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Long bone failure after intraosseous regional perfusion

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Keywords: horse; sepsis; antimicrobials; intraosseous regional perfusion; fracture

Summary

A horse that had sustained a penetrating injury to the proximal and medial aspect of the left fore MC3 was presented to the Charles Sturt University Veterinary Clinical Centre (CSUVCC). The horse had initially been treated with bandaging and systemic antimicrobials by the referring veterinarian but had failed to respond. Radiographs of the limb made at CSUVCC revealed bone lysis and a palisading new bone reaction on the medial splint bone that were thought to be characteristic of infection (**Fig 1**). As the associated soft tissue swelling extended above and below the carpus and therefore precluded intravenous regional perfusion (IVRP), a decision was made to use intraosseous regional perfusion (IORLP) to deliver antimicrobials to the limb. The horse was sedated with detomidine and butorphenol and an area of skin and subcutaneous tissue on the lateral diaphysis of MC3 was desensitised using an infiltration of 5 mL of lignocaine hydrochloride. A 4.5 mm bone portal was made in the lateral diaphysis of the left fore MC3 using the technique described by Scheuch *et al.* (2002). After a rubber tourniquet was applied to the limb above the carpus, one third of the calculated systemic dose of gentamicin sulphate was diluted with 30 mL of normal saline and injected slowly into the medullary cavity via a cannulated screw and a short i.v. extension tube. The tourniquet was removed after 30 minutes and the leg was supported with a padded bandage. The horse was maintained on systemic antimicrobials for 7 days and using the same portal, the IORLP procedure was repeated 3 times at 48-hour intervals.

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Approximately 13 weeks after the IORLP, the horse was re-presented to the CSUVCC for further assessment of lameness and a swelling centred over the LF cannon bone in the vicinity of the original IORLP access portal. Radiographs revealed a longitudinal fracture of MC3 (**Fig 2**) which appeared to have propagated from the drill hole and the swelling was identified as a bony callus. Fortunately, the horse had survived this fracture event and had undergone a healing process that stabilised the bone. At the time of writing, the horse is sound and the healing process has continued.

Key Points

- Intraosseous regional limb perfusion is a recognised method of delivering high concentrations of antimicrobials to infected tissues.
- Long bone fracture in the horse as a complication of this technique has not been reported.
- More work is needed to determine the safest site for making an IORLP access portal in MC3 and other equine bones.

Figure legends

Fig 1: DMPLO radiograph showing bone lysis (arrow head) and the palisading bone reaction on MC2 (block arrow).

Fig 2: LM radiograph showing the bone portal (block arrow) and the associated fracture lines (arrow heads).

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Summary

A five-year-old Australian Stock Horse gelding was initially referred to the Charles Sturt University Veterinary Clinical Centre (CSUVCC) for assessment of a penetrating laceration over the medial splint bone of the left foreleg. Clinical examination failed to reveal a communication with a synovial structure but radiographs showed a palisading bone reaction on the proximal aspect of the medial splint bone which was thought to be a characteristic of infection. Soft tissue swelling precluded access to a peripheral vein so a decision was made to use intraosseous regional limb perfusion. This was achieved through an access portal in the lateral diaphysis of MC3. Subsequently the horse developed lameness and further investigation revealed a non-displaced longitudinal fracture of MC3 propagating from the intraosseous regional perfusion access portal.

29 Introduction

30 Intraosseous (IORLP) and intravenous regional limb perfusion (IVRLP) have become popular
31 techniques for delivering high concentrations of antimicrobials to target tissues (Whitehair *et al.*
32 1992; Butt *et al.* 2001; Rubio-Martinez and Cruz 2006; Alkabes *et al.* 2011; Errico *et al.* 2012; Rubio-
33 Martinez *et al.* 2012). The hypothesis is that delivery of high concentrations of concentration-
34 dependant antimicrobials to infected tissues at or above the minimum inhibitory concentration
35 would be more likely to eliminate bacteria. It has been shown that regional limb perfusion in both
36 rabbits and humans will achieve high concentrations of antimicrobials even in poorly perfused and
37 chronically infected tissues (Finsterbush *et al.* 1970; Finsterbush and Weinberg 1972). The
38 techniques involve delivery of the perfusate (antimicrobial) directly into the limb through an
39 intravenous or intraosseous portal and distal to an occlusive tourniquet. Thus achieving a
40 concentration and pressure gradient across the intravascular and extravascular compartments
41 (Rubio-Martinez and Cruz 2006). Intraosseous regional limb perfusion is a more invasive technique
42 which involves the creation of a portal into the medullary cavity in the region of the septic site and
43 distal to the tourniquet. The technique is particularly useful when there is damage to and swelling of
44 soft tissues and access to a peripheral vein is challenging. After drilling a unicortical hole, a
45 cannulated, self-tapping bone screw is used as a conduit for the antimicrobial. The same hole can be
46 used for injection on consecutive days, thus avoiding damage to peripheral veins. In 2004, Mattson
47 *et al.* described an intraosseous technique for the delivery of gentamicin into the distal metacarpus
48 in standing, sedated horses. He used a 4.0 mm drill and a 5.5 mm tap for use with a 5.5 mm x 20 mm
49 cannulated bone screw. The portal was made in the distal metacarpus on the dorsolateral cortex 1
50 cm lateral to the common digital extensor tendon. From this work the authors concluded that IORLP
51 of gentamicin using the distal aspect of the limb should be considered for the treatment of
52 orthopaedic infections in horses. Keys *et al.* (2006) in an experimental study on cadaveric limbs
53 showed that after IORLP the pattern of distribution to the peripheral vessels was in a centrifugal
54 direction from the medullary cavity. However, they showed that this distribution included veins and
55 arteries and that these results supported the use of IORLP to deliver medications to the tissues of
56 the distal forelimb.

57 This report describes a potentially catastrophic complication of intraosseous regional limb perfusion
58 in a standing, sedated horse

59 Clinical History

In 2016, a horse that had sustained a penetrating injury to the proximal and medial aspect of the left fore MC3 was presented to the Charles Sturt University Veterinary Clinical Centre (CSUVCC). The horse had initially been treated with bandaging and systemic antimicrobials by the referring veterinarian, but had failed to respond. The horse was (AAEP) Grade 3/4 lame in the affected limb and palpation of the swelling over the medial splint bone elicited a marked pain response. Radiographs of the limb made at CSUVCC revealed bone lysis and a palisading new bone reaction on the medial splint bone that were thought to be characteristic of infection (**Fig 1**). As the associated soft tissue swelling extended above and below the carpus and therefore precluded intravenous regional perfusion (IVRLP), a decision was made to use IORLP to deliver antimicrobials to the limb. The horse was sedated with detomidine (Dozadine 0.01 mg/kg i.v.)^a and butorphanol (Butorgesic 0.02 mg/kg i.v.)^b and an area of skin and subcutaneous tissue on the lateral diaphysis of MC3 was desensitised using an infiltration of 5 mL of lignocaine hydrochloride (Lignocaine 20 20 mg/mL)^b. A 4.5 mm lateral bone portal was made in the diaphysis of the left fore MC3 using the technique described by Scheuch *et al.* (2002). After a rubber tourniquet was applied to the limb above the carpus, one third of the calculated systemic dose of gentamicin sulphate (Gentam 100 6.6 mg/kg bwt i.v.)^b was diluted with 30 mL of 0.9% sodium chloride solution and injected slowly into the medullary cavity via the 4.5 mm cannulated screw^d and a short i.v. extension tube (Discofix C)^c. The tourniquet was removed after 30 minutes and the leg was supported with a padded bandage. The horse was maintained on oxytetracycline^e (Engermycin 10 mg/kg i.v. q12h) and phenylbutazone^f (Bute Paste 2.2 mg/kg PO q12h) for 7 days and using the same portal, the IORLP procedure was repeated 3 times at 48-hour intervals. The skin incision for the IORLP was covered with a bandage and allowed to heal by second intention. The lameness gradually improved during the period of hospitalisation and the phenylbutazone was discontinued. Approximately 13 weeks after the IORLP, the horse was re-presented to the CSUVCC for further assessment of lameness and a swelling centred over the LF cannon bone in the vicinity of the drill hole. Radiographs revealed a longitudinal fracture of MC3 (**Fig 2**) which appeared to have propagated from the drill hole and the swelling was identified as a bony callus (**Fig 3**). Fortunately the horse had survived this fracture event and had undergone a healing process that stabilised the bone. At the time of writing, the horse is sound and the healing process has continued.

Discussion

91 This report describes a potentially catastrophic complication of IORLP. The horse developed a
92 fracture at the site of the IORLP at some time after the procedure. The only thing that alerted us to
93 the presence of a complication was that the owner reported a new swelling and lameness after the
94 initial treatment period. Although Mattson *et al.* (2004) alluded to the possibility of an iatrogenic
95 fracture occurring through an IORLP bone portal, to our knowledge such a long bone fracture in the
96 horse has not previously been reported.

97 In this case, for safety reasons and ease of access in the standing horse, the IORLP access hole was
98 made on the lateral side of the diaphysis of MC3. On reviewing the case, the question was asked if
99 choosing this particular site predisposed the MC3 to failure when placed under stress. In one study
100 that examined the effects of hole diameter on the torsional mechanical properties of isolated equine
101 cadaveric third metacarpal bones, it was found that the presence of lateral to medial 3/16 inch or
102 5/16 inch bicortical holes reduced all torsional structural properties (except stiffness) of the bone.
103 Failure torque and post yield properties were most dramatically reduced. The authors hypothesised
104 that these changes may prohibit affected horses from sensing pre-failure damage, enhancing the
105 likelihood of catastrophic failure when bone torsional load exceeds the yield point (Selzer *et al.*
106 1996).

107 The horse in this report was given free access to a small pasture after the IORLP procedure, so the
108 exercise was uncontrolled but limited in its intensity. In a treadmill exercise study, Davies (2005)
109 recorded the timing and distribution of surface strains from strain gauges distributed around the
110 periphery of the mid MC3 cortex of a 3-year-old Thoroughbred racehorse. That study showed that
111 peak compressive strains on the dorsal cortex of MC3 occurred in the first third of stance and were
112 probably related to deceleration of the limb as these strains were directly proportional to exercise
113 speed. Whereas, strains on the medial cortex were thought to be directly related to support of the
114 body weight. The strains on the lateral cortex occurred at variable times through stance and showed
115 a much larger variation than strains recorded on the other cortices. The lateral strains may be
116 associated with the maintenance of balance as well as the support of body weight. The lateral cortex
117 was not loaded significantly during exercise in a straight line on a flat treadmill surface in that study,
118 but exercise in circles significantly loads the lateral cortex of the inside limb in both compression and
119 tension (Davies and Merritt 2004). This inconsistent pattern of loading of the lateral cortex and the
120 increase in both tensile and compressive strains might help to explain the lateral condylar fractures
121 that occur in racehorses on the turns, and may have been significant in our case.

122 While these studies are helpful in explaining the strains imparted to MC3 in normal horses, more
123 work needs to be done to determine the safest site for making an IORP access portal in horses with
124 orthopaedic infections. As can be seen in Fig 3, the lateral cortex of MC3 in this case was much

narrower than the medial cortex which suggests that this horse may not have had sufficient residual strength in that region of bone to withstand sharp turns towards that limb following the surgery. It may be preferable to consider drilling into the medial cortex given that it is apparently more consistently loaded than the lateral cortex (Davies and Merritt 2004). Changes in bone density and loss of structural integrity associated with disease will have some effect on the outcome, but in most cases these will be unknowns. However, any evidence may help surgeons decide on the safest protocol, and it may also enable them to make better post-operative confinement and exercise plans.

Authors' declaration of interests

No conflicts of interest have been declared.

Ethical animal research

Research ethics committee oversight was not required. The owners gave their verbal consent to use the animal radiographs for educational purposes.

Source of funding

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Authorship

V. Slack-Smith and B. Hilbert undertook the clinical work. H.M.S. Davies provided expert biomechanical interpretation. All authors contributed to the production of the manuscript and approved the final version.

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Figure legends

Fig 1: DMPLO radiograph showing bone lysis (arrow head) and the palisading bone reaction on MC2 (block arrow).

Fig 2: LM radiograph showing the bone portal (block arrow) and the associated fracture lines (arrow heads).

Fig 3: DP radiograph showing the portal (block arrow) and the new bone reaction on MC3 (arrow head).



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