for the VL from the right lower limb (BARONI et al., 2015). A single experienced examiner performed all ultrasound scans. To ensure that all measurements were performed at the same position, the probe position as well as anatomical and skin references were drawn in a transparent paper which was used on the following evaluations. Three parallel images were taken from both RF and VL. Images were analysed in Image-J software (National Institutes of Health, USA). The superior and inferior muscle aponeurosis were identified and the distance between them was measured at three points of each image (left border, middle and right border). A mean value was calculated for each image and for

the three images for each assessment; the last one being used for analysis (BARONI et al., 2015). The method presented high interclass correlation coefficient (ICC): 0.97 for RF thickness and 0.98 for VL thickness.

Isokinetic dynamometry

Subjects were positioned seated according to the manufacturer's recommendations for knee flexion-extension assessment at the Biodex System 3 Pro equipment (Biodex Medical Systems, EUA). They performed a warm-up of two sets of ten submaximal repetitions with a one-minute rest interval between sets of knee flexion-extension movements at angular velocity of 90°s^{-1} . Thereafter, individuals performed three 5-sec maximal voluntary isometric contractions of knee extensors at 60° of knee flexion (0° = full knee extension). If there was more than 10% difference between tests, a fourth attempt was performed. Next, subjects performed two sets of three repetitions of isokinetic concentric knee flexion-extension movements at angular velocity of 60°s^{-1} (range of motion = 90° - 10°). A two-minute rest was given between each test to minimize possible fatigue effects (BARONI et al., 2015). The dynamometric tests were performed only with the right lower limb and the highest peak torque values obtained during isometric and isokinetic concentric tests were used for statistical analyses. The tests presented ICC values of 0.92 for isometric peak torque and 0.86 for isokinetic concentric peak torque.

1RM tests

Maximal dynamic strength was assessed by the one-repetition maximum test (1RM) on bilateral leg press and knee extension exercises. Subjects walked for 10 minutes on a treadmill and performed the specific movements as warm up before the tests. Each subject's maximal load was determined with no more than five attempts with a 4-minute recovery between tests. A 10-minute recovery was respected between leg press and knee extension tests. Performance time for each contraction (concentric and eccentric) was two seconds, controlled by the researchers using an electronic metronome (CADORE et al., 2013). The movement range of motion was also controlled, so that volunteers had to reach 90 degrees of knee and hip flexion in the 1RM leg press test as well as the maximal knee extension in the 1RM knee extension test. The tests presented had high test-retest reliability (ICC 0.96 for both leg press and leg extension 1 RM).

Functional tests

The Chair Rise to Standing (CRS) test was performed with the individuals seated in a hard-backed chair with a 43 cm height next to the wall with their arms across their chest. They were requested to rise to a full standing position and return to the initial sitting position as fast as possible for five times (MCCARTHY et al., 2004). They performed three attempts with a two-minute interval between them, being the lower value registered for statistical analysis. The test presented ICC value of 0.85.

The Timed up-and-go (TUG) test was performed with individuals seated in the same hard-backed chair with a 43 cm height next to the wall with their hands on their hips. They were requested to rise and walk as fast as possible without running until a cone positioned three meters away and return to the initial sitting position. Timing started with the command 'go' and ended when subjects touched their back on the back of the chair (PODSIADLO; RICHARDSON, 1991). Each individual performed three attempts with a two-minute interval and the lowest value was recorded for statistical analysis. The ICC of the test was 0.88.

Intervention procedures

Photobiomodulation Therapy (PBMT) / Placebo

Before each training session, PBMT/placebo were applied on both lower limbs. Treatments were delivered with a cluster probe composed by five 850 nm laser diodes, each one with a 100mW power output (Chattanooga Corp., Chattanooga, USA). Eight sites of the quadriceps muscle were defined by palpation and treated: two at vastus lateralis, three at vastus medialis, and three at rectus femoris (Figure 2). Each site was treated for 60s, leading to a dose of 6 J per diode, 30 J per site, and 240 J per leg (BARONI et al., 2015). The probe was held stationary in skin contact at 90° angle with light skin pressure. Placebo was applied on the same way, but with the device turned off. Subjects were in the supine position, blindfolded and with headphones listening to music of their own choice to prevent clues about the active or placebo application. As soon as the application was finished, they were instructed to start their training session.

<< Insert Figure 2 around here >>

Resistance Training

After the 10-minutes warm up on a treadmill, individuals started the resistance exercises. We standardized leg press and knee extension exercises as the first and the second exercises, respectively, for all volunteers in every training session to ensure a similar interval between the end of the PBMT/placebo application and the beginning of the goal exercises. After that, individuals performed other five exercises (seated supine, seated row, leg curl, abdominal curl and back extension exercises) so that they had a complete resistance training program. The periodization was performed as previously described (CADORE et al., 2013). Subjects performed two sets of 18–20 repetitions maximum (RM) during weeks 1 and 2, progressing to two sets of 15–17 RM at weeks 3 and 4, two sets of 12–14 RM in weeks 5–7, three sets of 8–10 RM in weeks 8–10 and finalizing with three sets of 6–8 RM in weeks 11 and 12. The recovery time between sets was of 90–120 seconds (CADORE et al., 2013). All sets were performed until failure and the workload was constantly adjusted to keep the number of repetitions within the established interval. The load and number of repetitions performed in each exercise were recorded so that instructors could control the training of each participant and their workload. These data were used to calculate mean workload applied in each mesocycle (Figure 3). The intervention was realized twice weekly for 12 weeks, leading to a total of 24 training sessions. The sessions were always performed on Mondays and Wednesdays afternoon with the supervision of one experienced instructor for each volunteer to ensure that exercises were performed correctly, within the interval of repetitions desired and until failure. It was established that if a volunteer missed one session, he could recover it on the following Friday. So, all the individuals completed all the training sessions.

Statistical analyses

Within-group analysis to verify training effects were performed with the following calculations: percent change ($\Delta\% = \text{post-training/pre-training}-1$); paired sample t-test, with a significance level set as 5% ($\alpha < 0,05$); effect size (ES) through the Cohen's d [$ES = (M_{\text{post}} - M_{\text{pre}})/SD_{\text{pooled}}$, where M_{post} is the mean post-training measure, M_{pre} is the mean pre-training measure, and SD_{pooled} is the pooled standard deviation of the pre- and post-measurement], and training effects were considered as “trivial” ($ES < 0.2$), “small” ($ES > 0.2$), “moderate” ($ES > 0.5$) or “large” ($ES > 0.8$) (COHEN, 1977).

Longitudinal percent changes (pre- to post-training) were used for between-group analysis. Data were analysed for practical significance using magnitude-based inferences because traditional statistical approaches often do not indicate the magnitude of an effect, which is typically more relevant to sports medicine than statistical

significance (HOPKINS et al., 2009). The chances of a possible substantial effect favourable to Placebo or PBMT groups were calculated [i.e., greater than the smallest worthwhile change (0.2 multiplied by the between-subject standard deviation)]. Quantitative chances of trivial effects and substantial effects for each group were assessed qualitatively as follows: <1%, almost certainly not; 1-5%, very unlikely; 5-25%, unlikely; 25-75% = possibly; 75-95% = likely; 95-99% = very likely; >99% = almost certain (BATTERHAM; HOPKINS, 2006). When the Placebo and PBMT values were both >5 %, the inference was classified as unclear (BATTERHAM; HOPKINS, 2006).

RESULTS

There were no differences ($p > 0.05$) between groups for age (Placebo 66.67 ± 5.84 ; PBMT 68.93 ± 7.47); height (Placebo 1.71 ± 0.05 ; PBMT 1.74 ± 0.06); weight (Placebo 76.64 ± 8.82 ; PBMT 80.81 ± 12.24) and body mass index (Placebo 26.25 ± 2.82 ; PBMT 26.78 ± 4.13). Moreover, the physical activity level of the individuals was also similar between groups: 75.0% from the Placebo and 66.7% from the PBMT were active, 12.5% from Placebo and 20.0% of the PBMT were sedentary and 12.5% from the Placebo and 13.3% from the PBMT were irregularly active.

There were no differences between the relative training load between groups across the training period and both groups showed similar pattern of mean load increases ($p > 0.05$) (Figure 3).

<< Insert Figure 3 around here >>

Table 1 shows the within-group analysis. The training programs increased VL muscle thickness in both groups with a small effect size, while RF muscle thickness achieved only trivial effect sizes. Both groups presented small and moderate effect sizes for isometric and concentric peak torques, respectively. Leg press 1RM, knee extension 1RM and the CRS had large effect sizes. TUG improved with a small effect size in Placebo group and a moderate effect size in PBMT group.

<< Insert Table 1 around here >>

Figure 4 presents the between-group analysis. Unclear nuclear differences were found for all outcomes, which support similar adaptations to resistance training in Placebo and PBMT groups for knee extensor muscle thickness, strength and functional capacity.

<< Insert Figure 4 around here >>

DISCUSSION

As expected, independently of PBMT or placebo therapy, resistance training increased knee extensor thickness, maximal strength and functional capacity of elderly men. The main finding of the current study is that the addition of PBMT prior to resistance training sessions did not add any benefit to resistance training plus placebo therapy in older men, since both training groups showed similar enhancements in all investigated outcomes. This finding means that practitioners should not apply PBMT before resistance training or rehabilitation session in elderly individuals, at least with the parameters utilized in the current study, as it does not provide additional effects on muscle mass, strength and physical function.

Resistance training programs are known to increase muscle strength and mass, as well as physical function in older adults (BORDE; HORTOBAGYI; GRANACHER, 2015; CSAPO; ALEGRE, 2016; RAYMOND et al., 2013). However, the training periodization (frequency, duration, volume of training, etc.) and subjects' characteristics (age, sex, physical activity level) interfere directly on the magnitude of the muscular adaptations (BORDE; HORTOBAGYI; GRANACHER, 2015; BROCCA et al., 2017). Comparison between studies should be made with caution and should always concern trials with similar study population and periodization (BORDE; HORTOBAGYI; GRANACHER, 2015; BROCCA et al., 2017). It means that the adaptations magnitude of the current study should not be directly compared to previous studies which associated resistance training and PBMT in young men (BARONI et al., 2015; FERRARESI et al., 2011, 2016; VANIN et al., 2016) and elderly women (TOMA et al., 2016) due to different training periodization, age and/or sex of the participants.

Although only small effect sizes have been observed on VL muscle thickness of both groups in our study, the percent increases (6-7%) are similar to the ones previously reported with the elderly population (CSAPO; ALEGRE, 2016; LIXANDRÃO et al., 2016). A recent systematic review found very modest training-associated gains in muscle thickness and cross sectional area in this population (CSAPO; ALEGRE, 2016). It was argued that the hypertrophic capacity of the elderly may be impaired due to age-related motor unit loss and concomitant accumulation of intramuscular non-contractile tissues (CSAPO; ALEGRE, 2016). However, the limited

hypertrophic response seen in elderly in relation to young is controversial, since studies have shown similar increases in muscle mass and strength in response to resistance training in both groups (HÄKKINEN et al., 2000; MAYHEW et al., 2009). Both groups present increments of approximately 10% and a high variability is seen between individuals (KRAEMER et al., 1999), what may limit the effect sizes observed in hypertrophic response.

The current study did not assess neuromuscular adaptations provided by resistance training program, but this is a typical response to resistance training and explains the larger relative increases found in 1 RM tests compared to those observed in muscle thickness (HÄKKINEN et al., 2000). The improvements found in leg press and knee extension 1RM (40% and 23-24%, respectively) are similar to the increases reported by studies which used the same periodization in concurrent trainings in elderly men (CADORE et al., 2012, 2013). Moreover, the increments found in isometric and isokinetic peak torques (6% and 7-10%, respectively) are in agreement with previous studies which also reported increments in isometric (CADORE et al., 2013; VAN ROIE et al., 2013) and isokinetic peak torques (VAN ROIE et al., 2013) in similar population. The smaller gains assessed by isokinetic dynamometry may be due to training specificity; isokinetic exercises were not performed along training program and there is no full transference of training-induced strength gains to that tests (ISSURIN, 2013).

The increments in the CRS and TUG tests (9% and 4-6%, respectively) are in agreement with previous trials analysing the effect of resistance training in functional capacity of elderly (DIAS et al., 2015; TIGGEMANN et al., 2016; VAN ROIE et al., 2013). They represent the training transfer, since the leg press and knee extension exercises led to an improvement in untrained tasks (ISSURIN, 2013). Although the within-group analyses showed different effect sizes for PBMT and placebo groups, suggesting an advantage for the PBMT group, the magnitude-based inference analyses supported the similar adaptations between groups, as well as for all other outcomes. Therefore, the current results further support findings of (TOMA et al., 2016) that there is no evidence of additional effect of adding PBMT to resistance training programs on the functional capacity of elderly.

Studies with young men have demonstrated that PBMT maximizes the strength gain and the hypertrophic response to resistance training (BARONI et al., 2015; FERRARESI et al., 2011; VANIN et al., 2016). Authors have justified the positive effects of PBMT mainly through the acute effects of PBMT, such as reduced muscle fatigue, enhanced exercise performance and optimized post-exercise recovery (BARONI et al., 2015; FERRARESI et al., 2011; VANIN et al., 2016). That resistance training studies and other previous investigations (see Vanin et al. 2018) have supported our initial hypothesis that PBMT group would have enhanced exercise performance during training sessions, leading to higher mechanical work supported by the muscle; as well as better

recovery between training sessions, also contributing to a higher performance in the following training day compared to placebo group. Consequently, we expected that PBMT group would have faster increments in training load than placebo group. However, training load progression was similar between PBMT and placebo groups (see Figure 3). These findings refuted our initial hypothesis about how PBMT would act as an ergogenic factor for elderly subjects engaged in a resistance training program.

It is important to note that almost all evidences supporting the ergogenic action of PBMT on human performance were provided by studies with young adults (VANIN *et al.*, 2018). Interestingly, trials involving older subjects have presented inconsistent results. While Toma *et al.* (2013) found an increase in the number of repetitions of knee extension exercise after PBMT application, Vassão *et al.* (2016) found no effect of its prior application on isokinetic exercise performance. At the same time, the only study so far with resistance training associated to PBMT in elderly found no additional benefits of this therapy compared to placebo application (TOMA *et al.*, 2016), which is further supported by our findings. Even though the exercise protocols and the PBMT parameters differed among the trials, it is not clear why PBMT seems to work for young but not for elderly individuals.

It is well known that PBMT acts on mitochondria improving the cytochrome c oxidase activity, the respiratory chain reactions, the ATP production and the cellular metabolism (KARU, 2010). It is also known that elderly people present muscle atrophy independently from being physically active or having a disease (BROCCA *et al.*, 2017). Furthermore, mitochondrial function presents deficits with age and it has been considered as a potential mediator of sarcopenia (CARTER; CHEN; HOOD, 2015). Aged muscles present mitochondria with less organelle content, reduced layer thickness, enzymes (being cytochrome oxidase one of them) and function, including protein synthesis, respiration and ATP production (CARTER; CHEN; HOOD, 2015). Therefore, if there are less enzymes to absorb the light photons, its energy may be lost leading to no clinical effect. Interestingly, PBMT seems to not affect the exercise performance of patients with cardiovascular diseases (BUBLITZ *et al.*, 2016; STEIN *et al.*, 2018), a population that also presents reduced mitochondrial activity and sarcopenia (MARZETTI *et al.*, 2013). However, as the present study did not investigate the mechanism of action of PBMT in the elderly, we do not have data to confirm or refute this hypothesis. We recommend that future studies should investigate the effects of PBMT in young in comparison to elderly to similar exercise protocol and/or training program to elucidate specific responses of each age group to PBMT.

PBMT presents a typical biphasic dose-response effect (HUANG et al., 2011) and there is still no consensus about the ideal parameters to optimize adaptations to resistance training in any population. Therefore, although the parameters used in our study have been proven effective in young people (BARONI et al., 2015), we cannot exclude the possibility that a different dosage may be needed to optimize training responses in older people. In fact, a few studies analysed the dose-response effect of PBMT in young people and found contradictory results (AVER VANIN et al., 2016; DELLAGRANA et al., 2018; LANFERDINI et al., 2018). The application of dosages of 270 J and 300 J were the most effective ones for increasing performance outcomes of recreational runners during running tests (DELLAGRANA et al., 2018) and soccer players during isokinetic tests (AVER VANIN et al., 2016), respectively. However, only the application of 135 J, and not 270 J or 405 J, improved the exercise performance of cycling athletes in incremental cycling tests until exhaustion (LANFERDINI et al., 2018). It remains unclear if the exercise protocol, the sample characteristics and/or the other parameters of the PBMT were responsible for that specific dose-response effects. Different dosages with different parameters of PBMT need to be tested to provide the best parameters to each population and exercise protocol before conclusions can be drawn.

A limitation of the current study is the lack of investigation on biochemical parameters, what limits the discussion about the mechanisms responsible for the absence of PBMT effect on this population. Moreover, the addition of experimental groups with different parameters of PBMT would have allowed us a broader view on the effects of this treatment in training responses of older men. To the best of our knowledge, there are only a few previous studies that investigated the effects of PBMT in muscular performance of the elderly population (TOMA et al., 2013, 2016; VASSÃO et al., 2016) and only one of them investigated the chronic effects of the therapy (Toma et al. 2016). One positive aspect of the current study is that its design was developed to be similar to the clinical practice and to provide results that could be easily replicated. Moreover, it provided reliable results on muscle strength, mass and physical function. The potential benefit of PBMT to counteract the age-related impairments in the musculoskeletal system and in functional capacity highlights the need for more studies to find how the older can also benefit from the therapy, what has been already shown for the young.

CONCLUSIONS

The current study did not find any additional effect of PBMT with the tested parameters compared to placebo treatment combined to resistance training on knee extensor muscle mass, strength and functional performance of elderly men. Given the potential benefits of PBMT on the fight against sarcopenia and its

deleterious effects, as well as the characteristic dose-response effect of PBMT, scientists are encouraged to perform further investigations on that issue.

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Authors' contributions

BB, EC, RP, MV conceived and designed research. CF, MD, JT and LS conducted experiments. CF and BB analyzed data. CF and BB wrote the manuscript. All authors read the manuscript, contributed to its improvement and approved its final version.

Authors declare that there is no professional relationships with companies or manufacturers who will benefit from the results of the present study.

Figure 1. Flowchart of the study

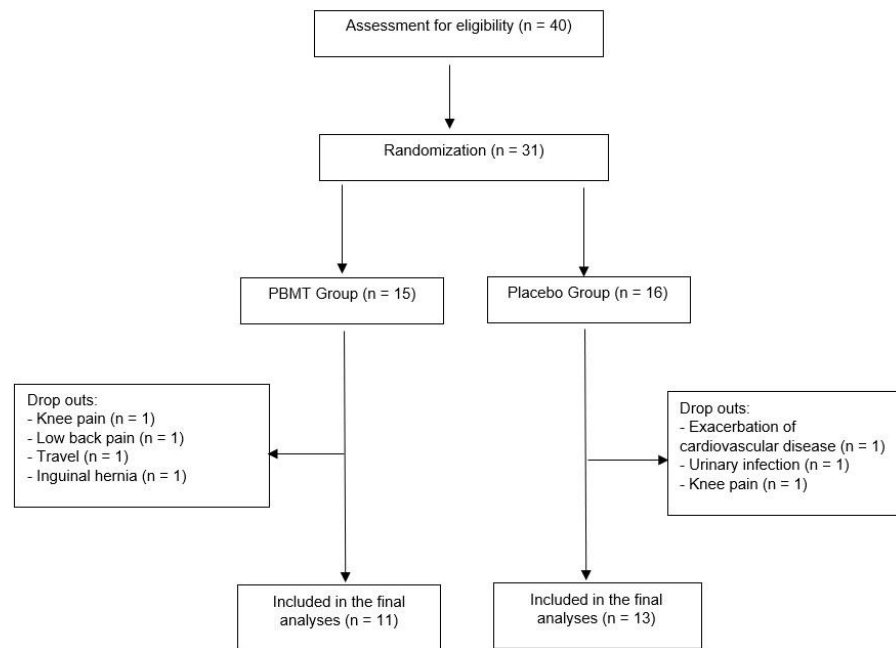


Figure 2. Photobiomodulation therapy (PBMT) parameters

PBMT PARAMETERS			
(per leg)	Number of points		8 points
	Treatment time		60 s
(per point)	Number of diodes used (cluster probe)		5 diodes
	Wavelength		850 nm
	Frequency		Continuous
	Spot size	(per diode)	0.029 cm ²
	Peak power	(per diode)	100 mW
	Power density	(per diode)	3.4 W/cm ²
	Energy density	(per diode)	206.9 J/cm ²
	Energy	(per diode)	6 J
	Energy	(per point)	30 J
	Energy	(per leg)	240 J

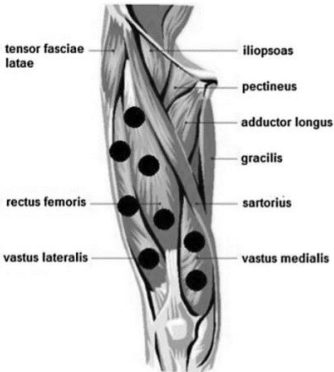
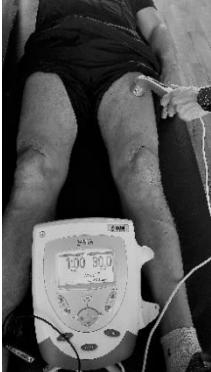



Figure 3. Leg press and leg extension mean training load relative to 1RM test on each mesocycle

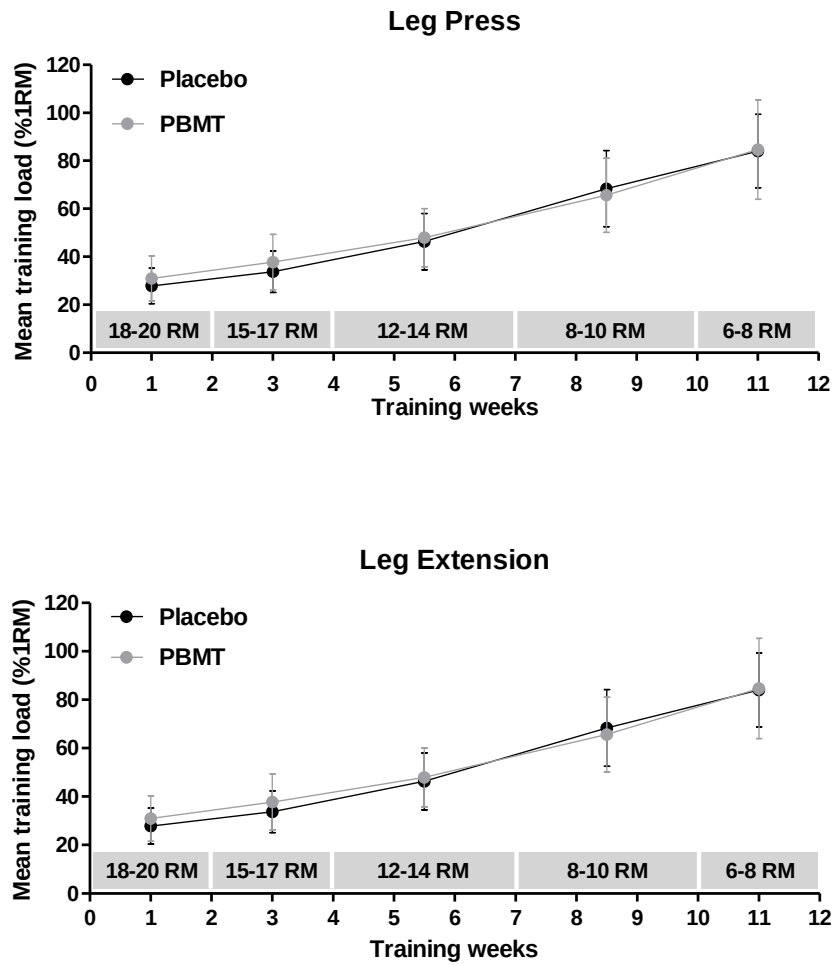
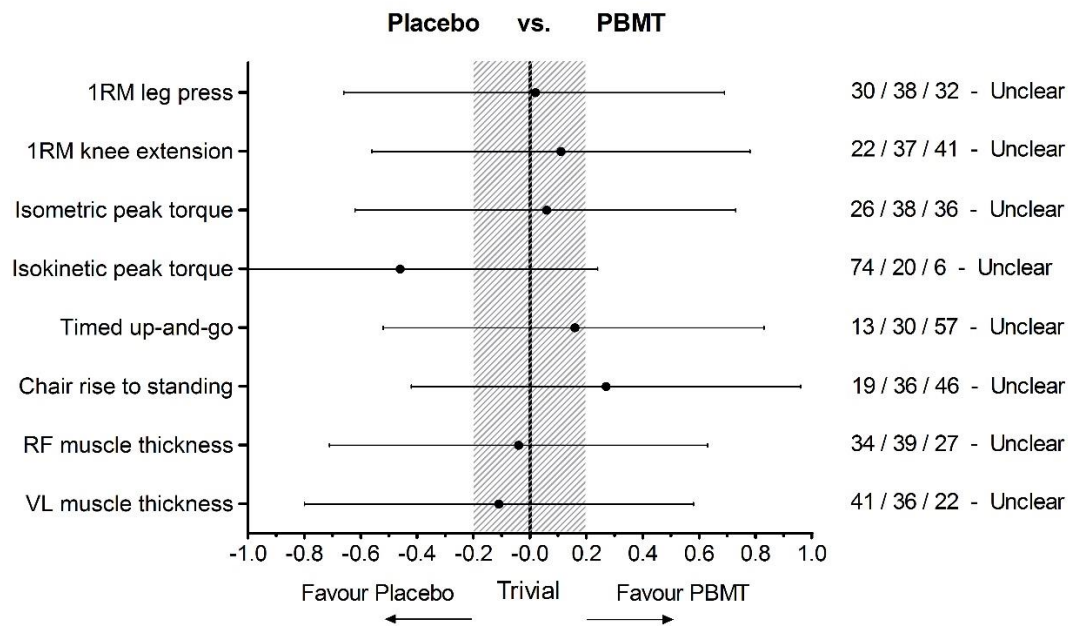


Figure 4. Magnitude-based inferences for between-group analyses



5 CONCLUSÃO GERAL

A terapia de fotobiomodulação com os parâmetros estudados neste trabalho não apresenta efeitos adicionais em relação à terapia placebo quando associada ao treinamento de força para ganhos de força e massa muscular de extensores de joelhos e capacidade funcional de idosos. Devido ao potencial desta terapia de combater a sarcopenia e à sua característica bifásica de dose e resposta, a comunidade científica deve ser encorajada a continuar investigando o efeito de diferentes parâmetros de fotobiomodulação sobre a otimização dos ganhos do treinamento de força nesta população.

ANEXOS

ANEXO A – Normas de formatação do periódico European Journal of Applied Physiology

Types of papers

The journal publishes Original Articles, Editorials, Invited Reviews, Letters to the Editor, and Replies.

- **Instructions for Invited Reviews**

The European Journal of Applied Physiology (EJAP) now publishes Invited Reviews only. The journal welcomes topics from physiological mechanics, biochemistry, endocrinology, ergonomics, immunology, motor control, neuroscience and nutrition.

Invitations are issued by the Reviews Editor Michael Lindinger (mi.lindinger@gmail.com) on behalf of EJAP.

Authors wishing to submit proposals for consideration are welcome to contact the Reviews Editor, providing a brief description of the review topic (i.e. title and one-paragraph summary). If a proposal is deemed to be relevant and appropriate after an initial evaluation, authors will receive a formal invitation to write the Review.

- Maximum manuscript size is 15 printed pages (4500-5000 characters/page)
- Colored illustrations highlighting critical concepts are strongly encouraged
- no limitation to number of references
- Abstracts should aim for 150-250 words
- Abstracts should contain the need for the review, primary objectives and logical conclusions
- Abstracts must not contain undefined abbreviations or references

- **Instructions for Letters to the Editor**

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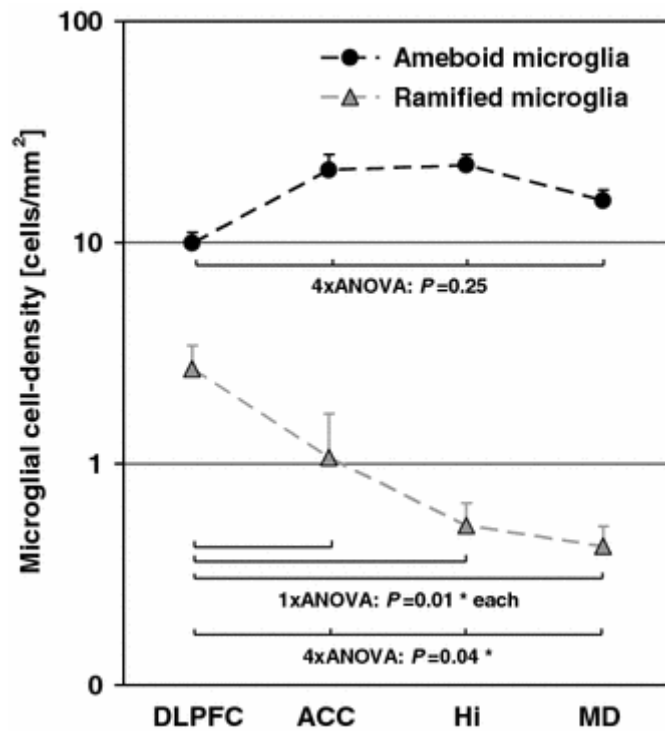
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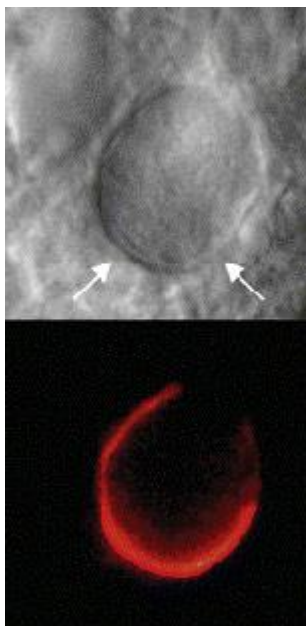
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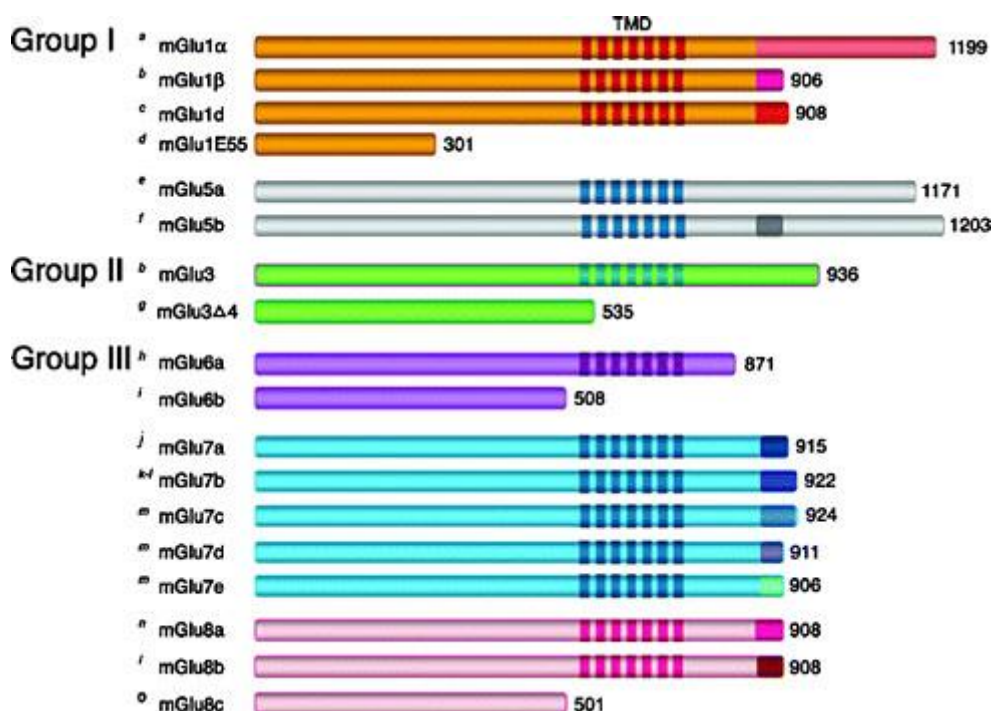
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ANEXO B – Parecer do CEP

PARECER CONSUBSTANCIADO DO CEP**DADOS DO PROJETO DE PESQUISA**

Título da Pesquisa: Fototerapia associada ao treinamento de força em idosos: Efeitos sobre a força, massa muscular e capacidade funcional **Pesquisador:** Bruno Manfredini Baroni **Área**

Temática:

Versão: 2

CAAE: 59328316.9.0000.5345

Instituição Proponente: Universidade Federal de Ciências da Saúde de Porto Alegre

Patrocinador Principal: Financiamento Próprio

DADOS DO**PARECER****Número do Parecer:**

1.783.982

Apresentação do Projeto:

Introdução: A Laserterapia de Baixa Potência (LBP), largamente utilizada na prática clínica de profissionais de fisioterapia, é uma modalidade de fototerapia com um robusto corpo de evidências acerca de seus efeitos analgésicos, anti-inflamatórios e regenerativos. Mais recentemente, a LBP vem sendo estudada com a finalidade de potencializar os efeitos do treinamento de força em indivíduos jovens e saudáveis. Entretanto, a literatura carece de evidências a cerca dos seus efeitos na população idosa, a qual poderá ser amplamente beneficiada devido ao declínio muscular apresentado. Objetivo: Verificar o efeito da fototerapia sobre as adaptações promovidas pelo treinamento de força em indivíduos idosos. Métodos: Este estudo se caracteriza como um ensaio clínico randomizado, duplo-cego e placebo-controlado. Serão incluídos 30 idosos com idades entre 60 e 80 anos, os quais serão randomizados entre Grupo Placebo (LBP placebo + treinamento de força) e Grupo Fototerapia (LBP ativa + treinamento de força). Todos os voluntários serão submetidos a um período controle (sem treinamento) de quatro semanas, seguido por um período de 12 semanas de treinamento de força de membros inferiores. As avaliações serão realizadas em três momentos do estudo: (1) quatro semanas antes de iniciar o período de treinamento (basal); (2) imediatamente antes de iniciar o período de treinamento (pré-treinamento); (3) após o término do período de treinamento (pós-treinamento). As sessões de

treinamento serão realizadas duas vezes por semana e será respeitada uma periodização progressiva pelo método de repetições máximas. A LBP será aplicada previamente às sessões de treinamento em oito pontos dos músculos extensores de joelho, totalizando uma dosagem de 240 J em cada quadríceps. A aplicação placebo será realizada da mesma forma, mas com o equipamento desligado. As avaliações contemplarão análises de: propriedades estruturais do quadríceps por meio de imagens ultrassonografia; capacidade de produção de força do quadríceps pela dinamometria isocinética; e capacidade funcional dos voluntários através do Teste de subir degraus, Teste de sentar e levantar em uma cadeira, Teste de caminhada de seis minutos e Teste “Timed Up-and-Go”.

Objetivo da Pesquisa:

Verificar o efeito da fototerapia sobre as adaptações promovidas pelo treinamento de força em sujeitos idosos.

Avaliação dos Riscos e Benefícios:

Adequados

Comentários e Considerações sobre a Pesquisa:

Sem comentários

Considerações sobre os Termos de apresentação obrigatória:

Adequados

Recomendações:

Sem recomendações

Conclusões ou Pendências e Lista de Inadequações:

Ajustadas a solicitação do CEP.

Considerações Finais a critério do CEP:

Dea acordo com o parecer do Relator.

Este parecer foi elaborado baseado nos documentos abaixo relacionados:

Tipo Documento	Arquivo	Postagem	Autor	Situação
Informações Básicas do Projeto	PB_INFORMAÇÕES_BÁSICAS_DO_PROJETO_753253.pdf	23/09/2016 10:25:36		Aceito
Outros	Carta_resposta_CEP.pdf	23/09/2016 10:25:03	Bruno Manfredini Baroni	Aceito

Continuação do Parecer: 1.783.982

TCLE / Termos de Assentimento / Justificativa de Ausência	TCLE_round2.pdf	23/09/2016 10:24:37	Bruno Manfredini Baroni	Aceito
Projeto Detalhado / Brochura Investigador	ProjetoCEP_round2.pdf	23/09/2016 10:23:57	Bruno Manfredini Baroni	Aceito
TCLE / Termos de Assentimento / Justificativa de Ausência	Termo_Anuencia_Lab_UFCSPA.pdf	29/08/2016 22:42:27	Bruno Manfredini Baroni	Aceito
TCLE / Termos de Assentimento / Justificativa de Ausência	Termo_Anuencia_ESEF_UFRGS.pdf	29/08/2016 22:42:11	Bruno Manfredini Baroni	Aceito
Declaração de Pesquisadores	Termo_Entrega_Relatorio.pdf	29/08/2016 22:41:48	Bruno Manfredini Baroni	Aceito
Folha de Rosto	Folhaderosto.pdf	25/07/2016 21:48:18	Carolina Gassen Fritsch	Aceito

Situação do Parecer:

Aprovado

Necessita Apreciação da CONEP:

Não

PORTO ALEGRE, 20 de Outubro de 2016

Assinado por:
Julia Fernanda Semmelmann Pereira Lima
(Coordenador)

ANEXO C – Termo de Consentimento Livre e Esclarecido

TERMO DE CONSENTIMENTO INFORMADO

	Dados
de Identificação do Participante:	
Nome:	
Identidade: Sexo: : Data de Nascimento: : Endereço:	
..... Cidade:	
..... Estado:..... País:..... Telefone:	
..... E-mail:	

	Dados
de Identificação do Pesquisador Responsável:	
Prof. Bruno Manfredini Baroni	
Universidade Federal de Ciência da Saúde de Porto Alegre (UFCSPA)	
Departamento de Fisioterapia	
Rua Sarmiento Leite, 245 – Centro Histórico – Porto Alegre, RS. CEP: 90050-170;	
Telefone: (51) 3303-8876	

	Termo
de Consentimento Informado:	

Este termo de consentimento, cuja cópia lhe foi entregue, é apenas parte de um processo de consentimento informado de um projeto de pesquisa ao qual você está sendo convidado a participar. Ele deve lhe dar uma ideia básica do que se trata o projeto, e o que sua participação envolverá. Se você quiser mais detalhes sobre algo mencionado aqui, ou informação não incluída aqui, sinta-se livre para solicitar. Por favor, leia atentamente esse termo, a fim de que você tenha entendido plenamente o objetivo desse projeto e o seu envolvimento nesse estudo como sujeito participante. O investigador tem o direito de encerrar o seu envolvimento nesse estudo, caso isso se faça necessário. De igual forma, você pode retirar o seu consentimento em participar no mesmo a qualquer momento.

O objetivo deste estudo é analisar o efeito da aplicação de Laser de Baixa Potência (LBP), um equipamento comumente utilizado na prática clínica de fisioterapia, sobre as adaptações decorrentes de um programa de treinamento de força de membros inferiores (exercício para os músculos da coxa). Todos os voluntários realizarão o mesmo programa de treinamento. A única diferença entre os participantes é que metade receberá aplicação de LBP antes das sessões de treino e a outra metade dos indivíduos receberá placebo (aplicação com o equipamento desligado). Salienta-se que a LBP é uma terapia totalmente segura, de aplicação não-invasiva, que consiste na emissão de uma fonte luminosa de natureza específica sobre a pele do indivíduo. Quando utilizada nas doses apropriadas (como no caso do presente estudo), tal terapia não provoca qualquer dano à pele ou tecidos subjacentes do indivíduo, sendo nulos os riscos de queimaduras, cicatrizes ou outros tipos de alterações estéticas no local das aplicações. Da mesma forma, a aplicação de LBP não causa nenhum tipo de sensação tátil, desconforto ou dor.

O estudo será realizado em dois locais: (1) Laboratório de Fisioterapia da UFCSPA, onde serão realizadas as avaliações; e (2) sala de musculação das ESEFID-UFGFRS, onde serão realizadas as sessões de treinamento. Os pesquisadores informarão como chegar a cada um desses locais.

As avaliações serão realizadas em três momentos: quatro semanas antes de começar o programa de treinamento, na semana anterior ao início do treinamento e na semana seguinte ao término do programa de treinamento. Essas avaliações incluem a massa muscular da região anterior da coxa (avaliada por exame de ultrassom), a capacidade de gerar força da musculatura anterior da coxa e o desempenho do voluntário em testes funcionais que envolvem movimentos do dia-a-dia.

O programa de treinamento terá duração de 12 semanas ininterruptas. As sessões de treino ocorrerão duas vezes por semana e em horários a serem acertados com cada voluntário. Em cada sessão, o participante executará um aquecimento em bicicleta ergométrica, seguido de um protocolo de exercícios para os músculos da coxa em equipamentos de musculação.

É importante destacar que a participação no presente estudo não acarretará despesas aos voluntários. Da mesma forma, não haverá qualquer espécie de remuneração financeira aos mesmos pela participação no estudo. Os benefícios para os participantes que fizerem parte dos grupos LBP e placebo se encontram nos incrementos de força que são esperados com o treinamento, assim como possíveis melhoras da capacidade de realização de tarefas do dia-a-dia que estejam relacionadas à força dos membros inferiores.

Os riscos à saúde dos participantes são mínimos, e consistem principalmente em dor ou desconforto nos músculos treinados nos dias subsequentes à realização das sessões de treino. Em caso de ocorrência de qualquer lesão musculoesquelética (estiramentos musculares, contraturas musculares e inflamações em tendões ou cartilagens) em função do treinamento, os pesquisadores se responsabilizam por fornecer reabilitação fisioterapêutica completa e gratuita nas dependências do Laboratório de Fisioterapia da UFCSPA. O tratamento será realizado por alunos do curso de Fisioterapia da instituição sob a supervisão do pesquisador responsável.

A sua assinatura nesse formulário indica que você entendeu satisfatoriamente a informação relativa à sua participação nesse projeto e você concorda em participar como sujeito. De forma alguma esse consentimento lhe faz renunciar aos seus direitos legais, e nem libera os investigadores, patrocinadores, ou instituições envolvidas de suas responsabilidades pessoais ou profissionais. A sua participação continuada deve ser tão bem informada quanto o seu consentimento inicial, de modo que você deve se sentir à vontade para solicitar esclarecimentos ou novas informações durante a sua participação. Se surgir qualquer dúvida referente a esta pesquisa, favor contatar os pesquisadores nos telefones indicados no início desse termo de consentimento ou o Comitê de Ética em Pesquisa com Seres Humanos da Universidade Federal de Ciências da Saúde de Porto Alegre, localizado na Rua Sarmento Leite, 245 (sala 407 – prédio III), telefone (51) 3303-8804 ou e-mail cep@ufcspa.edu.br, de segunda à sexta-feira, das 8h às 12h e das 13h às 17h.

Porto Alegre, ____ de _____ de 20__.

Pesquisador responsável

Participante