

1 **Unravelling the temporal association between lameness and body condition score in dairy**
2 **cattle using a multistate modelling approach**

3 P Y Lim ^{a,b}, J N Huxley^a, J A Willshire^c, M J Green^a, A R Othman^b, J Kaler^{a*}

4 1. PohYing Lim ^{a,b},

5 2. Jonathan N Huxley^a,

6 3. James A Willshire^c,

7 4. Martin J Green^a,

8 5. Abdul Rahman Othman^b,

9 6. Jasmeet Kaler^{a*}

10 ^aSchool of Veterinary Medicine and Science, University of Nottingham, Sutton Bonington
11 Campus, Sutton Bonington, Leicestershire, LE12 5RD, United Kingdom.

12 ^bUniversiti Sains Malaysia, 11800, PulauPinang, Malaysia.

13 ^c Endell Veterinary Group, 49 Endless Street, Salisbury, Wilts. SP1 3UH.

14

15 *Corresponding author: phone:+441159516564; email address: jasmeet.kaler@nottingham.ac.uk

16 **Complete correspondence address including email address to which the proofs should be**

17 **sent:** Dr Jasmeet,Kaler, School of Veterinary Medicine and Science, University of Nottingham,

18 Sutton Bonington Campus, Sutton Bonington, Leicestershire, LE12 5RD, United Kingdom,

19 jasmeet.kaler@nottingham.ac.uk

20 Email addresses:

21 PYL: pohyinglim@gmail.com

22 JNH: jon.huxley@nottingham.ac.uk

23 JAW: jim@endellfarmvets.co.uk

24 MJG: martin.green@nottingham.ac.uk

25 ARO: oarahman@usm.my

26 JK: jasmeet.kaler@nottingham.ac.uk

27

28 **Abstract (400 words)**

29 Recent studies have reported associations between lameness and body condition score (BCS) in
30 dairy cattle, however the impact of change in the dynamics of BCS on both lameness occurrence
31 and recovery is currently unknown. The aim of this study was to investigate in a longitudinal
32 study the effect of change in BCS on the transitions from the non-lame to lame, and lame to non-
33 lame states. A total of 731 cows with 6889 observations from 4 UK herds were included in the
34 study. Mobility score (MS) and body condition score (BCS) were recorded every 13-15 days
35 from July 2010 until December 2011. A multilevel multistate discrete time event history model
36 was built to investigate the transition of lameness over time. There were 1042 non-lame
37 episodes and 593 lame episodes of which approximately 50% (519/1042) of the non-lame
38 episodes transitioned to the lame state and 81% (483/593) of the lame episodes ended with a
39 transition to the non-lame state. Cows with a lower BCS at calving (BCS Group 1 (1.00-1.75)
40 and Group 2 (2.00-2.25)) had a higher probability of transition from non-lame to lame and a
41 lower probability of transition from lame to non-lame compared to cows with BCS 2.50-2.75 i.e.
42 they were more likely to become lame and if lame, they were less likely to recover. Similarly,
43 cows who suffered a greater decrease in BCS (compared to their BCS at calving) had a higher
44 probability of becoming lame and a lower probability of recovering in the next 15 days. An

45 increase in BCS from calving was associated with the converse effect i.e. a lower probability of
46 cows moving from the non-lame to the lame state and higher probability of transition from lame
47 to non-lame. Days of lactation, months of calving and parity was associated with both lame and
48 non-lame transitions and there was evidence of heterogeneity among cows in lameness
49 occurrence and recovery. This study suggests loss of BCS and increase of BCS could influence
50 the risk of becoming lame and the chance of recovery from lameness. Regular monitoring and
51 maintenance of BCS on farms could be a key tool for reducing lameness. Further work is
52 urgently needed in this area to allow a better understanding of the underlying mechanisms behind
53 these relationships.

54

55 **Keywords: Lameness, Body condition score, Dairy cow, Multilevel multistate model,**

56 **Discrete time event history model, transitions**

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58

59 **1. Introduction**

60 Lameness is one of the most challenging diseases for the dairy industry due to its serious welfare
61 impact and associated economic losses (Kossaibati and Esslemont, 1997; Cha et al., 2010).
62 Lameness in cows leads to discomfort and pain resulting in altered gaits (Whay et al., 1998).
63 There is a significant impact of lameness on milk production, reproductive performances and it
64 results in a higher culling rate (Rajala-Schultz and Grohn, 1999; Green et al., 2002; Bicalho et al.,
65 2007; Peake et al., 2011; Huxley, 2013). In the UK, the estimated average herd prevalence of
66 lameness is 36% (range: 0 to 79%) (Barker et al., 2010; Leach et al., 2010); similar prevalence
67 estimates have been reported from other locations around the world (Hernandez et al., 2005;
68 Dippel et al., 2009; Tadich et al., 2010).

69

70 Over the last decade a small number of studies have reported an association between lameness
71 and poor body condition score (BCS) (Espejo et al., 2006; Dippel et al., 2009; Hoedemaker et al.,
72 2009). The area has recently been reviewed (Huxley, 2013); historically it has been assumed
73 that lameness led to cows having a lower BCS because disease meant that cows were more likely
74 to have lower dry matter intakes, decreased feeding time and longer lying time (Bach et al., 2007;
75 Kilic et al., 2007). However, in a cross-sectional study, Bicalho et al. (2009) reported that lame
76 cows were more likely to have thinner digital cushions compared with non-lame cows and
77 reported a significant positive association between BCS and thickness of the digital cushion, i.e.
78 cows with low BCS had thinner digital cushions compared with cows with higher BCS. The
79 authors hypothesized that losing BCS could influence the cows to change from non-lame to lame
80 due to thinning of the digital cushion but could not test this due to the cross-sectional nature of
81 the data. In a longitudinal study conducted on one farm, Green et al. (2013) reported that cows

82 with low BCS ($BCS < 2.5$) in the previous 0-2 months and $> 2-4$ months had a higher risk of
83 treatment for lameness in a 30 day period. However, exploration of the dynamics of change of
84 BCS on both the occurrence and recovery from lameness was not investigated and the study was
85 conducted on a single farm which limited the generalizability of the findings.

86

87 Multilevel multistate discrete time event history models can be used to investigate the effects of
88 factors on the likelihood of transitions between states among animals (e.g. disease/healthy) in
89 longitudinal data. Their use is becoming more common in exploring the complex dynamics of
90 diseases on farms (Kaler et al., 2010; Reader et al., 2011; Nielsen et al., 2012) and understanding
91 the interplay of animal level factors. Moreover, they avoid the limitation of fitting a separate
92 model for each state transition (Steele et al., 2004) accounting for the correlation that may exist
93 between the transitions due to heterogeneity. The aim of this longitudinal study was to
94 investigate the temporal effect of changes in BCS in a cow, and other cow level factors, on the
95 transitions from non-lame to lame and lame to non-lame states in a 15-day risk period .

96

97 **2. Material and methods**

98

99 ***2.1. Dataset and study methodology***

100 The data was collected from a convenience sample of four dairy herds in the South West region
101 of the UK. Herds were selected based on their proximity and previous working relationship with
102 the observer (JAW), the quality of their records and their willingness to participate. Animals on
103 the study farms were predominantly Holstein Friesians. The number of animals in milk, average
104 yield and herd calving to conception intervals are outlined in Table 1. On three farms cows were

105 housed through the winter period and had access to pasture during the summer, cows in one herd
106 (Farm 3) were continuously housed throughout the year. All four herds, when housed, used
107 cubicles; herd 4 bedded cows on deep sand, herd 3 bedded cows on mats with straw and the
108 remaining two farms bedded animals on mats with sawdust. In all herds dry cows were loose
109 housed on straw and all farms had loose straw areas available for freshly calved and sick cows.

110 Animals calving between July 2010 and June 2011 were selected from each herd. One
111 trained observer (JAW) visited all the herds every 13-15 days from July 2010 until December
112 2011. At every visit, the body condition score (BCS) and mobility score (MS) of selected cows
113 were recorded i.e. the lameness state of selected cows was identified based on a visual
114 assessment every 13-15 days throughout lactation. No treatment interventions were instigated by
115 the observer, consequently no temporal information on the cause of the lameness association
116 with the elevation in mobility score was collected. Body condition score (BCS) was scored
117 according to Edmonson et al. (1989) using a scale of 1-5 with increments of 0.25. Mobility score
118 (MS) was scored according to Whay et al. (2003) on a four point scale (0 to 3). Other
119 information such as parity, age, days of lactation and month of latest calving of selected cows
120 were recorded.

121

122 ***2.2. Statistical analysis***

123 Cows with no information for parity, age, days of lactation, month of latest calving or BCS at
124 calving (0-15 days post-partum) were excluded from the dataset. Any missing observation at the
125 start of the study and end of the study was excluded and if cows had missing visits in the middle
126 of the observation period, the remaining observations following the missing visit were excluded.
127 Only cows with at least 5 observations were used in the analysis. The BCS was grouped into four

128 categories: group1 (BCS G1 1.00-1.75), group 2 (BCS G2 2.00-2.25), group 3 (BCS G3 2.50-
129 2.75) and group 4 (BCS G4 3.00-5.00), while MS was categorized into two groups: non-lame
130 (MS 0 and 1) and lame (MS 2 and 3). Descriptive analyses were conducted in Stata version 12
131 (StataCorp, USA).

132

133

134 ***2.3. Multilevel multistate discrete time event history model***

135 A multilevel multistate discrete time event history model (Steele et al., 2004) was set up to
136 investigate the effect of covariates on the probability of transitions between lameness states, in a
137 15 day risk period. Two origin states were defined for lameness in the multilevel multistate
138 model for each cow k : non-lame (MS 0 and 1; coded as 0) and lame (MS 2 and 3; coded as 1).
139 Data were censored at the end of the study. An episode was the continuous period of time (in
140 discrete time intervals) a cow spent in a state until a transition occurred or data was censored. For
141 each episode, the time interval was categorized in discrete periods (15 days) and the number of
142 time intervals (t) was measured as $t=1, 2, 3 \dots$ up to the maximum observed time. For each cow k ,
143 at the end of each discrete time interval t (15 day) the outcome y was two possible
144 transitions/event occurrences, non-lame to lame or lame to non-lame.

145 The response variable y was the binary indicator of event occurrence. The event was
146 when a cow transitioned from lame to non-lame or non-lame to lame. The event indicator was
147 coded as 1 or 0 depending on the occurrence of an event: if no event occurred, 0 was coded; 1
148 was coded if an event occurred (non-lame to lame state or vice versa).

149 The hazard of transition from origin state i to transition state r_i during discrete-time interval
150 t of episode j for cow k was denoted by $h_{ijk}^{(ri)}$, and the hazard of no transition was denoted by
151 $h_{ijk}^{(0)}$,

152

153 The multilevel multistate discrete time model took the form:

154 $\log(h_{ijk}^{(ri)} / h_{ijk}^{(0)}) = \beta_0^{(ri)} + \alpha_t^{(ri)} + \beta x_{ijk}^{(ri)} + \mu_{ik}^{(ri)}$ where $\beta_0^{(ri)}$ is a state specific intercept, $\alpha_t^{(ri)}$ is an effect of
 $r = 0, 1; i = 0, 1$

155 duration which is a piecewise constant step function of time interval with three categories, $\beta x_{ijk}^{(ri)}$

156 refer to the covariates and $\mu_{ik}^{(ri)}$ represents the random effect of cow level which were assumed to

157 follow a normal distribution with variance σ_u^2 and non-zero correlation between random effects.

158 The model was created in MLwiN version 2.25 (Centre for Multilevel Modelling, University of

159 Bristol) and estimation was carried out by quasi-likelihood methods and followed by Monte

160 Carlo Markov chain (MCMC) methods with 500,000 iterations and a burn in of 5,000. The chain

161 mixing and stability were then evaluated visually in the MCMC trajectories window and MCMC

162 diagnostic window (Browne, 2009).

163

164 **2.4. Covariates (Predictor variables) used in the model**

165 The covariates with fixed effects used in the analysis were: parity group (1st, 2nd, 3rd and >3rd),

166 herd (Herd 1, 2, 3 and 4), month of calving (January-March, April-June, July-September and

167 October-December) and BCS at calving (BCS G1: 1.00-1.75, BCS G2: 2.00-2.25, BCS G3: 2.50-

168 2.75 and BCS G4: 3.00-5.00). Days of lactation and changes of BCS at the current visit

169 compared with calving were included in the model as time-varying effects. Days of lactation

170 referred to the days which compared the current visit with the latest calving date and were
171 categorized into three categories: 0-90 days, 91-120 days and more than 120 days. Likewise, the
172 change in BCS at the current visit compared with BCS at calving was categorized into five
173 categories:

- 174 i) no change (C1)
- 175 ii) decrease in 2 or 3 categories of BCS compared with calving (C2) (e.g. BCS G4 drop
176 to BCS G2 or BCS G1)
- 177 iii) decrease in 1 category of BCS compared with calving (C3) (e.g. BCS G4 drop to
178 BCS G3)
- 179 iv) increase in 1 category of BCS compared with calving (C4) (e.g. BCS G2 increase to
180 BCS G3)
- 181 v) increase in 2 categories of BCS compared with calving (C5) (e.g. BCS G2 increase to
182 BCS G4).

183

184 **3. Results**

185 ***3.1. Descriptive results***

186 There were a total of 8351 observations obtained from 731 cows across the four herds. Of these,
187 a total of 1462 observations were excluded for reasons described in the methodology. The final
188 dataset had 6889 observations, these included 110 (1100 observations), 329 (2953 observations),
189 166 (1626 observations) and 126 (1210 observations) cows from Herd 1, 2, 3 and 4 respectively.
190 There were 214 (29%), 145 (20%), 113 (16%) and 259 (35%) cows in parity 1, 2, 3 and 4 or
191 greater respectively. A total of 166 (23%), 115 (16%), 258 (35%) and 192 (26%) cows calved

192 within Jan-March, April-June, July-Sep and Oct-Dec respectively. The frequencies of variables
193 at the cow level by herd are presented in Table 2.

194

195 Out of 1635 episodes, 1042 were non-lame episodes and 593 were lame episodes.
196 Approximately 50% (519/1042) of non-lame episodes transitioned to the lame state and 81%
197 (483/593) of lame episodes ended with a transition to the non-lame state. For transitions from the
198 non-lame to the lame state, approximately 65% occurred within 45 days, 84% within 90 days and
199 the remaining (16%) occurred after 90 days. Approximately 88% of transitions from the lame to
200 the non-lame state occurred within 45 days, 97% within 90 days and 3% after 90 days.

201

202 ***3.2. Multilevel multistate model***

203 The multistate model results are presented in Table 3 and 4. Cows in BCS G1 (1.00-1.75) and
204 BCS G2 (2.00-2.25) at calving had a significantly higher probability of transition from the non-
205 lame to the lame state in a 15 day period (OR: 7.73, CI: 3.37-17.71; OR: 1.54, CI: 1.11-2.14) and
206 a lower probability of transition from the lame to the non-lame state in a 15 day period (OR:
207 0.21, CI: 0.08-0.52; OR: 0.31, CI: 0.19-0.51) compared with the cows with BCS G3 (2.50-2.75)
208 respectively.

209

210 When comparing the BCS at the current visit with the BCS at calving; cows where the BCS had
211 decreased by 2 or 3 groups had a higher probability of transition from the non-lame to the lame
212 state over the next 15 day period (OR: 2.07, CI: 1.30-3.29) and a lower probability of transition
213 from lame to non-lame over the next 15 day period (OR: 0.44, CI: 0.23-0.82), compared with
214 cows with no change in BCS. Similarly, cows where BCS decreased 1 group had a lower

215 probability of transition from the lame to the non-lame state over the next 15 day period
216 compared with cows where BCS did not change (OR: 0.66, CI: 0.44-0.98); there was no
217 significant effect of this change in the transition from non-lame to lame.

218

219 Cows in BCS G1 (1.00-1.75), G2 (2.00-2.25) or G3 (2.5-2.75) that had increased one BCS
220 category at the current visit had a lower probability of transition from the non-lame to the lame
221 state (OR: 0.53, CI: 0.33-0.83) and a higher probability of transition from the lame to the non-
222 lame state (OR: 2.49, CI:1.37-4.53) over the next 15 day period, compared with the cows with no
223 change in BCS.

224

225 The probability of cows transitioning from the non-lame to the lame state was lower the
226 longer a cow remained non-lame (OR: 0.53, CI: 0.40-0.71; OR: 0.33, CI: 0.21-0.51). In contrast,
227 the longer the cow was in the lame state, the lower the probability (OR: 0.42, CI: 0.18-0.96) of
228 transition from the lame to the non-lame state (i.e. they were more likely to remain lame). Cows
229 between 91-120 days of lactation (OR: 1.64, CI: 1.17-2.29) and greater than 120 days of
230 lactation (OR: 2.11, CI: 1.58-2.82) were more likely to change from the non-lame to the lame
231 state compared with cows less than 91 days of lactation. Cows between 91-120 days of lactation
232 were less likely to change from the lame to the non-lame state compared with cows less than 91
233 days of lactation (OR: 0.67, CI: 0.44-1.00). Cows that calved between Oct-Dec had a higher
234 probability of transition from the lame to the non-lame state compared with cows that calved
235 during Jan-March (OR: 1.79, CI: 1.05-3.06); the month of calving did not have a significant
236 effect on transition from the non-lame to the lame state.

237

238 Cows in parity 3 and above had a higher probability (OR: 3.00, CI: 1.95-4.60; OR: 3.51,
239 CI: 2.39-5.17) of transition from the non-lame to the lame state and a lower probability (OR:
240 0.50, CI: 0.25-1.00; OR: 0.36, CI: 0.19-0.67) to transition from the lame to the non-lame state
241 compared with cows in parity 1. Cows in parity 2 did not show any significant differences in
242 both transitions compared with cows in parity 1.

243
244 Herd 2 (OR: 3.18, CI:1.98-5.11) and Herd 3 (OR: 1.94, CI:1.19-3.15) had a significantly higher
245 probability of transition from the non-lame to the lame state compared with Herd 1; Herd 2 had a
246 significantly lower probability of transition from the lame to the non-lame state compared with
247 Herd 1 (OR: 0.23, CI:0.10-0.53).

248
249 Estimated random effects covariance matrices for the models with duration effects only and
250 duration with all covariates are presented in Table 5. For the model with duration effects only,
251 there was significant moderate negative correlations (-0.76) between the probability of both
252 transitions from non-lame to lame and lame to non-lame. It indicated that the cows with a high
253 (low) probability of switching from non-lame to lame had a low (high) probability of changing
254 from the lame to the non-lame state. Equally, cows with a longer (shorter) duration of the non-
255 lame state were likely to have a shorter (longer) duration in the lame state. The correlation
256 between both transitions after adding covariates was not significant. The model converged
257 visually with stable chain mixing.

258

259 **4. Discussion**

260 The key finding from the study reported here is that loss of BCS increased a cow's probability of
261 becoming identifiably lame and decreased her likelihood of recovery, over the next 15 days; the
262 effect was apparent after controlling for body condition score at calving. Furthermore, cows with
263 a greater decrease of BCS had a higher (lower) probability of transition from non-lame to lame
264 (lame to non-lame) compared with cows that had a relatively lower decrease in BCS i.e. the
265 effect was greater in animals which lost more condition. Our findings demonstrate that in this
266 dataset loss of BCS preceded an animal being identified as lame by mobility scoring.

267 The findings from this longitudinal study shed no light on the causality of this
268 relationship, however they add further evidence in support of a previous cross-sectional study
269 that has demonstrated that BCS is associated with the thickness of the digital cushion (Bicalho et
270 al., 2009). The digital cushion is thought to absorb part of the load and protecting the corium
271 during locomotion, lowering the incidence of claw horn lesions (sole haemorrhage, sole ulcers
272 and white line disease) and lameness (Räber et al., 2004; Räber et al., 2006; Bicalho et al., 2009).
273 Results from the current study support the hypothesis that a greater loss of BCS leads to a higher
274 risk of lameness possibly due to the thinning effects on the digital cushion. They also suggest
275 that body condition score loss decreases an animal's chance of recovery i.e. animals which lose
276 weight are less likely to recover from any lameness events which do occur. Alternatively,
277 thinner cows may have lower social rank and therefore be less able to compete with other cows
278 for resources such as lying space and access to feed (Hohenbrink and Meinecke-Tillmann, 2012).
279 Being out competed in the environment could put them at greater risk of becoming lame and
280 make them less likely to recover once a lameness event occurs.

281

282 Our findings support the work of Green et al. (2013) who demonstrated that low BCS (<2.5) was
283 associated with an increased risk of treatment for lameness in the following four months.
284 Additionally they extend the previous findings by exploring the dynamics of changes in BCS and
285 lameness both in terms of lameness occurrence and recovery in a 15 day period. The work
286 described here suggests that a) in addition to low BCS per se being a risk for lameness, any
287 significant loss of BCS has an added detrimental effect on the occurrence and recovery from
288 lameness in a 15 day risk period, b) thinner cows at calving who gain BCS have a lower chance
289 of becoming lame and a higher chance for recovery compared with their counterparts who stay
290 thin. That is the interrelationship between BCS and BCS change and lameness are bidirectional.

291

292 Previous work has demonstrated that the thickness of the digital cushion is influenced by age,
293 parity and days of lactation (Räber et al., 2004; Bicalho et al., 2009). It is noteworthy that these
294 factors also had a significant effect on lameness transitions in the current study. The thickness of
295 the digital cushion increases to the third parity and then reduces after the third parity (Räber et al.,
296 2004; Bicalho et al., 2009). Cows in parity three and greater were more likely to become lame
297 and less likely to recover in the current study. Alternatively, previous work has demonstrated that
298 animals which have experienced a lameness event in previous lactations are at greater risk of
299 future cases of lameness (Hirst et al., 2002) The increased risk of lameness in older cows could
300 have been caused by their lameness history. For days of lactation, the thickness of the digital
301 cushion has been reported to decrease from early lactation until 120 days post calving and then
302 slowly increase in thickness after 120 days (Bicalho et al., 2009). Our results demonstrate that
303 cows were more likely to become lame and less likely to recover between 91 and 120 days of
304 lactation, this coincides with the period that the digital cushion is thinnest. Equally this finding

305 could be due to injuries to the corium around the time of parturition and in early lactation taking
306 a period of time to cause an identifiable lameness event.

307

308 The duration that cows spent in a particular lameness state influences the risk of lameness and
309 recovery (Reader et al., 2011). Our results suggest that cows that have been lame for longer are
310 less likely to recover. Equally, the longer that cows remain non-lame the less likely they are to
311 become lame. This was further confirmed by the significant moderate correlations observed
312 between lameness transitions. This could be due to the effect of cow heterogeneity or genetic
313 resistance to lameness, where cows with inherently thinner digital cushion will have a higher risk
314 of lameness and a lower chance of recovery compared with other animals. A recent study
315 (Oikonomou et al., 2013) reported that the heritability estimate for digital cushion thickness was
316 moderate (0.33) and it was genetically correlated with claw horn disruption lesions. Alternatively
317 the observed effects could be mediated via the impacts of lameness on nutrition. Lame cows
318 have shorter feeding times and suffer reduced feed intakes (Bach et al., 2007; Kilic et al., 2007)
319 which could further reduce the thickness of the digital cushion exacerbating the problem i.e.
320 lameness state is a vicious (or virtuous) cycle. Finally this increase in lameness risk could be due
321 to some failure in the efficacy of treatments. Since lameness was only measured by mobility
322 score, no information was captured on the causes of lameness or if, when and how animals were
323 treated on farm. Reader et al. (2011) reported that cows with a history of treatment for sole ulcer
324 and digital dermatitis were more likely to become lame compared with non-treated cows,
325 suggesting that current treatments may not be fully healing the lesion or leaving the animal with
326 changes to the foot which increase the risk of recurrence in the future.

327

328 In the current study, the chance of incorrect animal identification was minimised by identifying
329 cows whilst they were stationary in the milking parlour. Moreover, all the observations for BCS
330 and MS score were made by a single trained and experienced observer to reduce between
331 observer bias, although the possibility of intra-observer bias remains. We believe this approach
332 is likely to have reduced the chance of data misclassification and the authors have no reason to
333 suspect that what misclassifications did occur were not randomly distributed through the data set.
334 Although the 4 herds selected for the study were not randomly selected we have no reason to
335 believe that they are not representative of typical UK dairy herd w.r.t. range and change of body
336 condition and mobility scores. Given the consistency of our results with previous literature, we
337 consider that the results are likely to be generalisable to other similar cattle herds.

338

339

340 **5. Conclusion**

341 In conclusion, our results suggest that both a decrease and an increase in BCS influences the risk
342 of becoming lame and the chance of recovery, possibly due to the impact of body weight change
343 on the thickness of the digital cushion. Further longitudinal studies with detailed measurements
344 of the digital cushion alongside BCS and MS would help to explore these complex interactions.
345 Regular monitoring and maintenance of BCS on farms could be a key tool for managing the risk
346 of lameness on farm.

347

348 **6. Conflicts of interest**

349 The authors declare that they have no competing interests.

350

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355

356 **References**

357

358 Bach, A., Dinares, M., Devant, M., Carre, X., 2007. Associations between lameness and
359 production, feeding and milking attendance of Holstein cows milked with an automatic
360 milking system. *J. Dairy Res.* 74, 40-46.

361 Barker, Z.E., Leach, K.A., Whay, H.R., Bell, N.J., Main, D.C., 2010. Assessment of lameness
362 prevalence and associated risk factors in dairy herds in England and Wales. *J. Dairy Sci.*
363 93, 932-941.

364 Bicalho, R.C., Machado, V.S., Caixeta, L.S., 2009. Lameness in dairy cattle: A debilitating
365 disease or a disease of debilitated cattle? A cross-sectional study of lameness prevalence
366 and thickness of the digital cushion. *J. Dairy Sci.* 92, 3175-3184.

367 Bicalho, R.C., Vokey, F., Erb, H.N., Guard, C.L., 2007. Visual locomotion scoring in the first
368 seventy days in milk: impact on pregnancy and survival. *J. Dairy Sci.* 90, 4586-4591.

369 Cha, E., Hertl, J.A., Bar, D., Grohn, Y.T., 2010. The cost of different types of lameness in dairy
370 cows calculated by dynamic programming. *Prev. Vet. Med.* 97, 1-8.

371 Dippel, S., Dolezal, M., Brenninkmeyer, C., Brinkmann, J., March, S., Knierim, U., Winckler, C.,
372 2009. Risk factors for lameness in freestall-housed dairy cows across two breeds, farming
373 systems, and countries. *J. Dairy Sci.* 92, 5476-5486.

374 Edmonson, A.J., Lean, I.J., Weaver, L.D., Farver, T., Webster, G., 1989. A body condition
375 scoring chart for Holstein dairy cows. *J. Dairy Sci.* 72, 68-78.

376 Espejo, L.A., Endres, M.I., Salfer, J.A., 2006. Prevalence of lameness in high-producing
377 Holstein cows housed in freestall barns in Minnesota. *J. Dairy Sci.* 89, 3052-3058.

378 Galindo, F., Broom, D.M., 2000. The relationships between social behaviour of dairy cows and
379 the occurrence of lameness in three herds. *Res. Vet. Sci.* 69, 75-79.

380 Galindo, F., Broom, D.M., Jackson, P.G.G., 2000. A note on possible link between behaviour
381 and the occurrence of lameness in dairy cows. *Appl. Anim. Behav. Sci.* 67, 335-
382 341. Green, L.E., Hedges, V.J., Schukken, Y.H., Blowey, R.W., Packington, A.J., 2002.
383 The impact of clinical lameness on the milk yield of dairy cows. *J. Dairy Sci.* 85, 2250-
384 2256.

385 Green, L.E., Huxley, J.N., Banks, C., Green, M.J., 2013. Temporal associations between low
386 body condition, lameness and milk yield in a UK dairy herd. *Prev. Vet. Med.* 113, 63-71.
387

388 Hirst, W.M., Murray, R.D., Ward, W.R., French, N.P., 2002. A mixed-effects time-to-event
389 analysis of the relationship between first-lactation lameness and subsequent lameness in
390 dairy cows in the UK. *Prev. Vet. Med.* 54, 191-201. Hernandez, J.A., Garbarino, E.J.,
391 Shearer, J.K., Risco, C.A., Thatcher, W.W., 2005. Comparison of the calving-to-
392 conception interval in dairy cows with different degrees of lameness during the
393 prebreeding postpartum period. *J. Am. Vet. Med. Assoc.* 227, 1284-1291.

394 Hoedemaker, M., Prange, D., Gundelach, Y., 2009. Body condition change ante- and postpartum,
395 health and reproductive performance in German Holstein cows. *Reprod. Domest. Anim.*
396 44, 167-173.

397 Hohenbrink, S., Meinecke-Tillmann, S., 2012. Influence of social dominance on the secondary
398 sex ratio and factors affecting hierarchy in Holstein dairy cows. *J. Dairy Sci.* 95, 5694-
399 5701.

400 Huxley, J.N., 2013. Impact of lameness and claw lesions in cows on health and production.
401 *Livest. Sci.* 156, 64-70.

402 Kaler, J., Medley, G.F., Grogono-Thomas, R., Wellington, E.M., Calvo-Bado, L.A., Wassink,
403 G.J., King, E.M., Moore, L.J., Russell, C., Green, L.E., 2010. Factors associated with
404 changes of state of foot conformation and lameness in a flock of sheep. *Prev. Vet. Med.*
405 97, 237-244.

406 Kilic, N., Ceylan, A., Serin, I., Gokbulut, C., 2007. Possible interaction between lameness,
407 fertility, some minerals, and vitamin E in dairy cows. *Bull. Vet. Inst. Pulawy* 51, 425-429.

408 Kossaibati, M.A., Esslemont, R.J., 1997. The costs of production diseases in dairy herds in
409 England. *Vet. J.* 154, 41-51.

410 Leach, K.A., Whay, H.R., Maggs, C.M., Barker, Z.E., Paul, E.S., Bell, A.K., Main, D.C., 2010.
411 Working towards a reduction in cattle lameness: 1. Understanding barriers to lameness
412 control on dairy farms. *Res. Vet. Sci.* 89, 311-317.

413 Nielsen, B.H., Thomsen, P.T., Green, L.E., Kaler, J., 2012. A study of the dynamics of digital
414 dermatitis in 742 lactating dairy cows. *Prev. Vet. Med.* 104, 44-52.

415 Oikonomou, G., Banos, G., Machado, V., Caixeta, L., Bicalho, R.C., 2013. Short communication:
416 Genetic characterization of digital cushion thickness. *J. Dairy Sci.* 97, 1-5.

417 Peake, K.A., Biggs, A.M., Argo, C.M., Smith, R.F., Christley, R.M., Routly, J.E., Dobson, H.,
418 2011. Effects of lameness, subclinical mastitis and loss of body condition on the
419 reproductive performance of dairy cows. *Vet. Rec.* 168, 301.

420 Räber, M., Lischer, C.J., Geyer, H., Ossent, P., 2004. The bovine digital cushion – a descriptive
421 anatomical study. *Vet. J.* 167, 258-264.

422 Räber, M., Scheeder, M.R.L., Ossent, P., Lischer, C.J., Geyer, H., 2006. The content and
423 composition of lipids in the digital cushion of the bovine claw with respect to age and
424 location – A preliminary report. *Vet. J.* 172, 173-177.

425 Rajala-Schultz, P.J., Grohn, Y.T., 1999. Culling of dairy cows. Part III. Effects of diseases,
426 pregnancy status and milk yield on culling in Finnish Ayrshire cows. *Prev. Vet. Med.* 41,
427 295-309.

428 Reader, J.D., Green, M.J., Kaler, J., Mason, S.A., Green, L.E., 2011. Effect of mobility score on
429 milk yield and activity in dairy cattle. *J. Dairy Sci.* 94, 5045-5052.

430 Steele, F., Goldstein, H., Browne, W., 2004. A general multilevel multistate competing risks
431 model for event history data, with an application to a study of contraceptive use dynamics.
432 *Statistical Modelling* 4, 145-159.

433 Tadich, N., Flor, E., Green, L., 2010. Associations between hoof lesions and locomotion score in
434 1098 unsound dairy cows. *Vet. J.* 184, 60-65.

435 Whay, H.R., Main, D.C.J., Green, L.E., Webster, A.J.F., 2003. Assessment of the welfare of
436 dairy cattle using animal-based measurements: direct observations and investigation of
437 farm records. *Vet. Rec.* 153, 197-202.

438 Whay, H.R., Waterman, A.E., Webster, A.J., O'Brien, J.K., 1998. The influence of lesion type on
439 the duration of hyperalgesia associated with hindlimb lameness in dairy cattle. *Vet. J.* 156,
440 23-29.

441 **Tables**

442 **Table 1.** Descriptive data collected from 7/10/2010 to 12/11/2011 on 4 study farms located in
443 southwest region of UK

444

445 **Table 2.** The frequency of the covariates at the cow level by herd used in multilevel multistate
446 model

447

448 **Table 3.** Frequency of observations for each covariate/predictor variable in the multilevel
449 multistate model by state (Lame/Non-Lame)

450

451 **Table 4.** Multilevel multistate model of factors associated with transition of cows from the non-
452 lame to the lame and the lame to the non-lame state in dairy cows on 4 herds

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454 **Table 5.** Random effects covariance matrix for a multilevel multistate model looking into
455 transitions from the non-lame to the lame and the lame to the non-lame state in cows

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Table 1

	Herd 1	Herd 2	Herd 3	Herd 4
Number of cows in milk*	164	359	220	138
Milk Production (kg / 305d lactation)	5,889	8,057	9,171	9,005
Calving-Conception interval (d)	93	148	138	115

***Breed of cows: Predominantly Holstein Friesians**

Table 2.

Herd	1		2		3		4		Total
	Freq.	%	Freq.	%	Freq.	%	Freq.	%	Freq.
Cows	110	15.05	329	45.01	166	22.71	126	17.23	731
Parity group									
1	25	22.73	100	30.40	48	28.92	41	32.54	214
2	15	13.64	48	14.59	55	33.13	27	21.43	145
3	19	17.27	50	15.20	21	12.65	23	18.25	113
>3	51	46.36	131	39.81	42	25.30	35	27.78	259
Month of calving									
Jan-March			81	24.62	49	29.52	36	28.57	166
Apr-June			83	25.23	18	10.84	14	11.11	115
July-Sep	84	76.36	82	24.92	58	34.94	34	26.98	258
Oct-Dec	26	23.64	83	25.23	41	24.70	42	33.34	192
BCS at calving									
BCS G1 (1.00-1.75)			16	4.86	3	1.81	2	1.59	21
BCS G2 (2.00-2.25)	23	20.91	116	35.26	46	27.71	41	32.54	226
BCS G3 (2.50-2.75)	81	73.64	163	49.54	105	63.25	72	57.14	421
BCS G4 (3.00-5.00)	6	5.45	34	10.34	12	7.23	11	8.73	63

Table 3.

Covariates	Categories	State			
		Non-lame (%)		Lame (%)	
Duration spent in state (days)	≤45	2062	(37.50)	974	(70.07)
	46-90	1618	(29.42)	240	(17.27)
	≥90	1819	(33.08)	176	(12.66)
Herd	1	1028	(18.69)	72	(5.18)
	2	1989	(36.17)	964	(69.35)
	3	1402	(25.50)	224	(16.12)
	4	1080	(19.64)	130	(9.35)
Days of lactation	0-90	3136	(57.03)	684	(49.21)
	91-120	923	(16.78)	238	(17.12)
	>120	1440	(26.19)	468	(33.67)
Month of calving	Jan-March	1104	(20.08)	398	(28.63)
	Apr-June	671	(12.20)	251	(18.06)
	July-Sep	2211	(40.21)	484	(34.82)
	Oct-Dec	1513	(27.51)	257	(18.49)
Parity group	1	1818	(33.06)	145	(10.43)
	2	1239	(22.53)	159	(11.44)
	3	809	(14.71)	278	(20.00)
	>3	1633	(29.70)	808	(58.13)
BCS at calving	BCS G3 (2.50-2.75)	3413	(62.07)	549	(39.50)
	BCS G1 (1.00-1.75)	74	(1.35)	104	(7.48)
	BCS G2 (2.00-2.25)	1517	(27.58)	640	(46.04)
	BCS G4 (3.00-5.00)	495	(9.00)	97	(6.98)
BCS (Comparing current visit with calving)	No change	2860	(52.01)	574	(41.29)
	Decrease by 2,3 groups	341	(6.20)	210	(15.11)
	Decrease by 1 group	1801	(32.75)	517	(37.19)
	Increase by 1 group	467	(8.49)	87	(6.26)
	Increase by 2 groups	30	(0.55)	2	(0.14)

Table 4.

Transitions Covariates	Non-lame to lame ^{a,b,c}					Lame to non-lame ^{a,b,c}				
	Freq.	OR.	95%	C.I.	P-value	Freq.	OR.	95%	C.I.	P-value
Duration spent in state (days)										
≤45	339	Ref.				427	Ref.			
46-90	98	0.53	0.40	0.71	<0.001	43	0.64	0.40	1.04	0.07
≥90	82	0.33	0.21	0.51	<0.001	13	0.42	0.18	0.96	0.04
Herd										
1	55	Ref.				49	Ref.			
2	272	3.18	1.98	5.11	<0.001	247	0.23	0.10	0.53	<0.001
3	115	1.94	1.19	3.15	0.01	113	0.64	0.28	1.47	0.29
4	77	1.36	0.82	2.28	0.24	74	0.83	0.34	2.01	0.68
Days of lactation										
0-90	284	Ref.				267	Ref.			
91-120	84	1.64	1.17	2.29	<0.001	66	0.67	0.44	1.00	0.05
>120	151	2.11	1.58	2.82	<0.001	150	0.72	0.50	1.03	0.07
Month of calving										
Jan-March	132	Ref.				130	Ref.			
Apr-June	81	0.71	0.45	1.11	0.13	70	1.19	0.67	2.12	0.55
July-Sep	184	0.95	0.66	1.38	0.79	164	0.81	0.50	1.33	0.41
Oct-Dec	122	0.82	0.56	1.20	0.31	119	1.79	1.05	3.06	0.03
Parity group										
1	88	Ref.				80	Ref.			
2	78	1.26	0.83	1.91	0.28	76	1.08	0.52	2.25	0.83
3	109	3.00	1.95	4.60	<0.001	95	0.50	0.25	1.00	0.05
>3	244	3.51	2.39	5.17	<0.001	232	0.36	0.19	0.67	<0.001
BCS at calving										
BCS G3 (2.50-2.75)	266	Ref.				242	Ref.			
BCS G1 (1.00-1.75)	29	7.73	3.37	17.71	<0.001	24	0.21	0.08	0.52	<0.001
BCS G2 (2.00-2.25)	173	1.54	1.11	2.14	0.01	173	0.31	0.19	0.51	<0.001
BCS G4 (3.00-5.00)	51	0.92	0.56	1.51	0.74	44	1.51	0.73	3.10	0.27
BCS (Comparing current visit with calving)										
No change	253	Ref.				207	Ref.			
Decrease by 2,3 groups	64	2.07	1.30	3.29	<0.001	62	0.44	0.23	0.82	0.01
Decrease by 1 group	169	1.12	0.86	1.46	0.41	165	0.66	0.44	0.98	0.04
Increase by 1 group	31	0.53	0.33	0.83	0.01	48	2.49	1.37	4.53	<0.001
Increase by 2 groups	2	0.50	0.07	3.37	0.48	1	5.85	0.12	276.25	0.37

^a Intercept (coefficient (standard error)): Non-lame to lame: -3.80 (0.34); Lame to non-lame :2.44 (0.61)

^b Random variability between cows: Non-lame to lame : 0.80 (0.27) ; Lame to non-lame: 0.87 (0.36)

^c Covariance : -0.32(0.25); Freq.: frequency, OR.: odd ratios, C.I.: credible interval, P-value.: significant value, Ref.: reference

1

2 **Table 5.**

3 Model	Coefficient	SE
4 Duration effects only		
5 Non-lame to lame	2.45*	0.55
6 Lame to non-lame	1.63*	0.65
7 Covariance between both transitions	-1.51*	0.58
8 Correlation between random effects	-0.76	
9 Duration+covariates		
10 Non-lame to lame	0.80*	0.27
11 Lame to non-lame	0.87*	0.36
12 Covariance between both transitions	-0.32	0.25
13 Correlation between random effects	-0.38	

14 *Significant at 5% level, SE: standard error.

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