

Accepted Manuscript

Risk of Headache, Temporomandibular Dysfunction and Local Sensitization in Male Professional Boxers: A Case-Control Study

Miguel Mendoza-Puente, PT, PhD Ángel Oliva-Pascual-Vaca, PT, PhD Cleofás Rodríguez-Blanco, PT, PhD Alberto Marcos Heredia-Rizo, PT, PhD Daniel Torres-Lagares, DDS, PhD Francisco J. Ordoñez, MD, PhD

PII: S0003-9993(14)00468-7

DOI: [10.1016/j.apmr.2014.06.011](https://doi.org/10.1016/j.apmr.2014.06.011)

Reference: YAPMR 55876

To appear in: *ARCHIVES OF PHYSICAL MEDICINE AND REHABILITATION*

Received Date: 28 March 2014

Revised Date: 4 June 2014

Accepted Date: 13 June 2014

Please cite this article as: Mendoza-Puente M, Oliva-Pascual-Vaca Á, Rodríguez-Blanco C, Heredia-Rizo AM, Torres-Lagares D, Ordoñez FJ, Risk of Headache, Temporomandibular Dysfunction and Local Sensitization in Male Professional Boxers: A Case-Control Study, *ARCHIVES OF PHYSICAL MEDICINE AND REHABILITATION* (2014), doi: 10.1016/j.apmr.2014.06.011.

This is a PDF file of an unedited manuscript that has been accepted for publication. As a service to our customers we are providing this early version of the manuscript. The manuscript will undergo copyediting, typesetting, and review of the resulting proof before it is published in its final form. Please note that during the production process errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.



Running Head: Orofacial Sensitization in Boxing

Title: Risk of Headache, Temporomandibular Dysfunction and Local Sensitization in Male Professional Boxers: A Case-Control Study

Miguel Mendoza-Puente, PT, PhD,^a Ángel Oliva-Pascual-Vaca, PT, PhD,^b Cleofás Rodríguez-Blanco, PT, PhD,^b Alberto Marcos Heredia-Rizo, PT, PhD,^b Daniel Torres-Lagares, DDS, PhD,^c Francisco J Ordoñez MD, PhD,^d

^a University Center of Health Sciences San Rafael-Nebrija, University Antonio de Nebrija, Madrid, Spain; ^b Department of Physical Therapy, Faculty of Nursing, Physiotherapy and Podiatry, University of Sevilla, Sevilla, Spain; ^c Department of Stomatology, Faculty of Odontology, University of Sevilla, Sevilla, Spain; ^d Department of Anatomy and Human Embryology, School of Physical Education and Sports Medicine, University of Cádiz, Cádiz, Spain

The study was performed at an University-based physical therapy research clinic.

No commercial party having a direct financial interest in the results of the research supporting this article has or will confer a benefit on the authors or on any organization with which the authors are associated.

No conflicts of interest are reported for this study.

Address correspondence to Dr. Angel-Oliva-Pascual-Vaca. Departamento de Fisioterapia, Facultad de Enfermería, Fisioterapia y Podología, Universidad de Sevilla, Sevilla, España. c/ Avicena s/n, 41009 Sevilla, Spain.

E-mail: angeloliva@us.es

Tlf: 00 34 954486528 Fax: 00 34 954486527

No reprints are available for this study.

The study protocol was approved by the Ethical and Research Committee of the University of Sevilla, Spain.

1 **Title:** Risk of Headache, Temporomandibular Dysfunction and Local Sensitization in
2 Male Professional Boxers: A Case-Control Study

3

4 **Abstract**

5 **Objective:** To evaluate the differences in the incidence of headache, trigeminal
6 nerve mechanosensitivity, and temporomandibular functionality, in professional male
7 boxers (exposed to repetitive cranio-facial trauma) who were actively training and
8 without severe previous injuries, compared to handball players.

9 **Study Design:** Case-control study.

10 **Setting:** University-based physical therapy research clinic.

11 **Participants:** Eighteen boxers as the cases group and twenty handball players as
12 the comparison group (age, ± 2 years, and sex matched), were included (23 ± 4.61
13 years). All participants completed the assessment protocol.

14 **Interventions:** Not applicable.

15 **Main Outcome Measures:** Measurements were taken of the headache impact
16 (Headache Impact Test-6) and the pressure pain threshold (PPT) over the trigeminal
17 nerve sensory branches, the masseter and tibialis anterior muscles. Secondary
18 outcome measure included the temporomandibular function (Helkimo Clinic index).

19 **Results:** The boxers showed slight mandibular function impairment, local muscular
20 and neural sensitization and a higher impact from headaches compared to the
21 handball players. The between-group comparison found significant differences in all
22 outcome measures ($p < .05$), except in the tibialis anterior muscle PPT on the
23 dominant ($p = .958$) and the non-dominant sides ($p = .453$).

24 **Conclusion:** Professional male boxers seem to suffer a greater headache impact
25 and local sensitization of the craniomandibular region when compared to

26 professional handball players. It cannot be determined if these findings are transient,
27 as a result of the training activity, or more permanent.

28

29 **Key Words** Boxing; Case-control study; Headache; Sensitization; Sport.

30

31 ***List of abbreviations***

32 TMD temporomandibular disorders

33 MTP myofascial trigger points

34 TMJ temporomandibular joint

35 PPT pressure pain threshold

36 TA tibialis anterior muscle

37 HIT-6 headache impact test-6

38 V1 supraorbital nerve

39 V2 infraorbital nerve

40 V3 mental nerve

41 HCI helkimo clinic index

42 BMI body mass index

43

44 Boxing is one of the 'contact sports' that generates more controversy.¹ Up to about
45 34% of professional fights lead to later injury.^{2,3} The most common problems occur at
46 the cranio-facial, cervical and hands level.¹ Craniomandibular trauma in boxers can
47 cause concussion, jaw fractures, facial lacerations and dental contusion.⁴ Head and
48 face injuries account for about 20% of medical consultations in this sport.⁵ The
49 nervous system may also suffer mechanical stress due to the repeated blows.⁶ At
50 least 17% of boxers develop chronic traumatic encephalopathy after repetitive
51 concussion,⁷ with headache being one of the most common symptoms.⁸ Even
52 though headache usually appears right after the injury, it seems to be long lasting.⁹

53
54 Increased pain experience among high-level athletes has been correlated to
55 reduced sensitivity for pain.¹⁰ Competitive sport practice has also been linked to
56 increased endogenous opioid activity and altered hormones, which could result in
57 changes of the neural matrix and nociceptive pathways.¹⁰

58
59 Orofacial pain is present in 96.1% of subjects with temporomandibular disorders
60 (TMD),¹¹ and it can lead to an alteration of the nociceptive modulation in the Central
61 Nervous System, starting from afferent impulses from the cranio-cervical region.¹² As
62 a consequence, a central sensitisation process may take root, manifesting as muscle
63 tissue tenderness, a decreased mechanosensitive threshold and an extension of the
64 areas of referred pain.¹³ For this process to become established, there must be prior
65 peripheral sensitisation, prolonged over time, and induced mainly by the release of
66 algogenic substances that irritate the peripheral nerve endings, particularly in
67 muscles.¹⁴ Peripheral sensitisation has been described as the local sensation of hurt
68 and pain resulting from an enhanced responsiveness of nociceptive pathways to

69 peripheral natural stimuli.^{15,16} Therefore, it is a nociceptive pain that reflects the
70 perception of noxious stimuli.¹⁷ It seems to induce changes in the posterior horn of
71 the spinal cord associated with the afferent nerve fibers of the affected tissues,¹⁶ and
72 to contribute to central sensitisation.¹⁷ The presence of myofascial trigger points
73 (MTP) can generate nociceptive impulses which, when sustained over time, may
74 lead to a state of central sensitisation.¹² In addition, myofascial pain has been
75 purported to play a key role in the establishment of tension-type headaches.¹⁸

76

77 The purpose of the study was to observe the differences between sports-people
78 exposed to repeated cranio-facial trauma (boxers) versus high-level sportsmen
79 (handball players) not exposed to such continual trauma, with regards to neural and
80 muscular mechanosensitivity at a cranio-cervical level, the impact of headache, and
81 the temporomandibular function.

82

83

84 **Methods**

85

86

87 **Study Design**

88

89

90 An observational, and blinded case-control study was carried out. All the participants
91 were divided into two groups taking into account the exposure (boxers) or non-
92 exposure (handball players) to a risk factor (cranio-facial trauma).

93

94

95 Sample Size

96

97

98 Sample size calculation was made considering a 20% difference on
99 mechanosensitivity in the between-groups comparison,¹⁹ an estimated interindividual
100 coefficient of variation for pain threshold measures of 20%, an alpha value of 0.05
101 and a 80% power (Tamaño de la Muestra 1.1®).^a Sixteen subjects per group were
102 necessary to complete the study. A loss to follow-up rate of 10% was taking into
103 account.

104

105

106 Study Population

107

108

109 Based on a non-probability convenience sample, from April to August 2012, 18 male
110 elite boxers, and 20 handball players (age, within ± 2 years, and sex matched)²⁰ as
111 the comparison group, were selected (figure 1). To minimize sampling bias, all
112 eligible participants were drawn from the population of the same source (a high-
113 performance sports centre), in the same sex- and age-groups, and were included in
114 the final analysis.

115

116 Sixteen of the eighteen boxers were part of the national team and all were
117 awarded scholarships by the national sports council. The boxers had competed
118 previously in European and world championships, and some of them in the Olympic

119 games. The active training of the boxers consisted of physical training (running,
120 skipping, and gym) and training in skills. The latter was divided in different routines of
121 “shadow-boxing”, and practices with the punch bag and the punch ball to improve
122 speed and accuracy. Training with sparring was practiced with heavier and better
123 padded boxing gloves to soften the punches, and it was only performed occasionally
124 for those who were in the four weeks prior to competition. At the time of study, the
125 handball players were playing in honour division B, and some of them had previously
126 participated in international championships.

127

128 The inclusion criteria were: (a) active sports male; (b) aged between 18-35
129 years; (c) receiving regulated training at least 5 days/week; (d) no previous fight for
130 at least 45 days prior to the study for the boxers.²¹ The exclusion criteria were: (a) a
131 medical record of fractures and/or surgery at any vertebral, cranial vault, facial bones
132 or temporomandibular joint (TMJ) level; (b) central and/or peripheral degenerative
133 diseases; (c) a history of suffering from active migraine, based on a positive
134 response to any of the following questions “Have you suffered from migraine over
135 the year before data collection?” and “Do you have a medical diagnosis of migraine
136 from a consulting physician, e. g., a neurologist?”,^{22,23} (d) a previous history of
137 osteitis, rheumatic diseases or tumours; (e) current use of any medication (e.g., anti-
138 inflammatories, analgesics) which might interfere with the results; and (f) subjects
139 undergoing orthodontic treatment.²⁴

140

141 The study protocol complied with the ethical precepts of research involving
142 human subjects. It was designed and conducted according to the Helsinki
143 Declaration and was approved by the Institutional Review Board.

144

145

146 Data Collection Protocol

147

148

149 Upon confirming that the participant fulfilled the inclusion/exclusion criteria, the
150 subjects provided written informed consent. Then, the individuals received the
151 evaluation protocol together in a single session that lasted around forty-five
152 minutes. Boxers were asked about their fighting history (age when boxing started as
153 a competitive sport).²¹ The measurements were performed by an evaluator
154 previously trained in handling the measurement tools, and with 10 years of clinical
155 experience. The evaluator was blinded to the aims of the study and to the
156 case/control status of the participants.

157

158

159 Outcome Measures

160

161

162 Pressure Pain Threshold (PPT)

163 The PPT is the minimum force needed to induce pain or discomfort.²⁵ The
164 measurements were made using a digital algometer,^b model FPX 25, with a contact
165 probe size of 1cm². Pressure was applied with a depth of 5N/cm²/sec. The PPT was
166 measured on both sides in: (a) emerging trigeminal nerve sensory branches. The
167 supraorbital nerve (V1) was found in the superior orbital margin in the frontal bone.²⁶
168 The infraorbital nerve (V2) was found below the inferior orbital rim and in line with the

169 pupil.²⁷ The mental branch (V3) was located below the corner of the mouth and
170 slightly medial;²⁷ (b) masseter muscle. The reference point was located 1 cm cranial
171 and 2 cm anterior to the mandibular angle considering the minimum criteria outlined
172 for MTP (presence of a palpable taut band with a tender spot within it, referred pain
173 pattern, and local twitch response provoked by palpation of the MTP)²⁸; and (c)
174 tibialis anterior muscle (TA). The TA was located halfway between the most superior
175 attachment to the tibia and its tendon in the upper one third of the muscle belly.²⁹
176 The TA has been previously used as a remote location on the study of central or
177 peripheral sensitisation in chronic-whiplash associated disorders,²⁹ myofascial
178 TMD,³⁰ shoulder impingement syndrome,³¹ and tension-type headache.³²

179
180 Three measurements were performed with a 30 seconds interval, and the mean
181 was taken as the reference value.³³ Pressure algometry has proven to be valid and
182 has shown a high inter-examiner reliability, ICC = 0.91 (95% CI 0.82-0.97).³⁴ The
183 standard error of measurement range from 5.06 N/cm² to 6.27 N/cm².³⁴

186 **Headache Impact**

187 The headache impact test 6 (HIT-6) is a self-administered questionnaire that
188 evaluates the general and the previous four-weeks headache impact in the functional
189 capacity of the individual at a work and social level.³⁵ A final score below 50 points
190 implies that the headache is not functionally disabling. Above this score, the impact
191 is classified as mild (50-55), moderate (56-59) and severe (> 59).³⁵ In adolescent
192 athletes, the HIT-6 has observed good internal consistency ($\alpha = .90$), and test-retest
193 reliability ($r = .72$).³⁶

194

195

196 Mandibular functionality

197 The Helkimo clinic index (HCI) analyses the presence of signs and/or symptoms of
198 TMD. This index evaluates mandibular mobility, the presence of joint sounds and/or
199 muscle pain (in the TMJ itself or during its movements) and deviations in the jaw
200 movements.³⁷ Subjects may present a negative (<5), moderate (5-9 points) or severe
201 (≥ 10 points) dysfunction in the HCI.³⁷ The HCI has shown a good inter-
202 examiner reliability, ICC between 0.744 and 0.838.³⁸

203

204

205 Statistical Analysis

206

207

208 The data were processed using the PASW advanced statistics 18.0 programme.^c
209 The results are expressed as mean, with the corresponding standard deviation (SD)
210 and/or 95% confidence intervals (95% CI). The Shapiro-Wilk test was used to
211 analyse the normality of the variables. All outcome variables followed a normal
212 distribution ($p > .05$) and were analyzed with parametric tests. The baseline
213 characteristics of the study groups (age and body mass index, BMI) were compared
214 with the Student t test. The frequency distribution of the subjects in regard to the
215 HIT-6 and HCI scores was assessed with the Chi-square test (X^2).

216

217 To minimize the risk of type 1 error, the between-groups comparison of the mean
218 differences in the PPT, HIT-6 and HCI was performed using a multivariate analysis

219 of variance (MANOVA test). Post-hoc testing was assessed with Bonferroni
220 correction. Effect sizes (partial eta square) for these comparisons were used
221 considering that a small effect has a value ≥ 0.2 and < 0.5 , whereas a moderate
222 effect is considered to be a value ≥ 0.5 and < 0.8 , and a large effect must be ≥ 0.8 .³⁹

223

224 Finally, the impact of age and BMI was assessed as possible confounding
225 variables in the outcome measures, by means of contingency tables (2x2) (Chi-
226 square). The statistical analysis was conducted considering significance at a p value
227 $< .05$.

228

229

230 Results

231

232

233 The baseline characteristics of the sample are displayed in table 1. The BMI found
234 statistical significance in the between-groups comparison ($p < .05$), although both
235 groups showed values within the average range (18.5 - 24.99 points). Boxers had
236 been fighting for 7.5 ± 2.64 years, whereas handball players had been playing for 9.1
237 ± 3.43 years.

238

239 According to the MANOVA overall analysis, Wilks' Lambda value was 0.228, F-
240 ratio was 6.246 and p-value $< .001$. Table 2 lists the between-groups mean
241 differences (MANOVA) in the post-hoc comparison of the outcome measures. The
242 boxing group showed an increased mechanosensitivity in all the locations and on
243 both sides, with statistical significance ($p < .05$), except for the TA on the dominant

244 side ($p=.958$) and on the non-dominant side ($p=.453$). Considering the average value
245 of the measurements in both sides (table 2), the effect size was close to moderate
246 for V1 [$p<.001$; $F_{1,36}=24.12$; $R^2=0.40$] and V2 [$p<.001$]; $F_{1,36}=23.55$; $R^2=0.38$], and
247 low for V3 [$p=0.015$; $F_{1,36}=9.59$; $R^2=0.21$]. For the masseter muscle, the effect size
248 was also moderate in the dominant [$p<.001$; $F_{1,36}=47.90$; $R^2=0.57$] and non-dominant
249 sides [$p<0.001$; $F_{1,36}=46.00$; $R^2=0.56$].

250

251 Table 3 presents the frequency distribution of the study subjects with regard to
252 the severity of the HIT-6 and HCI results. The HIT-6 values were lower in the
253 handball players, with significant differences in the between-groups comparison, and
254 a small effect size [$p=0.028$; $F_{1,36}=5.28$; $R^2=0.12$]. Although 55.5% (10/18) of the
255 boxers revealed a mild or moderate impact in the HIT-6 (value >50), compared to
256 only 20% (4/20) of the handball players (table 3), the average score in both groups
257 (value below 50) showed that the headache was not functionally disabling in any of
258 them. Similar results were observed from the HCI. The mean value in the boxers
259 (7.44 ± 4.00) indicated a moderate TMD compared to an absence of TMD in the
260 handball players (3.81 ± 2.25) (table 2). The inter-group mean difference was also
261 significant [$p=0.008$; $F_{1,36}=8.02$; $R^2=0.18$].

262

263 The between-groups comparison using contingency tables showed no statistical
264 significance between age and BMI and the measured variables ($p>.05$).

265

266

267 Discussion

268

269

270 This study has assessed the differences in headache impact, orofacial
271 mechanosensitivity and TMJ functionality between sportsmen exposed to repetitive
272 craniofacial trauma, as an associated risk factor, and high-level handball players.

273

274 With regard to the PPT, the results seem to indicate that professional boxers in
275 training showed increased excitability in the trigeminal nerve sensory branches and
276 in the masseter muscle, but not in a distant location, the TA, when compared to the
277 handball players. The capacity to withstand pain is a key factor in competitive sports,
278 and pain tolerance appears to differ between athletes and non-sports people.¹⁰
279 Therefore, the comparison group included high-level athletes that were selected from
280 a single sport to homogenize the sample and increase internal validity.

281 Mechanosensitivity thresholds were similar in a distal muscle, the TA, which would
282 help to assume that both groups were suspected to have similar pain tolerance.

283

284 The present findings may suggest a local sensitization in the craniocervical
285 region of the boxers. Local sensitization can increase the presence of pain in chronic
286 injuries in high-level sports people.⁴⁰ Whether these differences are only transient as
287 a result of the training activity remains unknown. For a sensitization phenomenon to
288 occur, a maintained nociceptive input is required.³⁰ Boxing carries risk of facial and
289 head injuries that are self-evident, and the incidence of contact injuries increases in
290 professional boxing,² compared to other sports such as American football, wrestling
291 and soccer.^{3,4} .

292

293 The impact of headache was also higher in the boxers. The use of a self-report
294 headache scale, like the HIT-6, as part of a testing protocol for concussion, appears
295 to help clinicians to identify those with existing headache disorders.³⁶ A difference ≥ 5
296 points in the HIT-6 has been reported as clinically meaningful, whereas a 3-point
297 change is considered to be remarkable.⁴¹ According to this, the between-group
298 difference in the present study (4.82 points) could be considered close to clinical
299 relevance. Among the boxers, only 45.5% (8/18) revealed no functionally disabling
300 impact in the HIT-6, compared to 80% (16/20) of the handball players (table 3).
301 Nonetheless, HIT-6 scores in both groups were within the average values for non-
302 functionally disabling headache. Hence, results must be interpreted with caution.
303 Sensitization of the cranial nerves may contribute to the establishment and severity
304 of headache.²⁶ Trigeminal nerve increased excitability, such as that found in the
305 boxers, has been linked to increased prevalence of migraine,¹² and to chronic-
306 tension or cervicogenic headache.²⁶ Different headache subgroups have been
307 established when considering the management of headache in sport, including the
308 "*headache arising from mechanisms that occur during exertion*", i.e. headaches
309 resulting from trauma, impact and/or cervical dysfunction.⁴² This subgroup could be
310 linked to boxing, although no direct relationship has been found between the severity
311 of the impact and the resulting headache.⁴² Changes in the vascular pressure and
312 heart rate may also be related to headache in sport.⁴² However, these aspects have
313 not been monitored in this study.

314

315 The trigeminal caudal nucleus is responsible for innervating the masticatory
316 muscles.⁴³ Therefore, the increased mechano-excitability over the masseter muscles
317 in the boxers is consistent with the hypersensitivity recorded in the trigeminal nerve

318 branches in these subjects. A sensory impairment in the trigeminal nerve has been
319 purported to be associated with TMD,³⁰ and trigeminal neuropathic pain is frequently
320 caused by trauma.⁴⁴ A sensitization process (central or peripheral) may also be
321 involved in the development of these disorders.³⁰ Pain in the muscles of mastication
322 has been found as the most prevalent sign of TMD among young athletes who play
323 basketball.⁴⁵ For the HCI, 77.7% of the boxers (14/18) had moderate or severe TMD
324 (table 3), and the mean value in this group was 7.44 ± 4.00 , which confirms the
325 presence of signs and symptoms of a moderate disorder. Between 44% and 99% of
326 TMDs are secondary to trauma, which may help to explain our findings.⁴⁶ Moreover,
327 the absence of immediate symptoms or injury is common in contact sports that
328 involve direct trauma to the jaw.⁴⁵

329
330 Despite the observed differences, all the boxers in the sample used a head
331 protector and a mouth guard, both in sparring training and in actual competition
332 events. The use of foam head guards has shown to be effective in preventing
333 superficial head injuries, although their effect on severe head injuries is unknown.⁴⁷
334 The mouth guard prevents tooth loss and minimises the impact of trauma directly
335 upon the oral cavity,⁴⁸ although scientific evidence on its effect in preventing
336 contusions is lacking. Its use is considered a priority in contact sports because it
337 decreases the risk of orofacial injury between 1.6-1.9 times.⁴⁹

340 **Limitations of the study**

341

342

343 The prevalence of injury is higher in male boxing than in female boxers.¹ Hence,
344 the study only evaluated male subjects. It would be interesting to assess whether
345 there are gender-related effects in regard to the local sensitization, the headache
346 impact and the TMJ functionality. Future studies should include men and women,
347 and other sports with high risks of cranio-facial impacts and potential for injury, such
348 as rugby, martial arts, and basketball.⁵⁰ It also seems necessary to compare the
349 present findings with a control group of non-sports subjects, as in previous studies.⁴⁵
350 Handball players are physically quite different from boxers, which makes blinding
351 difficult. Nevertheless, the evaluator was unaware of the aims of the study and the
352 groups to be tested.

353
354 Boxers were included in the study if they had not fought in a competition within
355 45 days prior to the study. Nonetheless, training habits regarding sparring schedules
356 before data collection were not controlled, which may have influenced the results.²¹ It
357 is plausible to state that the present findings may only reflect transient changes
358 because of the ongoing sparring training instead of permanent changes. The results
359 suggest the need to assess boxers later in life to observe the possible impairments
360 and the duration of changes. In addition, those boxers with a severe prior injury were
361 excluded, yet these are the ones most likely to have an orofacial problem.

362

363

364 **Conclusion**

365

366

367 Male professional boxers who are actively training seem to show a local muscular
368 and neural sensitization in the orofacial region, a higher impact from headache and
369 slight TMD, when compared to high-level handball players. It cannot be determined if
370 these findings are transient, as a result of the training activity, or more permanent.

371

372

373 **References**

374

375

- 376 1. Gambrell RC. Boxing: Medical care in and out of the ring. *Curr Sports Med Rep*
377 2007;6:317–21.
- 378 2. Bledsoe GH, Li G, Levy F. Injury risk in professional boxing. *South Med J*
379 2005;98:994-8.
- 380 3. Zazryn T, Cameron P, Mc Crory P. A prospective cohort study of injury in
381 amateur and professional boxing. *Br J Sports Med* 2006;40:670-4.
- 382 4. Bianco M, Pannoizzo A, Fabbricatore C, Sanna N, Moschetti M, Palmieri V,
383 Zeppilli P. Medical survey of female boxing in Italy in 2002-2003. *Br J Sports*
384 *Med* 2005;39:532-6.
- 385 5. Timm KE, Wallach JM, Stone JA, Ryan EJ. Fifteen years of amateur boxing
386 injuries / illness at the United States Olympia training center. *J Athl Train*
387 1993;28:330-4.
- 388 6. Topp KS, Boyd BS. Structure and biomechanics of peripheral nerves: nerve
389 responses to physical stresses and implications for physical therapist practice.
390 *Phys Ther* 2006;86:92-109.

- 391 7. Galetta KM, Barrett J, Allen M, Madda F, Delicata D, Tennant AT, Branas CC,
392 Maguire MG, Messner LV, Devick S, Galetta SL, Balcer LJ. The King-Devick test
393 as a determinant of head trauma and concussion in boxers and MMA fighters.
394 Neurology 2011;76:1456-62.
- 395 8. Ledic D, Sosa I, Linic IS, Cvijanovic O, Kovacevic M, Desnica A, Banicek I.
396 Vomiting as a reliable sign of concussion. Med Hypotheses 2012;78:23-5.
- 397 9. Jagoda A, Riggio S. Mild traumatic brain injury and the postconcussive
398 syndrome. Emerg Med Clin North Am 2000;18:355-63.
- 399 10. Manning EL, Fillingim RB. The influence of athletic status and gender on
400 experimental pain responses. J Pain 2002;3:421-8.
- 401 11. Cooper BC, Kleinberg I. Examination of a large patient population for the
402 presence of symptoms and signs of temporomandibular disorders. Cranio
403 2007;25:114-26.
- 404 12. Fernández-de-las-Peñas C, Arendt-Nielsen L, Cuadrado ML, Pareja JA.
405 Generalized mechanical pain sensitivity over nerve tissues in patients with
406 strictly unilateral migraine. Clin J Pain 2009;25:401-6.
- 407 13. Merrill RL. Central mechanisms of orofacial pain. Dent Clin North Am
408 2007;51:45-59.
- 409 14. Mork H, Ashina M, Bendtsen L, Olesen J, Jensen R. Experimental muscle pain
410 and tenderness following infusion of endogenous substances in humans. Eur J
411 Pain 2003;7:145-53.
- 412 15. Lechner SG, Lewin GR. Peripheral sensitisation of nociceptors via G-protein-
413 dependent potentiation of mechanotransduction currents. J Physiol
414 2009;587:3493-503.

- 415 16. Ramiro-González MD, Cano-de-la-Cuerda R, De-la-Llave-Rincón AI,
416 Miangolarra-Page JC, Zarzoso-Sánchez R, Fernández-de-Las-Peñas C. Deep
417 tissue hypersensitivity to pressure pain in individuals with unilateral acute
418 inversion ankle sprain. *Pain Med* 2012;13:361-7.
- 419 17. Woolf CJ. Central sensitization: implications for the diagnosis and treatment of
420 pain. *Pain* 2011;152(3 Suppl):S2-15.
- 421 18. Bendtsen L. Central sensitization in tension-type headache--possible
422 pathophysiological mechanisms. *Cephalalgia* 2000;20:486-508.
- 423 19. La Touche R, Fernández-de-Las-Peñas C, Fernández-Carnero J, Díaz-Parreño
424 S, Paris-Alemany A, Arendt-Nielsen L. Bilateral mechanical-pain sensitivity
425 over the trigeminal region in patients with chronic mechanical neck pain.
426 *J Pain* 2010;11:256-63
- 427 20. Torelli P, Manzoni GC. Clinical observations on familial cluster headache.
428 *Neurol Sci* 2003;24:61-4.
- 429 21. Bernick C, Banks S, Phillips M, Lowe M, Shin W, Obuchowski N, Jones S, Modic
430 M. Professional fighters brain health study: rationale and methods. *Am J*
431 *Epidemiol* 2013;178:280-6.
- 432 22. Fernández-de-Las-Peñas C, Hernández-Barrera V, Carrasco-Garrido P, Alonso-
433 Blanco C, Palacios-Ceña D, Jiménez-Sánchez S, Jiménez-García R. Population-
434 based study of migraine in Spanish adults: relation to socio-demographic factors,
435 lifestyle and co-morbidity with other conditions. *J Headache Pain* 2010;11:97-
436 104.
- 437 23. Schürks M, Buring JE, Kurth T. Agreement of self-reported migraine with ICHD-II
438 criteria in the Women's Health Study. *Cephalalgia* 2009;29:1086-90.

- 439 24. Branco LP, Santis TO, Alfaya TA, Godoy CH, Fragoso YD, Bussadori SK.
440 Association between headache and temporomandibular joint disorders in
441 children and adolescents. *J Oral Sci* 2013;55:39-43.
- 442 25. Fredriksson L, Alstergren P, Kopp S. Pressure pain thresholds in the craniofacial
443 of female patients with rheumatoid arthritis. *J Orofac Pain* 2003;17:326-32.
- 444 26. Fernández de las Peñas C, Coppieters MW, Cuadrado ML, Pareja JA. Patients
445 with chronic tension-type headache demonstrate increased mechano-sensitivity
446 of the supra-orbital nerve. *Headache* 2008;48:570-7.
- 447 27. Song WC, Kim SH, Paik DJ, Han SH, Hu KS, Kim HJ, Koh KS. Location of the
448 infraorbital and mental foramen with reference to the soft-tissue landmarks. *Plast*
449 *Reconstr Surg* 2007;120:1343-7.
- 450 28. Simons DG, Travell J, Simon LS. Myofascial pain and dysfunction: the trigger
451 point manual. Volume 1 (2nd ed) Baltimore: Williams & Wilkins; 1999.
- 452 29. Scott D, Jull G, Sterling M. Widespread sensory hypersensitivity is a feature of
453 chronic whiplash-associated disorder but not chronic idiopathic neck pain. *Clin J*
454 *Pain* 2005;21:175-81.
- 455 30. Fernández de las Peñas C, Galán del Río F, Fernández Carnero J, Pesquera J,
456 Arendt-Nielsen L, Svensson P. Bilateral widespread mechanical pain sensitivity
457 in women with myofascial temporomandibular disorder: evidence of impairment
458 in central nociceptive processing. *J Pain* 2009;10:1170-8.
- 459 31. Albuquerque-Sendín F, Camargo PR, Vieira A, Salvini TF. Bilateral myofascial
460 trigger points and pressure pain thresholds in the shoulder muscles in patients
461 with unilateral shoulder impingement syndrome: a blinded, controlled study. *Clin*
462 *J Pain* 2013;29:478-86.

- 463 32. Ashina S, Bendtsen L, Ashina M, Magerl W, Jensen R. Generalized
464 hyperalgesia in patients with chronic tension-type headache. *Cephalalgia*
465 2006;26:940-8.
- 466 33. Heredia-Rizo AM, Oliva-Pascual-Vaca A, Rodríguez-Blanco C, Torres-Lagares
467 D, Albornoz-Cabello M, Piña-Pozo F, Luque-Carrasco A. Craniocervical posture
468 and trigeminal nerve mechanosensitivity in subjects with a history of orthodontic
469 use: a cross-sectional study. *Cranio* 2013;31:252-9.
- 470 34. Chesterton LS, Sim J, Wright CC, Foster NE. Interrater reliability of algometry in
471 measuring pressure pain thresholds in healthy humans, using multiple raters.
472 *Clin J Pain* 2007;23:760-6.
- 473 35. Kosinski M, Bayliss MS, Bjorner JB, Ware JE Jr, Garber WH, Batenhorst A,
474 Cady R, Dahlöf CG, Dowson A, Tepper S. A six-item short-form survey for
475 measuring headache impact: the HIT-6. *Qual Life Res* 2003;12:963-74.
- 476 36. Piebes SK, Snyder AR, Bay RC, Valovich McLeod TC. Measurement properties
477 of headache-specific outcomes scales in adolescent athletes. *J Sport Rehabil*
478 2011;20:129-42.
- 479 37. Helkimo M. Studies on function and dysfunction of the masticatory system. II.
480 Index for anamnestic and clinical dysfunction and occlusal state. *Sven Tandlak*
481 *Tidskr* 1974;67:101-21.
- 482 38. Fu K, Ma X, Zhang Z, Tian Y, Zhou Y, Zhao Y. Study on the use of
483 temporomandibular joint dysfunction index in temporomandibular disorders.
484 *Zhonghua Kou Qiang Yi Xue Za Zhi* 2002;37:330-2. [Chinese]
- 485 39. Cohen J. *Statistical Power Analysis for the Behavioral Sciences*. 2nd ed.
486 Hillsdale, NJ (USA): Lawrence Erlbaum; 1988.

- 487 40. van Wilgen CP, Keizer D. Neuropathic pain mechanisms in patients with chronic
488 sports injuries: a diagnostic model useful in sports medicine? *Pain Med*
489 2011;12:110-7.
- 490 41. Bayliss MS, Batenhorst AS. The HIT-6™: a user's guide. Lincoln, RI (USA):
491 Quality Metric Inc; 2002.
- 492 42. Kernick DP, Goadsby PJ. Guidance for the management of headache in sport on
493 behalf of The Royal College of General Practitioners and The British Association
494 for the Study of Headache. *Cephalalgia* 2011;31:106-11.
- 495 43. Schünke M, Schulte E, Schumacher U. Text and atlas of anatomy general
496 anatomy and musculoskeletal system. Bogota (Colombia): Ed Médica
497 Panamericana; 2008.
- 498 44. Zakrzewska JM. Multi-dimensionality of chronic pain of the oral cavity and face. *J*
499 *Headache Pain* 2013;14:37
- 500 45. Weiler RM, Vitalle MS, Mori M, Kulik MA, Ide L, Pardini SR, Santos FM.
501 Prevalence of signs and symptoms of temporomandibular dysfunction in male
502 adolescent athletes and non-athletes. *Int J Pediatr Otorhinolaryngol*
503 2010;74:896-900.
- 504 46. American Academy of Pediatric Dentistry University of Texas Health Science
505 Center at San Antonio Dental School. Treatment of temporomandibular
506 disorders in children: summary statement and recommendations. *J Am Dent*
507 *Assoc* 1990;120:265-9.
- 508 47. Jones SJ, Lyons RA, Evans R, Newcombe RG, Nash P, McCabe M, Palmer SR.
509 Effectiveness of rugby headgear in preventing soft tissue injuries to the head: a
510 case-control and video cohort study. *Br J Sports Med* 2004;38:159-62.

- 511 48. Lesić N, Seifert D, Jerolimov V. Orofacial injuries reported by junior and senior
512 basketball players. Coll Antropol 2011;35:347-52.
- 513 49. Knapik JJ, Marshall SW, Lee RB, Darakjy SS, Jones SB, Mitchener TA,
514 delaCruz GG, Jones BH. Mouthguards in sport activities: history, physical
515 properties and injury prevention effectiveness. Sports Med 2007;37:117-44.
- 516 50. Soporowski NJ, Tesini DA, Weiss AI. Survey of orofacial sports-related injuries. J
517 Mass Dent Soc 1994;43:16-20.
- 518

519 **Suppliers**

520 a. Pontificia Universidad Javeriana, 40-62, Carrera 7, Bogotá (Colombia),

521 11001000.

522 b. Wagner Instruments, 17 Wilmot Ln, Riverside, CT, 06878.

523 c. SPSS Inc, 233 S Wacker Dr, 11th Fl, Chicago, IL, 60606.

524 d.

525

ACCEPTED MANUSCRIPT

526 **Figure Legends**

527

528 **Figure 1** Flowchart Diagram of the Study Subjects.

ACCEPTED MANUSCRIPT

Table 1 Baseline characteristics of the subjects in the study groups *

	Total Sample (N=38)	Boxing Group (n=18)	Control Group (n=20)	p value
Mean Age (yrs)	23 ± 4.61	22 ± 4.06	23 ± 5.04	.299
BMI (kg/m ²)	22.71 ± 2.83	20.76 ± 2.27	24.47 ± 2.02	.001

Abbreviations: BMI, Body Mass Index; p-value, statistical significance of the difference between groups.

*Data are reported as mean ± standard deviation

Table 2 Between-groups differences of the headache impact, mandibular function (Helkimo test) and muscular and neural mechanosensitivity*

		Mean value ± SD	Between-group mean differences (95% CI)	p- value
HIT-6	Boxing Group	48.77 ± 5.77	4.82	.028
	Handball Group	43.95 ± 7.02	(0.56 / 9.08)	
PPT-V1 D	Boxing Group	13.94 ± 4.57	- 7.06	<.001
	Handball Group	21.00 ± 4.44	(-10.22 / -3.90)	
PPT-V1 ND	Boxing Group	14.77 ± 4.42	- 7.09	<.001
	Handball Group	21.87 ± 4.82	(-10.32 / -3.86)	
PPT-V2 D	Boxing Group	20.78 ± 6.79	- 13.15	<.001
	Handball Group	33.94 ± 8.06	(-17.12 / -7.14)	
PPT-V2 ND	Boxing Group	21.63 ± 6.72	- 8.22	<.001
	Handball Group	29.86 ± 6.56	(-12.87 / -3.57)	
PPT-V3 D	Boxing Group	20.86 ± 7.39	- 6.40	.003
	Handball Group	27.27 ± 5.60	(-11.03 / -1.77)	
PPT-V3 ND	Boxing Group	20.85 ± 6.51	- 6.77	.005
	Handball Group	27.63 ± 7.49	(-11.42 / -2.13)	
PPT-M D	Boxing Group	16.04 ± 4.88	- 11.16	<.001
	Handball Group	27.21 ± 5.39	(-14.75 / -7.57)	
PPT-M ND	Boxing Group	16.52 ± 4.90	- 10.22	<.001

	Handball Group	26.74 ± 5.22	(-13.76 / -6.67)	
PPT-TA D	Boxing Group	82.25 ± 17.16	- 1.49	.958
	Handball Group	83.75 ± 16.52	(-10.67 / 13.65)	
PPT-TA ND	Boxing Group	81.79 ± 17.25	- 2.23	.453
	Handball Group	84.03 ± 17.22	(-14.68 / 10.20)	
Helkimo test	Boxing Group	7.44 ± 4.00	3.63	.008
	Handball Group	3.81 ± 2.25	(1.32 / 5.88)	

* Data are expressed as mean ± standard deviation (SD) or as mean (95% CI, confidence interval). PPT values are expressed in N/cm²

Abbreviations: HIT-6, Headache Impact Test-6; PPT, pressure pain threshold; V1, supraorbital nerve; V2, infraorbital nerve; V3, mental nerve; M, masseter muscle; TA, tibialis anterior muscle; D, dominant side; ND, non-dominant side

Table 3 Frequency distribution of the study subjects based on the scores of the HIT-6 and the HCI*

Observed events (N=38)	Boxers (n=18)	Controls (n=20)	p value
HIT-6 not disabling (<50)	8 (21.0)	16 (42.1)	.023
HIT-6 slight (50-55)	8 (21.0)	2 (5.2)	.016
HIT-6 moderate (56-59)	2 (5.2)	2 (5.2)	.911
HIT-6 severe (>59)	0 (0)	0 (0)	1.00
HIT-6 > slight	10 (26.3)	4 (10.5)	.023
HCI slight (<5)	4 (10.5)	11 (28.9)	.139
HCI moderate (5-9)	8 (21.0)	8 (21.0)	.516
HCI severe(\geq 10)	6 (15.7)	1 (2.6)	.024
HCI > slight	14 (36.8)	9 (23.6)	.039

* Data are reported as number of subjects and (frequency, %, of total sample, N); p-value, sourced from the Chi Square test (X^2).

Abbreviations: HIT-6, Headache Impact Test-6; HCI, Helkimo Clinic Index

