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Immediate effects of dry needling on pain sensitivity and pain modulation in patients with chronic idiopathic neck pain : a single-blinded randomized clinical trial

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1 Immediate effects of Dry Needling on Pain Sensitivity and Pain  
2 Modulation in Patients With Chronic Idiopathic Neck Pain: A single-  
3 blinded randomized clinical trial

4 Running title: Effects of Dry Needling on Pain Sensitivity and Pain Modulation.

5 Marjolein Chys<sup>1</sup>, Jente Bontinck<sup>1,2</sup>, Lennard Voogt<sup>2,3</sup>, Gracia María Gallego Sendarrubias<sup>4</sup>, Barbara  
6 Cagnie<sup>1</sup>, Mira Meeus<sup>1,2,5</sup>, Kayleigh De Meulemeester<sup>1,2</sup>

7 <sup>1</sup>Spine, Head and Pain Research Unit Ghent, Department of Rehabilitation Sciences, Faculty of  
8 Medicine and Health Sciences, Ghent University, Belgium

9 <sup>2</sup>Pain in Motion International Research Group, , [www.paininmotion.be](http://www.paininmotion.be)

10 <sup>3</sup>Research Centre for Health Care Innovations, Rotterdam University of Applied Sciences, Rotterdam,  
11 The Netherlands

12 <sup>4</sup>Department of Physical Therapy, Camilo José Cela University, Madrid, Spain

13 <sup>5</sup>MOVANT Research group, Department of Rehabilitation Sciences and Physical Therapy, Faculty of  
14 Medicine and Health Sciences, University of Antwerp, Belgium

15

16 Corresponding author: Marjolein Chys, [Marjolein.Chys@Ugent.be](mailto:Marjolein.Chys@Ugent.be), Ghent University, Corneel  
17 Heymanslaan 10, 9000 Gent, Belgium

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## 24 Abstract

25 **Background:** Dry needling is frequently used for the treatment of neck pain but knowledge about its  
26 neurophysiological central effects is scarce.

27 **Objectives:** To compare the immediate effects of a single session of dry needling (DN) and sham needling  
28 (SN) on local and distant pressure pain thresholds and conditioned pain modulation in patients with  
29 chronic idiopathic neck pain.

30 **Method:** Participants with chronic idiopathic neck pain were randomly allocated to a DN or SN group.  
31 The primary outcome measure was the pressure pain threshold (PPT) at one peripheral location:  
32 quadriceps muscle (Q). Secondary outcome measures were local PPTs at the treated (most painful) (tUT)  
33 and non-treated upper trapezius muscle (ntUT), absolute and relative conditioned pain modulation  
34 (CPM) effects and pain during hot water immersion. Patients were assessed at baseline and immediately  
35 post intervention. Linear mixed models were used to examine interaction effects as well as between-  
36 and within-group differences.

37 **Results:** Fifty-four participants were included for statistical analysis. Linear mixed model analyses  
38 showed no significant “group X time” interaction effects for any of the outcome measures. The relative  
39 CPM effect at the Q was significantly higher post-intervention, compared to baseline within the DN  
40 group (mean difference= 13.52%; 95% CI: 0.46, 26.59).

41 **Conclusion:** The present study shows no superior effect of DN, compared to SN, in the immediate effect  
42 on local and distant PPTs and CPM in patients with chronic idiopathic neck pain.

43 **Keywords:** chronic neck pain, dry needling, pain modulation, pain sensitivity.

44

## 45 Highlights

- 46 • It is hypothesized that dry needling improves pain sensitivity and pain modulation
- 47 • There was no difference on pain sensitivity or pain modulation between dry needling and  
48 sham needling
- 49 • Dry needling shows better antinociceptive pain modulation after treatment

## 50 Introduction

51 Over the last few years, the number of studies suggesting myofascial pain syndrome (MPS) as one of  
52 the possible underlying causes of chronic idiopathic neck pain (CINP) has increased.<sup>1-4</sup> CINP can be  
53 associated with (referred) muscle pain caused by active or latent myofascial trigger points (MTrPs).<sup>5</sup> The  
54 prolonged presence of MTrPs may lead to altered peripheral and central pain processing, also referred  
55 to as peripheral and central sensitization (CS).<sup>6-8</sup> Peripheral primary sensory neurons and pain-  
56 processing neurons in the spinal cord and brain become more sensitive due to neuronal plasticity caused  
57 by continuous nociceptive afferent information coming from the MTrP to spinal cord neurons and supra-  
58 spinal structures of the central nervous system.<sup>6</sup> Nevertheless, the presence and clinical importance of  
59 CS in CINP is still under discussion.<sup>9-13</sup>

60 Although there is no gold standard to diagnose CS, multiple screening and diagnostic tools have already  
61 been established.<sup>8</sup> A screening questionnaire that identifies self-reported signs of CS is the Central  
62 Sensitization Inventory (CSI).<sup>14</sup> Another option is the use of Quantitative Sensory Testing (QST).<sup>15</sup> This  
63 testing includes, amongst others, the determination of local and distant pain sensitivity or hyperalgesia  
64 as assessed by pressure pain thresholds (PPTs) and endogenous pain inhibition efficiency as assessed by  
65 conditioned pain modulation (CPM) paradigms.<sup>16, 17</sup> Changes in central nociceptive processing may  
66 explain persistent and recurrent symptoms in CINP and failure of treatments to obtain long-lasting  
67 relief.<sup>11-13</sup>

68 A common intervention for treatment of MTrPs is dry needling (DN). Although several local and  
69 mechanical effects have already been established, more research is needed on the unclear underlying  
70 central neurophysiological effects of DN. Preliminary experimental evidence shows that the application  
71 of DN may be able to reduce the excitability of the central nervous system in patients with chronic  
72 pain.<sup>18, 19</sup> Niddam et al. found in an MRI study that pain mediation after DN happens through the  
73 periaqueductal gray substance in the brainstem, possibly indicating that DN may activate  
74 enkephalinergic inhibitory dorsal horn interneurons.<sup>18</sup> Stieven et al.<sup>20</sup> found that a single application of  
75 DN in CINP resulted in higher local and distant PPTs, compared to sham needling (SN).<sup>20</sup> However, only  
76 a paucity of trials about the effect of DN on PPTs and CPM have been performed to date.<sup>19-21</sup>

77

78 Consequently, the aim of this randomized controlled trial was to compare the immediate effects of a  
79 single DN or SN session on distant and local PPTs and CPM in patients with CINP. It was hypothesized  
80 that DN would have immediate positive effects resulting in higher distant and local PPTs (reflecting a  
81 decrease in pain sensitivity) and higher CPM (reflecting more efficient pain inhibition).

## 82 Methods

### 83 Protocol and registration

84 This study design was approved by the Ethics and Research Committee of Ghent University (project  
85 number EC2019/0980) and prospectively registered at Clinicaltrials.gov (registration number:  
86 NCT04725825). This trial was reported according to the recommendations of the Consolidated  
87 Standards of Reporting Trials (CONSORT) statement.<sup>22, 23</sup>

88

### 89 Study Population

90 Between February 2021 and July 2021, patients with CINP were recruited for this study. Patients were  
91 recruited by flyers at the waiting rooms for physical medicine and rehabilitation of the Ghent University  
92 Hospital and on social media. Before participating, patients were asked to complete an online  
93 questionnaire concerning their current neck complaints and general health. After completing the online  
94 questionnaire, all participants were selected based on inclusion and exclusion criteria as stated in Table  
95 1. All eligible individuals provided informed consent and were informed about the study procedures  
96 before the trial started.

97

98 Table 1: Inclusion and exclusion criteria.

Inclusion criteria	Exclusion criteria
Age between 18 and 65 years old	Patients with a specific cause of their neck pain (such as cervical radiculopathy and/or myelopathy, severe osteoarthritis, fractures)
Chronic neck pain, present for more than three months	Major depression or any other psychiatric condition
An average NPRS of three or more during the past month	Life-threatening metabolic diseases (such as diabetes mellitus and any symptoms of restless legs, etc.),
Presence of a clinically relevant trigger point in the upper trapezius muscle using the following criteria: “(A) a palpable taut band of skeletal muscle, (B) exquisite (unusual) local muscle tenderness in the taut band, (C) patient pain recognition and (D) patient pain referral”. <sup>47</sup>	Transmittable diseases (such as Hepatitis, HIV, etc.)
	Cardiovascular, neurological, and systemic diseases
	Pregnancy or given birth in the past year
	Fear of needles and/or presence of other conditions that preclude dry needling
	Clotting disorders, use of blood thinning medication
	A history of head, neck, or shoulder surgery
	Fibromyalgia or chronic fatigue syndrome
	BMI > 30 kg/m <sup>2</sup>

	Whiplash within the past 10 years, current neck pain associated to this whiplash and/or whiplash associated disorder
	Inability to read or understand Dutch
	Being in treatment for neck pain during the study
	Skin abnormalities at the treatment region

99 BMI, body mass index; NPRS, numeric pain rating scale

100

### 101 Randomization procedure and blinding

102 All procedures were performed at the Department of Rehabilitation Sciences, Ghent University.  
 103 Participants were randomly allocated to one of the 2 study groups (DN or SN) by an independent  
 104 researcher, using an internet-based randomization website ([www.randomizer.org](http://www.randomizer.org)) with an allocation  
 105 ratio of 1:1. Allocation concealment was guaranteed by using sealed opaque envelopes. All participants  
 106 were informed that they would be randomly assigned to one of the two study groups and were blinded  
 107 for treatment allocation. All outcome measures were assessed at baseline and immediately post-  
 108 intervention by assessors blinded to treatment allocation.

109

### 110 Sample size determination and pilot

111 A total sample size of at least 36 subjects had to be recruited based on an a priori sample size calculation  
 112 (G\*Power 3.1.9.2). This calculation was determined for the primary outcome measure “PPT Quadriceps”  
 113 and was based on pilot data, which showed an effect size of 0.28 for the difference between a DN group  
 114 (n=9) and a SN group (n=9) post intervention. The a priori sample size calculation was performed for the  
 115 within-between interaction in a repeated-measures analysis of variance with two groups and two  
 116 measurements, a minimum power of 0.90, an effect size of 0.28, and an  $\alpha$  level of 0.05.  
 117 The PPT data from this pilot study were pooled with the data from the present study, which resulted in  
 118 a total sample size of 54 participants for the PPT data and 36 participants for CPM data.

119

### 120 Interventions

121 Both groups received one single needling intervention at the upper trapezius (UT) of the (most) painful  
 122 side, there was no follow-up treatment. All interventions were performed by one of the three trained  
 123 physical therapists with at least 4 years of experience in the treatment of MPS and manual therapy. All  
 124 therapists performed both interventions. Prior to the intervention, therapists provided the same  
 125 standardized information to all participants about MTrPs, the intervention and possible post-  
 126 intervention effects. The interventions were performed with a solid filiform needle (0.30x0.40 mm C-  
 127 Type acupuncture needle). Participants were placed in a prone position with their arms comfortably  
 128 supported in 90° shoulder abduction.

129

### 130 *Dry needling*

131 The DN was applied unilaterally at the (most) painful UT. First, the skin was cleaned with alcohol and a  
132 relevant MTrP was identified. Second, the skin was pierced subcutaneously at the MTrP location,  
133 followed by piercing into the muscle tissue in a poster-anterior direction (from therapist's thumb to  
134 index), while the muscle belly was held in a pincer palpation. The "fast in, fast out" method was used,  
135 for this technique the needle was quickly moved up- and downwards into the muscle fibers of the taut  
136 band with the aim of provoking local twitch responses (LTRs) until extinction. In case no LTRs were  
137 elicited, the needle was moved up and downwards for 10 times in 3 slightly different directions and was  
138 then withdrawn from the muscle.

139

### 140 *Sham needling*

141 The same procedure as for the DN group was implemented to replicate an authentic clinical experience  
142 and maintain credibility and participants' blinding.<sup>24</sup> The needle was inserted into the subcutaneous  
143 layer and went up and down 10 times on the MTrP location without penetrating the deep muscle fascia  
144 while the therapist pretended to change the direction of the needle 3 times. Because the needle did not  
145 penetrate the muscle fascia, no LTRs were provoked.<sup>21, 25-27</sup> Contextual clues associated with DN such as  
146 skin's cleaning, needle insertion, and manipulation (simulation in sham needling), and haemostatic  
147 compression after procedure were identical in both interventions.<sup>24</sup>

148

### 149 *Outcome measures*

150 The outcome measurements were performed by three independent assessors who were blinded to  
151 treatment allocation. Baseline and post-intervention measurements for each participant were always  
152 performed by the same assessor. During the testing, the patient was placed in a seated position with a  
153 neutral spine and the feet flat on the ground. First, each patient was asked to score their NP at that  
154 moment on a numeric pain rating scale (NPRS). Second, PPTs were measured on both UT muscles  
155 (treated and non-treated side) and quadriceps muscle for the treated side. The sequence of PPT muscle  
156 testing was randomly selected via the online tool Randomizer ([www.randomizer.org](http://www.randomizer.org)). Third, for the  
157 CPM protocol, the function of the descending pain inhibitory pathways was evaluated by examining the  
158 effect of a conditioning stimulus of the non-dominant hand (hot water immersion) on the PPTs.  
159 Additionally, pain intensity caused by the hot water immersion was assessed on a NPRS. After  
160 implementation of the intervention (DN or SN of the (most) painful UT), the same testing protocol was  
161 repeated.

162

163 *Primary outcome measures:*

164 *Distant PPTs – Quadriceps (distant pain sensitivity/ hyperalgesia) – Fig. 2A.*

165 PPTs were measured at a standardized location with a hand-held pressure algometer (Wagner FPX 25  
166 Force Gage). The quadriceps muscle on the painful side was assessed at the middle of the distance  
167 between the anterior superior iliac spine and the base of the patella.<sup>28, 29</sup> The probe (1cm<sup>2</sup>) was placed  
168 perpendicular to the test surface. The pressure was expressed in Newton (N) and the average was taken  
169 of two measurements with a 30-second interval between each application. Pressure was increased by  
170 1N/s until the participant reported this feeling as unpleasant.<sup>9</sup> Digital algometry performed at the Q  
171 muscle is shown to have a good intrarater reliability (intraclass correlation, 0.74-0.85).<sup>30</sup>

172

173 *Secondary outcome measures*

174 *Local PPTs – treated and non-treated upper trapezius (local pain sensitivity/ hyperalgesia) – Fig. 2B.*

175 PPTs were measured at the treated (tUT) and non-treated upper trapezius (ntUT). The reported treated  
176 side was the most painful side indicated by the patient. The average of two measurements at the middle  
177 between the processus spinous of C7 and the centre of the acromion was calculated.<sup>9</sup> Digital algometry  
178 is shown to have sufficient intrarater reliability in measuring the PPT on the trigger point of the UT  
179 muscle in patients with CINP.<sup>31</sup> The interrater reliability of PPT measurements has shown to be  
180 excellent.<sup>32</sup> In a study of Walton et al., PPT at the UT showed a significant ability to detect global change  
181 (AUC=0.76), using minimal clinically important difference (MCID) change scores within a clinically  
182 reasonable range (between approximately 5 and 22 N/cm<sup>2</sup>).<sup>33</sup> Minimal detectable change (MDC) values  
183 at the UT site ranged between approximately 4.45 and 11.12 N/cm<sup>2</sup>; intrarater reliability was almost  
184 perfect (ICC = 0.94-0.97).<sup>34</sup>

185

186 *Conditioned pain modulation (Efficacy of pain inhibition) – Fig. 2C.*

187 The conditioning stimulus in this study was a 1-minute hot water immersion (45.5°C) of the non-  
188 dominant hand (up to the most distal point of the ulnar styloid process) in a VersaCool Circulating Bath  
189 (Thermo Fisher Scientific).<sup>35, 36</sup> PPTs were used as test stimulus, which are shown to be a valid tool to  
190 measure CPM.<sup>37</sup> For analysis of CPM efficacy, absolute CPM effects were calculated: the mean PPT  
191 measured before the hot water immersion was subtracted from the mean PPT after hot water  
192 immersion (PPT post - PPT pre). Hence, a lower CPM value reflects a less efficient endogenous pain  
193 inhibition, whereas a higher CPM value reflects a more efficient endogenous pain inhibition.  
194 Additionally, the relative CPM effect (CPM efficacy expressed in percent change) was calculated: ((PPT  
195 post – PPT pre)/PPT pre) \* 100. This resulted in either a pronociceptive value (CPM value less than or  
196 equal to zero, indicating a less efficient endogenous pain inhibition: no CPM effect) or antinociceptive



197 value (CPM value more than zero, indicating a more efficient endogenous pain inhibition: CPM effect).  
198 No information about MCID has been found.

199

#### 200 [NPRS during hot water immersion \(heat hyperalgesia\)](#)

201 After placing the hand in the VersaCool for one minute, the patient was asked to score the pain caused  
202 by the hot water on an 11-point NPRS. The MDC and MCID are 2.1 and 1.3 points, respectively, in  
203 patients with mechanical neck pain.<sup>38</sup>

204

#### 205 [Statistical analysis](#)

206 Data analysis was performed based on an intention-to-treat principle with IBM SPSS Statistics version  
207 27.0 (IBM, Armonk, NY, USA) for all outcome measures. Data normality was assessed by means of the  
208 Shapiro-Wilk test, histograms and Q-Q plots. Boxplots were used as quality control to find any outliers  
209 and extreme values. Patients' characteristics, baseline and post-intervention values between groups,  
210 were evaluated with the independent T-test, the Mann-Whitney U test (for the non-normally distributed  
211 data), and the Chi Square test (for categorical variables sex and affected side). Means and standard  
212 deviations were calculated for all demographic data. Linear mixed model analyses were used to  
213 determine the differences of all outcomes between and within the intervention groups over time for  
214 the PPTs, as well as for the absolute and relative CPM effects at the tUT, ntUT, and Q, and for heat  
215 hyperalgesia. Participant number was used as random intercept and residuals were checked for  
216 normality. Fixed factors were 'intervention' (DN and SN group), 'time' (baseline and post-intervention)  
217 and 'intervention x time'. Sex was included as covariate in the linear mixed model analyses. All PPT  
218 analyses were performed on the entire group (DN; N=26 and SN;N=28). CPM data were only available  
219 from a subgroup (DN; N=17 and SN; N=19). Statistical significance was accepted at the 0.05  $\alpha$ -level.

## 220 [Results](#)

### 221 [Participants](#)

222 Fifty-four patients with CINP were randomly allocated to the DN group (n=26) or the SN group (n=28)  
223 (Figure 1). Demographic features of both groups are presented in Table 2. Patients' characteristics  
224 between groups (except for sex) and outcome measures were comparable at baseline (Tables 1 and 2).  
225 The mean NDI and CSI-scores in both groups were considered to represent mild disability levels and  
226 presence of mild features of CS. The mean CSI-score did not reach the clinically relevant cutoff value of  
227 40/100, although some participants reached higher CSI-scores on an individual level.<sup>14</sup>

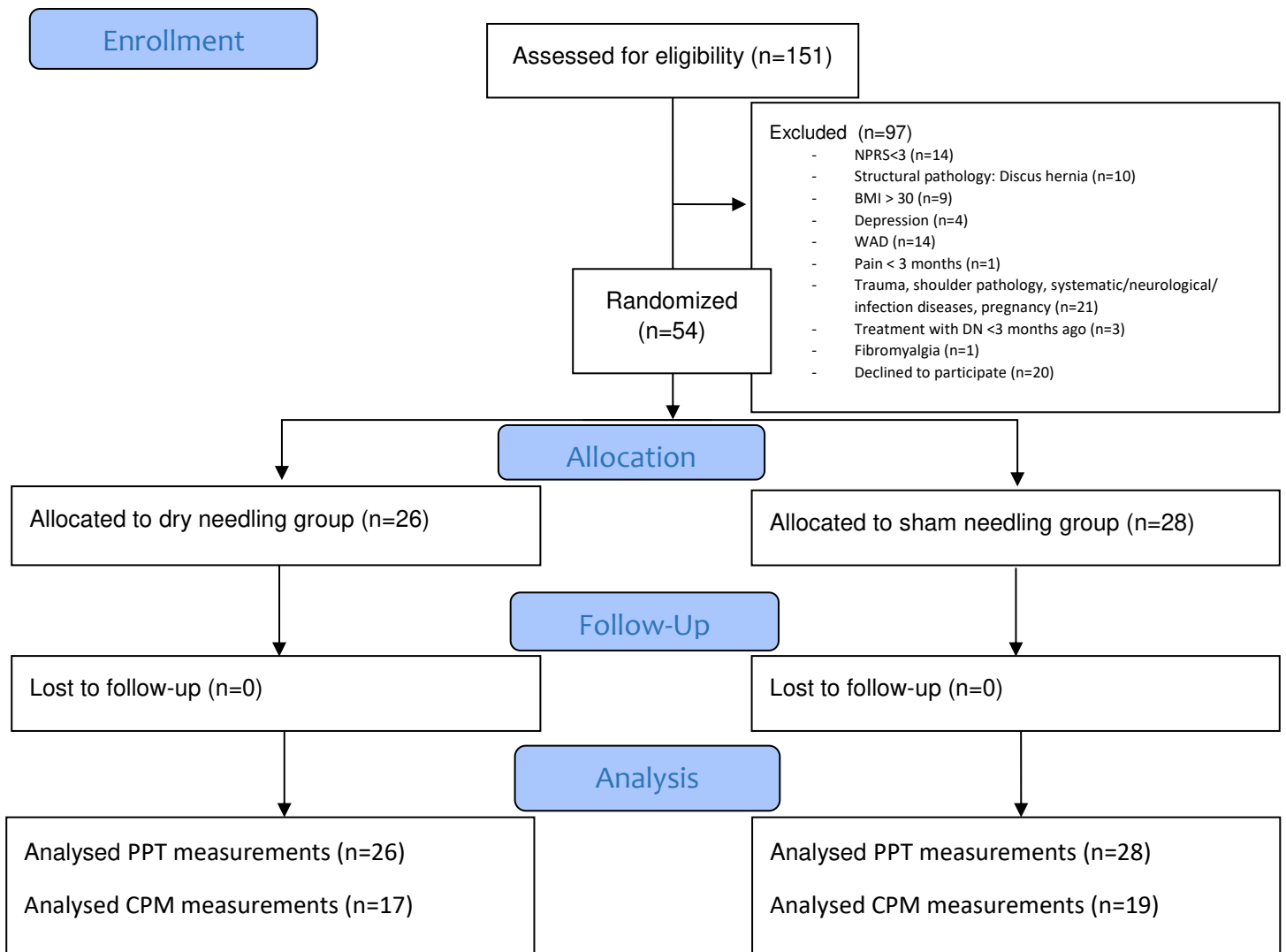


Figure 1: CONSORT Flow Diagram

230 **Table 2.** Patients’ characteristics of the DN and SN Group.

Demographics	Dry needling (n=26)	Sham needling (n=28)
Age, year	33 ± 13.5	32 ± 11.8
Sex, n (%)		
Male	2 (7.7%)	6 (21.4%)
Female	24 (92.3%)	22 (78.6%)
Height, cm	170 ± 6.3	169 ± 10.9
Weight, kg	65.6 ± 9.4	66.6 ± 12.3
BMI, kg/m <sup>2</sup>	22.3 ± 2.7	23.1 ± 2.5
NDI (0-50)	11.2 ± 5.1	10.1 ± 4.0
CSI (0-100)	33.6 ± 14.6	33.9 ± 10.8
Duration, months	(n=23) 89.83 ± 75.9	(n=25) 64.8 ± 44.4
Treated (most painful) side		
Right	17 (65.4%)	14 (50%)
Left	9 (34.6%)	14 (50%)
NPRS treated UT, range 0-10	4.8 ± 1.6	4.1 ± 1.8
Values are expressed as means ± standard deviation for continuous variables and absolute frequency (%) for categorical variables. Abbreviations: n, number of participants; BMI, body mass index; NDI, Neck Disability Index; CSI, Central Sensitization Inventory; NPRS, Numeric Pain Rating Scale; UT, upper trapezius muscle.		

231

232 **Primary and secondary outcome measures**

233 *Data for all outcome measures are provided in Table 3.*

234 *Pressure Pain Thresholds*

235 The linear mixed-models revealed no significant “group x time” interaction effect for PPTs at the tUT,  
 236 ntUT, or Q. No post hoc pairwise comparisons for between-group or within group-comparisons showed  
 237 any significant results. No difference in PPTs between DN and SN groups was found for tUT (mean  
 238 difference [MD]=-1.38; 95% CI: -8.13, 5.37); ntUT (MD= -2.15; 95% CI:-9.52, 5.23); and Q (MD= -0.13;  
 239 95% CI:-9.78 , 9.52). Between-group differences in mean changes from baseline to post-intervention  
 240 were smaller than the reported MDC and MCID.

241 *Absolute CPM effect*

242 The linear mixed-models revealed no significant “group x time” interaction effects for tUT, ntUT, or Q. ,  
 243 No post hoc pairwise comparisons for between-group or within group-comparison show any significant  
 244 results. No difference in absolute CPM effect between DN and SN groups was found for tUT (MD= -0.96;  
 245 95% CI: -3.22, 1.30); ntUT (MD= -0.38; 95% CI:-4.08, 3.33); and Q (MD= 3.44; 95% CI: -0.51, 7.40).

246 *Relative CPM effect*

247 No significant “group x time” interaction effect for tUT, ntUT, or Q was found. There were no significant  
248 results for the between-group or within group-comparison at the tUT and ntUT. The within group  
249 difference in the DN group for the Q indicated that the CPM efficiency was significantly higher post-  
250 intervention compared to baseline (MD= 13.52%; 95% CI: 0.46, 26.59). In the SN group, no significant  
251 differences were found. No between-group mean differences were found for the tUT, ntUT, and Q.

252 *NPRS during hot water immersion*

253 No significant “group x time” interaction effect for NPRS water temperature were found, reflecting the  
254 absence of heat hyperalgesia. The post hoc pairwise comparisons for between-group or within group-  
255 comparison also showed no significant results. Between-group differences in mean changes from  
256 baseline to post-intervention were smaller than the reported MDC and MCID.

257 *Adverse events*

258 During the trial, no adverse events were registered.

259

260

261 **Table 3:** Descriptive statistics and within-group change scores (post-pre intervention) for pressure pain  
 262 thresholds and conditioned pain modulation.

Outcome	Dry Needling	Sham Needling	Between-group difference
<b>PPT tUT (N/cm<sup>2</sup>)</b>			
Baseline	18.52 ± 10.59	19.79 ± 12.54	
Post-intervention	19.55 ± 12.40	20.93 ± 13.65	
Within-group difference	1.03 (-1.06, 3.12)	1.15 (-0.87, 3.16)	0.12 (-2.78, 3.02)
<b>PPT ntUT (N/cm<sup>2</sup>)</b>			
Baseline	19.05 ± 10.53	22.90 ± 17.24	
Post-intervention	19.62 ± 10.59	21.77 ± 14.18	
Within-group difference	0.58 (-1.78, 2.94)	-1.13 (-3.40, 1.15)	-1.70 (-4.98, 1.58)
<b>PPT Q (N/cm<sup>2</sup>)</b>			
Baseline	29.25 ± 15.77	32.80 ± 19.18	
Post-intervention	30.08 ± 15.94	30.21 ± 19.24	
Within-group difference	0.83 (-1.89, 3.54)	-2.59 (-5.21, 0.03)	-3.42 (-7.19, 0.36)
<b>Absolute CPM effect tUT (N/cm<sup>2</sup>)</b>			
Baseline	1.72 ± 3.19	1.31 ± 3.64	
Post-intervention	-0.35 ± 3.22	0.61 ± 3.47	
Within-group difference	-2.08 (-4.40, 0.25)	-0.70 (-2.90, 1.49)	1.37 (-1.82, 4.57)
<b>Absolute CPM effect ntUT (N/cm<sup>2</sup>)</b>			
Baseline	1.38 ± 4.02	-0.33 ± 6.77	
Post-intervention	1.83 ± 4.44	2.20 ± 6.27	
Within-group difference	0.44 (-3.36, 4.25)	2.53 (-1.07, 6.13)	2.09 (-3.15, 7.33)
<b>Absolute CPM effect Q (N/cm<sup>2</sup>)</b>			
Baseline	-0.56 ± 4.88	-0.05 ± 7.66	
Post-intervention	3.34 ± 5.72	-0.10 ± 4.92	
Within-group difference	3.90 (-0.19, 7.98)	-0.05 (-3.91, 3.81)	-3.95 (-9.54, 1.64)
<b>Relative CPM effect tUT (%)</b>			
Baseline	8.74 ± 14.91	5.06 ± 24.32	
Post-intervention	3.68 ± 25.57	4.40 ± 16.02	
Within-group difference	-5.07 (-19.26, 9.12)	-0.66 (-14.08, 12.77)	4.41 (-15.14, 23.96)
<b>Relative CPM effect ntUT (%)</b>			
Baseline	4.17 ± 14.68	3.50 ± 22.87	
Post-intervention	6.34 ± 20.16	7.24 ± 17.67	
Within-group difference	2.17 (-11.03, 15.37)	3.74 (-8.74, 16.22)	1.57 (-16.49, 19.63)
<b>Relative CPM effect Q (%)</b>			
Baseline	-2.95 ± 16.87	-1.52 ± 24.93	

Post-intervention	10.58 ± 17.94	0.29 ± 14.67	
Within-group difference	<b>13.52* (0.39, 26.66)</b>	1.81 (-10.62, 14.23)	-11.72 (29.70, 6.26)
<b>NPRS water temperature</b>			
Baseline	5.65 ± 2.19	4.37 ± 2.73	
Post-intervention	4.97 ± 2.06	3.74 ± 2.73	
Within-group difference	-0.68 (-1.54, 0.18)	-0.63 (-1.45, 0.18)	0.045 (-1.14, 1.23)

Data are expressed as mean ± standard deviation and mean difference (95% confidence interval).

Within-group difference (Baseline – post-intervention)

\*= statistically significant.

**Abbreviations:** CI, confidence interval; CPM, conditioned pain modulation; DN, dry needling; NPRS, numeric pain rating scale; ntUT, non-treated upper trapezius muscle; PPT, pressure pain threshold; Q, Quadriceps muscle; SN, sham needling; tU, treated (most painful) upper trapezius muscle.

263

## 264 Discussion

265 No significant differences between DN and SN were found for PPTs at the local and remote locations  
266 and change in PPT values did not exceed the SE, MDC, or MCID, as identified by Walton et al.<sup>33,34</sup> Walton  
267 et al. stated that local PPTs appears to be a useful tool for measuring change over time, but remote  
268 (measured at the tibialis anterior muscle in their study) PPT is not useful for this purpose.<sup>34</sup> Our result  
269 contradicts the findings of Stieven et al., who found that a single session of DN or manual release, but  
270 not SN, resulted in an increase in PPTs at the UT bilaterally and at the ipsilateral and contralateral  
271 proximal head of the radius in patients with CINP.<sup>20</sup> Pecos-Martin et al. also reported a significant  
272 increase in PPT over the lower trapezius after one DN session performed in an active MTrP, immediately  
273 and up to at least one month after the treatment session, compared to DN on another location of the  
274 same muscle (but not a MTrP).<sup>39</sup> Mejuto-Vázquez et al. found superior effects of DN, compared to no  
275 treatment on local (C5-C6 zygapophyseal joint) and remote (second metacarpal and tibialis anterior  
276 muscle) PPTs in patients with acute neck pain 10 minutes and one week after intervention.<sup>40</sup> However,  
277 because no control group was included, placebo effects cannot be ruled out. Although our results show  
278 a general increase in PPTs for all DN locations, in contrast to the SN group where only a local increase  
279 was seen at the treated location, the results were statistically non-significant. A possible explanation is  
280 that DN is often accompanied by local post needling soreness, lasting up to 48 hours.<sup>41,42</sup> This soreness  
281 is the result of a direct local hyperalgesia response at the treated area and thus can mask effects  
282 immediately after intervention.<sup>43,44</sup>

283 This study found no differences in absolute CPM effects within or between groups. To our knowledge,  
284 there are no other studies evaluating CPM effect after DN in patients with neck pain, which makes  
285 comparing results difficult. One study evaluated the CPM effect of DN in patients with knee

286 osteoarthritis and found no larger effect of DN on central pain processing, compared to SN, immediately  
287 and 3 days postintervention.<sup>21</sup> Nevertheless, it may be hypothesized that eliciting LTRs during DN, which  
288 is mostly experienced as ‘painful’, may be considered an extra painful conditioning stimulus, which may  
289 have influenced the CPM protocol. In this case, the LTR might blur the CPM response, as this acts as a  
290 third pain stimulus besides the test and conditioning stimulus.<sup>45</sup> This contrasts with SN, which may not  
291 have influenced the testing protocol in a similar way since no LTRs were elicited and less pain was  
292 present.

293 When considering the relative CPM effects; there was a significant increase in the percentage of change  
294 at the Q location in the DN group, indicating a possible amelioration of the antinociceptive pain  
295 modulation. This may be caused by activation of descending inhibitory pain mechanisms.<sup>5, 18</sup> This is in  
296 line with the generalized, however not significant, increase in PPTs after DN.

297 The sample in this study included patients with CINP with mild disability and mild features of CS.  
298 Generalizability of the results to other patient groups (eg. whiplash, patients with cervical radiculopathy  
299 or generalized musculoskeletal complaints) is not applicable because previous research has shown that  
300 QST-features differ in these patient groups.<sup>9, 46-48</sup>

301 Considering the complexity of blinding in physical intervention research, two Delphi studies have been  
302 performed to evaluate the most important elements of shams for DN research.<sup>49</sup> Experts placed high  
303 importance on the entire intervention experience for active and sham protocols. Sham credibility may  
304 be maintained using cognitive strategies, potentially relinquishing the need for indistinguishable shams  
305 that have proved problematic to design.<sup>24</sup>

### 306 Strengths

307 This study is to our knowledge one of the first studies investigating pain modulatory effects of DN. The  
308 combination of evaluating PPT measurements and both absolute and relative CPM effects on local and  
309 remote locations is an added value to the insights on central neurophysiological effects of DN. The  
310 intervention was performed by three experienced DN therapists, who actively searched for trigger  
311 points instead of needling a predetermined point. All outcome assessors were blinded for the  
312 intervention. Therapists were trained to give identical verbal and non-verbal communication to both  
313 groups to maximize blinding of the participants.

### 314 Limitations

315 First, only one measurement was performed, making it impossible to evaluate and discuss the long-term  
316 effects and the possible influence of muscle soreness on the results immediately post-intervention.  
317 Second, patients who experienced DN in the past were not excluded. Although participants were not  
318 aware of the group allocation and were blinded to their treatment, expectations and previous

319 experience with DN may have influenced the results. Nevertheless, in a recent study, evaluating the  
320 effects of previous experience with DN therapy on blinding effectiveness and pain outcomes in people  
321 with neck pain, participants with previous experience were 22% more accurate at identifying their group  
322 allocation than those without experience, but the difference was not significant. Previous experience  
323 did not influence most clinical outcomes, except for pain intensity after real DN, although the difference  
324 was not clinically relevant.<sup>50</sup> Lastly, SN may have provoked neurophysiological effects as well, resulting  
325 in the comparison of two interventional groups instead of comparing an intervention to a control  
326 group.<sup>51, 52</sup> The insertion of a needle in the skin is interpreted as a noxious stimulus that can result in the  
327 activation of central inhibitory mechanisms and cause the excitation of A $\delta$  and A $\beta$  nerve fibers, which  
328 automatically provokes an analgesic effect.<sup>53, 54</sup> Nevertheless, there are no high-quality alternatives that  
329 may counter this possible effect.

### 330 Implications for clinical practice & future research

331 Future trials are needed to examine the effects of DN on central pain processing, after recovering from  
332 the associated post needling soreness. A follow-up period of more than 48 hours post-intervention  
333 should therefore be indicated. Because there is no widely accepted sham protocol for DN research,  
334 researchers should incorporate cognitive influences that extend beyond mimicking of tactile sensations  
335 to create a believable simulation of active dry needling. Assessment of blinding, using a blinding index  
336 might provide more robustness to the results.<sup>24</sup>

### 337 Conclusion

338 Based on the results of this study, we cannot conclude that DN has better effects on pain sensitivity and  
339 central pain modulation immediately post-intervention, compared to SN. Future trials are needed to  
340 examine the effects after post-needling soreness is resolved.

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344

### 345 Conflict of interest: none

346



347



Fig. 2. A: Measurement of pressure pain threshold on the quadriceps muscle.



Fig. 2. B: Measurement of pressure pain threshold on the upper trapezius muscle



Fig. 2. C: Measurement of conditioned pain modulation with hand in hot water bath

348

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