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Intra-osseous heat generation during implant bed preparation with static navigation: Multi-factor *in vitro* study

Running head (abbreviated title)

Intra-osseous heat generation during bone drilling

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52

53 J.G-A: Drafting of the article, critical revision, approval of the submitted and final versions.
54 O.S-C: Data collection, drafting of the article, approval of the submitted and final versions.
55 N.L-C: Data collection, drafting of the article, approval of the submitted and final versions.
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58

59 **Abstract**

60 **Objectives:** To compare the intra-osseous temperature reached during bone drilling for dental implant
61 placement using open versus closed static surgical guides and evaluate the influence of bone density,
62 osteotomy drilling depth, and irrigation fluid temperature.

63 **Material and methods:** 960 osteotomies were performed with 2 mm pilot drills in 16 solid rigid
64 polyurethane foam blocks. Two main variables were considered: the guide type (open or closed guide) and
65 bone density (hard (D1) or soft (D4)). The blocks were divided into four groups according to the type of
66 surgical template and bone density as follows: group one: closed guide and hard bone; group two: open guide
67 and hard bone; group three: closed guide and soft bone; and group four: open guide and soft bone. A
68 combination of different experimental conditions was used, including different bone osteotomy depths (6 or
69 13 mm) and irrigation fluid temperatures (5 °C or 21 °C).

70 **Results:** The highest mean temperature was found in group one (28.29±4.02 °C). In the soft bone groups
71 (three and four), the mean maximum temperature decreased compared to groups one and two (dense bone)
72 and was always higher with closed guides (23.38±1.92 °C) compared to open guides (21.97±1.22 °C)
73 (p<0.001). The osteotomy depth and irrigation fluid temperature also significantly influenced the bone
74 temperature (p<0.001), especially in hard bone.

75 **Conclusions:** The greatest heat generation was observed in high-density bone. The final intra-bone
76 temperature was about 1 °C higher with a closed static surgical guide than with an open guide. The heat
77 generation in osteotomy sites was substantially reduced by cooling the irrigation fluid to 5 °C.

78
79 **Keywords:** Computer-aided surgery, Computer-assisted surgery, Dental implants, Osteotomy, Bone tissue.

80 81 **Introduction**

82 Adequate three-dimensional (3D) positioning of dental implants has been revealed as a key factor for
83 aesthetic outcomes, easy maintenance, stability of peri-implant soft and hard tissues, and long-term success
84 (Buser, Martin, & Belser, 2004; D'Haese, Ackhurst, Wismeijer, De Bruyn, & Tahmaseb, 2017; Linkevicius,
85 Puisys, Vindasiute, Linkeviciene, & Apse, 2013; Tarnow, Cho, & Wallace, 2000). Guided surgery has been
86 widely applied to achieve ideal 3D implant positioning (Bover-Ramos, Viña-Almunia, Cervera-Ballester,
87 Peñarrocha-Diago, & García-Mira, 2018; Colombo et al., 2017; Moraschini, Velloso, Luz, & Barboza, 2015;
88 Raico Gallardo et al., 2017; Schneider, Marquardt, Zwahlen, & Jung, 2009) as it can reduce human error
89 (Amorfini, Migliorati, Drago, & Silvestrini-Biavati, 2017). Freehand implant surgery is less accurate in
90 transferring 3D pre-surgical planning to the patient than guided surgery (Farley, Kennedy, McGlumphy, &
91 Clelland, 2013; Pozzi, Tallarico, Marchetti, Scarfò, & Esposito, 2014; Vercruyssen et al., 2015; Younes et
92 al., 2018). The application of static surgical templates for bone drilling and implant placement is a popular
93 implant surgery navigation system. Closed or open static implant guides are currently available. The open
94 guide has an open access located on the buccal side of the template that allows a buccal view of the surgical
95 field and better contact of the irrigation fluid with the implant drill. However, this open design may affect the
96 stent stability during drilling (Fauroux, De Boutray, Malthiéry, & Torres, 2018). In contrast, closed guides
97 tend to cover the entire surgical field and do not allow for direct visualization of the surrounding soft and
98 hard tissues. In addition, closed guides limit irrigation, which is usually used to cool down the temperature of
99 the drill. Consequently, it has been shown that the use of a closed surgical guide for implant site preparation
100 instead of a surgical template generates more heat (Liu, Wu, Zhang, Peng, & Liao, 2018; Misir, Sumer,
101 Yenisey, & Ergiöglu, 2009).

102
103 Thermal changes during bone drilling can cause bone alterations such as osteonecrosis (ON), which is
104 defined as the *in situ* loss of a bone segment and is characterized by avascular necrosis of the bone (Bolland,
105 Hood, Bastin, King, & Grey, 2004). Thermally induced osteonecrosis is a traumatic type of necrosis caused
106 by high temperatures that can result in micro-damage to the osteocytes with cell apoptosis, followed by
107 osteoclast activation and bone resorption (Augustin et al., 2012; Jeong et al., 2014; Noble, 2003). Therefore,

108 bone temperature monitoring during osteotomy drilling using thermocouple methods has been used to
109 prevent implant integration complications (Delgado-Ruiz et al., 2018; Gehrke et al., 2018; Jeong et al., 2014;
110 Liu et al., 2018; Misir et al., 2009; Möhlhenrich, Modabber, Steiner, Mitchell, & Hölzle, 2015; Sumer,
111 Keskiner, Mercan, Misir, & Cankaya, 2014).

112

113 In summary, static navigation for implant bed preparation appears to generate greater internal bone heat than
114 using a freehand technique. Open static surgical guides seem to reduce heat generation compared to closed
115 guides because there is more contact between the irrigation fluid and drill. Consequently, the influence of
116 surgical guide designs and other factors such as bone density, irrigation fluid temperature, and osteotomy
117 depth must be considered when planning guided implant surgery procedures (Raico Gallardo et al., 2017;
118 Younes et al., 2018)

119

120 This *in vitro* study aimed to evaluate the maximum bone temperature reached during implant bed preparation
121 using two different types of surgical guides (open and closed) and assess the influence of bone density,
122 osteotomy drilling depth, and irrigation fluid temperature on the bone thermal changes.

123

124 **Material and methods**

125 **Study design**

126 The present *in vitro* study was carried out on eight solid rigid polyurethane foam blocks measuring 130 x 160
127 x 200 mm in size and distributed into four blocks simulating type I bone (D1) and four blocks simulating
128 type IV bone (D4) according to the bone density classification of Misch and Degidi (Misch & Degidi, 2003)
129 (Sawbones, Pacific Research Laboratories, USA). Each bone block was then cut into ten smaller pieces,
130 resulting in 40 smaller blocks of D1 and D4 density, respectively, measuring 130 x 16 x 200 mm in size. To
131 allow for more osteotomies, each block was drilled on both sides. The D1 and D4 foam blocks'
132 characteristics were a density, compressive strength, and compressive modulus of 0.08 g/cm³ and 0.48 g/cm³;
133 0.6 MPa and 18 MPa; and 16 MPa and 445 MPa, respectively. This study was approved by the Research
134 Ethics Committee of the Universitat Internacional de Catalunya (Ref. CIR-ELM-2017-02).

135

136 In partially edentulous patients, canine, premolar, and molar acrylic resin teeth were inserted in the blocks
137 and fixed with triad gel (TriadGel® Dentsply Sirona, USA) to simulate the remaining teeth for better guiding
138 stabilization. A space was left between the inserted simulated teeth to allow for six osteotomies per block,
139 resulting in a total of 960 osteotomies.

140

141 The blocks were divided into four groups according to the type of surgical template (open or closed guide)
142 and bone density: group one: closed guide and hard bone (D1); group two: open guide and hard bone (D1);
143 group three: closed guide and soft bone (D4); and group four: open guide and soft bone (D4).

144

145 The closed guide (Figure 1) covered the entire surgical field and did not allow for the visualization of the
146 simulated bone during the drilling sequence. In contrast, the open guide (Figure 2) had an open access
147 located on the buccal side of the template that offered buccal visibility of the surgical field, direct visual
148 control of the bone drilling, and better contact with the irrigation fluid.

149 The blocks in each group were subjected to four different experimental conditions, combining two factors
150 with two magnitudes each: 1) the depth of the bone osteotomy (6 or 13 mm), and 2) the temperature of the
151 irrigation fluid (fridge temperature 5 °C or room temperature 21 °C).

152 Figure 3 shows the distribution of the samples, specifying the number of blocks (n), number of osteotomies
153 (m), and number of temperature measurements (k).

154

155 **Surgical templates**

156 The solid blocks with the inserted acrylic resin teeth were digitally scanned (REV 100, Optical Reven
157 Dental, 3DBiotech®, Spain). The scanned image of each block was reconstructed with built-in software and
158 analyzed on a desktop computer with specialized implant planning software (Limaguide V01.2, Innovación
159 Dental®, Spain). The surgical templates were designed and finally printed with a 3D printer (M300,
160 Zortrax®, Poland) according to the design involved (open or closed). Each surgical template provided six
161 osteotomy drilling sites, prepared with six open or closed plastic sleeves to allocate 2 mm diameter drills.

162

163 **Drilling procedure**

164 A specially designed machine with an attached implant motor that could produce a continuous drilling
165 movement with a predetermined position and load was used. The device was coupled to a 20:1 reduction
166 speed handpiece with a predetermined load of 2 kg (SurgicPro, NSK Nakanishi Inc., Japan). A thermometer
167 with two thermocouples (DEM 106, Velleman®, Belgium) was added to the block to register the maximum
168 temperature reached during the osteotomies (Figure 4).

169 Two different 2 mm diameter drills (Limaguide V01.2, Innovación Dental®, Spain), with mechanical
170 stoppers at 6 mm and 13 mm, respectively, were used to perform the osteotomies through the surgical guide
171 to ensure the correct depth and minimize any human errors (LimaGuide system, Barcelona, Spain). The drill
172 speed was kept at 900 rpm constantly. Each set of drills was only used for five osteotomies to prevent drill
173 wear from influencing the results.

174

175 Saline solution (Braun, GmbH, Germany) was continually used at 50 ml/min, at two different temperatures:
176 room temperature (21 °C) and cooled in the fridge (5 °C). The saline solution's temperature was recorded
177 every 5 min and was replaced if the temperature changed by 1 °C.

178

179 **Data retrieval**

180 For the temperature measurements, two type K thermocouple devices were coupled to a digital thermometer
181 with an accuracy of 0.1 °C and inserted into two holes placed at each side of the osteotomy, 1 mm lateral to
182 the osteotomy level (6 and 13 mm, respectively). The maximum temperature obtained (in °C) in each

183 perforation drilling procedure in the two positions (buccal and lingual) was the primary variable of the study.
184 The temperatures were recorded in a Microsoft Excel Office® 2011 spreadsheet (Microsoft Corporation,
185 Redmond, USA). We waited until the temperature had returned to baseline (21 °C) before commencing the
186 next drilling procedure after each bone perforation.

187

188 **Statistical analysis**

189 The statistical analysis was performed using Statgraphics®Plus version 5.1 (Statpoint Technologies, Inc.
190 Virginia, USA). A descriptive analysis of the different variables was made. The Kolmogorov-Smirnov test
191 showed that the temperature data exhibited a non-normal distribution. However, the large sample size meant
192 that a parametric approach could be used. Regarding the inferential analysis, linear models of the generalized
193 estimation equations (GEE) were used, based on the hierarchical design of the observations (two
194 measurements per bed and six beds per block). The dependent variable was always the maximum
195 temperature reached, and the mean was compared for different levels of the independent factors. The chi-
196 squared statistic of Wald was considered for the significance of the factors, with a significance level of 5%
197 ($\alpha = 0.05$). Multiple comparisons were corrected according to Bonferroni's criteria.

198

199 **Results**

200 **Bone temperature assessment**

201 The maximum drilling temperature was reached in group one, with a mean value of 28.29 ± 4.02 °C,
202 approximately 1 °C above the value recorded in group one (27.16 ± 3.39 °C). The mean maximum
203 temperature was lower in the soft bone groups (groups three and four) than in the hard bone groups (groups
204 one and two). Furthermore, the mean maximum temperature was always higher with a closed guide than
205 with an open guide (23.38 ± 1.92 °C vs. 21.97 ± 1.22 °C). Table 1 details the temperatures reached in each
206 group. The mean maximum temperature was not equal in the four groups ($p < 0.001$). Bonferroni's multiple
207 comparison test showed that there were significant differences between any two of the groups. The detailed
208 results are listed in Table 2.

209

210 **Influence of bone density**

211 Bone density was found to introduce great variability in heat generation, as seen in Figure 5. The effect of
212 bone type on temperature was the same regardless of whether a closed or open guide was used. Linear
213 models of GEE were established. First, in hard bone (D1), the temperature was significantly higher (p
214 < 0.001). Second, the temperature was significantly higher with a closed guide ($p < 0.001$). Third, the effect of
215 bone type did not appear to be influenced by the type of template ($p = 0.436$). Fourth, the effect of the type of
216 guide did not appear to be influenced by the type of bone density ($p = 0.436$). However, the chi-squared
217 statistic was much greater for the bone factor than the guide factor, implying that the observed variability
218 was mostly due to bone density. The detailed results are presented in Table 2.

219

220 The maximum threshold temperature of ≥ 47 °C, as described by Eriksson and Albrektsson (1983, 1984),
221 was only reached in the hard bone groups. In contrast, the 47 °C threshold was never reached in the soft bone
222 groups.

223

224 **Influence of osteotomy drilling depth and cooling irrigation**

225 Apart from the observation that the combination of hard bone (D1) and closed guide yielded the highest
226 temperatures ($p < 0.001$), the 13 mm drilling depth in groups one and two (dense bone) was associated with
227 higher mean temperatures than with a 6 mm drilling depth. In contrast, in group three, the mean temperature
228 at a depth of 6 mm was higher than at 13 mm. Besides, in group four, no difference was observed in the
229 mean temperature between osteotomy depths (Tables 1 and 2).

230

231 Lastly, the irrigation fluid temperature also significantly influenced the heat generated during bone drilling (p
232 < 0.001). Chilled fluid at 5 °C reduced the temperature peaks for all groups to similar degrees, which was not
233 the case with room temperature irrigation (Table 1). For the 6 mm drilling depth, the reduction of the bone
234 temperature in relation to bone density and irrigation fluid temperature proved to be more linear across the
235 four groups (Figure 6). The detailed results are shown in Table 2.

236

237 Most importantly, maximum temperatures above the threshold of 47 °C (Eriksson & Albrektsson, 1983;
238 Eriksson & Albrektsson, 1984) were only observed in the high-density bone groups when room temperature
239 irrigation was used. The maximum temperatures never reached the 47 °C threshold when chilled irrigation
240 fluid was used.

241

242 **Discussion**

243 Our findings indicate that high-density bone and closed surgical guides contribute to increased intra-osseous
244 temperature during bone drilling compared to low-density bone and open guides. Furthermore, high-density
245 bone and greater drilling depths lead to increased heat generation. Using chilled irrigation fluid helps to
246 reduce heat generation and avoid bone temperatures above the 47 °C threshold.

247

248 Several factors have been reported to influence bone temperature during bone drilling, including bone
249 density, cortical bone thickness, aspects related to bone drilling parameters, different surgical guides, and
250 irrigation fluid temperature (Augustin et al., 2012; Boa et al., 2016; Bolland et al., 2004; Delgado-Ruiz et al.,
251 2018; Jeong et al., 2014; Lee, Gozen, & Ozdoganlar, 2012).

252

253 Bone density has been previously reported as the primary influencing factor behind increased temperature
254 variability (Karaca, Aksakal, & Kom, 2011). It has been argued that the duration of bone drilling depends on
255 the hardness of cortical bone, which correlates to bone mineral density, and these parameters are closely
256 related to increased bone temperature (Karaca et al., 2011). Furthermore, the present study found that the
257 depth of the bone preparation was seen to significantly influence the bone temperature reached in hard

258 density bone, in agreement with Misir et al. (Misir et al., 2009). They recorded an increase in heat generation
259 associated with deeper osteotomy drillings (3, 6, and 9 mm). However, they did not take bone density into
260 account. Our study did not find a significant increase in the intra-bone temperature associated with deeper
261 osteotomies in soft bone. Interestingly, the mean bone temperature was higher at a depth of 6 mm than at 13
262 mm. This may be due to the short duration of the shallow drilling procedure in soft bone and the lesser effect
263 of the irrigation fluid over this short period.

264
265 The use of surgical templates to achieve better results in implant positioning has several shortcomings.
266 Surgical templates can reduce the surgeon's visibility and prevent the irrigation fluid from cooling the drill
267 during bone preparation, thereby generating more heat than with freehand implant placement surgery
268 (Fauroux et al., 2018; Misir et al., 2009). However, not all surgical template designs have the same effect in
269 terms of bone temperature changes (Amorfini et al., 2017; Farley et al., 2013; Pozzi et al., 2014; Younes et
270 al., 2018). An open guide design enhances surgical visibility and allows for the proper irrigation of the drills
271 while transferring the pre-surgical implant positioning planning to the surgical sites (Fauroux et al., 2018). In
272 the present study, open surgical guides reduced the bone temperature during drilling by approximately 1 °C
273 compared to closed guides in hard and soft density bone. Nevertheless, this degree of difference did not
274 imply a variation that exceeded the recommended temperature limit.

275
276 The mean temperature reached in our study was far from the temperature threshold of 47 °C reported by
277 Eriksson and Albrektsson in 1983 and 1984 (Eriksson & Albrektsson, 1983; Eriksson & Albrektsson, 1984),
278 and is in agreement with the observations of Di Fiore, Sivoletta, Stocco, Favero and Stellini (2018).

279
280 Nonetheless, this study revealed that bone temperatures above 47° C during implant bed preparation might be
281 reached in high-density bone when room temperature irrigation fluid is used. This finding highlights the
282 importance of using a chilled irrigation fluid to avoid excess intra-bone temperatures. Irrigation during bone
283 drilling has two main functions, that is, to reduce the bone temperature (Liu et al., 2018) and enhance the
284 removal of bone debris (Augustin et al., 2012). Boa et al. found that external irrigation during flapless guided
285 surgery could reduce 50% of the bone temperature generated compared to non-irrigation (Boa et al., 2015).
286 Interestingly, studies involving implant static navigation surgery have reported no differences in terms of the
287 bone temperature reached between continuous and intermittent drilling (Di Fiore et al., 2018) or between
288 flapless and flap surgery (Jeong et al., 2014). Furthermore, according to the results of Boa et al. (2016) and
289 Di Fiore et al. (2018), the use of pre-cooled irrigation fluid (10 °C and 6 °C, respectively) is recommended to
290 control bone temperature, especially when using a closed guide. This is in agreement with our findings,
291 where chilled irrigation fluid (5 °C) reduced the intra-bone temperature compared to room temperature
292 irrigation fluid (21 °C) and prevented intra-bone temperatures from rising above 47° C during implant bed
293 preparations in high-density bone.

294

295 This study has several limitations. Polyurethane foam blocks (Möhlhenrich et al., 2015) may differ from
296 living bone in quality and elasticity. However, they have been found to have physical and mechanical
297 characteristics similar to cortical and cancellous bone and are easy to standardize to different bone densities.
298 Nevertheless, the results obtained could differ from those recorded in clinical practice due to differences
299 between patients (Vilani, Ruellas, Mattos, Fernandes, & Elias, 2015). Additionally, to standardize this study,
300 the drilling speed and drill diameter were not modified throughout the investigation, and only external
301 irrigation was employed. The use of other conditions could have yielded different results. Drilling
302 parameters such as the drill design (e.g., helix angle, chisel edges, helical or straight flutes), drill diameter,
303 and the type of material have all been shown to influence bone temperature during drilling (Matthews &
304 Hirsch, 1972). Moreover, drill wear and sterilization cycles leading to drill dullness also contribute to heat
305 production (Matthews & Hirsch, 1972; Möhlhenrich et al., 2015). Additionally, drilling speed, the torque
306 applied, and the depth of the preparation have all been shown to modify bone temperature during bone
307 drilling (Cordioli & Majzoub, 1997; Matthews & Hirsch, 1972). Delgado-Ruiz et al. (Delgado-Ruiz et al.,
308 2018) reported significantly lower temperatures when drilling at a slow-speed (50 rpm; 22.11 ± 0.8 °C)
309 compared to the drilling speeds of 150 rpm (24.75 ± 1.1 °C) and 300 rpm (25.977 ± 1.2 °C). Furthermore,
310 the bone temperatures in cortical bone were found to be significantly higher while working at higher speeds.
311 No irrigation with lower speeds was revealed to avoid excessive heat generation (Salomó-Coll et al., 2020).
312 Salomó-Coll et al. (2020) also revealed that the mean temperatures attained when using 2 or 3.5 mm
313 diameter drills were similar, indicating little variability with different drill diameters under similar conditions
314 (Salomó-Coll et al., 2020). This contrasts with other studies that showed that a decrease in drill diameter is
315 related to an elevation in bone temperature due to increased energy transfer to the bone (Boa et al., 2016;
316 Matthews & Hirsch, 1972).

317 Furthermore, internal or double irrigation systems were not used. However, their benefits remain
318 controversial. Boa et al. (2016) described that double irrigation during osteotomies could reduce the heat
319 generated in the bone. Misir et al. (2009) observed no differences in bone heat generation with external or
320 external and internal irrigation during implant osteotomies with surgical guides. For standardization
321 purposes, external irrigation was selected in the present study to achieve clear results and more specific
322 clinical guidance for this commonly used irrigation system.

323
324 Further studies should involve an upgraded mechanism to assess the intra-bone temperature during implant
325 bed preparations. For example, a device that does not require the insertion of thermocouple electrodes inside
326 the apical zone of the bone preparations would be preferable.

327
328 When interpreting the results of this study, it is important to recognize that the mean maximum temperature
329 reached the 47 °C threshold in none of the tested scenarios. However, when looking at the maximum values
330 measured in each group, a bone temperature of >47 °C was reached in several samples when irrigating with
331 room temperature fluid in dense bone sites. Using a chilled irrigation fluid (5° C) was found to be effective in
332 maintaining the intra-bone temperature below the 47 °C threshold.

333

334 **Conclusions**

335 Within the limitations of this *in vitro* study, it can be concluded that in high-density bone (D1), a closed
336 guide for a deep osteotomy will lead to the highest intra-bone temperatures. With a closed static surgical
337 guide, the final intra-bone temperature was about 1 °C higher than with an open guide. The osteotomy depth
338 significantly affected the maximum temperature reached in high-density bone samples, especially when
339 room temperature irrigation fluid was used. This study also confirmed the value of using chilled irrigation
340 fluid (~5 °C) to avoid heat generation above 47 °C in dense bone.

341

342

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344

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453 **Tables legends**

454 Table 1. The temperature recorded (°C) according to each group, osteotomy drilling depth, and irrigation
455 fluid.

456

457 Table 2. Beta coefficients (mean temperature differences) and 95% CI for different effects from GEE models.
458 Corrected p-values of Wald's Chi² test using Bonferroni's criteria (*p<0.05; **p<0.01; ***p<0.001)

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460

461 **Figure legends**

462 Figure 1. Closed implant guide.

463 Figure 2. Open implant guide.

464 Figure 3. Flowchart distribution of the study sample.

465 Figure 4. Handpiece, thermometer, and thermocouple.

466 Figure 5. Boxplot of bone temperature distribution according to each group.

467 Figure 6. Mean maximum temperature (°C) according to each group, osteotomy drilling depth, and irrigation
468 fluid.

Table 1

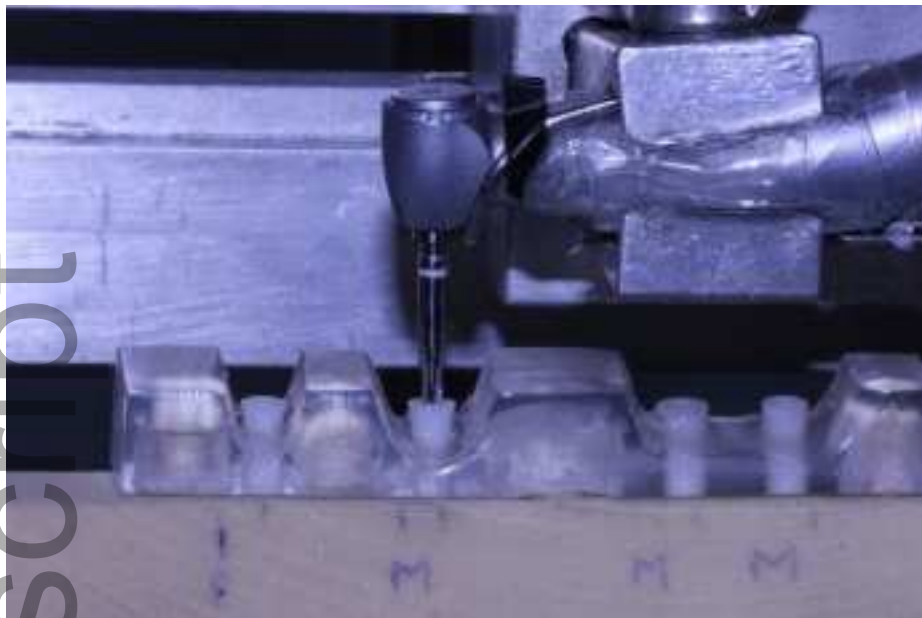
GROUP	DRILLING DEPTH (mm)	IRRIGATION FLUID (°C)	N	MEAN	SD	MINIMUM	MAXIMUM	MEDIAN
1	6	5	120	27,58	3,85	21,60	37,90	26,30
		21	120	28,17	5,26	22,30	50,00	26,50
	13	5	120	28,26	3,26	22,00	35,60	27,55
		21	120	29,17	3,25	23,20	35,40	28,70
2	6	5	120	26,05	1,81	21,80	32,10	25,95
		21	120	27,10	4,31	22,00	47,80	26,55
	13	5	120	27,14	2,90	22,30	35,60	26,90
		21	120	28,34	3,64	22,30	36,70	28,00
3	6	5	120	23,95	1,39	19,80	27,30	24,05
		21	120	24,77	2,33	21,10	30,10	24,20
	13	5	120	22,04	,98	20,30	27,10	22,00
		21	120	22,77	1,43	20,10	26,80	22,45
4	6	5	120	21,28	,85	19,50	23,20	21,30
		21	120	22,82	1,24	20,10	26,20	22,70
	13	5	120	21,48	,91	19,70	23,70	21,50
		21	120	22,29	1,16	20,10	26,40	22,10

Table 1. Temperature recorded (°C) according to group, drilling depth and irrigation fluid.

Table 2

Comparison between GROUPS	MEAN TEMPERATURE DIFFERENCE (° C)	95% CI	p-value
1 - 2	1.13	0.51 – 1.77	0.003**
1 - 3	4.91	4.31 – 5.51	<0.001***
1 - 4	6.33	5.74 – 6.91	<0.001***
2 - 3	3.78	3.28 – 4.27	<0.001***
2 - 4	5.19	4.66 – 5.72	<0.001***
3 - 4	1.42	1.18 – 1.66	<0.001***
D1 – D4	4.90	4.30 – 5.50	<0.001***
Closed – Open guide	1.42	1.18 – 1.66	<0.001***
in D1: Closed - Open	1.14	0.51 – 1.77	0.003**
in D4: Closed - Open	1.42	1.18 – 1.66	<0.001***
in Closed: D1 – D4	4.91	4.31 – 5.51	<0.001***
in Open: D1 – D4	5.19	4.66 – 5.72	<0.001***
Fluid temp: 5 – 21°	0.96	0.62 – 1.29	<0.001***
Depth: 6 – 13 mm	-0.03	-0.19 – 0.13	0.717
In Group 1: Depth 6 – 13mm	-0.83	-1.29 - -0.37	0.010*
In Group 2: Depth 6 – 13mm	-1.16	-1.96 - -0.36	0.124
In Group 3: Depth 6 – 13mm	1.95	1.47 – 2.43	<0.001***
In Group 4: Depth 6 – 13mm	0.16	0.03 – 0.29	0.455

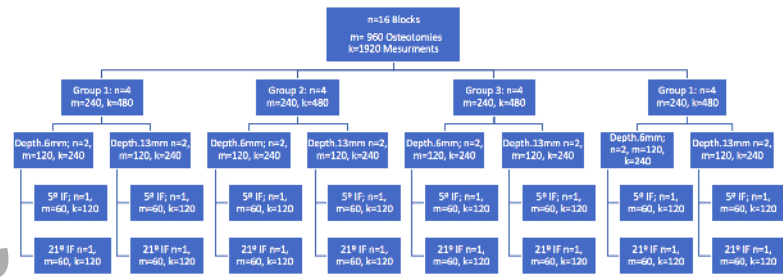
Table 2. Beta coefficients (mean temperature differences) and 95% CI for different effects from GEE models. Corrected p-values of Wald's Chi² test using Bonferroni's criteria (*p<0.05; **p<0.01; ***p<0.001).



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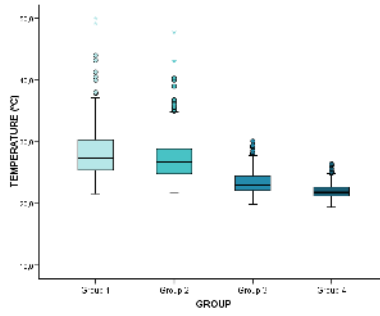


n: blocks; m: osteotomies; k: temperature measurements; IF: irrigation fluid.

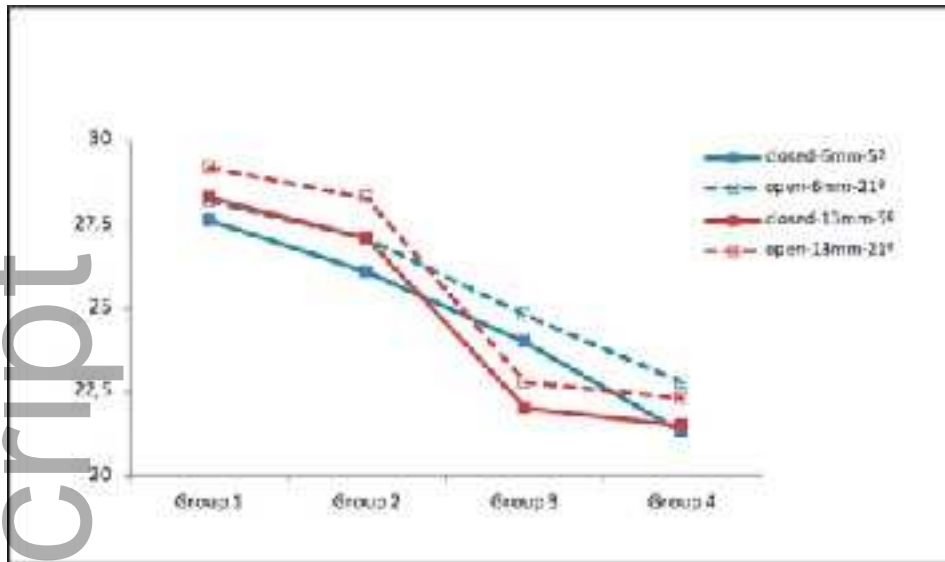
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