

1 **Immediate effects of two, isometric calf muscle exercises on mid-portion Achilles tendon pain.**

2 **Word Count**

3 Abstract : 1414 (Characters with spaces)

4 Manuscript : 18534 (Characters with spaces)

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27 **Abstract**

28 The objectives of this pilot, randomised, cross-over study were to determine if isometric
29 plantarflexion exercises resulted in an immediate change in Achilles tendon pain during a loading
30 task and if this differed in knee extension or flexion. Eleven participants with mid-portion Achilles
31 tendinopathy were recruited from NHS community physiotherapy services and local running clubs.
32 Participants were then randomised to complete an isometric calf muscle exercise with the knee fully
33 extended or flexed to 80 degrees. Participants switched to the alternate exercise after a minimum
34 seven-day period. Achilles tendon pain during a specific, functional load test was measured on a 11-
35 point numeric pain rating scale (NPRS) pre and post-intervention. There was a small, immediate,
36 mean reduction in pain following isometric plantar flexion performed in both knee extension (1.6,
37 95%CI 0.83 to 2.45, $p=0.001$) and knee flexion (1.3, 95%CI 0.31 to 2.19, $p=0.015$). There were no
38 significant differences between the two positions. A non-significant, potentially clinically relevant
39 finding was a 20% larger reduction in symptoms in knee extension versus flexion ($p=0.110$). In
40 conclusion, isometric plantarflexion holds gave an approximately 50% immediate reduction in
41 Achilles tendon pain with a functional load test. There were no significant differences between the
42 two positions and both were well tolerated.

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44 **Key Words**

45 Tendinopathy; Physiotherapy; Exercise; Pain

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53 **Introduction**

54 Achilles tendinopathy is a musculoskeletal disorder characterised by pain and loss of function [1, 2].

55 Exercise is the cornerstone of management of tendinopathy, but pain experienced during or

56 following exercise may present a barrier to adherence [3]. Exercise is more effective than wait-and-

57 see in the management of mid-portion Achilles tendinopathy [4] and by 12 weeks appears to result

58 in clinically meaningful improvements in tendon-pain related disability [5]. Thus, improving

59 adherence to exercise and reducing transition to less efficacious, invasive procedures should be a

60 focus. For in-season athletes, interventions that target immediate reductions in symptoms may also

61 be important, however few interventions demonstrate an immediate, beneficial effect on tendon

62 pain [6, 7]. Studies that investigate the immediate effects of calf muscle exercises and whether they

63 are tolerable (and beneficial) to participants with Achilles tendinopathy may assist with exercise

64 prescription and provide options for clinicians when faced with those who have irritable symptoms

65 or fear of exercise, particularly in the early phases on rehabilitation [8].

66 Isometric exercise has been investigated in a small case series in people with Achilles tendon pain

67 with variable effects on pain during a functional load test [9]. Currently most of the research on

68 isometrics is based in patellar tendinopathy with variable results. Improvements were first

69 demonstrated in a small laboratory based-cross over study [7] and then a 4-week case series of 20

70 athletes [10]. Further, both isometric and isotonic exercises were shown to be beneficial in a 4-week

71 RCT [11] indicating two potential options for clinicians treating athletes with patellar tendon pain in-

72 season. It has also been shown that short-duration isometric contractions were as effective as longer

73 duration contractions in relieving patellar tendon pain, potentially giving clinicians greater options in

74 the prescription of isometric exercise [12].

75 Research suggests that of the two main calf muscles forming the Achilles tendon, soleus weakness,

76 more than gastrocnemius weakness, may be a feature of chronic Achilles tendinopathy [13, 14].

77 Evidence suggests that activities with the knee flexed at 80-90 degrees inhibits the force production

78 capabilities of the gastrocnemius muscle [15] and may be considered a suitable method of biasing

79 the soleus muscle during the assessment of muscle strength and subsequent exercise prescription
80 [16]. Conversely, in full knee extension, the entire Triceps Surae (Gastrocnemius and Soleus) are
81 recruited, and maximal force production may differ from that of the flexed knee position [17].
82 Therefore, one position may be superior to the other with respect to patient tolerance of load and
83 result in increased adherence to rehabilitation. The primary objectives of this pilot study were to
84 determine if isometric plantar flexion resulted in an immediate change in Achilles tendon pain
85 during a loading task and investigate if differences exist between isometrics performed in a position
86 of knee extension, versus knee flexion.

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Methods

This study was a participant-blinded, randomized, cross-over pilot trial. We confirm that we have read and understood the journal’s ethical standards document [18] and confirm that the study meets the ethical standards of the journal. Ethical approval was obtained from the National Research Ethics Service (NRES) in September 2018.

Participants

Participants were recruited from patients referred to the Norfolk Community Health and Care NHS Trust (NCH&C) physiotherapy services by their GP and in addition, from local running clubs. Participants wishing to take part provided written informed consent at their initial assessment. Participants were eligible for inclusion if they were over 18, had mid-portion Achilles tendon pain for greater than 3 months (Supplementary File A: participant selection of Image B or D) had a score of less than 80 points on the Victorian Institute Sport Assessment – Achilles (VISA-A) and reported only mid-portion Achilles tendon pain during at least one of the following tests performed at the initial physiotherapy appointment: I) double leg heel raise (II) single leg heel raise (III) double leg hop on spot (IV) single leg hop on spot. All four tests were performed consecutively to establish the test that reproduced the patient’s pain between 2 - 7/10 on the NPRS (Supplementary File B). This range was pre-defined as our “clinical tolerance range” based on previous research [19, 20]. The test with the highest level of pain was documented as these tests represent increasing Achilles tendon load. Participants were excluded if they had pain outside the “clinical tolerance range” during any of the four tests above, a history of inflammatory systemic disease or fluoroquinolone(s) usage and pain in any area other than the mid-portion of the Achilles tendon. Participants completed both interventions but were randomized to one of two intervention groups, isometric plantar flexion with the knee in flexion (FLX) or extension (EXT), at their first session of data collection. The order of completion for the two interventions were randomized for all participants by the drawing of sealed, opaque envelopes containing the allocation sequence.

Baseline Strength Assessment.

116 Isokinetic dynamometry was used to measure participant maximal voluntary isometric contraction
117 (MVIC) torque (Nm) for the two test positions. The participant was positioned in long sitting, with
118 the back rest and hip joint range of movement standardised to 60 degrees. This position
119 demonstrates test retest reliability, accuracy and reproducibility [21, 22, 23]. The lateral malleolus
120 was aligned with the rotational axis of the dynamometer lever arm and the ankle positioned in 10
121 degrees of dorsiflexion. This range of movement was selected because prior pilot testing
122 demonstrated peak MVIC torque between neutral and 20 degrees of ankle dorsiflexion. The
123 participant completed a five-minute stationary cycle and three sub-maximal isometric contractions
124 as a warm-up. The highest recorded value from three, maximal efforts was then recorded as the
125 MVIC for that position. The guidance and encouragement given during the procedure was
126 standardized to reduce the risk of performance bias.

127 **Intervention**

128 Each participant completed two exercises for this study: isometric ankle plantarflexion (at 10
129 degrees of dorsiflexion) with an extended knee (EXT) and knee flexed to 80 degrees (FLX). The two
130 test positions are shown in Figure 1 below. To ensure recovery of the calf muscle(s) there was a
131 minimum of five minutes rest following the MVIC testing procedure. Each participant was positioned
132 so that they could monitor the torque vs time graph displayed on the computer monitor. The
133 participant completed five isometric, ankle plantarflexion contractions at 70% of their MVIC each
134 lasting 45 seconds. 70% of MVIC was chosen as the target exercise intensity because in a previous
135 laboratory study, an isometric quadriceps exercise performed at 70% of MVIC reduced patella
136 tendon pain immediately and for up to 45 minutes during a previously painful functional load test [7,
137 10]. The participant was advised that it was acceptable to maintain the isometric contraction within
138 a range 5% above and below this value. Each set of 45 second contractions was separated by a two-
139 minute rest period [7, 24]. The guidance given during each phase of data collection was standardized
140 to reduce the risk of bias influencing performance (Supplementary File C). Participants were blinded
141 to the hypotheses of the study but supervision of the exercise(s) and outcome measure assessment

142 were completed by the principal investigator (BB). Participants switched to the alternate exercise
143 after a minimum of one week.

144 **Outcome Measure**

145 The primary outcome measure was the level of patient-reported Achilles tendon pain during one of
146 four specific, functional load test(s) on a Numeric Pain Rating Scale (NPRS). The intention was to
147 establish which of the four progressive Achilles tendon tests reproduced the patients Achilles tendon
148 pain between 2 and 7/10 on the NPRS. In the event that more than one test reproduced pain, the
149 test with the highest level of pain was used as the pre and post-intervention outcome measure as
150 these tests represent increasing Achilles tendon load. The guidance given during performance of
151 each Achilles tendon test was standardized (Supplementary File C) to reduce the risk of performance
152 bias. The NPRS is an 11-point verbal, pain-rating scale ranging from 0 (no pain) to 10 (worst pain
153 ever) and has been shown to be a reliable, valid and responsive measurement of pain for
154 musculoskeletal pain conditions [25].

155 Previous research in the patellar tendon have shown large effect sizes (>0.8) for within-group
156 change when performing isometrics [7, 10] and between group change when comparing isometrics
157 to isotonics [10]. However, the within-group change in the only paper examining Achilles isometrics
158 displays a small effect size (0.27). As the diagnostic criteria for this pilot study are more similar to
159 those of the patellar tendon it was hypothesised that participants would experience a large effect
160 size (>0.8). Therefore, a minimum sample size of 12 was determined for paired t-tests using G.Power
161 version 3.1.9.4 based on detecting an effect size of 0.9 with 80% power and two-tailed significance
162 set to 0.05.

163 **Statistical Methods**

164 Data analysis was conducted using SPSS v25 (IBM Corp. released 2017). Participant age, sex, height,
165 weight, body mass index (BMI), symptom duration (months), baseline VISA-A and which side the
166 participant had symptoms were described using count, mean and standard deviation (SD), where
167 appropriate. The value for the NPRS at each timepoint are reported as mean (SD). Changes in the

168 NPRS which reduced pain are reported as a negative value. The effect of isometrics on tendon pain,
169 for both FLX and EXT were reported as absolute mean (SD), percentage mean (SD). Within-group
170 differences were determined using paired t-tests. A generalised estimating equation (GEE) was used
171 to determine if there was a difference between plantar flexion isometrics performed with FLX versus
172 EXT when controlling for confounding/ influencing factors. It has been suggested that clinically,
173 people with Achilles tendon pain who have larger baseline NPRS may have a larger absolute value
174 reduction in the NPRS following isometrics. (Cook, Personal Communication, 2020). Therefore, in
175 additional to testing order being included as a factor, baseline pain was reported as a covariate. To
176 account for baseline pain as a possible influence, the relative effect (percentage change in the NPRS)
177 with isometrics was assessed in an additional GEE to determine if there was a difference between
178 plantar flexion isometrics performed with the knee straight versus the knee bent when controlling
179 for testing order as a factor. Significance was set as <0.05.

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Results

194 Between 01 January and 01 October 2019, 42 patients with Achilles tendon pain were screened for
195 eligibility. 17 out of 42 patients were not eligible to participate (7 patients had pain for <12 weeks, 7
196 patients self-reported pain other than the mid-portion of Achilles tendon and 3 patients had a
197 rheumatological condition.) 14 participants declined to participate. 11 participants with chronic,
198 mid-portion Achilles tendinopathy were recruited to study. There were no differences between the
199 groups allocated to FLX first or EXT first in terms of age, height, weight, BMI and baseline VISA-A
200 score with all baseline data presented in Table 1 below. The difference in the mean symptom
201 duration was likely due to the randomisation to FLX first of a participant who had had episodic
202 Achilles tendon pain for 6 years.

203 **Within-group differences**

204 Individual patient data are presented in Supplementary File D. Isometric ankle plantarflexion in FLX
205 and EXT resulted in a significant reduction of pain when performing a load test (Table 2). Isometric
206 ankle plantar flexion with EXT resulted in a 52% mean reduction in symptoms ($p=0.001$) whereas
207 isometric ankle plantar flexion with FLX resulted in a 47% mean reduction in symptoms ($p=0.016$).

208 **Between group differences**

209 The GEE using the absolute change value on the NPRS (Table 3) showed that there was an additional
210 0.47 point reduction in pain on the NPRS during a functional loading task for every 1-point increase
211 of the baseline NPRS ($p=0.002$). Due to this influence a subsequent GEE was performed using the
212 relative change score (percentage change) instead of the absolute score (raw-value) (Table 4). This
213 subsequent GEE showed that there was no significant effect of the knee position or testing order on
214 the percentage of pain change when completing isometric plantar flexion. However, whilst not
215 significant ($p=0.110$) isometric plantar flexion in knee extension appeared to give a 20% larger
216 reduction in pain when compared to isometric plantar flexion in knee flexion.

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Discussion

This pilot study demonstrated that two positions for isometric exercise were well tolerated for people with mid-portion Achilles tendinopathy, resulting in pain reduction of approximately 50% in a functional task. For this small study, a 20% reduction in tendon pain was observed with EXT compared to FLX but this was not statistically significant. However, the clinical significance of these changes is uncertain and this warrants further investigation in a larger RCT. Complete rest is detrimental for musculoskeletal capacity so ensuring the musculoskeletal system is getting as much load as tolerated, as soon as possible is vital for effective rehabilitation [26]. Isometric holds may provide a well-tolerated start point for rehabilitation, especially if patients have a fear of movement/ loading which has been demonstrated in tendinopathy [27]. The immediate pain relief observed following isometric plantar flexion in this study may assist the management of the patient with Achilles tendon pain in several ways. Firstly, it may allow completion or participation in a previously painful task or activity. Secondly, as a non-pharmacological treatment, exercise is not associated with the side effects of medications that may typically be taken for tendon pain (e.g. non-steroidal anti-inflammatory medications)[28]. Furthermore, exercise is likely to have many other, general positive impacts for the patient that will not occur with medication use [29]. Finally, if patients had strategies to independently manage their pain, it may improve adherence with any further rehabilitation [30].

This study differs from a previous case series study [9] investigating the efficacy of isometric exercise in Achilles tendinopathy in several respects. Only 9 of the 16 participants in O’Neil’s study (2019) [9] reported pain during a bilateral or unilateral heel raise or subsequent hopping test, the loading test(s) used for inclusion within our study. O’Neil et al [9] placed a stronger emphasis on palpatory pain. Other differences between the two studies include how MVIC torque was assessed, the apparatus used for assessment and treatment and the range of dorsiflexion in which isometrics were performed. The diagnostic criteria for tendinopathy are not universal [31]. It is not clear from the current literature if there is a heterogenous response to isometric (or any) exercise or alternatively if

243 there is heterogeneity in the included participants and procedures. In this study participants
244 demonstrated increasing pain with increasing load that remained localised and the clinical take
245 home is that isometric exercise is well tolerated in people with that phenotype. Whether this can be
246 applied to people that have pain with Achilles tendon palpation and imaging change (common
247 inclusion criteria used) is not known [32].

248 Although we primarily sought to recruit participants through the NHS, we struggled to recruit the
249 numbers expected. Therefore, any interpretation of our data must acknowledge that the sample size
250 was small. Despite these difficulties, it was felt that the robustness of the study was improved by the
251 implementation of strict eligibility criteria to ensure that the final sample was representative of the
252 patient with chronic, mid-portion Achilles tendon pain [33]. The number of participants that were
253 included in the final analysis are comparable to other studies that have examined the use of
254 isometric exercise in the management of tendon pain [7,11] including those that did not find any
255 effect from isometric exercise on tendon pain [9, 34]. Finally, it is acknowledged that due to the
256 absence of a placebo intervention, and as blinding of the outcome assessor was not possible, this
257 does introduce the potential for bias. However, to negate the possibility of performance bias
258 influencing the participants response to the exercises, we chose to standardise our methods of data
259 collection, including the use of standardised explanations of the study intervention(s) and all
260 procedures for data collection (Supplementary File C).

261 Whilst recognising the methodological limitations of this pilot study, the isometric exercises
262 investigated were well tolerated and regardless of the position of the knee joint offered small but
263 statistically significant reductions in pain during a functional load test. The protocol investigated two
264 options for isometric contractions and provides a viable option for clinicians to use in the
265 management of Achilles tendon pain. For example, it may be that clinicians look to implement
266 isometric exercise as the starting point to rehabilitation, to demonstrate to the fearful patient that
267 loading the tendon with exercise is safe or as a means of promoting self-efficacy and adherence with
268 any ongoing strengthening programme. Finally, isometric exercise should never be expected to

269 provide comprehensive rehabilitation and restoration of function in those with Achilles
270 tendinopathy and immediate analgesia is not the only reason to provide an exercise to a patient
271 with Achilles tendinopathy. If exercise is truly going to be used as medicine then studies should seek
272 to identify what exercise suits which patient, and with what dosage and progression. This requires
273 adequate description of participants to identify sub-groups of responders and non-responders to aid
274 clinical interpretation [31].

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295 **Legend for Tables and Figures**

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297 **Table 1: Baseline Data.**

298 **Legend:** EXT= extension, FLX= flexion, BMI= body mass index, VISA-A= Victorian Institute of Sport

299 Assessment- Achilles ¹data are presented as number, ²data are presented as mean (SD)

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301 **Table 2: Within-group differences of pain with isometric plantar flexion.**

302 **Legend:** * Significance of $p < 0.05$

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304 **Table 3: Generalised estimating equation of absolute change in pain with isometric plantar flexion.**

305 **Legend:** β = beta, SE= standard error, 95%CI= 95% confidence interval

306 ^a Knee flexion set to 0, ^bIsometric plantar flexion in knee flexion first set to 0

307 * Significance of $p < 0.05$

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309 **Table 4: Generalised estimating equation of relative change in pain with isometric plantar flexion .**

310 **Legend:** β = beta, SE= standard error, 95%CI= 95% confidence interval,

311 ^a Knee flexion set to 0, ^bIsometric plantar flexion in knee flexion first set to 0,

312 * Significance of $p < 0.05$

313

314 **Figure 1: EXT and FLX Isometric Exercise Position(s)**

315 **Legend:** EXT – the knee joint positioned in terminal extension and secured with non-elastic

316 strapping. FLX – the knee joint positioned at 80 degrees of flexion and secured against the thigh

317 stabilizer with non-elastic strapping.

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321 **Reference List**

322

323 1. van Dijk CN, van Sterkenburg MN, Wiegerinck JI et al. **Terminology for Achilles tendon related**
324 **disorders.** *Knee Surg Sports Traumatol Arthrosc* 2011; 19: 835-841.

325

326 2. Scott A, Squier K, Alfredson H, et al. **ICON 2019: International Scientific Tendinopathy**
327 **Symposium Consensus: Clinical Terminology.** *Br J Sports Med* 2020; 54:260-262.

328

329 3. Alfredson H, Pietilä T, Jonsson P et al. **Heavy-load eccentric calf muscle training for the treatment**
330 **of chronic Achilles tendinosis.** *Am J Sports Med* 1998; 26: 360-366.

331

332 4. Murphy MC, Travers MJ, Chivers P et al. **Efficacy of heavy eccentric calf training for treating mid-**
333 **portion Achilles tendinopathy: a systematic review and meta-analysis.** *Br J Sports Med* 2019;
334 bjsports-2018-099934. doi:10.1136/bjsports-2018-099934

335

336 5. Murphy M, Travers MJ, Gibson W et al. **The rate of improvement of pain and function in mid-**
337 **portion Achilles tendinopathy with loading protocols: A Systematic Review and Longitudinal Meta-**
338 **Analysis.** *Sports Med* 2018; 48: 1875-1891.

339

340 6. Kearney RS, Parsons N, Metcalfe D et al. **Injection therapies for Achilles tendinopathy.** *Cochrane*
341 *Database Syst Rev* 2015; CD010960. Published 2015 May 26.
342 doi:10.1002/14651858.CD010960.pub2.

343

344 7. Rio E, Kidgell D, Purdam C et al. **Isometric exercise induces analgesia and reduces inhibition in**
345 **patellar tendinopathy.** *Br J Sports Med* 2015; 49: 1277-1283.

346

- 347 8. Mc Auliffe S, Synott A, Casey H et al. **Beyond the tendon: Experiences and perceptions of people**
348 **with persistent Achilles tendinopathy.** *Musculoskelet Sci Pract* 2017; 29: 108-114.
349
- 350 9. O'Neill S, Radia J, Bird K et al. **Acute sensory and motor response to 45-s heavy isometric holds**
351 **for the plantar flexors in patients with Achilles tendinopathy.** *Knee Surg Sports Traumatol Arthrosc*
352 2019; 27: 2765-2773.
353
- 354 10. Rio E, van Ark M, Docking S et al. **Isometric Contractions Are More Analgesic Than Isotonic**
355 **Contractions for Patellar Tendon Pain: An In-Season Randomized Clinical Trial.** *Clin J Sport Med*
356 2017; 27: 253-259.
357
- 358 11. van Ark M, Cook JL, Docking SI et al. **Do isometric and isotonic exercise programs reduce pain in**
359 **athletes with patellar tendinopathy in-season? A randomised clinical trial.** *J Sci Med Sport* 2016;
360 19: 702-706.
361
- 362 12. Pearson SJ, Stadler S, Menz H et al. **Immediate and Short-Term Effects of Short- and Long-**
363 **Duration Isometric Contractions in Patellar Tendinopathy [published online ahead of print, 2018**
364 **Aug 8].** *Clin J Sport Med.* 2018; 10.1097/JSM.0000000000000625.
365 doi:10.1097/JSM.0000000000000625
366
- 367 13. Wyndow N, Cowan SM, Wrigley TV, Crossley KM. **Triceps surae activation is altered in male**
368 **runners with Achilles tendinopathy.** *J Electromyogr Kinesiol* 2013; 23: 166-172.
369
- 370 14. O'Neill S, Barry S, Watson P. **Plantarflexor strength and endurance deficits associated with mid-**
371 **portion Achilles tendinopathy: The role of soleus.** *Phys Ther Sport.* 2019; 37: 69-76.
372

- 373 15. Lenhart RL, Francis CA, Lenz AL, Thelen DG. **Empirical evaluation of gastrocnemius and soleus**
374 **function during walking.** *J Biomech.* 2014; 47: 2969-2974.
375
- 376 16. Cronin NJ, Avela J, Finni T, Peltonen J. **Differences in contractile behaviour between the soleus**
377 **and medial gastrocnemius muscles during human walking.** *J Exp Biol* 2013; 216: 909-914.
378
- 379 17. Al-Uzri M, O'Neil S, Watson P, Kelly C. **Reliability of isokinetic dynamometry of the**
380 **plantarflexors in knee flexion and extension.** *Physio Pract. & Res* 2017; 38: 49–57.
381
- 382 18. Harriss DJ, Macsween A, Atkinson G. **Ethical standards in sport and exercise science research:**
383 **2020 update.** *Int J Sports Med* 2019; 40: 813 - 817
384
- 385 19. Silbernagel KG, Thomeé R, Eriksson BI, Karlsson J. **Continued sports activity, using a pain-**
386 **monitoring model, during rehabilitation in patients with Achilles tendinopathy: a randomized**
387 **controlled study.** *Am J Sports Med* 2007; 35: 897-906.
388
- 389 20. Smith BE, Hendrick P, Smith TO et al. **Should exercises be painful in the management of chronic**
390 **musculoskeletal pain? A systematic review and meta-analysis.** *Brit J of Sports Med* 2017; 51: 1679-
391 1687.
392
- 393 21. Moller M, Lind K, Styf J, Karlsson J. **The test-retest reliability of concentric and eccentric muscle**
394 **action during plantar flexion of the ankle joint in a closed kinetic chain.** *Isokinetic Exercise Science*
395 2000; 8: 223- 228.
396
- 397 22. Moller M, Lind K, Styf J, Karlsson J. **The reliability of isokinetic testing of the ankle and a heel-**
398 **raise test for endurance.** *Knee Surg Sports Traumatol Arthrosc* 2005; 13: 60-71

399

400 23. Harbo T, Brincks J, Andersen H. **Maximal isokinetic and isometric muscle strength of major**
401 **muscle groups related to age, body mass, height, and sex in 178 healthy subjects.** *Euro J Applied*
402 *Phys* 2012; 112: 267–275.

403

404 24. Ahtiainen JP, Pakarinen A, Kraemer WJ, Häkkinen K. **Acute hormonal and neuromuscular**
405 **responses and recovery to forced vs maximum repetitions multiple resistance exercises.** *Int J*
406 *Sports Med* 2003; 24: 410-418.

407

408 25. Ferreira-Valente MA, Pais-Ribeiro JL, Jensen MP. **Validity of four pain intensity rating**
409 **scales.** *Pain* 2011; 152: 2399-2404.

410

411 26. Docking SI, Cook J. **How do tendons adapt? Going beyond tissue responses to understand**
412 **positive adaptation and pathology development: A narrative review.** *J Musculoskelet Neuronal*
413 *Interact* 2019; 19: 300-310.

414

415 27. Corrigan P, Cortes DH, Pontiggia L, Silbernagel KG. **The degree of tendinosis is related to**
416 **symptom severity and physical activity levels in patients with midportion Achilles tendinopathy.**
417 *Int J Sports Phys Ther* 2013; 13: 196-207

418

419 28. Aminoshariae A, Kulild JC, Donaldson M. **Short-term use of nonsteroidal anti-inflammatory**
420 **drugs and adverse effects: An updated systematic review.** *J Am Dent Assoc* 2016; 147: 98-110.

421

422 29. Warburton DER, Bredin SSD. **Health benefits of physical activity: a systematic review of current**
423 **systematic reviews.** *Curr Opin Cardiol* 2017; 32: 541-556.

424

425 30. Silbernagel KG, Hanlon S, Sprague A. **Current Clinical Concepts: Conservative Management of**
426 **Achilles Tendinopathy.** *J Athl Train* 2020; 55: 438-447.
427

428 31. Rio EK, Mc Auliffe S, Kuipers I et al. **ICON PART-T 2019–International Scientific Tendinopathy**
429 **Symposium Consensus: recommended standards for reporting participant characteristics in**
430 **tendinopathy research (PART-T)** *Br J Sports Me.* 2020; 54: 627-630.
431

432 32. Vicenzino B, de Vos RJ, Alfredson H et al. **ICON 2019-International Scientific Tendinopathy**
433 **Symposium Consensus: There are nine core health-related domains for tendinopathy (CORE**
434 **DOMAINS): Delphi study of healthcare professionals and patients.** *Br J Sports Med.* 2020; 54: 444-
435 451.
436

437 33. Martin RL, Chimenti R, Cuddeford T et al. **Achilles Pain, Stiffness, and Muscle Power Deficits:**
438 **Midportion Achilles Tendinopathy Revision 2018.** *J Orthop Sports Phys Ther* 2018; 48: A1-A38.
439

440 34. Riel H, Vicenzino B, Jensen MB et al. **The effect of isometric exercise on pain in individuals with**
441 **plantar fasciopathy: A randomized crossover trial.** *Scand J Med Sci Sports* 2018; 28: 2643-2650.
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