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1 **Recognition and Management of Pain in Cattle**

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6 **Abstract**

7 ATTITUDES towards pain and its control in farm animals have lagged behind those
8 in companion animal species. However, a considerable amount of work over the past
9 15 years has focused on the perception of pain in cattle based on objective and
10 subjective assessment by clinicians working with this species. A recent large-scale
11 survey of cattle practitioners revealed that over half of the respondents felt their
12 knowledge of pain and analgesia in cattle was inadequate or could be improved, and
13 the majority of these identified a lack of readily available information on the subject
14 as being a contributory factor. This article reviews current knowledge on pain
15 assessment in cattle in a clinical setting, and discusses some protocols for pain
16 management in specific conditions.

17 **Introduction**

18 Pain in humans has been described as “an unpleasant sensory and emotional
19 experience with actual or potential tissue damage”. It is reasonable to suppose that
20 animals experience pain in a similar way to humans because experimental work has
21 demonstrated that the neural pathways of pain sensation are similar in human and
22 other mammals. Application of the “precautionary principle” would also suggest that
23 this is the safest assumption unless strong experimental evidence proves otherwise.
24 Attitudes towards pain and its control in farm animals have lagged behind those in
25 companion animal species. However, a considerable amount of work in recent years

26 has focussed on perception of pain in cattle by clinicians working with the species,
27 and on its subjective and objective assessment.

28

29 In a recent large-scale survey of cattle practitioners (Huxley and Whay 2006), over
30 half of respondents felt that their knowledge of pain and analgesia in cattle was
31 inadequate or could be improved, and the majority of these identified a lack of readily
32 available information on the subject as a contributory factor. This article reviews
33 current knowledge on pain assessment in cattle in a clinical setting before discussing
34 methods to prevent and alleviate it.

35

36 **The Physiology of Pain**

37 Pain results from by chemical, mechanical or thermal stimulation of free nerve
38 endings containing nociceptors. Injury to cells in tissues causes release of
39 inflammatory mediators (e.g. prostaglandins, histamine and bradykinin), which
40 stimulate nociceptors in nearby nerve endings. This is an amplification process; a
41 stimulus affecting a relatively small number of nerve endings stimulates many more.
42 Impulses resulting from this stimulation are conducted via the ventrolateral part of the
43 spinal cord to the brainstem and thalamus. There is further amplification at this level
44 (centrally); this is known as “wind-up”. Conscious perception of pain is a result of
45 activation of certain areas of the cerebral cortex (via the thalamus). Theoretically, pain
46 is a central “experience” that occurs as a result of nociception in peripheral nerves.

47

48 Tissue injury results in acute pain, which stimulates muscular action to avoid the
49 noxious stimulus (either as a result of reflex limb flexion or via conscious
50 mechanisms) and causes sympathetic autonomic nervous system activation and a
51 heightened state of arousal. Increased sympathetic tone can become persistent if the
52 insult is prolonged or severe. In chronic pain, the presence of high levels of
53 inflammatory mediators around the site of injury and persistent activation of pain
54 fibre pathways in the spinal cord leads to a decrease in pain threshold, so that stimuli
55 are perceived as more painful than would be normal for the individual concerned.
56 This is known as hyperalgesia. Another phenomenon associated with chronic pain is
57 allodynia, whereby similar mechanisms lead to perception of normally non-painful
58 stimuli as painful. Prevention or modulation of hyperalgesia and allodynia is one of
59 the main objectives of analgesia. For example, a chronically lame cow may over time
60 perceive the lesion as more painful than it was initially (hyperalgesia) and perceive
61 pain in undamaged surrounding tissues on touch (allodynia).

62

63 As well as implications for welfare, pain is also significant in terms of disease
64 progression, potentially having a major effect on the physiological state of the animal
65 (Otto and Short 1998). This may interfere with wound healing.

66

67 **Assessment and Recognition of Pain in Cattle**

68 A large volume of research has been conducted in the past 15 years in the field of pain
69 assessment in ruminants. A number of methodologies have been employed
70 experimentally to assess or quantify levels of pain experienced by animals. These can
71 be broadly categorised as objective and subjective. Objective methods measure

72 physiological stress responses (e.g. plasma cortisol levels), changes in levels of
73 biochemical markers (e.g. acute phase proteins) or the incidence of clearly defined
74 behaviour patterns (e.g. vocalisation). Subjective methods are value judgements made
75 by the human observer. These will become more repeatable and reliable with
76 appropriate experience and training. Subjective pain assessment relies on the
77 evaluation of behaviour, posture and other cues. The degree of pain is then either
78 described using a verbal descriptor (e.g. mild, moderate, severe), assigned a numerical
79 value (e.g. zero to 10) or described using a visual analogue scale (e.g. placement of a
80 mark somewhere on a line between no pain and the worst pain imaginable).

81

82 In a practical situation, a variety of subjective indicators can be employed to assess
83 pain, and this should form part of a standard clinical examination. The following
84 indicators are useful in cattle:

- 85 • Decrease in movement/locomotion
- 86 • Decreased interaction with other animals in the group
- 87 • Decreased feed intake (e.g. “hollow” left flank caused by an empty rumen)
- 88 • Changes relevant to the source of the pain being experienced (e.g. altered
89 locomotion, flank watching or kicking, ear twitching)
- 90 • Level of mental activity/responsiveness (animals in severe pain often show reduced
91 responsiveness to stimuli)
- 92 • Changes in normal postures associated with pain (e.g. lateral recumbency, standing
93 motionless, drooping of the ears)

- 94 • Easily measurable indicators of physiological stress (e.g. increased heart rate,
95 increased pupil size, altered rate and depth of respiration, trembling)
- 96 • Bruxism (tooth grinding)
- 97 • Poor coat condition (e.g. rough, dusty or unkempt) caused by decreased grooming

98 As with all types of clinical examination, it is important to have a consistent approach
99 to pain assessment, and ensure that the same behavioural and physiological signs are
100 assessed in each animal.

101

102 It is important to remember that cattle are stoical by nature because, as a species, they
103 have been subject to a strong evolutionary pressure to mask pain and its implied
104 weakness from predators. As a result they often do not demonstrate appreciable
105 definite signs of pain until the stimulus is severe. Often, particularly in adult cattle,
106 unwillingness to move may be the predominant indicator. This means that the
107 precautionary principle should be applied, i.e. the clinician should err on the side of
108 treating or preventing pain, as the cost of unnecessary treatment is relatively less
109 severe than the cost of failing to manage animals that are suffering.

110

111 **Barriers to Treatment**

112 A survey by Huxley and Whay (2006) examined the reasons why practitioners tended
113 to under-use analgesia in cattle. Over 90% of respondents to the survey considered
114 that cattle benefited from analgesics as part of their treatment and that they recovered
115 faster if they were administered; however, two thirds of respondents considered that
116 the cost of analgesia was a major issue to their clients. Whilst the financial constraints

117 of the industry must always be considered, there are a number of reasons why they
118 need not always preclude effective analgesia:

119 • Many analgesic protocols (such as local anaesthetic techniques) are inexpensive to
120 perform. Local anaesthetic drugs are economical and volumes required are
121 generally low. Time spent in performing these techniques is also usually low, and
122 will decrease as the experience of the clinician grows.

123 • Financial benefits are often an unexpected outcome of analgesic treatment.
124 Increases in parameters such as growth rate after calf disbudding (Faulkner and
125 Weary 2000) and milk yield after lameness cases (O'Callaghan-Lowe and others
126 2004) have been reported after analgesic therapy was combined with standard
127 treatments. Whilst this increased performance may not cover the total cost of
128 analgesic treatment, in the majority of situations it will partially offset the cost.

129 • Prices of the most commonly used non-steroidal anti-inflammatory drugs
130 (NSAIDs) may fall in the future as more generic products become available.

131

132 Clinicians too often assume that farmers are unwilling to carry costs associated with
133 improvements to the welfare of their animals. In the case of many farm animal owners
134 (especially “hobby” farmers or owners of small herds), the client may be more
135 prepared to pay than the clinician realises. A recent survey of commercial UK cattle
136 farmers (with over 1,000 respondents) has demonstrated that for the majority of
137 owners the cost of analgesics remains a significant issue; however this was not true
138 for all respondents. When asked to state how much they would be prepared to pay for
139 analgesics during and following the treatment of a range of conditions and procedures,
140 the answers varied considerably. For all 13 conditions considered, a small minority of

141 respondents stated they would consider between £35 and £50 (the highest bracket) an
142 acceptable cost for analgesic treatment, but a more significant number were prepared
143 to meet a lower cost. For example, when considering surgical castration of calves,
144 32% of respondents stated they would pay between £5 and £10 and eight percent
145 stated £11 or more. When considering caesarean section surgery, 28% of respondents
146 stated they would pay £11 to £20, 13% stated £21 to £35 and 6% stated £36 to £50
147 and when considering disbudding, 21% stated they would pay £5 to £10 and 4%
148 stated £11 or more. This work suggests that for some owners cost is not the issue
149 practitioners may initially believe and therefore it is important to offer a variety of
150 costed analgesic treatment protocols for painful procedures and conditions.

151

152 One of the most noteworthy findings in the practitioner survey (Huxley and Whay
153 2006), was that respondents who did not use any analgesic agents during treatment
154 estimated significantly lower pain scores for the condition or procedure in question
155 (Table 1). This suggests that one of the key motivators for analgesic usage is the
156 attending clinician's own perceptions of the patient's suffering. Therefore, one of the
157 barriers for the provision of appropriate analgesia in cattle is, in some cases, an
158 unwillingness or inability to consider or identify the level of pain cattle are suffering.

159

160 **Misconceptions about Analgesia and Cattle**

161 A number of common misconceptions also emerged from the survey. These are
162 summarised and discussed below:

163 • **Age of the animal:** Young animals are often assumed to feel less pain than adults.

164 A good example of this is the lack of analgesia used in castrating calves and lambs

165 using rubber ring techniques. There is no evidence to show that young animals
166 perceive pain to a lesser extent than adults. In the authors' opinion, young animals
167 should be considered in exactly the same manner as adult animals.

168 • **Pain restricts movement which may be potentially damaging to the animal's**
169 **condition:** If movement is likely to be damaging to an animal's condition
170 appropriate analgesia should be provided and movement should be restricted by
171 penning the animal tightly, rather than relying on the animal's suffering.

172 • **Analgesia may mask deterioration in the condition of the animal:** This view is
173 likely to have stemmed from received wisdom in equine practice, where it has been
174 considered that use of potent anti-endotoxic drugs such as flunixin may hamper
175 monitoring of colic cases. Farm animals are rarely monitored on such a short-term
176 basis, and other clinical signs can be used in these species to monitor progression
177 of a disease.

178 • **Corticosteroids are effective analgesic agents:** Although corticosteroids are
179 potent anti-inflammatories and act on the same pathway as NSAIDs, at therapeutic
180 dose rates they are likely to produce less profound analgesia than NSAIDs.

181

182 **Techniques for Alleviating Pain**

183 An important concept in terms of alleviation of pain in cattle is that of pre-emptive
184 versus reactive analgesia. Where pain is predictable (e.g. surgical procedures), it is
185 preferable to provide pre-emptive analgesia. By ensuring that effective analgesia is in
186 place before the onset of pain, phenomena such as wind-up, hyperalgesia and
187 allodynia can be reduced or prevented. Obviously this is not always possible, but
188 provision of analgesia as soon as possible after the onset of pain will minimise these

189 effects. It should always be remembered that pre-emptive analgesia is likely to be
190 more effective than reactive analgesia.

191

192 Multimodal analgesia is also an important concept. It is well recognised in human and
193 companion animal medicine that the most effective analgesia is provided by using a
194 combination of agents that act on different pathways, but this is an often-neglected
195 strategy in relation to cattle, where it could frequently be gainfully employed (e.g. the
196 use of an epidural containing local anaesthetic and xylazine, combined with systemic
197 NSAID to provide analgesia for dystocia).

198

199 There are several routes that can be used to provide analgesia to cattle. Systemic
200 treatment involves parenteral administration of systemically active analgesic agents,
201 while local techniques such as epidural anaesthesia, local nerve blocks and
202 intravenous regional anaesthesia provide analgesia to specific areas.

203

204 **Systemic Analgesic Techniques**

205 The main groups of analgesic drugs available for use in animals are NSAIDs, α_2 -
206 agonists and opioids. Licensing is a major issue in prescribing for food producing
207 animals (see Box Figure 1), and this places a major restriction on the agents that can
208 be used in cattle. A variety of NSAIDs are licensed, along with the α_2 -agonist
209 xylazine.

210 • **NSAIDs:** This class of drugs works by inhibition of inflammatory mediators (see
211 Box Figure 2). They provide effective analgesia for mild to moderate pain, and are

212 administered by a variety of routes (see Table 2). They also have anti-endotoxic
213 effects, which provide major benefits in terms of morbidity and mortality in some
214 disease states. Duration of activity is generally in the range 24-72 hours per dose.
215 Some products are licensed for repeated administration (up to a maximum of five
216 days of treatment), but have been used for longer periods with few reports of side-
217 effects (although abomasal ulceration has been reported anecdotally).

- 218 • **α_2 -agonists:** These agents work by activation of α_2 -adrenoreceptors in the central
219 and peripheral autonomic nervous system. These have a negative effect on
220 sympathetic activity and release of noradrenaline, leading to sedation and
221 analgesia. They can provide deep sedation and effective analgesia for moderate
222 pain in cattle. As the sedative and analgesic effects of this class of drug go
223 together, they are more useful during some types of surgery and are not used for
224 provision of longer-term analgesia. Xylazine is the only licensed drug in this class.
- 225 • **Opioids:** Opioids are very potent analgesics, and are an important component of
226 multimodal analgesia protocols in other species. However, under current
227 legislation, no opioid agents are available for use in cattle.

228

229 **Regional and Local Techniques**

230 The main techniques for providing local analgesia are epidural analgesia, intravenous
231 regional anaesthesia and nerve blocks.

- 232 • **Epidural analgesia:** This technique involves injection of analgesic agents into the
233 epidural space, to provide desensitisation of nerves leaving the spinal cord.
234 Although outside the scope of this article, a full description of the technique, which
235 is quick and straightforward to perform in cattle, is provided by Holden (1998).

236 Low-volume (4-6ml of injectate for an adult bovine) epidural anaesthesia is most
237 commonly performed, providing anaesthesia of the genital tract, rectum and
238 perinaeal area and abolition of tenesmus. High volume (up to 100ml per adult
239 bovine) techniques are also described, and may be used to provide anaesthesia for
240 the entire abdomen. High volume techniques will involve loss of motor control to
241 the hindlimbs, so will result in the patient becoming recumbent. Local anaesthetic
242 is the most commonly used agent, but xylazine has also been extensively used.
243 Xylazine provides a longer duration of action compared to local anaesthetic alone,
244 and the two agents are often used in combination (Grubb and others 2002). A
245 number of dosage regimes are described, including a xylazine dosage of 0.05mg/kg
246 (1.25ml of 2% xylazine per 500kg) with the remainder of the injectate made up of
247 local anaesthetic. This use of xylazine is not licensed.

248 • **Intravenous regional anaesthesia (IVRA):** This is another quick and easy
249 technique (see Box Figure 3 for description). It provides desensitisation of the limb
250 distal to the tourniquet, and so is very useful for painful procedures in the foot
251 (both foot surgery and treatment of severe claw horn lesions). It should be
252 remembered that the effects of the IVRA will quickly wear off once the tourniquet
253 is released, and no ongoing analgesia is provided. It is therefore usually advisable
254 to use NSAIDs in combination, to provide a longer duration of effective pain relief.
255 This use of local anaesthetic is off-licence.

256 • **Local nerve blocks:** A number of nerve block techniques are described in cattle.
257 Again, specific description of the techniques is outside the scope of this article, but
258 have been described previously (Edwards 2001). These techniques are summarised
259 in Table 3. Local anaesthetics (procaine is now the only licensed product in food
260 producing animals) are the most commonly used agents, providing 30-90 minutes

261 of effective anaesthesia. Local anaesthetic/xylazine combinations may also be used
262 for these techniques, and are thought to provide an extended duration of analgesia.
263 Again, this is off-licence, and there is little research evaluating the combination. It
264 has been suggested that α_2 -agonists are unlikely to have local effects and that any
265 clinical differences seen may be due to systemic absorption. With this in mind, the
266 possibility of sedation as a side-effect should be considered.

267

268 Ring blocks are a method for blocking distal appendages (e.g. teats and distal
269 limbs). Local anaesthetic is introduced at various points and depths around the
270 circumference of the appendage in order to block nerve supply distal to the
271 location of injection. These require multiple injections, and are generally less
272 effective. In addition, as procaine with adrenaline is the only licensed product, the
273 potential vasoconstrictive effect of infiltrating adrenaline around a small
274 appendage (e.g. a teat) should also be considered, as necrosis may result.
275 Infiltration of local anaesthetic around the area to be desensitised is also useful in
276 some situations.

277

278

279 **Suggested Standard Operating Procedures for Management of** 280 **Pain in Specific Situations**

281 **Standing Flank Laparotomy in Adult Cattle**

- 282 • Systemic NSAID before the start of surgery.
- 283 • Systemic xylazine could be used, but extreme care would be needed with dose to
284 ensure the animal does not become recumbent. May be useful to provide extra

285 short-term analgesia if a very painful procedure is anticipated. May be necessary in
286 very fractious patients.

287 • Paravertebral nerve blocks to provide effective anaesthesia of the flank area using
288 procaine.

289 • Epidural anaesthesia may be used in the case of a caesarean section, to abolish
290 tenesmus that may hinder surgery.

291

292 **Castration and Disbudding of Calves**

293 • Cornual nerve block using local anaesthetic (possibly in combination with low
294 dose perineural xylazine in some cases - although this use of xylazine is off-
295 licence, the consequences of which would have to be explained to the owner).

296 • Local infiltration of local anaesthetic in the skin of the distal scrotum (surgical
297 castration) and over the neck of the scrotum to provide analgesia to the spermatic
298 cord (surgical and burdizzo castration). Injection of local anaesthetic into the testes
299 themselves may or may not be used.

300 • Where economically acceptable, pre-emptive use of NSAIDs is a desirable
301 addition to the protocol. Several researchers have found welfare benefits as a result
302 (Earley and Crowe 2002; Ting and others 2003). Clinicians too often fail to offer
303 this option to clients, who may well be happy to the relatively small cost of
304 extended analgesia.

305

306 **Foot surgery/ treatment of severe claw horn lesions e.g. severe sole ulcers**

307 • Systemic NSAID before start of treatment.

308 • IVRA for short-term anaesthesia of the foot.

309

310 **Other potentially painful conditions where NSAID use should be**
311 **considered**

312

313 • Joint ill and navel ill in calves: Septic arthritis in particular is considered to be an
314 extremely painful condition in humans and companion animals, but analgesia is too
315 often neglected in cattle.

316 • Mastitis: NSAIDs should be considered for use in all cases of mastitis involving
317 udder or systemic signs (as opposed to mastitis where signs are restricted to milk
318 changes) (Milne and others 2003).

319 • Lameness: In addition to foot surgery and radical treatment of severe claw horn
320 lesions, benefits are also seen as a result of NSAID use in less severe lameness
321 cases (O'Callaghan-Lowe and others 2004).

322 • Dystocia: NSAID use is relatively common following dystocia. It is worth
323 considering providing analgesia to the calf as well as the dam.

324 • Uveitis and keratoconjunctivitis: These are both relatively common ocular
325 conditions in cattle (“silage eye” and “New Forest eye”), and the underlying
326 pathology (uveitis and corneal ulceration respectively) is considered to produce
327 severe pain in humans and companion animals. As well as providing analgesia,
328 NSAIDs may increase speed of response to treatment by decreasing inflammation
329 in cases of uveitis.

330

331 **Conclusions**

332 Despite an increase in research into and awareness of pain in cattle over the last
333 fifteen years, management of painful conditions in cattle is still too rarely considered
334 in practice. This article provides information on the methods by which analgesia may
335 be provided, as well as suggesting standard protocols for pain management in specific
336 conditions.

337

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386

387 Box figure 1 – place near reference in main text (line 193)

388 **Licensing of Veterinary Medicines**

389 Licensing issues have a major effect on product choices in farm animal medicine. The
390 Veterinary Medicines Regulations (2005) provide for administration of products
391 outside the terms of the product’s marketing authorisation under the prescribing
392 “cascade”. However, they state that “any pharmacologically active substances
393 included in a medicinal product administered to a food-producing animal under the
394 cascade must be listed in Annex I, II or III to Council Regulation (EEC) No.
395 2377/90.” Annex I lists substances for which a definitive maximum residue limit
396 (MRL) has been established. These are generally found in products which have an
397 authorisation for use in food producing animals. Annex II lists substances which,
398 following initial evaluation by the European Medicines Agency (EMA), were
399 deemed to pose sufficiently little risk to public health not to necessitate the
400 determination of a MRL. Annex III lists substances which are undergoing MRL
401 determination at the current time and have been given provisional MRLs as there are
402 considered to be no outstanding ongoing safety issues.

403

404 In practical terms, this means that only pharmacologically active substances listed in
405 Annexes I, II and III can be used in veterinary medicinal products for use in food
406 production animals. The only potentially useful compounds listed in Annex II for food
407 producing animals are ketamine and thiopentone (butorphanol, isoflurane and
408 lidocaine are also listed but the listing is restricted to equidae only). This restricts
409 drugs for use as analgesics to a variety of NSAIDs, procaine and xylazine (with
410 ketamine and thiopentone the only options for general anaesthesia). Further and

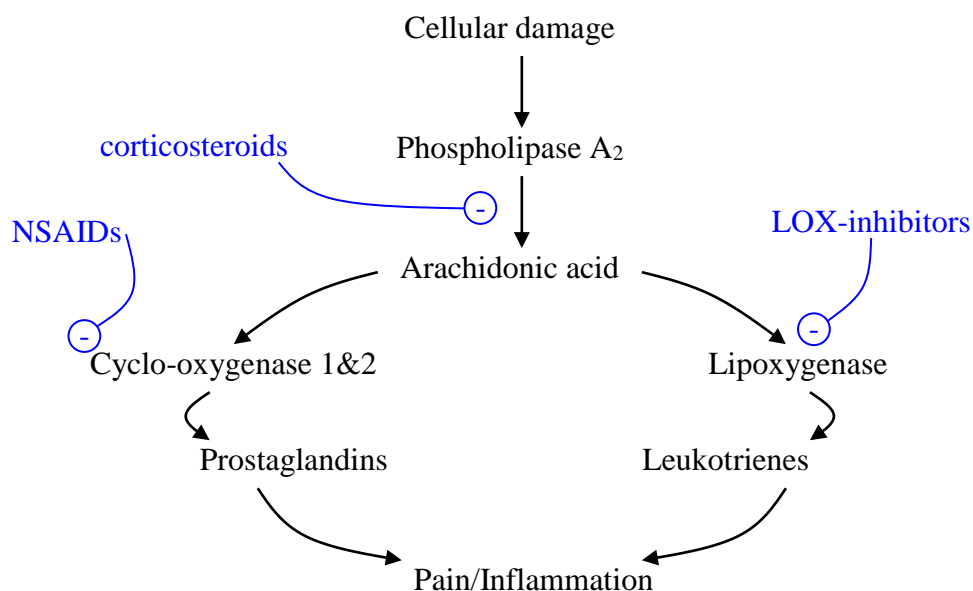
411 frequently updated information is available from

412 <http://www.emea.europa.eu/index/indexv1.htm>.

413

414 Box figure 2 – place near reference in text (line 197)

415 Mode of Action of NSAIDs



416

417 A simplified mode of action for NSAIDs is given in the diagram above. Inhibition of
418 cyclo-oxygenase (COX) enzymes decreases prostaglandin synthesis, thereby
419 decreasing pain and inflammation. As well as mediating inflammation, some COX-1
420 enzymes have alternative “housekeeping” functions and are induced in the absence of
421 cellular damage. Their roles include gastroprotection and maintenance of renal
422 bloodflow. Inhibition of these enzymes can bring about the side-effects of NSAIDs,
423 notably gastric ulceration and renal disease. These are not well recognised in farm
424 animals, although there have been anecdotal reports of abomasal ulceration in calves
425 after treatment with NSAIDs. One possible reason for the apparent lack of side effects
426 is the short term nature of the vast majority of NSAID use in these species. NSAIDs
427 that preferentially inhibit COX-2 enzymes to a greater degree than COX-1 may be
428 less likely to produce side effects.

429

430 It is clear from the diagram that production of leukotrienes by induction of the
431 enzyme lipoxygenase (LOX) is also an important inflammatory pathway. LOX
432 inhibitors are available in human and companion animal medicine (e.g. tepoxalin), but
433 as yet none are licensed for farm animals. However, there is evidence that tolfenamic
434 acid has some inhibitory activity against LOX. Some NSAIDs are also thought to have
435 a direct inhibitory action on leukotrienes.

436

437 A third COX enzyme has been identified (COX-3). Inhibition of this enzyme is
438 thought to be responsible for some of the activity of paracetamol (a NSAID-like agent
439 which is considered to have relatively little COX-1 and -2 inhibition). While
440 paracetamol is not clinically relevant to cattle, the activity of carprofen may also be
441 partly due to COX-3 inhibition. Other factors, such as how effectively the agent
442 crosses the blood-brain barrier, may also be significant in determining the
443 effectiveness of a NSAID.

444

445 Box figure 3 – place near reference in text (line 235)

446 **Intravenous Regional Anaesthesia for the Bovine Hindlimb: A**

447 **Standard Operating Procedure**

448

449 IVRA is a quick and simple technique to perform, and is underused in the treatment of

450 lame cows.

451 *Equipment required:*

- 452 • Appropriate handling facilities (i.e. foot trimming crush)
- 453 • Clippers/scissors
- 454 • Chlorhexidine surgical scrub
- 455 • Surgical spirit
- 456 • Tourniquet (a bicycle tyre inner tube is a good choice)
- 457 • 20-30ml local anaesthetic in syringe (procaine is now the only legal option),
- 458 depending on the size of the animal
- 459 • 18 gauge, 1.5 inch needle

460 *Procedure:*

- 461 • The animal is restrained in the crush, with the affected limb raised. The procedure
- 462 is easier to perform if the limb is not tied to the upright of the crush (as this will get
- 463 in the way of the injection site).
- 464 • The dorsolateral aspect of the metatarsus is clipped and surgically prepared.

465 • A tourniquet is applied to the limb, either below or above the hock. If the
466 tourniquet is above the hock, rolls of bandage or similar may be required to fill the
467 spaces either side of the gastrocnemius tendon. For this reason, the authors prefer
468 to apply it below the hock. The tourniquet must be applied sufficiently tightly and
469 secured.

470 • The lateral saphenous vein is palpated running directly up the dorsolateral aspect of
471 the metatarsus. The needle is then placed in the vein (directed distally), with the
472 entire length of the needle in the lumen of the vessel. A good needle placement
473 increases stability of vascular access while the local anaesthetic is injected.

474 • Blood is allowed to drain through the needle until the pressure drops so that blood
475 is dripping rather than running out of the hub.

476 • The syringe is connected and the local anaesthetic slowly injected.

477 • After five to ten minutes, desensitisation of the foot can be checked by pricking the
478 skin of the interdigital space with a sterile needle.

479 • When the procedure is finished, the tourniquet should be removed gradually (to
480 prevent the theoretical possibility of a bolus of local anaesthetic entering the
481 circulation). This is more important if the procedure has been very short.

482

483 The procedure is easily adapted to use in a forelimb (although restraint can be more
484 difficult in this situation, and it is worth considering casting the animal).

485

486 Box figure (no reference) – place anywhere in second half of text (preferably around
487 “Techniques for alleviating pain” section, lines 168-264)

488 **General Anaesthesia in Farm Animals**

489 In many respects, general anaesthesia can be thought of as the “gold standard” in
490 terms of pain management. However, it is important to remember that induction of
491 and recovery from general anaesthesia are stressful processes and that general
492 anaesthesia only provides pain relief for the duration of the anaesthetic. Some agents
493 have very poor analgesic properties, and multimodal pain relief should be employed.

494

495 Detailed descriptions of anaesthetic techniques are outside the scope of this article,
496 but general anaesthesia (either in the field or in a hospital setting) is a useful
497 procedure, particularly in young animals (the weight of the gastrointestinal tract
498 makes it more dangerous in adults). Licensing restrictions affect which products may
499 be used, but anaesthesia may be induced using a xylazine and ketamine combination,
500 and maintained with incremental doses of ketamine. A side-benefit of the use of
501 ketamine is that, as an NMDA antagonist, it is thought to interrupt central pain
502 amplification processes (wind-up). Endotracheal intubation (with or without oxygen
503 supplementation) is recommended in all cases, even though volatile agents cannot be
504 used for maintenance. Obviously, this is only suitable for shorter procedures.
505 Isoflurane could be used for maintenance to make longer procedures practicable, but
506 under current licensing rules this is not allowed.

507

508 Box figure (no reference) – place anywhere in first half of text (preferably near
509 “Barriers to treatment” and “Misconceptions about analgesia and cattle” sections,
510 lines 97-167)

511 **Legal Aspects of Analgesia**

512 The major legislation relevant specifically to this area is summarised below:

513 • **Protection of Animals (Anaesthetics) Act 1954** (as amended) states that
514 anaesthetic must be used for “any operation, with or without the use of
515 instruments, which involves interference with the sensitive tissues or the bone
516 structure of an animal” with the exception of injection or extraction by means of a
517 hollow needle. Some specific exclusions apply (e.g. life-saving or emergency first
518 aid treatment or minor procedures customarily performed without anaesthetic).

519 With regard to routine husbandry procedures in calves, anaesthetic must be used
520 when:

- 521 ○ Castrating calves over two months of age
- 522 ○ Disbudding or dehorning cattle of any age (with the exception of
523 chemical cautery, which is only permitted during the first week of life)
- 524 ○ Removing supernumerary teats from calves of over three months of
525 age

526 • More generally, the **Agriculture (Miscellaneous Provisions) Act 1968** states that
527 it is an offence to cause unnecessary pain or unnecessary distress to any livestock
528 on agricultural land.

529 • Similarly, the **Welfare of Farmed Animals (England) Regulations 2000** state
530 that owners and keepers of animals shall take all reasonable steps:

- 531 ○ To ensure the welfare of the animals under their care; and
- 532 ○ To ensure that the animals are not caused any unnecessary pain,
- 533 suffering or injury.
- 534 • The most relevant issues raised by the new **Animal Welfare Act 2006** are the
- 535 requirement for the person responsible for an individual animal to comply with
- 536 “good practice” in order satisfy the animals need to be “protected from pain”, and
- 537 the concept that duty of care passes from the owner/keeper to the veterinary
- 538 surgeon during treatment of the animal.
- 539

540 Box figure (no reference) – place anywhere in second half of text, preferably near to
541 “Techniques for alleviating pain” section (lines 168-264)

542 **Provision of Long Term Analgesia to Farm Animals**

543 Providing long-term pain management to farm animals with chronically painful
544 conditions is currently difficult. This is largely due to licensing restrictions and cost.
545 In the past, oral phenylbutazone has been used for this purpose, but use of
546 phenylbutazone in food producing species is now illegal.

547

548 The alternative approach is the use of repeated doses of injectable NSAID. This is off-
549 licence beyond five days, although it could be justified under the cascade system (as
550 authorised NSAIDs all have established MRL values, the standard withdrawal periods
551 of seven days milk withdrawal and 28 days meat withdrawal would apply). The safety
552 of long-term NSAID treatment in farm animals has not been extensively researched,
553 although there were anecdotal reports of long term phenylbutazone use being well
554 tolerated. The major barrier to long-term use of parenteral NSAID is cost: ketoprofen
555 given daily would cost approximately £1.20 - £1.70/100kg/day, while meloxicam
556 given every three days would cost £0.57/100kg/day (at list price). This level of
557 expenditure may only be justified in a limited number of cases, so use of NSAIDs is
558 usually restricted to coverage of episodes of acute pain. Euthanasia should always be
559 seriously considered in cases where an animal is likely to experience long-term pain.

560

561 Table 1 – place near reference in text (line 141)

562

563

564 **Caption:** A recent survey of UK cattle practitioners with over 500 respondents asked
 565 clinicians to judge the severity of pain associated with a range of procedure and
 566 conditions on a ten point scale (1 = No pain at all; 10 = The worst pain imaginable).
 567 The results are outlined in the table below. The median pain score is the score
 568 assigned by the middle clinician if all the scores are arranged in ascending order and
 569 the modal pain score was the most frequently given answer.

570

571

	Median		Modal	
	Pain	Range	Range	Pain
	Score	- Min	- Max	Score
Treatment of a sole ulcer	6	1	10	7
Claw amputation	10	2	10	10
Caesarean section	9	1	10	10
Adult Cattle Procedures Dystocia ¹	7	2	10	8
De-horning ²	8	2	10	10
Debriding a digital dermatitis lesion	6	1	10	5
LDA surgery	9	2	10	10

Uveitis	6	1	10	7
Fracture of tuber coxae	7	2	10	8
LDA	3	1	10	3
Digital dermatitis	6	2	10	5
Adult Cattle Conditions Acute Metritis	4	1	10	3
Swollen hock	5	1	10	5
Hock with hair loss	3	1	10	2
Acute toxic <i>Escherichia coli</i> mastitis	7	1	10	7
Mastitis (clots in milk only)	3	1	10	2
Neck calluses	2	1	7	2

	White line disease with Sub-				
	sole abscess	7	1	10	7
<hr/>					
Calf Procedures	Calf castration (Surgical)	6	2	10	5
	Calf castration (Rubber ring)	6	1	10	5
	Calf castration (Burdizzo)	7	2	10	8
	Umbilical hernia surgery	8	2	10	10
	Disbudding	7	2	10	8
<hr/>					
Calf Conditions	Distal limb fracture	8	2	10	8
	Following dystocia ¹	4	1	10	3
	Umbilical abscess	5	1	10	4
	Joint ill	7	1	10	8
	Pneumonia	6	1	10	5

572

Footnote: Respondents were asked to estimate the severity of pain assuming *NO* analgesic drugs were administered

573 ¹Fetal-maternal disproportion requiring traction alone

574 ²Horns >8cm/3"

575 Reproduced from Huxley and Whay (2006).

576

577

Place near to reference in text (line 198)

Table 2: Costs of commonly used drugs and techniques.

	Products	Licensed route(s)	Milk/meat withdrawal	Dose rate	Cost /ml	Cost /100kg	Licensed indications
Systemic							
NSAIDs							
Carprofen	Rimadyl™ (Pfizer)	i/v, s/c	n-1/21d	1ml/35kg	121p	£3.44 ²	Respiratory disease
Flunixin	Binixin™ (Bayer) Cronyxin™ (Bimeda) Finadyne™ (Schering Plough) Meflosyl 5% (Fort Dodge)	i/v	12-36h/ 5-8d	2ml/45kg	12- 42p	51p – £1.86	Respiratory disease, mastitis
Ketoprofen	Comforion™ (Janssen) Ketofen™ (Merial)	i/v, i/m	0/1-4d	1ml/33kg	39- 61p	£1.18- £1.85	Parturient paresis, respiratory disease, mastitis, udder oedema ¹
Meloxicam	Metacam™ (Boehringer Ingelheim)	i/v, s/c	5d/15d	2.5ml/100kg	75p	£1.88 ²	Respiratory disease, mastitis, diarrhoea in youngstock
Tolfenamic acid	Tolfine™ (Vetoquinol)	i/v, s/c (s/c limited)	24h/3-7d	1ml/10kg or 1ml/20kg ³	29p	£1.44 - £2.87	Mastitis, respiratory disease
α₂-agonists							
Xylazine ⁴	Chanazine™ 2% (Chanelle) Rompun™ 2% (Bayer) Sedaxylan™ (CEVA) Virbaxyl™ 2% (Virbac) Xylacare™ 2% (Animalcare) Xylapan™ (Vetoquinol)	i/v (Sedaxylan only), i/m	0d-24h, some n-1 / 1d – 14d	0.15 – 1.5ml /100kg	91p - £1.39	18p - £2.09	Sedation, muscle relaxation and analgesia. See data sheets for specific indications.
Local techniques							
Procaine	Willcain™ (Arnolds)		0/0	Depends on	3.5p/		

¹ Indications vary between the two licensed products.

² Meloxicam and carprofen may have longer durations of action than the other NSAIDs.

³ Dose rate for tolfenamic acid varies with indication

⁴ Xylazine should be used with caution in potentially pregnant animals – see data sheet for further information.

			technique:	ml		
			epidural ⁵		18p	
			cornual		35p	
			paravertebral		£2.80	
			IVRA ⁵		88p	

i/v = intravenous, i/m = intramuscular, s/c = subcutaneous

n-l = not licensed for use in lactating cattle

This is not a comprehensive list – it is restricted to those products listed in the NOAH data sheets compendium. Other NSAIDs for animal use are available. Drug costs are given at list price. Costs and dose rates are intended as guides only – the product datasheet should always be consulted before administering a medicine. The authors accept no liability for costs arising due to errors in this material.

⁵ Use of procaine by this route is off-licence.

Place near to reference in text (line 245).

Table 3: Nerve blocks commonly used in cattle.

Nerve block	Area of analgesia	Notes
Paravertebral	Flank	Quick and simple way to provide anaesthesia for flank surgery
Cornual	Horn and surrounding skin in calves	Less effective in adults
Retrobulbar	Eye and adnexa	May result in damage to adnexal structures, usually reserved for enucleation
Peterson	Eye and adnexa except eyelids	Less destructive than retrobulbar, need to anaesthetise eyelids separately for enucleation.
Auriculopalpebral	Eyelids (<i>motor function only</i>)	Provides paralysis but not desensitisation of eyelids
Common peroneal and tibial	Hindlimb distal to tarsus	A good alternative to IVRA, although technically more difficult.