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Comparison of epidural versus intrathecal anaesthesia in dogs undergoing pelvic limb orthopaedic surgery

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(Article begins on next page)

1 **Comparison of epidural versus intrathecal anaesthesia in dogs undergoing pelvic limb** 2 **orthopaedic surgery**

3 **Abstract and Keywords**

4 **Objective:** To compare the procedural failure rate (PFR), intraoperative rescue analgesia (iRA)
5 probability and postoperative duration of motor block after epidural and intrathecal anaesthesia in
6 dogs undergoing pelvic limb orthopaedic surgery.

7 **Study design:** Prospective, randomized clinical trial.

8 **Animals:** 92 client-owned dogs.

9 **Methods:** Dogs were assigned randomly to receive either lumbosacral epidural anaesthesia (EA),
10 injected through a Tuohy needle (0.1 mg kg⁻¹ bupivacaine 0.5% and 0.1 mg kg⁻¹ morphine 1%), or
11 intrathecal anaesthesia with the same drugs in a hyperbaric solution (HIA) injected through a
12 Quincke needle at the L5-L6 level, with dosage based on body mass and spinal cord length.
13 Inaccurate positioning of the Tuohy needle, assessed by radiographic imaging, and lack of cerebral
14 spinal fluid outflow were considered procedural failures (PFs) of EA and HIA, respectively.
15 Fentanyl (1 µg kg⁻¹ IV) was provided for intraoperative rescue analgesia. Its use was recorded as a
16 sign of intraoperative analgesic failure. Fentanyl was administered when either the heart rate or the
17 mean arterial pressure increased by 30% above the pre-stimulation value. The motor block
18 resolution was postoperatively evaluated.

19 **Results:** The PFRs in the EA and HIA groups were 15/47 (32%) and 3/45 (7%), respectively
20 ($p=0.003$). Differences in iRA were analysed in 26 and 30 subjects in the EA and HIA groups
21 respectively, using Kaplan-Meier survival analysis. The iRA probability within the first 80 minutes
22 of needle injection (NI) was higher in the EA group ($p= 0.045$). The incidence of dogs walking
23 within 3 hours of NI was significantly higher in the HIA group (8/20, 40%) than in the EA group
24 (0/17) ($p = 0.004$).

25 **Conclusions and Clinical Relevance:** HIA was found to have lower PF, lower intraoperative
26 analgesic failure and faster motor block resolution. In this study HIA was shown to provide some
27 advantages over EA in dogs undergoing commonly performed pelvic limb orthopaedic surgery in a
28 day-hospital regime.

29 *Keywords:* dogs, epidural anaesthesia, orthopaedic surgery, spinal anaesthesia.

30 **Main text**

31 **Introduction**

32 Single injection epidural anaesthesia (EA) is the most performed neuraxial technique in dogs
33 undergoing common orthopaedic pelvic limb surgery and its use is supported by abundant
34 veterinary literature (Troncy et al. 2002; Valverde 2008; Campoy et al. 2012; Caniglia et al. 2012).
35 Some studies have shown that EA can decrease postoperative pain and rescue analgesic
36 requirements (Hendrix et al. 1996; Kona-Boun et al. 2006). In contrast, intrathecal anaesthesia is
37 usually the preferred neuraxial technique in humans undergoing pelvic limb surgery in a day-
38 surgery regime (Korhonen 2006). The feasibility, incidence of side effects and quality of intrathecal
39 nervous block in dogs undergoing anaesthesia for pelvic limb orthopaedic surgery have been
40 recently reported (Sarotti et al. 2011, 2013); however, studies that compare these two neuraxial
41 techniques in a clinical setting, allowing for an evidence-based choice, are still lacking. Ideally,
42 regional anaesthesia should be able to achieve intraoperative muscle relaxation, a prolonged
43 sensitive block in order to limit systemic drugs administration, and rapid postoperative motor
44 function recovery.

45 The aim of this study was to prospectively compare the use of epidural and intrathecal
46 anaesthesia using bupivacaine and morphine. The procedural failure rate (PFR), intraoperative
47 rescue analgesia (iRA) requirement and time to postoperative motor block resolution was evaluated
48 in dogs anaesthetized for pelvic limb orthopaedic surgery.

49 **Materials and methods**

50 This study was approved by the Ethical Committee of the University of Padua (Prot. N. 49574) and
51 all owners gave their informed consent.

52 *Animals and inclusion criteria*

53 Dogs, older than 6 months presenting to the Centro Veterinario Fossanese, from January 2011 to
54 January 2013, for various scheduled surgical procedures involving the pelvic limbs were enrolled in

55 this prospective study. All animals underwent a preoperative physical examination. Blood test
56 analyses, such as packed cell volume, plasma total protein, urea, creatinine and electrolyte
57 concentrations, were performed for all the animals. Dogs were fasted 8 hours prior to surgery, while
58 water was freely available.

59 Dogs were excluded from the study when, based on clinical and laboratory examination, the
60 American Society of Anesthesiologists (ASA) Physical Status class of assignment was III or
61 superior and when there were reasons to consider neuraxial anaesthesia as being either absolutely
62 (infection of the puncture site, uncorrected hypovolemia, bleeding disorders, degenerative central or
63 peripheral diseases, anatomical abnormality of the lumbosacral region, the disapproval of the
64 owner) or relatively (bacteremia and neurologic disorders) contraindicated. Dogs undergoing
65 surgical procedures distal to the knee joint were excluded from iRA and postoperative evaluation.
66 Motor block resolution was not evaluated in dogs that were unable to walk before surgery or that
67 had postoperative bandaging of the leg.

68 *Randomization*

69 The estimated sample size to detect a difference in the primary endpoint (rate of iRA probability at
70 60 minutes) with a power of 80% and an alpha error of 5% using a two group study design has been
71 performed with an effect size (w) of 0.61. It resulted in a minimum number of subjects of 21 in each
72 group. The effect size (w) was calculated with a hypothesis of iRA probability in the HIA group of
73 10% and in the EA group of 40%. The dogs were assigned to one of the two treatment groups
74 according to a computer-generated randomization sequence: epidural anaesthesia (EA) group or
75 hyperbaric intrathecal anaesthesia (HIA) group, using a stratified randomization for type of surgery
76 in order to obtain homogenous groups. All anaesthetic procedures were performed by the same
77 experienced operator (DS), who was not blinded to the assigned technique.

78 *Anaesthesia*

79 All the dogs received a fentanyl bolus ranging from 1 to 3 $\mu\text{g kg}^{-1}$ intravenously (IV) two minutes
80 before anaesthesia induction through a previously inserted catheter. The dosing of fentanyl was

81 decided by the anaesthetist, based on the dog's temperament, in order to achieve stress-free
82 anaesthesia induction. General anaesthesia (GA) was induced by administering propofol to effect
83 and was maintained using a variable rate of propofol titrated to maintain a sluggish palpebral reflex.
84 All patients were allowed to breathe spontaneously during anaesthesia, unless end-tidal CO₂
85 (P_{E'}CO₂) exceeded 6 kPa, in which case intermittent positive pressure ventilation (Alpha-Delta,
86 Siare, Italy) was imposed to restore and maintain normocapnia. Lactated Ringer's solution (Ringer
87 Lattato; Fresenius Kabi, Italy) was administered IV at 10 mL kg⁻¹ hour⁻¹ during anaesthesia in all of
88 the dogs. The oscillometric technique (Viridia C26, HP, Germany) was used to measure systemic
89 arterial blood pressure every 2-3 minutes using an appropriately sized cuff placed on the distal third
90 of the left forelimb, while the fraction of inspired oxygen (FiO₂), P_{E'}CO₂, electrocardiogram,
91 arterial oxygen saturation, heart rate (HR), respiratory rate and oesophageal temperature were
92 monitored continuously (Viridia C26, HP, Germany). In order to perform the regional technique,
93 anaesthetized dogs were positioned in lateral recumbency and the skin over the L₃-S₁ vertebrae was
94 aseptically prepared with chlorhexidine (4%) and alcohol (70%), after clipping the hair. The body
95 temperature was maintained above 35°C during the perioperative period using an active heating
96 system (Bair Hugger Warmer Model 505, Augustine Biomedical Design, MN, USA). Thirty
97 minutes before the end of surgery, all dogs received 0.2 mg kg⁻¹ of meloxicam (Metacam 0.5%;
98 Boehringer Ingelheim, Spain) subcutaneously (SC). At the end of surgery, the urinary bladder was
99 manually voided. Two different experienced operators evaluated postoperative pain 30 minutes
100 after extubation and then every 2 hours until discharge using the short form of the Glasgow
101 composite pain scale (Reid et al. 2007). If Glasgow composite pain scale score was \geq 6 methadone
102 0.1 mg Kg⁻¹ was administered IM as a test dose, when it was $>$ than 6 methadone 0.2 mg kg⁻¹ was
103 administered, as rescue analgesia..

104 The following perioperative events were recorded: bradycardia (HR<60 beats minute⁻¹),
105 hypotension (mean arterial pressure (MAP) < 60 mmHg for at least 5 min or any MAP value lower
106 than 55 mmHg), vomiting, pruritus and neurologic deficit. Hypotension was treated by reducing the

107 administration of general anaesthetic and by giving a bolus of fluids (Lactate's Ringer) at 3 mL kg⁻¹
108 IV. If MAP increased after the first bolus, an additional 2 mL kg⁻¹ of fluid was administered. If the
109 hypotension persisted, the dogs were treated with a bolus of ephedrine (50-100 µg kg⁻¹) and/or
110 norepinephrine CRI (0.05-0.3 µg kg⁻¹ min⁻¹). Urinary retention was defined as the inability to
111 spontaneously void in the presence of bladder over-distension. Bladder over-distention was
112 evaluated by abdominal palpation and ultrasonography in all patients that did not spontaneously
113 urinate within 12 hours after discharge from the veterinary clinic. Owners were instructed to
114 monitor their dog's micturition and to report episodes of prolonged sedation and marked lameness.
115 If the owner noted that the dog had not urinated for at least 12 hours, he or she was asked to return
116 the dog to the veterinary practice. Inaccurate positioning of the Tuohy needle, as assessed by
117 radiographic imaging (Manchikanti et al. 2004) and lack of cerebral spinal fluid (CSF) outflow
118 were considered procedural failures, respectively, of EA and HIA. An iRA (fentanyl 1 µg kg⁻¹ IV)
119 was administered when HR or/and MAP was raised by more than 30% of the pre-incisional value,
120 which was defined as the mean value of the parameter during the 5 minutes prior to skin incision.
121 One µg kg⁻¹ of fentanyl IV was repeated every two minutes until HR and MAP were below 30% of
122 the pre-incisional value. The ability to walk was tested by the same operator (DS). If necessary,
123 dogs could be helped to get up, but they had to walk on their own. Recovery of ambulation was
124 tested at 3, 4, 5 and 8 hours after performing the neuraxial technique, unless the dog had already
125 been sent home because it was able to walk.

126 *EA Group*

127 An epidural injection was administered with a Tuohy needle (Perican 22, 20 or 18 G; B. Braun,
128 Brazil), with the dogs in lateral recumbency, using a median approach at the level of the
129 lumbosacral intervertebral space (L₇-S₁). The needle was advanced perpendicularly into the skin
130 until an increase in resistance was felt, indicating the *ligamentum flavum*. The epidural space was
131 then identified using the following clinical signs: 1) loss of resistance (LOR), assessed using an air-
132 filled LOR syringe (Perifix, B. Braun; Germany) and 2) sudden LOR to needle advancing. The

133 correct positioning of the needle was confirmed by a radiograph, with the animal in lateral
134 recumbency. If the radiograph showed incorrect positioning, a further two radiographically assessed
135 attempts were made to reach the epidural space. An isobaric solution of bupivacaine 0.5%
136 (Bupivacaina Angelini 5 mg mL⁻¹; Angelini, Italy) at 1 mg kg⁻¹ (maximum dose allowed 30 mg, 6
137 mL) and morphine 1% (Morfina Cloridrato; Molteni, Italy) at 0.1 mg kg⁻¹ was administered over at
138 least 20 seconds. Surgery started between 25 and 50 minutes after the epidural injection.

139 *HIA Group*

140 The intrathecal injection was administered using a paramedian approach at the level of the
141 intervertebral space between L₅ and L₆. Three attempts were allowed to reach the subarachnoid
142 space. A 75-mm-long 25 G Quincke needle (Aghi spinali; Pic, Italy) was used to administer all the
143 intrathecal injections reported in this study. Once the CSF outflow became visible in the hub of the
144 needle, the intrathecal solution was injected over 20-40 seconds. The needle bevel always faced
145 cranially during administration of the intrathecal solution. Once the injection was complete, dogs
146 were maintained in lateral recumbency with the pelvic limb to be operated lowermost for at least 12
147 minutes (Hocking & Wildsmith 2004).

148 The bupivacaine (Bupisen iperbarica 0.5%; Galenica Senese, Italy) dose calculations were
149 based on body mass (BM) and spinal cord length (SCL). SCL was determined as being the distance
150 between the caudal part of the L7 spinal process and the occipital bone in accordance with the
151 following equation (Sarotti et al. 2013).

$$152 \text{ Bupivacaine 0.5\% (mg): } 0.21 \text{ BM (kg)} + 0.035 \text{ SCL (cm)}$$

153 The formula used to calculate the bupivacaine dose resulted in a dosing regimen 30% less than that
154 suggested by Sarotti et al. (2013). The 1% morphine dose was 0.3 mg in dogs less than 10 kg, 0.5
155 mg in dogs between 11 and 20 kg, and 1 mg in dogs over 20 kg.

156 *Statistical Analysis*

157 Categorical variables were reported as frequencies and percentages and differences between groups
158 were analysed using exact fisher's test. Continuous variables were checked for normal distribution

159 with visual inspection of bar graphs, histograms and using the Shapiro–Wilk test. Data not normally
160 distributed were reported as the median and the range (minimum–maximum) and differences
161 analyzed with Mann-Whitney test. The significance level was set at 5% for all statistical methods.
162 The intraoperative time-to-event probability of iRA was analysed using Kaplan-Meier survival
163 analysis. The survival curves were analysed using the log-rank test and Hazard ratio statistic
164 (MedCalc Software for Windows version 12.5 Belgium).

165 **Results**

166 The types of pelvic limb orthopaedic surgery and the demographic data are listed, respectively, in
167 Tables 1 and 2. Procedural data concerning the local anaesthetic (LA) dose related to BM and SCL,
168 the morphine dose related to BM, the fentanyl induction bolus, the propofol induction bolus, the
169 median propofol dose consumption in the first hour, the time between the intrathecal injection and
170 the beginning of surgery and between the intrathecal injection and the end of surgery are reported in
171 Table 3. Intermittent positive pressure ventilation was provided to 24/26 (92%) dogs in the EA
172 group and 25/30 (83%) in the HIA group.

173 Epidural and intrathecal anaesthesia were attempted on 92 dogs, with an overall PFR of
174 15/47 (32%) and 3/45 (7%), respectively ($p = 0.003$). The Tuohy needle was correctly repositioned
175 and an epidural injection administered in eight out of the 15 cases in the EA group, using
176 radiographic imaging to redirect the needle. Considering that 14 dogs were excluded because they
177 had surgery distal to the knee joint in the EA group and 12 were excluded using the same criteria in
178 the HIA group, a total of 26 and 30 anaesthetic periods, respectively, were analysed for
179 intraoperative and postoperative evaluation. A CONSORT diagram is shown in Figure 1.

180 Procedural data concerning the local anaesthetic (LA) dose related to BM and SCL, the
181 morphine dose related to BM, the fentanyl induction bolus, the propofol induction bolus, the

182 median propofol dose consumption in the first hour, the time between the intrathecal injection and
183 the beginning of surgery and between the intrathecal injection and the end of surgery are reported in
184 Table 3.

185 The iRA survival probability at 60 minutes after needle injection (NI) was 64% (16/25) and
186 90% (26/29), respectively, in the EA and HIA groups, and 52% (11/21) and 68% (13/19),
187 respectively, in the EA and HIA groups 80 minutes after NI. The iRA probability during surgical
188 stimulation within the first 80 minutes of NI was higher in the EA group ($P=0.045$ log-rank test,
189 Hazard Ratio 0.355, 95% CI 0.127 to 0.991) (Fig. 2). The occurrence of iRA was not related to the
190 LA or morphine dose, BM, SCL, age or type of surgery ($p > 0.05$). There was no difference in the
191 incidence of perioperative side effects between the groups ($p > 0.05$). The incidence of hypotension
192 was 6/26 (23%) and 7/30 (23%), while bradycardia was found in 1/26 (4%) and 2/30 (7%) cases,
193 respectively, in the EA and HIA groups. Two cases in the EA group and five in the HIA group
194 needed norepinephrine CRI to maintain normotension. In the HIA group, one case of postoperative
195 pruritus, focused in the back area, was recorded 3 hours after the intrathecal injection and was
196 treated with a bolus of propofol ($1 \text{ mg kg}^{-1} \text{ IV}$) followed by 30 minutes of CRI ($10 \text{ mg kg}^{-1} \text{ h}^{-1}$). One
197 case of urinary retention was recorded in the EA group and was treated by urinary catheterization
198 20 hours after the local injection. No nervous deficit or sign of paraesthesia was recorded 24 hours
199 after the loco-regional technique in any patient. All patients included in this study scored less than
200 six on the Glasgow composite pain scale, during the postoperative observation period (at least 5
201 hours after NI) and did not receive postoperative rescue analgesia. The return of ambulation at 3
202 hours and at 4 hours after NI was significantly higher in the HIA group ($p = 0.004$ and $p = 0.045$,
203 respectively). **There was not a statistical difference in the motor block resolution at 5 and 8 hours**
204 **($p > 0.05$)**, (see Table 4).

205 Discussion

206 This study demonstrates that a single HIA injection provides a lower PFR, a higher intraoperative
207 analgesia efficacy and faster motor block resolution compared with a single EA injection in dogs
208 undergoing pelvic limb orthopaedic surgery. According to our results LOR technique to identify
209 epidural space can be inaccurate, even when executed by an expert operator in about one-third of
210 cases. In this study, the PFR in the HIA group was five times lower than in the EA group ($p =$
211 0.003).

212 One reason to explain this finding may simply be the different end point for needle
213 positioning between the two techniques. While correct positioning of the needle during IA can be
214 ascertained by CSF outflow, the LOR technique is prone to operator subjectivity and experience.
215 The PFR found in the EA group, using the LOR technique to identify the epidural space, was higher
216 than previously reported for dogs (Iff & Moens 2010). In the veterinary literature, to the authors'
217 best knowledge, this is the first work that has studied the PFR, as determined by radiographic
218 assessment, of EA performed using the LOR technique in dogs. To evaluate procedural failure by
219 just clinically assessing the presence of nervous block can be misleading for at least three reasons.
220 First, the needle can correctly reach the epidural space through a different spinal segment than
221 planned; second, as reported in the literature, the LA can, though correctly injected into the epidural
222 space, not produce a consistent nervous block (Curatolo et al. 1995); and, third, the tip of the needle
223 can be positioned paravertebrally next to the lumbar plexus, thus producing a nervous block of the
224 posterior limb. Other techniques have been proposed to monitor the correct positioning of the
225 needle in the epidural space, such as electrical stimulation, ultrasonography, fluoroscopy and
226 detection of pressure changes on entering the epidural space (Read 2005; Iff et al. 2007; Naganobu
227 & Hagio 2007; Carvalho 2008) but unfortunately they add to the complexity of the procedure and
228 often require expensive equipment.

229 In this study, dogs undergoing EA had a significantly higher iRA probability. This finding is
230 in accordance with what is well established for humans. Inadequate surgical analgesia, after spinal
231 anaesthesia, has been reported as being less than 1% (Fettes et al. 2009) while it ranges between 9

232 and 15% after epidural anaesthesia (Curatolo et al. 1995; Kinsella 2008). In contrast to the
233 subjective experience of many anaesthetists, failure of epidural anaesthesia is a frequent clinical
234 problem in humans: a recent heterogeneous cohort review of 2140 surgical patients undergoing
235 lumbar epidural anaesthesia reported a failure rate as high as 27% (Hermanides et al. 2012).
236 However, it is difficult to meaningfully compare our data with those presented for humans for many
237 reasons, not least because single-injection epidural anaesthesia is rarely used in daily practice in
238 adult subjects. The use of this technique, which does not allow top-up dosing, could result in a
239 higher incidence of inadequate control of surgical stimulation. One of the reasons that can explain
240 the difference in efficacy between the neuraxial techniques discussed in this study arises from
241 anatomical differences relative to the site of injection of the LA. Intrathecal anaesthesia provides a
242 dense block due to the deposit of LA and morphine next to the spinal nerve roots, while during an
243 epidural block the solution is injected into the extra-meningeal layer, which produces a less dense
244 nervous block. The higher morphine concentration, around the spinal cord, produced by intrathecal
245 injection could also have produced a faster onset time and more intense analgesic effect in the HIA
246 group.

247 Another reason to explain the poorer efficacy of EA could be incorrect dosing of the LA
248 used to perform the epidural block. The use of BM as the only predictive variable of the LA dosing
249 might not be adequate in all canine subjects. In the authors' clinical experience and as suggested by
250 a recent study (Otero et al. 2009) a linear correlation does not exist between the appropriate LA
251 dosage and BM. The use of a fixed ratio between the volume of LA and BM can increase the risk of
252 underdosing in small subjects and overdosing in heavier ones, even though the use of 6 mL as a
253 maximum volume can limit this problem. In the HIA group, the LA dose was calculated by
254 including the SCL in the formula along with BM. In a recent study the SCL was found to be a
255 predictive variable of LA to control cardiovascular response in dogs undergoing pelvic limb surgery
256 (Sarotti et al. 2011).

257 Inadequate surgical analgesia in the HIA group started to become apparent 80 minutes after
258 the subdural injection, as shown by the Kaplan-Meier curve, probably due to the offset of the
259 nervous block. A similar characteristic of intrathecal nervous block in dogs has been previously
260 reported in veterinary literature (Sarotti et al. 2013). Hyperbaric bupivacaine solution was used to
261 perform intrathecal anaesthesia. Baricity refers to the density of a substance compared with the
262 density of CSF. A local anaesthetic is commonly made hyperbaric by adding dextrose to the
263 mixture, such solutions will flow in the direction of gravity and settle in the most dependent areas of
264 the intrathecal space. The main advantage of using hyperbaric solutions is to produce a profound
265 unilateral spinal block with a reduced haemodynamic impact (Di Cianni et al. 2008).

266 This study included dogs undergoing different orthopaedic procedures. As a consequence,
267 surgical stimulation may have been variable in terms of intensity and nerves involved in the
268 nociceptive transmission. In order to reduce this source of variability, dogs undergoing surgery
269 below the knee joint were excluded from iRA and postoperative analysis. The vast majority of
270 nociceptive nerves involved in the surgical stimulation on the distal part of the rear leg belong to the
271 sciatic nerve.

272 The major study limitation of this work is that the operators collecting the data were aware
273 of the group assignment of the dogs enrolled. This limitation may have influenced our findings.
274 Subjects belonging to HIA group may have been involuntarily kept at a deeper level of anaesthesia by the
275 operators, introducing a bias in the study. The use of propofol, to maintain anaesthesia, may have led
276 this error to be more frequent compared to the use of a volatile anaesthetic agent, due to the lack of
277 monitoring of anaesthetic concentration.

278 No subjects enrolled in this study required rescue analgesia during the postoperative
279 observation period (maximum 8 hours). The limited observation time of our study did not allow a
280 true comparison between the analgesic properties of EA and HIA. The motor block resolution time
281 was faster in the HIA group, with a significant difference at 3 hours ($p=0.004$) and 4 hours ($p=$
282 0.045) after the LA injection even if no difference was found at 5 hours. This finding is in

283 accordance with what is reported in humans (Atef et al. 2010). Various factors influence the offset
284 of motor block produced by neuraxial techniques. A possible explanation of the slower motor block
285 recovery produced by EA can be the much higher dose of LA used in comparison with spinal
286 anaesthesia and the persistence of the LA within the vertebral canal absorbed by the epidural fat,
287 which acts as a reservoir for the drug (Reina et al. 2009). Hyperbaric intrathecal anaesthesia is the
288 most commonly used neuraxial technique in day surgery procedures in human patients. It has been
289 proven to provide recovery and discharge times comparable with short-acting general anaesthetics
290 such as propofol, desflurane and sevoflurane (Ben-David et al. 2001; Lennox et al. 2002).
291 Continuous, and not single-shot, epidural anaesthesia using short-acting LAs has also successfully
292 been used in day surgery procedures; however, considering the technical difficulties, the longer
293 preparation time and the onset time of the nerve block, this technique is not normally used (Michael
294 et al. 2003).

295 There were no significant differences in perioperative side effects between the HIA and EA
296 groups. Hypotension was the most frequent effect in both groups and was generally resolved by
297 fluid bolus administration, a lighter plane of anaesthesia and, in some cases, the use of vasoactive
298 drugs in both groups.

299 No sign of dural puncture was found in the over 40 epidural punctures successfully
300 performed in this study. Iff and Moens (2010) reported an incidence of 4% of subarachnoid
301 puncture over 98 epidural blocks. In two different papers by Bosmans and colleagues a fatality
302 (Bosmans et al. 2011) and an incidence of Horner's syndrome (Bosmans et al. 2009) potentially
303 caused by an accidental intrathecal injection were reported. In the authors' opinion, the use of a
304 Quincke spinal needle to perform EA may play a role in increasing the risk of a dural puncture. The
305 use of a Tuohy needle makes it easier to perceive a LOR to needle advancement and a dural
306 puncture less probable due to its blunt curved cutting tip (Hadzic [19972007](#)).

307 This study shows that the use of HIA in dogs undergoing common pelvic limb surgeries
308 provides a lower PFR, a lower risk of intraoperative cardiovascular response to surgical stimulation

309 for at least 80 minutes, and earlier motor block resolution, with a similar incidence of side effects as
310 the single-shot EA technique. Epidural and intrathecal anaesthesia have different characteristics
311 and, consequently, different indications for use; however, in this study, HIA has been shown to
312 provide some advantages over EA in dogs undergoing commonly performed pelvic limb
313 orthopaedic surgery in a day-hospital regime.

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Figure legends

Fig. 1 The CONSORT diagram on patient recruitment, inclusion, and exclusion is shown. EA (Epidural anaesthesia), HIA (Hyperbaric intrathecal anaesthesia), iRA (intraoperative rescue analgesia).

388 **Fig. 2.** The time-to-event probability of intraoperative rescue analgesia (iRA) was analysed using
389 Kaplan-Meier survival analysis for epidural anaesthesia (EA) and hyperbaric intrathecal anaesthesia
390 (HIA). The iRA survival probability was not significantly different between groups for the entire
391 study period ($p = 0.0536$ log-rank test, Hazard ratio 0.418, 95% CI 0.168 to 1.041), but the EA
392 group required more iRA during the first 80 minutes ($P=0.045$ log-rank test, Hazard ratio 0.355,
393 95% CI 0.127 to 0.991). "Censored" refers to subjects for which no events (iRA) were observed
394 during surgery period. Censored data in the graph were marked with a small vertical line.

395 **Table 1.**

396 Types of surgical procedures and anaesthesia (EA: epidural anaesthesia and HIA: intrathecal
397 anaesthesia with hyperbaric solution) included in the study. TPLO (Tibial Plate Levelling
398 Osteotomy); ESCCL (Extracapsular Stabilization of Cranial Cruciate Ligament); DPO (Double
399 Pelvic Osteotomy); FHNO (Femoral Head and Neck Osteotomy).

	EA (26)	HIA (30)
TPLO	10	9
ESCCL	5	7
FHNO	2	3
Luxating patella	4	3
DPO	2	2
Femoral fracture	1	2
Others	2	4

400 **Table 2.**

401 Demographic data of dogs that met the inclusion criteria for intraoperative and postoperative
 402 evaluation. EA (epidural anaesthesia) and HIA (intrathecal anaesthesia with hyperbaric solution);
 403 BM: body mass; SCL: spine cord length; ASA, American Society of Anesthesiologists.

	EA (n =26)	HIA (n =30)
Type of breed	Mongrel (16), Labrador (2), Yorkshire (2), Beagle (1), West Highland White Terrier (1), Collie (1) German Shorthaired Pointer (1), Scottish Shepard (1), Bernese Mountain Dog (1)	Mongrel (12), Labrador (5), Yorkshire (4), Cane Corso (2), American Staffordshire Bull Terrier (1), Poodle (2), Boxer (1), German Shepard (1), Italian Spitz (1), Neapolitan Mastiff (1)
Median (range) age (Years)	6 (0.5-13)	7 (0.6-14)
Median (range) BM (kg)	8.2 (4-33)	13 (1.8-56)
Median (range) SCL (cm)	45 (33-76)	47.5 (30-85)
ASA	ASA I (22) ASA II (4)	ASA I (26) ASA II (4)

404 **Table 3.**

405 Procedural data according to EA (epidural anaesthesia) or HIA (intrathecal anaesthesia with
 406 hyperbaric solution); BM: body mass; SCL: spine cord length; Ii, intrathecal injection.

	EA	HIA	<i>p</i> -value
Median (range) bupivacaine dose related to BM (mg kg ⁻¹)	1	0.35 (0.26-0.85)	
Median (range) bupivacaine dose related to SCL (mg cm ⁻¹)	0.21 (0.11-0.48)	0.1 (0.18-0.05)	
Median (range) morphine dose	0.1	0.04 (0.02-0.16)	
Median (range) fentanyl bolus (µg kg ⁻¹ IV)	2 (1-3)	2 (1-3)	<i>p</i> > 0.05
Median (range) propofol induction bolus (mg kg ⁻¹ IV)	5 (3-6)	5 (3-6)	<i>p</i> > 0.05
Median (range) propofol dose consumption in the first hour (mg kg ⁻¹ IV)	25 (15-38)	23 (15-35)	<i>p</i> > 0.05
Median time (range) between Ii and beginning of surgery (min)	30 (25-50)	27.5 (15-39)	<i>p</i> > 0.05
Median time (range) between Ii and end of surgery (minutes)	88 (50-140)	94 (50-150)	<i>p</i> > 0.05

407 **Table 4.**

408 Resolution time of motor block. At 3 and 4 hours after local anaesthetic injection there is a
 409 statistical difference in the motor block resolution between the EA (epidural anaesthesia) and HIA
 410 (intrathecal anaesthesia with hyperbaric solution) groups.

Dogs able to walk	EA (26)	HIA (30)	<i>p</i> -value
At 3 h	0/17	8/20 (40%)	<i>p</i> = 0.004
At 4 h	4/17 (24%)	12/20 (50%)	<i>p</i> = 0.045
At 5 h	11/17 (65%)	17/20 (85%)	<i>p</i> > 0.05
At 8 h	17/17 (100%)	20/20 (100%)	<i>p</i> > 0.05

