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1	Interpretive summary: Impact of mobility score on milk yield and activity. Reader
2	The hypothesis tested was that delay in treatment of lame cows explains the reduction
3	in milk yield before treatment. Delay in treatment was one likely explanation for a reduction
4	in milk yield. Reduced yield occurred before cows were visibly lame; one explanation is that
5	mobility scoring in less than 100% sensitive. An alternative hypothesis is that reduced body
6	condition caused both reduced milk yield and lameness as the digital cushion became thin.
7	LAMENESS AND MILK YIELD
8	
9	Impact of mobility score on milk yield and activity in dairy cattle
10	
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19 ABSTRACT

20 Previous studies have indicated that lame cows have a reduced milk yield both before 21 and after they are treated. One explanation for the reduction in yield before treatment is that 22 there is a delay to treatment, that is, cows have impaired mobility for some time before they 23 are treated. The aim of this study was to test this hypothesis by investigating temporal 24 associations between change in milk yield and change in mobility score. Mobility score (MS, 25 on a scale 0 to 3), milk yield, treatments for lameness and cow activity were recorded on 312 cows in a dairy herd in Somerset, UK for 1 yr. The MS was scored every 2 wk and 26 27 compared with the daily yield and activity (steps/h) averaged over the previous 16 d. 28 Approximately 44 % of MS changed within 14 d, usually by 1 score. Overall, milk yields of 29 cows with MS 1 were higher than those of cows with other scores. Cows with MS 2 and 3 30 produced 0.7 (0.35 - 0.97) kg and 1.6 (0.98 - 2.23) kg less milk / d, respectively, compared 31 with cows with MS 1. In addition, cows with MS 1 were slightly but significantly more 32 active than cows with MS 0, 2 or 3. Cows with MS 2 and 3 were 0.0.02 (0.01 - 0.03) and 33 0.03 (0.01 - 0.05) mean log steps less active than cows with MS 1.

34 There was a reduction in yield from 6 - 8 wk before becoming MS $2\ 0.5\ (0.12 - 0.47)$ 35 or 3 0.9 (0.16 - 1.65) to 4 wk after recovering from MS 2 0.42 (0.09 - 0.75) and non-36 significantly, score 3. The activity of cows was significantly less but quantitatively small 37 (mean log steps 0.01) with increasing MS; the associations between activity and parity 38 (mean 0.03 - 0.11) and month of lactation (mean 0.03 - 0.36) were quantitatively larger. 39 Results from a multistate model indicated that once cows were lame they remained lame or 40 become lame again despite treatment. We conclude that cows started to reduce milk 41 production before their mobility is visibly impaired. One explanation for this is that MS is not 100% sensitive. An alternative hypothesis, using evidence from other studies, is that reduction 42 43 in milk yield and development of lameness are on a common causal pathway most likely

linked to loss in body condition and reduced digital cushion thickness as a result of thedemands from producing high milk yields.

Key words Dairy cow, Milk yield, Lameness, Treatment, Multistate model

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INTRODUCTION

The prevalence and incidence of lameness in dairy cows in intensive systems is
unacceptably high with estimates of prevalence in the UK ranging from 21 % (Clarkson et al.,
1996) to 36 % (Leach et al., 2010). Lame cows are in pain and their welfare is compromised
(Whay et al., 1997).

53 Lameness is associated with a reduction in milk yield (Juarez et al., 2003; Archer et 54 al., 2010). This reduced milk yield is present before and after a treatment event, but varies by 55 the type of lesion (Green et al., 2002; Amory et al., 2008; Bicalho et al., 2008). The reduction in yield detected before a treatment event with non infectious horn lesions (Amory et al., 2008; 56 57 Green et al., 2010) might occur because of a long pathogenesis in disease before cows become 58 lame or because of delayed treatment. There is less evidence that infectious claw conditions 59 are associated with reduced milk yield before cows are observed lame, although Warnick et al. 60 (2001) reported that interdigital phlegmon was associated with reduced yield before treatment, 61 possibly because the time to lameness from infection is rapid. For both types of disorders, 62 delay in treatment would probably lead to reduced milk yield because of the increased 63 metabolic demands from pain and reduced feed intake. The treatment of lame cows depends 64 on the ability of farmers to recognize a lame cow and to treat affected cows promptly and appropriately. Most dairy cow farmers underestimate the prevalence of lameness on their 65 66 farms (Whay et al., 2003) and do so inconsistently compared with a trained researcher 67 (Leach et al., 2010), suggesting that most dairy cow herdsmen do not have a logical way 68 of detecting lameness, in contrast to sheep farmers (King and Green, in press).

Mobility scoring has been developed to help farmers improve detection of mild lameness and stimulate treatment and prevention as part of a herd health program. The currently accepted system used in the UK is a 4 point mobility scoring (**MS**, on a scale 0 to 3) system (Whay et al., 2003). This system is used by many researchers and veterinary practitioners, but has not been evaluated for repeatability. Some authors have reported that daily activity levels are lower in cows with reduced mobility (O'Callaghan et al., 2003; Mazrier et al., 2006; Walker et al., 2008).

The current study was designed to test the hypothesis that the reduction in milk yield that occurs before lame cows are treated is as a result of delayed treatment. This was tested by investigating the temporal association between change in milk yield and change in locomotion and time to treatment. The MS, milk yield, and activity in cattle from 1 farm was observed every 2 wk for 1 yr to estimate precise relationships between MS and changes in MS, milk yield, and cow activity.

82

MATERIALS AND METHODS

83 A dairy herd that calved all year round, located in Somerset UK, with a milking herd of 84 200 Holstein cows, producing approximately 9,000 kg milk/cow per year was used for the 85 study. The study started on October 24, 2007 and finished on November 5, 2008. Calving was all yr around; The numbers of cows in milk ranged from 168 (November 5, 2008) to 217 86 (April 23, 2008) with a mean of 197 and median of 200. The herd was divided into 2 groups of 87 88 about equal size based on milk yield, both housed in 1 building with a floor of concrete and 230 89 free stalls fitted with mattresses and bedded with sawdust. Milking cows had access to pasture 90 in summer with high yielding cows only on pasture for a limited period each day. Non-91 lactating cows were kept in a separate building and their locomotion was not scored. The 92 herd was milked twice daily through an 18/36 Westfalia herringbone parlor. Milking cows 93 walked through a 5% formalin footbath as they exited the parlor once each week.

94 Cows were selected for foot trimming by the herdsman. Approximately 35 cows 95 were trimmed per month; foot trimming was carried out by a paraprofessional foot trimmer 96 from Kingfisher Veterinary Practice (Synergy Farm Health, West Hill Barns, Evershot, 97 Dorset, England. DT2 0LD). The selection criteria for foot trimming were cows that were 98 clinically lame (MS 2 or 3) or cows that were due to be dried off. The farmer intended to 99 trim feet of each cow at least once each year, but this was not cross checked. Lesions were 100 defined using the definitions in the EU Lamecow Project (Barker et al., 2007) and all foot 101 trimming and lameness were recorded on lameness scoring sheets designed by the EU 102 Lamecow project. Cases of lameness treated by the herdsman or veterinarian (who treated 103 severe cases) were recorded in the same way.

104 All cows were individually identified and fitted with pedometers (Westfalia Dairy 105 Plan C21 (GEA Farm Technologies Australia Pty. Ltd. PO Box 39816 Trade Park Drive 106 Tullamarine VIC 3043). Activity readings for each cow were automatically downloaded to 107 the farm computer in the parlor twice daily and onto a lap top once weekly. The mobility of 108 lactating cows was scored (Table 1) every 2 wk after evening milking by JDR using the 109 system described by Whay et al., (2003). The identity of each cow was recorded as she 110 entered the parlor and mobility was scored and recorded on standardized sheets as the cow 111 exited the parlor. The MS was transferred to an Excel 2003 spreadsheet (Microsoft Corp., 112 Redmond, WA). Milk vield, activity (mean steps/hr), health records, lameness records, and 113 group were downloaded from the farm computer into the spreadsheet.

114 Data analysis

The mean proportion of cows with each MS by stage of lactation (1 to 90 d, 91 to 180 d, >180 d), mean milk yield, and mean activity over 16 d previously were calculated. The probability of transition between MS from time t to time t + 1, 14 d later, was estimated. Two multilevel statistical models were constructed, using conventional methods (Goldstein,

119 1995). In the first model the outcome variable was mean milk yield in the 16 d before a MS and 120 the impact of MS before and after this outcome was investigated. In the second model log¹⁰ 121 mean activity score for the previous 16 d was the outcome and the impact of MS on activity 122 was investigated.

123 The models took the form:

124
$$Y_{ij} = \alpha + \beta_1 X_{ij} + \beta_2 X_j + v_j + e_{ij} v_j \sim N(0, a^2 v)$$

125
$$eij \sim N(0,a^2_e)$$

where the subscripts i, and j denote the ith observation of the jth cow, respectively; α is the 126 regression intercept; X_{ij} is the vector of covariates associated with each observation; β_1 the 127 128 coefficients for covariates X_{ij} ; X_i the vector of covariates associated with each cow; β_2 the 129 coefficients for covariates X_i; v_i a random effect to reflect residual variation between cows which is normally distributed with mean = 0 and variance = σ^2 ; and e_{ii} a random effect to 130 131 reflect residual variation between MS which is normally distributed with mean = 0 and variance = σ^2 . The analysis was carried out using MLwiN 2.02 with penalized quasi-132 133 likelihood for parameter estimation (Rasbash et al., 2005). Covariates were left in the model 134 when the significance probability was P < 0.05 based on the Wald Test. When mean milk yield was the outcome, DIM, the exponential DIM ^{0.05} (Wilmink, 1987) and parity 1, 2, 3, and > 135 136 3, and first or second lactation in the study were forced into the model. Then the discrete 137 variable MS (0, 1, 2, and 3) at time t was added. The impact of MS at time t - 1, t - 2,..., t - 5 138 and t + 1, t + 2, ..., t + 5, where each time interval i was 14 d, was tested in the model. When 139 log mean activity was the outcome, parity 1, 2, 3, and > 3, second lactation in the study and 140 month in milk were forced into the model and then the mobility score at times t, t - 1, ..., t - 5 and t + 1, ..., t + 5, where each time interval t was 14 d, were tested in the model. Missing 141 142 observations were random and so were fitted in the model as discrete variables to minimize 143 loss of data. The model fit was checked.

144 Finally, a multistate model was set up to test the factors associated with cows 145 becoming lame, remaining lame, becoming sound, and remaining sound. Mobility score was 146 categorized into 2 states: not lame (scores 0 and 1) and lame (scores > 1). A cow was in 1 of 147 2 states, not lame or lame. An episode was defined as the continuous period of time a cow 148 spent in either state until a transition to the other state occurred. For each episode j for cow k149 there was an original state i (0 (not lame), 1 (lame)) the duration spent in that state was 150 categorized into discrete time intervals of 14 d, t_i (measured as $t = 1, 2, \dots, n$ with n being the 151 maximum duration of an episode) and an outcome event at the end of the discrete time 152 interval, y, with 0 = no change in state, and 1 = occurrence of a change in state. A logit link 153 function was used to express the ratio of probability of a change in state to probability of no 154 change in the state and took the form:

155
$$\operatorname{logit}[\pi_{ik(t)}] = \beta_{0_i} + \alpha_i(t) + \beta x_{ik(t)} + u_k^{(i)}$$

where β_{0_i} is a state specific intercept, $\alpha_i(t)$ a set of dummy variables for the discrete time interval *t* depicting duration of state, $\beta_{x_{ik(t)}}$ covariates include a vector of explanatory variables varying by time or cow with a dummy variable for original state. The model was run in MlwiN 2.02 (Rasbash et al., 2005) using Markov chain Monte Carlo estimation. The first 5,000 iterations were discarded and then 500,000 iterations until the chains were visually stable.

162

RESULTS

163	Mobility was scored on 28 occasions, 312 cows (allowing for additions and removals)
164	were scored with 168 to 217 at each observation, the number of scores arranged from 5 to 28 /
165	cow. The percent of scores 0, 1, 2, and 3 were 23, 45, 27, and 5, respectively, with 1, 20,
166	48, and 31% of cows with maximum scores of 0, 1, 2, and 3, respectively. The mean number
167	of observations with MS 2 or 3 was 32%, ranging from 24% in October 2008 to 40% in July

168 2008. The mean duration of lameness was 5.5 [s.e. 3] wk (median 4 wk, interquartile range 2 169 to 7 wk). Only 48% of scores remained unchanged from 1 score to the next, but cows were 170 unlikely to move more than 1 score in a 2-wk period. Once cows were a certain MS for 2 171 observations they were more likely to remain at that MS than change score. Patterns of scores 172 are in Table 2.

The milk yield was highest in cows with MS 1 (Table 3). Cows produced 0.7 kg/d and 1.6 kg/d less milk when MS 2 or 3, respectively, compared with cows with MS 1 (P <0.05). There was a reduction in yield from t – 3 before becoming MS 2 (0.47 CI (0.11 – 0.82) or MS 3 (0.9 (0.15 – 1.65) and t + 2 after recovering from MS 2 (0.85 CI 0.5 – 1.2).

177 First, second and third lactation cows were 58, 48, and 19%, respectively, less active (took fewer steps) than cows parity >3 (P < 0.05; Table 3). Cows were less active in early 178 179 lactation (mean log 1.38 steps/hr in month 1) and became more active as lactation progressed 180 (mean log 1.74 steps/hr in month 10), e.g., cows that were 9 months into lactation were 42% 181 more active than those in the first month of lactation (P < 0.05). Cows with MS 0 were 1% 182 less active than a cow with MS 1 (P < 0.05). Cows with MS 2 and 3 were 3 and 5 % less active than a cow with MS 1 (P < 0.05). Cows had a decreased activity for 42 d before being 183 184 MS 2 (mean 0.02 (CI 0.01 - 0.03)): they were 3% less active 2 wk before and 2% less active 185 4 wk before they became MS 2 compared with a cow with MS 1 (P < 0.05). Cows with MS 186 3 were less active from 28 d before they developed MS 3 (-0.02 CI (0.00 - 0.04)). Similarly, 187 cows that were MS 2 were less active by 3 to 4 % for the following 5 recordings and cows 188 that had MS 3 were less active by 3 to 6 % for the following recordings (P < 0.05).

A total 444 lesions (185/100 cows per yr) with 385 primary lesions on 258 feet were recorded by the herdsman, veterinarian, and foot trimmer. Over the 12 mo study period 178 cows (74%) were treated for at least 1 lesion; 72 (30%) cows had more than 1 foot with a lesion and 81 (31%) feet were treated more than once. The lesions recorded were digital

dermatitis (39%) sole ulcer (25%), white line disease (WLD) (12%), interdigital growth (9%),
and other (15%).

195	From the multistate model (Table 4) the longer the period a cow was not lame (i.e.,
196	not MS 2 or 3) the less likely she was to make a transition to being lame and the longer a cow
197	was lame the less likely she was to recover from being lame. Cows < 90 DIM were less likely
198	to become lame than cows \geq 90DIM (Odds Ratio (OR) = 0.66) and cows with milk yield >
199	15 to \leq 35 kg in the previous 16 d were less likely to recover from lameness (OR = 0.73) than
200	cows with milk yield > 35 kg.
201	Cows in parity 1 (OR = 0.49) or 2 (OR = 0.79) were less likely to become lame and
202	they were more likely to recover ($OR = 1.26$ and 1.32 , respectively) once they had become

they were more likely to recover (OR = 1.26 and 1.32, respectively) once they had become lame compared with cows of parity >2. Lame cows with 'other' lesions that were treated were less likely to recover from being lame (OR = 0.58) than untreated lame cows. Cows treated with a sole ulcer (OR = 1.35), digital dermatitis (OR = 1.51) or 'other' lesions (OR =1.39) were more likely to become lame again in comparison with non lame cows that had not been treated (Table 4).

208

DISCUSSION

209 In the current study, milk yield was reduced in cows with MS 2 or 3 for up to 4 to 8 wk 210 before their locomotion moved from MS 1. This period of time was considerably less than the 211 reduction in yield seen 3 to 4 mo before treatments reported by Green et al. (2002) and Amory 212 et al. (2008) and suggests that there was a delay in treatment in these 2 studies. If MS was used 213 to identify lame cattle and they were treated promptly the duration of both lameness and milk 214 loss might be reduced (Green et al., 2010). From the multistate model and patterns of MS 215 (Tables 4 and 2), treatment in the current study herd was not successful, with treated cattle 216 either not recovering (digital dermatitis) or being more likely to become lame again (sole ulcer 217 and other diseases). Note that WLD was not associated with lameness (Table 4) as in other

studies (Tadich et al., 2010). Repeated occurrences of lameness might indicate meager treatment strategy or efficacy, but might also indicate that treatment cannot address intrinsic factors such as a thin digital cushion. Treatment was added to the milk yield model; however, it did not alter the associations between yield and MS and so was excluded.

222 That cows with MS 1 had a lower milk yield for 4 to 8 wk before there was a change in 223 mobility score from MS 1 to MS 2 or 3 suggests that the reduction in yield occurred before 224 lameness was detectable. One possible explanation for the reduction in yield before MS changed is that MS was not sufficiently sensitive to detect the initial stages of disease. In 225 226 other studies of dairy cow lameness authors have reported lesions on sound cows (Manske et 227 al., 2002; Tadich et al., 2010; Bicalho et al., 2008). One hypothesis, drawing evidence from 228 Bicalho et al. (2009), is that lameness and foot lesions are positively associated with a thin 229 digital cushion which is associated with low body condition, this might cause sub clinical 230 disease that is not detectable externally or by MS, but is sufficiently painful to reduce food 231 intake, increase metabolic rate and so reduce milk yield. Low body condition per se could 232 also lead to reduced milk yield. It is unfortunate that we did not score the body condition of 233 the cattle in the current study but one could speculate that the cattle that moved from MS 1 to 234 MS 2 or 3 lost body condition before the transition whilst those that remained at MS 1 did not. 235 The fact that high yielding cattle at greater risk of lameness (Green et al., 2002; 236 Amory et al., 2008; Green et al., 2010) might help explain why cows with MS 1 produced 237 more milk than cows with scores 0, 2 or 3. These cows are producing high yields and their 238 locomotion is impaired (they are marginally lame). Over time, a proportion remain at MS 1 239 (Tables 2 and 5) and continue to produce high yields (Table 3) but some move to MS 2 or 3 240 and the pattern of lower yield and higher mobility score ensues. Once a cow is lame, she 241 might continue to have a further reduction in yield because extra energy is required to cope 242 with the pain of the foot lesion and energy is directed to this rather than milk production.

Depending on farm layout, lame cows might also feed less frequently and so reduce feed intake, exacerbating the disease process. If this was so, then successful treatment might increase mobility and stabilize milk yield, as seen in Green et al. (2010).

246 A large numbers of transitions in MS were seen between fortnightly scores for 247 individual cows in our study. In the UK farmers often MS their cattle annually or biannually 248 to comply with assurance scheme standards e.g. Tesco scheme, the current results suggest that 249 infrequent MS would give a snap shot of prevalence, but have little value in management of 250 lameness. Cows that had a MS of 2 or 3 had a high probability of remaining a 2 or a 3 (Table 251 2) and becoming lame again (Tables 2 and 4). The effects of this may be seen in terms of milk 252 production, but the effects on cow welfare are not so easy to quantify, although these cows did 253 have lower activity.. This suggests that prevalence, incidence, and repeat cases should be standard 254 recordings.

The results demonstrate that it is not only the MS on the day of recording that is important, but that the length of time that a cow has been at a particular MS is highly relevant. Our examples demonstrate that a cow that had been MS 2 for 6 wk lost 4.5 kg of milk per d while at MS 3 lost 6 kg/d of milk . These results support Juarez et al. (2003) who demonstrated a drop in milk yield of 4 kg/d for a lame cow. Extrapolating these results to a cow that is lame for 12 wk equates to 610 kg milk lost, supporting Amory et al. (2008).

Results from this herd suggest that activity data may not play a useful role in early identification of lameness because the absolute changes were so small: parity and stage of lactation had a much greater effect on activity than MS (Table 3). Cows became steadily more active as lactation progressed and with increasing parity, contrary to the findings of O'Callaghan et al. (2003) who reported a decreased level of activity as lactation progressed. The average change in activity associated with mobility score was less than 1%/d in our study, while they reported that cows that were lame were 24% less active than non lame cows.

Comment [FCG1]: Over what period of

There might be large variations in activity between herds, this might depend on the farm layout, and this might be very important when considering the necessary and unnecessary activity of cows.

271 The results suggest that a decrease in milk yield could have a role as an early indicator of 272 lameness, while change in activity is a less sensitive measure. In order to be practically 273 applied on farms, algorithms for milk yield, correcting for parity and stage of lactation, would 274 need to be incorporated into on-farm software alongside daily milk recording. In conjunction 275 with fortnightly MS this could alert the farmer that cows need early intervention. Before this 276 could be achieved, research needs to be repeated across many farms and systems to validate 277 the findings. In addition, unexpected reduction in milk yield might indicate that a cow is not 278 metabolically stable (Bicalho et al., 2009) and lameness is only one of the risks for such 279 cattle.

280 The advantage of this study was the large amount of detailed data that were collected. 281 This farm was chosen because it was similar to many farms in the UK with Holsteins 282 producing large quantities of milk under intensive conditions; the patterns within cow are 283 useful additions to our understanding of the associations between milk yield, MS, activity, and 284 lameness. A disadvantage of this study was that the data were from only 1 farm. It is not 285 possible to generalize prevalence, incidence, and transitions between MS. Whatever the 286 factors initiating lameness it appears that changing external management (Barker et al., 287 2007, 2009) is likely to be only part of the story to prevent lameness in dairy cows, possibly 288 explaining part of the limited success of intervention studies (Bell et al., 2007; Barker 2007). 289 Further work is required to elucidate when biochemical and pathological changes occur in 290 the development of lameness. If these changes can be identified, then we can move forward 291 in preventing lameness in dairy cows.

292

CONCLUSIONS

293	A reduction in mobility occurred 4 to 8 wk after cows had started to reduce milk
294	yield and an increase in milk yield occurred approximately 6 wk after a cow returned to
295	MS 0 or 1, suggesting that either mobility scoring is insufficiently sensitive to detect
296	lameness, that cattle mask lameness despite being diseased, or that a lameness and
297	reduction in yield are linked by a common intrinsic event. Once lame, cows were likely
298	to remain lame or become lame again, suggesting that either treatment was unsuccessful
299	or that the internal origin of lameness overrode treatment. Further work investigating
300	body condition, biochemical profiles, mobility, and lameness longitudinally could have a
301	huge impact on our understanding of the etiology of lameness.
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371

Mobility score	Definition	Description of cow mobility		
0	Good mobility / sound Walks with even weight bearing and rhythm on with a flat back. Long fluid strides possible.			
1	Imperfect mobility	Steps unevenly or shortened strides. Affected limbs not immediately identifiable.		
2	Impaired mobility	Uneven weight bearing on limb immediately identifiable and/or obviously shortened stride. Usually arched back.		
3	Severely impaired mobility	Unable to walk as fast as brisk human pace plus signs of score 2.		

Table 1. Definitions of mobility scores (Whay et al., 2003)

	376	Table 2.	. Transitions	in mobility	score from	time $t - 3$	to time t	where $t =$	14 d intervals
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377 illustrating that 50 - 60% of cows remain at a score for 8 weeks but that 40 - 50% cows move

- 378 mobility score

· · · · ·					4
t' - 3	t - 2	t - 1	t	Probability of	N^2 sequence
				score at t	observed
0	0	0	0	0.57	244
0	0	0	1	0.41	244
1	0	0	1	0.51	182
1	0	0	0	0.44	182
1	1	1	1	0.65	665
1	1	1	0	0.19	665
2	2	2	2	0.64	390
3	3	3	3	0.67	54
3	3	3	1	0.02	54
3	3	3	2	0.31	54
3	3	3	3	0.67	54
3	3	2	1	0.16	31
2	3	2	1	0.09	54
1	3	2	1	0.20	10

 1 t = time, t +/- i = time from / to t in 2 wk intervals 2 N = number of occasions,

Table 3. Random effects model of mean 16 d yield and 16 d mean log activity in 312 cows

384 from 1 dairy herd in Somerset, UK

	Mean	lower	upper	mean Log	lower 95%	upper 95%
	yield	95% CI ³	95% CI	activity	CI	CI
intercept	41.9	40.685	43.115	1.384	0.972	1.796
parity >3	referen	reference	reference	reference	reference	reference
	ce					
parity 1	-5.78	-7.113	-4.447	0.113	0.072	0.154
parity 2	-2	-3.078	-0.922	0.039	0.006	0.072
parity 3	-2.4	-3.282	-1.518	0.072	0.047	0.097
2nd lactation	-0.7	-1.366	-0.034	0.237	0.217	0.257
DIM	-0.05	-0.052	-0.048			
Wilmink	-15.7	-17.013	-14.387			
month in milk 1				reference	reference	reference
month in milk 2				0.033	0.011	0.055
month in milk 3				0.065	0.043	0.087
month in milk 4				0.078	0.056	0.100
month in milk 5				0.113	0.089	0.137
month in milk 6				0.125	0.101	0.149
month in milk 7				0.157	0.132	0.182
month in milk 8				0.205	0.180	0.230
month in milk 9				0.244	0.217	0.271
month in milk 10				0.304	0.277	0.331
month in milk 11				0.361	0.330	0.392
at t ¹						
MS^2 1	referen	reference	reference	reference	reference	reference
	ce					
MS 0	-0.45	-0.764	-0.136	-0.004	-0.014	0.006
MS 2	-0.66	-0.974	-0.346	-0.016	-0.026	-0.006
MS 3	-1.61	-2.237	-0.983	-0.025	-0.045	-0.005
at t+1						
MS 1	referen	reference	reference	reference	reference	reference
	ce					
MS 0	-0.76	-1.093	-0.427	-0.007	-0.017	0.003
MS 2	-0.43	-0.763	-0.097	-0.012	-0.022	-0.002
MS 3	-0.5	-1.147	0.147	-0.011	-0.031	0.009
at t+2						
MS 1	referen	reference	reference	reference	reference	reference
	ce		o 45 -	0.65-		
MS 0	-0.85	-1.203	-0.497	-0.005	-0.015	0.005

388 Table 3. Two level random effects model of mean 16 d milk yield and log activity in 312

389 cows from one herd in Somerset, UK continued

390

387

	Mean	lower 95%	upper	mean Log	lower 95%	upper 95%
	yield	CI	95% CI	activity	CI	CI
MS 2	-0.42	-0.753	-0.087	-0.002	-0.012	0.008
MS 3	0.26	-0.387	0.907	0.002	-0.018	0.022
at t+3						
MS 1	reference	reference	reference	reference	reference	reference
MS 0	-0.84	-1.212	-0.468	0.001	-0.011	0.013
MS 2	-0.26	-0.613	0.093	0.007	-0.003	0.017
MS 3	0.47	-0.196	1.136	0.009	-0.011	0.029
at t+4						
MS 1	reference	reference	reference			
MS 0	-0.65	-1.022	-0.278			
MS 2	-0.1	-0.453	0.253			
MS 3	0.28	-0.406	0.966			
at t-1		0.000	0.000			
MS 1	reference	reference	reference	reference	reference	reference
MS 0	-0.4	-0.733	-0.067	-0.005	-0.015	0.005
MS 2	-0.95	-1.283	-0.617	-0.015	-0.025	-0.005
MS 3	-2.67	-3.336	-2.004	-0.031	-0.051	-0.011
at t-2						
MS 1	reference	reference	reference	reference	reference	reference
MS 0	-0.44	-0.773	-0.107	-0.010	-0.020	0.000
MS 2	-0.69	-1.043	-0.337	-0.170	-0.180	-0.160
MS 3	-1.39	-2.096	-0.684	-0.019	-0.041	0.003
at t-3						
MS 1	reference	reference	reference	reference	reference	reference
MS 0	-0.25	-0.603	0.103	-0.013	-0.023	-0.003
MS 2	-0.47	-0.823	-0.117	-0.015	-0.025	-0.005
MS 3	-0.9	-1.645	-0.155	0.010	-0.225	0.245
at t-4						
MS 1	reference	reference	reference			
MS 0	0.09	-0.282	0.462			
MS 2	-0.41	-0.782	-0.038			
MS 3	0.31	-0.474	1.094			

391 I = time, t +/- i = time from / to t in 2-wk intervals

 $^{2}MS = mobility score$

 3 CI = confidence interval

Table 4: Multivariable multistate model of transitions between lame (mobility score 2 or 3)

and non lame (mobility score 0 or 1) states in 312 cows from 1 dairy herd observed for 1 yr in

396 Somerset, UK

Non lame to lame Lame to non lame variables intercept -5.15 0.21 -4.58 0.37 OR CI OR CI Duration spent in state - - - $\leq 2 wk$ 4.06 2.96-5.55 3.63 1.90-6.94.89 > 2-4wk 3.16 2.22-4.49 2.51 1.29-4.89 > 4-18 wk 1.80 1.32-2.47 1.93 1.01-3.69 > 18 wk reference reference - - DIM -			Tran	sition	
variables intercept -5.15 0.21 -4.58 0.37 OR CI OR CI Duration spent in state ≤ 2 wk 4.06 2.96-5.55 3.63 1.90-6.94 > 2.4 wk 3.16 2.22-4.49 2.51 1.29-4.89 > 4-18 wk 1.80 1.32-2.47 1.93 1.01-3.69 > 18 wk reference reference 0.90 0.66 0.57-0.78 1.25 0.93-1.67 91-180 1.00 0.79-1.26 1.15 0.91-1.46 0.84 0.69-1.02 92 wes 1.35 1.11-1.64 0.84 0.69-1.02 reference Past treatments Sole ulcer yes 0.51 1.29-1.76 0.86 0.72-1.03 no reference reference reference 0.65-1.05 0.65-1.05 no reference 0.91-1.46 0.83 0.65-1.05 0.45-0.74 no reference reference reference reference 0.45-0.		Nor	n lame to lame	Lan	ne to non lame
intercept -5.15 0.21 -4.58 0.37 OR CI OR CI Duration spent in state 2 $0R$ CI 2 wk 4.06 2.96-5.55 3.63 1.90-6.94 > 2-4wk 3.16 $2.22-4.49$ 2.51 $1.29-4.89$ > 4-18 wk 1.80 $1.32-2.47$ 1.93 $1.01-3.69$ > 18 wk reference reference 0.90 0.66 $0.57-0.78$ 1.25 $0.93-1.67$ 91-180 1.00 $0.79-1.26$ 1.15 $0.91-1.46$ 0.81 $0.69-1.02$ Past treatments Sole ulcer yes 1.35 $1.11-1.64$ 0.84 $0.69-1.02$ poistal dermatitis yes 1.51 $1.29-1.76$ 0.86 $0.72-1.03$ no reference reference $reference$ $reference$ Vhite line disease yes $0.51-1.05$ $0.45-0.74$ no reference $reference$ $reference$ Yes 0.87 $0.70-1.08$ 1.67 $1.34-2.07$ </td <td>variables</td> <td></td> <td></td> <td></td> <td></td>	variables				
OR CI OR CI OR CI Duration spent in state $\leq 2 \text{ wk}$ 4.06 2.96-5.55 3.63 1.90-6.94 $\geq 2.4\text{wk}$ 3.16 2.22-4.49 2.51 1.29-4.89 > 4-18 wk 1.80 1.32-2.47 1.93 1.01-3.69 > 18 wk reference reference 0.93-1.67 0-90 0.66 0.57-0.78 1.25 0.93-1.67 91-180 1.00 0.79-1.26 1.15 0.91-1.46 >180 reference reference 0.91-1.46 Past treatments Sole ulcer yes 1.51 1.29-1.76 0.86 0.72-1.03 no reference reference reference 0.65-1.05 0.65-1.05 no reference reference 0.65-1.05 0.45-0.74 no reference reference 0.65-1.05 0.45-0.74 no reference reference reference 0.72-1.03 No reference 0.58	intercept	-5.15	0.21	-4.58	0.37
Duration spent in state $\leq 2 wk$ 4.06 2.965.55 3.63 1.90-6.94 >2-4wk 3.16 2.224.49 2.51 1.29-4.89 >4-18 wk 1.80 1.32-2.47 1.93 1.01-3.69 > 18 wk reference reference 0.90 0.66 0.57-0.78 1.25 0.93-1.67 91-180 1.00 0.79-1.26 1.15 0.91-1.46 0.84 0.69-1.02 >180 reference reference reference 0.69-1.02 0.69-1.02 no reference reference 0.69-1.02 0.65-1.05 0.65-1.05 Digital dermatitis yes 1.51 1.29-1.76 0.86 0.72-1.03 no reference reference 0.65-1.05 0.65-1.05 0.65-1.05 no reference reference 0.65-1.05 0.65-1.05 no reference reference 0.65-1.05 No reference reference 0.51-0.74 no reference reference 0.51-0.74 no reference reference		OR	CI	OR	CI
$ \begin{array}{cccccccccccccccccccccccccccccccccccc$	Duration spent in state				
> 2.4 wk 3.16 2.22-4.49 2.51 1.29-4.89 > 4-18 wk 1.80 1.32-2.47 1.93 1.01-3.69 > 18 wk reference reference DIM 0-90 0.66 0.57-0.78 1.25 0.93-1.67 91-180 1.00 0.79-1.26 1.15 0.91-1.46 > 180 reference reference Past treatments Sole ulcer yes 1.35 1.11-1.64 0.84 0.69-1.02 no reference network reference 0.69-1.02 no reference reference 0.69-1.02 no reference 0.69-1.02 no reference 0.60-1.02 no reference 0.60-1.0	$\leq 2 \text{ wk}$	4.06	2.96-5.55	3.63	1.90-6.94
> 4-18 wk $> 1.80 $ $> 1.32-2.47 $ $1.93 $ $reference $ $Past wk $ $reference $ $Past treatments $ $Sole ulcer $ $yes $ $1.35 $ $1.11-1.64 $ $0.84 $ $0.69-1.02 $ $no $ $reference $ $1.29-1.76 $ $0.86 $ $0.72-1.03 $ $no $ $reference $ $0.91-1.46 $ $0.84 $ $0.69-1.02 $ $no $ $reference $ $0.91-1.46 $ $0.84 $ $0.69-1.02 $ $0.91-1.46 $ $0.86 $ $0.72-1.03 $ $0.91-1.46 $ $0.83 $ $0.65-1.05 $ $0.91-1.46 $ $0.83 $ $0.45-0.74 $ $0.91-1.16 $ $0.50-2.70 $ $1.5 $ $1.15 $ $0.89-1.48 $ $0.73 $ $0.54-1.22 $ $1.5-35 $ $1.15 $ $0.89-1.48 $ $0.73 $ $0.55-0.98 $ $35 $ $1.15 $ $0.89-1.48 $ $0.73 $ $0.55-0.98 $ $1.55 $ $1.15 $ $0.89-1.48 $ $0.73 $ $0.55-0.98 $ $1.55 $ $1.15 $ $0.89-1.48 $ $0.73 $ $0.55-0.98 $ $1.55 $ $0.55-0.98 $ $1.55 $ $0.55-0.98 $ $1.55 $ $0.55-0.98 $ $1.55 $ $0.85 $ $0.$	> 2-4wk	3.16	2.22-4.49	2.51	1.29-4.89
> 18 wk reference reference DIM 0-90 0.66 0.57-0.78 1.25 0.93-1.67 91-180 1.00 0.79-1.26 1.15 0.91-1.46 >180 reference reference 0.69-1.02 Past treatments Sole ulcer yes 1.35 1.11-1.64 0.84 0.69-1.02 no reference 1.29-1.76 0.86 0.72-1.03 no reference 0.91-1.46 0.83 0.65-1.05 No reference 0.58 0.45-0.74 0.45-0.74 No reference 0.87 0.70-1.08 1.67 1.34-2.07 Nean milk yield in	>4-18 wk	1.80	1.32-2.47	1.93	1.01-3.69
$\begin{array}{cccccccccccccccccccccccccccccccccccc$	>18 wk	reference		reference	
$\begin{array}{cccccccccccccccccccccccccccccccccccc$	DIM				
91-180 1.00 $0.79-1.26$ 1.15 $0.91-1.46$ >180 reference reference reference Past treatments Sole ulcer $0.91-1.46$ 0.84 $0.69-1.02$ no reference $1.11-1.64$ 0.84 $0.69-1.02$ Digital dermatitis yes 1.51 $1.29-1.76$ 0.86 $0.72-1.03$ no reference $91-1.46$ 0.83 $0.65-1.05$ No reference $92-1.48$ $0.45-0.74$ $0.65-1.05$ No reference $92-1.48$ 1.1	0-90	0.66	0.57-0.78	1.25	0.93-1.67
>180referencereferencePast treatments Sole ulcer yes1.351.11-1.640.840.69-1.02noreference1.351.11-1.640.840.69-1.02Digital dermatitis yes1.511.29-1.760.860.72-1.03noreference0.91-1.460.830.65-1.05Noreference0.91-1.460.830.65-1.05Noreference0.91-1.460.580.45-0.74Other yes1.391.10-1.760.580.45-0.74noreference1.671.34-2.07noreferencereferencereferencePregnant yes0.870.70-1.081.671.34-2.07noreferencereferencereferenceMean milk yield in previous 16 d0.900.46-1.801.160.50-2.70≤151.220.84-1.770.810.54-1.22>15-351.150.89-1.480.730.55-0.98	91-180	1.00	0.79-1.26	1.15	0.91-1.46
Past treatments Sole ulcer yes 1.35 1.11-1.64 0.84 0.69-1.02 no reference reference 0.69-1.02 Digital dermatitis yes 1.51 1.29-1.76 0.86 0.72-1.03 no reference 0.91-1.46 0.83 0.65-1.05 No reference 0.91-1.46 0.83 0.65-1.05 White line disease yes 1.15 0.91-1.46 0.83 0.65-1.05 Other yes 1.39 1.10-1.76 0.58 0.45-0.74 No reference reference 0.45-0.74 No reference 1.39 1.10-1.76 0.58 Other yes 0.87 0.70-1.08 1.67 1.34-2.07 No reference reference reference 1.34-2.07 No reference 0.87 0.70-1.08 1.67 1.34-2.07 No reference reference reference 1.34-2.07 No reference 0.81 0.65 0.50-2.70 ≤15 1.22 0.84-1.77 0.81 0.54-1.22 <	>180	reference		reference	
Sole ulcer .1.35 1.11-1.64 0.84 0.69-1.02 no reference reference 0.69-1.02 Digital dermatitis .1.51 1.29-1.76 0.86 0.72-1.03 no reference 0.91-1.46 0.83 0.65-1.05 No reference 0.91-1.46 0.83 0.65-1.05 White line disease 0.91-1.46 0.83 0.65-1.05 no reference 1.10-1.76 0.58 0.45-0.74 Other	Past treatments				
yes 1.35 $1.11-1.64$ 0.84 $0.69-1.02$ noreferencereference $reference$ $0.69-1.02$ Digital dermatitisyes 1.51 $1.29-1.76$ 0.86 $0.72-1.03$ noreference $reference$ $0.91-1.46$ 0.83 $0.65-1.05$ White line diseaseyes 1.15 $0.91-1.46$ 0.83 $0.65-1.05$ noreference $0.91-1.46$ 0.83 $0.65-1.05$ Otheryes 1.39 $1.10-1.76$ 0.58 $0.45-0.74$ noreferencereference $reference$ $0.45-0.74$ Pregnantyes 0.87 $0.70-1.08$ 1.67 $1.34-2.07$ noreference $reference$ $reference$ $1.34-2.07$ noreference 1.15 $0.84-1.80$ 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $0.54-1.22$ >15-35 1.15 $0.89-1.48$ 0.73 $0.55-0.98$ >35referencereference $reference$	Sole ulcer				
no reference reference Digital dermatitis 1.51 1.29-1.76 0.86 0.72-1.03 no reference reference 0.86 0.72-1.03 White line disease reference 0.91-1.46 0.83 0.65-1.05 yes 1.15 0.91-1.46 0.83 0.65-1.05 no reference reference 0.45-0.74 Other 1.39 1.10-1.76 0.58 0.45-0.74 no reference reference 0.45-0.74 no reference reference 0.45-0.74 no reference 0.58 0.45-0.74 no reference reference 0.51 Pregnant yes 0.87 0.70-1.08 1.67 1.34-2.07 no reference reference reference 1.34-2.07 no reference reference 1.05 1.20 No reference reference 1.10-1.76 1.34-2.07 no reference reference 1.10-1.76 1.10-1.76 1.10-1.76 <td< td=""><td>yes</td><td>1.35</td><td>1.11-1.64</td><td>0.84</td><td>0.69-1.02</td></td<>	yes	1.35	1.11-1.64	0.84	0.69-1.02
Digital dermatitis yes1.511.29-1.760.860.72-1.03noreferencereference0.72-1.03White line disease yes1.150.91-1.460.830.65-1.05noreference0.91-1.460.830.65-1.05Other yes1.391.10-1.760.580.45-0.74Noreferencereference0.45-0.74Pregnant yes0.870.70-1.081.671.34-2.07NoreferencereferencereferenceMean milk yield in previous 16 d0.900.46-1.801.160.50-2.70≤151.220.84-1.770.810.54-1.22>15-351.150.89-1.480.730.55-0.98≥35referencereferencereference	no	reference		reference	
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White line disease yes1.15 $0.91-1.46$ 0.83 $0.65-1.05$ noreference $1.10-1.76$ 0.83 $0.65-1.05$ Other yes 1.39 $1.10-1.76$ 0.58 $0.45-0.74$ noreference $1.10-1.76$ 0.58 $0.45-0.74$ Pregnant yes 0.87 $0.70-1.08$ 1.67 $1.34-2.07$ noreferencereference 1.67 $1.34-2.07$ noreference 1.67 $1.34-2.07$ noreference 1.67 $1.34-2.07$ noreference 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $>15-35$ 1.15 $0.89-1.48$ 0.73 >35 referencereference	no	reference		reference	
yes 1.15 $0.91-1.46$ 0.83 $0.65-1.05$ noreferencereference0Otheryes 1.39 $1.10-1.76$ 0.58 $0.45-0.74$ noreferencereference0Pregnantyes 0.87 $0.70-1.08$ 1.67 $1.34-2.07$ noreferencereferencereferenceMean milk yield inreferencereferencemissing 0.90 $0.46-1.80$ 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $0.54-1.22$ >15-35 1.15 $0.89-1.48$ 0.73 $0.55-0.98$ >35referencereference $reference$	White line disease				
noreferencereferenceOther yes1.39 $1.10-1.76$ 0.58 $0.45-0.74$ noreferencereference $0.45-0.74$ Pregnant yes 0.87 $0.70-1.08$ 1.67 $1.34-2.07$ noreferencereference $reference$ Mean milk yield in previous 16 d 0.90 $0.46-1.80$ 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $0.54-1.22$ >15-35 1.15 $0.89-1.48$ 0.73 $0.55-0.98$ >35referencereference	yes	1.15	0.91-1.46	0.83	0.65-1.05
Other yes1.391.10-1.760.580.45-0.74noreferencereference0.45-0.74Pregnant yes0.870.70-1.081.671.34-2.07noreferencereference1.34-2.07Mean milk yield in previous 16 d0.900.46-1.801.160.50-2.70 ≤ 15 1.220.84-1.770.810.54-1.22>15-351.150.89-1.480.730.55-0.98>35referencereference1.160.50-2.70	no	reference		reference	
yes 1.39 $1.10-1.76$ 0.58 $0.45-0.74$ noreferencereference $0.45-0.74$ Pregnantyes 0.87 $0.70-1.08$ 1.67 noreferencereferenceMean milk yield inprevious 16 dmissing 0.90 $0.46-1.80$ 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $0.54-1.22$ $>15-35$ 1.15 $0.89-1.48$ 0.73 $0.55-0.98$ >35 reference	Other				
noreferencereferencePregnant yes 0.87 $0.70-1.08$ 1.67 $1.34-2.07$ noreferencereferencereferenceMean milk yield in previous 16 d 0.90 $0.46-1.80$ 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $0.54-1.22$ >15-35 1.15 $0.89-1.48$ 0.73 $0.55-0.98$ >35referencereference	yes	1.39	1.10-1.76	0.58	0.45-0.74
Pregnant yes 0.87 $0.70-1.08$ 1.67 $1.34-2.07$ noreferencereferenceMean milk yield in previous 16 d 0.90 $0.46-1.80$ 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $0.54-1.22$ >15-35 1.15 $0.89-1.48$ 0.73 $0.55-0.98$ >35referencereference	no	reference		reference	
yes 0.87 $0.70-1.08$ 1.67 $1.34-2.07$ noreferencereferenceMean milk yield inprevious 16 dmissing 0.90 $0.46-1.80$ 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $0.54-1.22$ >15-35 1.15 $0.89-1.48$ 0.73 $0.55-0.98$ >35referencereference	Pregnant				
noreferencereferenceMean milk yield in previous 16 d $$	yes	0.87	0.70-1.08	1.67	1.34-2.07
Mean milk yield in previous 16 d 0.90 $0.46-1.80$ 1.16 $0.50-2.70$ ≤ 15 1.22 $0.84-1.77$ 0.81 $0.54-1.22$ $> 15-35$ 1.15 $0.89-1.48$ 0.73 $0.55-0.98$ > 35 referencereference	no	reference		reference	
$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	Mean milk yield in				
$\begin{array}{c ccccccccccccccccccccccccccccccccccc$	missing	0.00	0 /6 1 90	1 16	0 50 2 70
>15 1.22 0.04-1.77 0.01 0.34-1.22 >15-35 1.15 0.89-1.48 0.73 0.55-0.98 >35 reference reference	<15	1.20	0.40-1.00 0.94 1.77	1.10 A Q1	0.50-2.70
>15-55 1.15 0.07-1.40 0.75 0.03-0.96 >35 reference reference reference	<u></u> 15_35	1.22	0.04-1.//	0.01	0.54-1.22
	×15-55 ×35	1.1J	0.07-1.40	U.75 reference	0.33-0.98

Parity				
1	0.49	0.39-0.62	1.26	1.00-1.59
2	0.79	0.63-0.98	1.32	1.05-1.67
3	0.94	0.74-1.19	1.15	0.89-1.48
>3	reference	r	eference	