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# Footbathing, formalin and foot trimming

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1	Original article
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4	Footbathing, formalin and foot trimming: The 3Fs associated with granulomas and
5	shelly hoof in sheep
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#### 21 Abstract

Granulomas and shelly hoof (SH), are lesions of sheep feet. Our objective was to use data from four questionnaires on lameness sent to English sheep farmers in 2004, 2013, 2014 and 2015 to further understanding of the risks and aetiologies of both lesions.

25

26 Granulomas were more likely in flocks where routine foot trimming (odds ratio [OR] 27 = 3.17; 95% confidence intervals [CI] 1.11 - 11.47) and routine footbathing (OR = 2.38; 95%) 28 CI (1.19 - 4.83) were practised than where these management protocols were not. SH was 29 more likely in flocks that were footbathed in formalin compared with not footbathing (OR = 30 1.65; 95% CI 1.19 - 2.30), and was less common in flocks that stocked ewes at more than 31 eight vs. four per acre (OR = 0.34; 95% CI 0.17 - 0.68). There were weak associations 32 between SH and foot trimming. In 2004 only, SH was more likely in flocks where therapeutic 33 foot trimming was practised than not practised (OR = 2.24; 95% CI 1.12 – 4.68). In 2014 34 only, SH was marginally less likely in flocks where no feet bled during trimming, compared 35 with flocks not routinely trimmed (OR = 0.55; CI 0.30- 1.00); SH was not related to foot trimming once severe footrot was included. We propose that flocks with granulomas and SH 36 37 would decrease if farmers stopped footbathing in general, in particular with formalin, and 38 avoided foot trimming whether as a therapeutic or routine practice. Further work is needed to 39 understand the role of stocking density.

40

*Keywords:* Contagious ovine digital dermatitis; Footrot; Lameness prevalence; Multivariable
models; Sheep flock management

#### 43 Introduction

44 Lameness in sheep is one of the most important farm animal welfare concerns in the UK (Goddard et al., 2006). It is estimated to cost the sheep industry £24 to £80 million per 45 46 annum (Nieuwhof and Bishop, 2005; Wassink et al., 2010b; Winter and Green, 2017) due to 47 its significant impact on productivity (Marshall et al., 1991; Glynn, 1993; Wassink et al., 48 2010b). Footrot (both interdigital dermatitis and severe footrot) is the most common cause of lameness in the United Kingdom. It is present in >95% flocks and causes approximately 70% 49 50 of lameness (Winter et al., 2015). Contagious ovine digital dermatitis (CODD), another 51 infectious cause of lameness, is present in >50% flocks and causes approximately 30% - 35% 52 of lameness in affected flocks (Angell et al., 2014; Winter et al., 2015). 53 54 Granulomas and shelly hoof (SH) are less prevalent causes of lameness in sheep. 55 They are present in 50% - 60% flocks and farmers generally attribute <3% lameness/lesions 56 to these conditions (Kaler and Green, 2008a; Winter et al., 2015). A granuloma is a 57 proliferation of highly vascularised connective tissue that protrudes through disrupted hoof horn at the wall horn junction or through sole horn (Fig.1). It is very painful and sheep are 58 59 severely lame. Granulomas cause chronic lameness and sheep that have been lame for a long 60 period of time, and not bearing weight on the affected foot, often present with the wall horn 61 grown over the sole with the granuloma hidden from view. Granulomas are very difficult to 62 cure and often lead to premature culling of sheep (Winter, 2008). A professional review 63 suggests that the proliferation is a pathological response to trimming into the sensitive tissue 64 of the foot or as a result of untreated severe footrot (Winter, 2008). 65

66 Shelly hoof (SH) presents grossly as detachment of the hoof horn wall, typically the 67 abaxial wall, from the underlying epidermis, leaving a cavity between the wall and the sole

68 horn (Fig.1). SH per se does not cause lameness (Winter et al., 2015), however the impaction 69 of debris in the cavity formed by the detached horn, followed by penetration of a foreign 70 body, such as a thorn or stone, into sensitive tissue can lead to formation of an abscess and 71 then sheep become severely lame (Winter, 2008). Conington et al. (2010a) reported that wall 72 horn in cases of SH presents histologically as poorly keratinised epithelial horn cells with 73 microfissures and 'unzipping' of cell membranes that create microscopic crevices leading 74 deep into the horn. The authors hypothesised that this characteristic structure makes the hoof 75 physically weak, more permeable to moisture and highly vulnerable to entry of water-soluble 76 solutions and organisms that could further contribute to hoof breakdown. Conington et al. 77 (2010a) also reported a genetic component to SH. A professional review hypothesised that 78 physical damage, opportunistic bacteria/fungi or nutritional deficiency may cause SH (Winter 79 and Arsenos, 2009). While many farmers report low SH prevalence, some farmers report 80 high prevalence (Conington et al., 2010b) and the disease is of considerable concern for some 81 farmers (personal observation).

82

83 There is a body of published evidence that routine foot trimming and routine 84 footbathing are associated with a higher prevalence of all lameness. Treating individual sheep 85 lame with footrot rather than incorporating foot trimming and footbathing into routine flock 86 management protocols reduces the prevalence of footrot (Kaler and Green, 2009; Winter et 87 al., 2015; Witt and Green 2018). The only evidence of benefit from footbathing is for the 88 treatment of interdigital dermatitis (Clifton and Green, 2016). Which factors are effective in 89 the control of CODD are not well understood. However, recent studies indicate that CODD is 90 typically introduced by newly introduced sheep and that once it is endemic in flocks, CODD 91 can be controlled effectively using the same management protocols that control footrot

92	(Angell et al., 2016; Dickins et al., 2016). To the authors' knowledge, there are no published
93	studies reporting risk factors for granulomas and SH.
94	
95	The aim of the current study was to use data from four questionnaires sent to English
96	sheep farmers in 2004, 2013, 2014 and 2015 to investigate associations between management
97	of lameness and farm characteristics with the presence of granulomas and SH to generate
98	hypotheses on the risks for, and possible aetiologies of, these lesions.
99	
100	Materials and methods
101	Ethical approval
102	All questionnaire data used in this paper originates from studies that had ethical
103	approval from when this was required by the University of Warwick's Biomedical and
104	Scientific Research Ethics Committee (BSREC). For the 2013-2014 questionnaires this was
105	BSREC 159-01-12; approved 07 December, 2011. For the 2015 questionnaire this was
106	BSREC REG0-2014-620; approved 02 April, 2014. Ethical approval was not required in
107	2004.
108	
109	Questionnaires and responses
110	In 2004, a postal questionnaire was sent to a random sample of 3000 English sheep
111	farms selected from lists using the RAND function in Microsoft Excel and obtained from the
112	Agriculture and Horticulture Development Board (AHDB) stratified by region of England
113	(South west, South east, Central, North west and North east) and by flock size within each
114	region (Kaler and Green, 2008a, 2009). The questionnaire contained a photograph and a
115	description of granuloma and shelly hoof (Fig.1). Farmers were asked to identify each lesion

and estimate its prevalence in their flock, as well as answering questions on the prevalence of

117 lameness and management of lameness; approximately 6% of the target population was 118 surveyed (Kaler and Green, 2008a, 2009). In 2013, similar questions were posed in a 119 questionnaire sent to 4000 lowland sheep farmers in England with >199 ewes; lists were 120 obtained from the Department for Environment, Food and Rural Affairs (DEFRA) and 121 AHDB who selected flocks randomly stratified by county and size (Winter et al., 2015). In 122 2004 and 2013, randomization to select farms was achieved by assigning each farm a number 123 using the number autofill function in Microsoft Excel and then generating a second random 124 list of numbers in the same number range and corresponding to the number of farms required 125 using the RAND function in Microsoft Excel. Farms were assigned to the questionnaire 126 group if the number assigned to them was generated in the random number list. The lists for 127 the 2013 questionnaire may have included some of the 809 respondents from 2004. In 2014, 128 an abridged version of the 2013 questionnaire was sent to 1355/4000 compliant farmers who 129 responded to the 2013 questionnaire (Grant et al., 2018). Finally, in 2015, 722 compliant 130 farmers who responded to the 2014 questionnaire were invited to participate in a study which 131 included completing a questionnaire; 192/722 agreed to participate and 144 (75%) completed the questionnaire (Prosser et al., unpublished data). A further 18 farmers from the 2014 study 132 133 who participated in a clinical trial (Witt and Green, 2018) and an additional five farmers from 134 another clinical trial (Monaghan et al., unpublished data) also completed the questionnaire, 135 resulting in 167 responses to the 2015 questionnaire. Consequently, farmers who responded 136 to the 2015 questionnaire had also responded to the questionnaires in 2014 and 2013 and 137 farmers who responded to the 2014 questionnaire had also responded to the 2013 138 questionnaire. Data from 2004 were from a separate random sample of English sheep farmers 139 selected from lists using the RAND function in Microsoft Excel with unknown overlap in 140 responses from farmers in 2004 and other years. Data were collected on granulomas in 2004 141 and 2015 and SH in 2004, 2013 and 2014. Farmer responses to questions in Fig.1 were used

to identify flocks where granulomas and SH were present and to estimate the percentage of
granulomas and SH that farmers observed. Farmers were asked whether they practised
therapeutic and routine foot trimming in all four questionnaires; in addition, for 2013-2015,
the proportion of sheep that bled when trimmed was requested. Careful trimming is defined
here, as in Winter et al. (2015), as trimming where no sheep in a flock bled when routinely
trimmed.

- 148
- 149 Data preparation and analysis

150 All four datasets were clean because they had been used in previous analyses (Kaler 151 and Green, 2008a, 2009; Winter et al., 2015, Grant et al., 2018; Prosser et al., unpublished 152 data). The data were stored in Microsoft Excel. Responses were excluded if there were no 153 data on the presence of granulomas or SH. Explanatory variables were selected from the 154 questionnaires to test the following hypotheses. For granulomas, treatment variables e.g. 155 treated footrot with foot trimming, were used to test the hypothesis that management 156 protocols used for footrot were associated with presence of granulomas. Farm characteristics, 157 e.g. stocking rate, were used to test the hypothesis that certain types of farms had a higher 158 risk of granulomas. In addition, for SH, biosecurity variables e.g. the flock mixed with other flocks, were used to test the hypothesis that SH is an infectious disease (Supplementary 159 160 Tables 1 and 2, respectively).

161

The global mean period prevalence of granulomas, SH and all lameness were estimated and prevalence of lameness in flocks with and without granulomas and SH were calculated. Previous work has reported that farmers recognise lameness (Kaler and Green, 2008b; King and Green, 2011) and foot lesions, but do not necessarily identify lesions correctly (Kaler and Green, 2008a). Consequently, the prevalence of granulomas and SH by

167	correct (identifying the lesion in the photograph; Fig.1) and incorrect (identifying the lesions
168	as anything else, but most commonly footrot) identifying the lesions were investigated using
169	Mann Whitney tests. Binomial logistic regression models in R version 3.4.1 <sup>1</sup> (R Core Team
170	2018) were used to investigate univariable and multivariable associations between the
171	presence of granulomas and SH and management protocols and farm characteristics. The data
172	from 2004 and 2015 were used to investigate granulomas and data from 2004, 2013 and 2014
173	were used to investigate SH; each year was modelled separately. The models took the
174	following form:

....

175  $y_i \sim \alpha + \beta_i X_i + e_i$ 

176 where  $y_j$  is the probability that a flock has granulomas / SH, ~ is a logit link function,  $\alpha$  is 177 the intercept and  $\beta_j$  represents the series of vectors of coefficients of explanatory variables 178 for  $X_j$  that vary by farm *j* and  $e_j$  is the residual random error that follows a binomial 179 distribution.

180

The results were assessed using Wald's test for significance, that is 95% confidence 181 182 intervals did not include unity. Variables significant at P < 0.05 in the univariable analysis 183 were tested in the multivariable model using manual forward stepwise selection. All variables 184 were then retested using manual forward stepwise selection regardless of their significance in 185 the univariable analysis to check for residual confounding (Cox and Wermuth, 1996) and 186 were included in the model if significant. Where two variables were highly correlated, the 187 most biologically relevant was kept in the model. Presence of other lesions (interdigital 188 dermatitis, severe footrot and CODD) in the flock were tested in the models to investigate 189 residual confounding. Correlations between the variables in the multivariable models and

<sup>&</sup>lt;sup>1</sup> See: R Core Team, 2018. R: A language and environment for statistical computing. R Foundation for Statistical Computing. <u>https://www.R-project.org</u> (accessed 30 May 2019).

190 between lesions were investigated using chi-square tests. The model fit was investigated by

191 visually comparing the ranked expected values against the observed values.

192

193 **Results** 

194 Farmer response proportions

195 Granulomas

196 Of the 809 (27%) and 922 (68%) farmers that responded to questionnaires in 2004 197 and 2015, 89-95% answered questions regarding granulomas and flock size. Granulomas 198 were reported in 54-63% flocks (Supplementary Table 3). The median flock size was 220 199 (interquartile range [IQR], 90-450) in 2004 and 400 (IQR, 223-600) in 2015. The geometric 200 mean (95% confidence intervals [CI]) farmer reported prevalence of lameness was 8.1% (8.0-201 8.1%) in 2004 and 4.1% (3.7-4.7%) in 2015. The prevalence of granulomas was right skewed 202 meaning the distribution was positive on the x axis with a long tail to the right. Flock 203 prevalence in affected flocks ranged from 0.0-8.0% (upper quartile range, 1.0-8.0% in 2004; 204 1.2-7.0% in 2015; Supplementary Table 3). 205

206 Shelly hoof (SH)

207 There were 809 (27%), 1354 (34%) and 922 (68%) responses from farmers who 208 received a questionnaire in 2004, 2013 and 2014, respectively. Of these, 84-86% answered 209 questions on shelly hoof and flock size, with 58-76% reporting shelly hoof in their flock 210 (Supplementary Table 3). The median flock size ranged from 225 ewes (IQR, 95-460) in 211 2004, to 340 ewes (IQR, 220-540) in 2013 and 320 ewes (IQR, 220-518) in 2014. The 212 geometric mean (95% CI) farmer reported prevalence of lameness was 6.9% (6.4-7.3%), 213 3.7% (3.3-3.9%) and 3.3% (3.1-3.5%) in 2004, 2013 and 2014, respectively. In flocks with 214 SH, the prevalence of lesions was mostly <5%, however, in each year, some flocks had very

high prevalence. The upper quartile flock prevalence range was 3.8-37.5% in 2004, 5.070.0% in 2013 and 5.0-95.0% in 2014 (Supplementary Table 3).

217

218 Farmer identification and prevalence of granuloma and shelly hoof

219 Granuloma

In 2004 and 2015, 56% (n=288/516) and 98% (n=155/158) of farmers identified granulomas correctly from a photograph (Fig. 1). There was no significant difference in median flock prevalence of lameness by farmer identification of granuloma (Table 1).

223

224 Shelly hoof

In 2004, 2013 and 2014, 28% (*n*=134/477), 73% (*n*=787/1079) and 76% (*n*=605/800) farmers, respectively, identified SH correctly from the photograph (Fig.1). Only 55% (*n*=356/642) farmers correctly identified SH in both 2013 and 2014; 7% (*n*=43/642) identified SH incorrectly in both years, and 37% farmers identified SH correctly once in 2013 or 2014. However, the median prevalence of SH was not significantly different between flocks where farmers correctly and incorrectly identified SH and identification of SH was not significantly associated with flock prevalence of lameness (Table 1).

232

233 Univariable and multivariable binomial logistic regression models of factors associated with
234 the presence of granulomas and SH

235 Granulomas

The univariable associations between management variables and presence of granulomas in 2004 and 2015 are listed in Supplementary Table 1 and the multivariable models are presented in Table 2. In 2004, granulomas were more likely to be present in flocks where routine footbathing (odds ratio [OR] = 1.71; 95% CI 1.26 – 2.32) was practised 240 compared with those where routine footbathing was not practised and in flocks where farmers 241 practised therapeutic foot trimming of lame sheep (OR = 2.07; 95% CI 1.03 – 4.32) compared with flocks where farmers did not practise therapeutic foot trimming. In 2015, 242 243 granulomas were more likely to be present in flocks where routine footbathing (OR = 2.38; 244 95% CI 1.19 – 4.83) and routine trimming (OR = 3.17; 95% CI 1.11 – 11.47) were practised 245 compared with flocks where these management protocols were not practised. When other 246 foot lesions were added to the models, only CODD in 2015 was significantly associated with 247 the presence of granulomas; all management variables remained significant in this model 248 (data not shown).

249

250 Shelly hoof

251 The univariable associations between management variables and the presence of SH 252 in 2004, 2013 and 2014 are listed in Supplementary Table 2 and the multivariable models are 253 in Table 3. Approximately 94% farmers practised therapeutic foot trimming in 2004 (Table 254 4) and SH was significantly more likely to be present in flocks where farmers practised 255 therapeutic foot trimming compared with flocks where this was not practised (OR = 2.24; 256 95% CI 1.12 – 4.68). Approximately 65% farmers practised routine footbathing of sheep (Table 4) and SH was more common in flocks where footbaths were used compared to where 257 258 they were not used (OR = 1.81; 95% CI 1.33 - 2.46). In 2013, SH was more likely to be 259 present in flocks where sheep were routinely footbathed, and this was significantly associated 260 with flocks footbathed in formalin (OR = 1.65; 95% CI 1.19 – 2.30) compared with flocks never footbathed or footbathed with other products. The likelihood of SH was lower as 261 262 stocking density increased from fewer than four to more than eight ewes per acre (OR = 0.34; 95% CI 0.17 - 0.68). In 2014, routine foot trimming was associated with increased 263 likelihood of SH (data not shown). However, SH was less likely to be present (Wald's P =264

265 0.047; OR = 0.55; 95% CI 0.30 – 1.00) in flocks where farmers reported that no feet bled 266 during trimming (these were carefully, but not excessively, trimmed), compared with flocks 267 that were not routinely trimmed. When other foot lesions were tested in the models, ID, 268 CODD and severe footrot were significantly associated with the presence of SH in 2004, 269 2013 and 2014. The only change in the management variables was in 2014 when careful foot 270 trimming became non-significant (P = 0.44). 271

272 Correlations between explanatory variables in the multivariable models and with footrot and273 CODD

There were significant ( $P \le 0.05$ ) correlations between variables in the multivariable models. Typically, there were positive correlations between routine and therapeutic foot trimming and footbathing. Stocking density was positively associated with footbathing and positively and negatively associated with foot trimming in different models (Supplementary Table 4). All lesions that were recorded in a single year were positively correlated with each other.

280

281 Model fit

The model fits were reasonable, with predicted values following the pattern of observed for both granulomas and SH (Supplementary Figs.1-5).

284

### 285 Discussion

This is the first study to provide evidence that the associations between therapeutic foot trimming and routine footbathing and all lameness (Kaler and Green, 2009; Winter et al., 2015) are associated with specific lesions, granulomas and SH, in addition to footrot

(Wassink et al., 2003; Kaler and Green, 2009) and CODD (Dickins et al., 2016). These
findings are discussed in detail below.

291

292 Granuloma data was available from 2004 and 2015. These data came from two 293 independent randomly selected samples of English sheep farmers, the degree of overlap in 294 farmers between these two samples is unknown, but it is likely to be low given a population of >35,000 sheep farmers in England<sup>2</sup>. Additionally, given the time interval of 13 years, even 295 296 if some of the farmers were the same, practices changed (Winter et al., 2015), so it seems 297 reasonable to assume that the datasets were independent. Consequently, the consistency 298 between results for both years (Table 2) provides confidence in their robustness. We 299 hypothesised that foot trimming could damage the foot structure and so lead to proliferation 300 of connective tissue i.e. a granuloma. The results indicate that routine and therapeutic foot 301 trimming are associated with the presence of granulomas (Table 2) and support the 302 hypothesis that damaging connective tissue in the foot is associated with trimming. In both 303 questionnaires, therapeutic and routine foot trimming were positively correlated and so the 304 presence of one or the other in the final models is in part because the degree of correlation 305 was sufficient to exclude the other. There is insufficient statistical power in the 2015 dataset 306 to explore whether excessive routine foot trimming vs. all routine foot trimming, was 307 associated with granulomas. Despite this, we conclude that granulomas are associated with 308 the practice of foot trimming. Granulomas were also more likely to be present in flocks that 309 were footbathed in both datasets. It is not clear why this might be. However, if formalin was 310 the main footbathing product in 2004 and 2015, as it was in 2013 (73% farmers; Winter et al., 311 2015), and formalin can cause neoplasia and damage cells (see below), then either reduced

<sup>&</sup>lt;sup>2</sup> See: AHDB, 2018. UK Sheep Yearbook 2018. Agriculture and Horticulture Development Board. <u>http://beefandlamb.ahdb.org.uk/wp-content/uploads/2018/10/UK-Sheep-Yearbook-2018.pdf</u> (Accessed 14 March 2019).

312 regrowth of excessively trimmed hoof horn, or proliferation of digit connective tissue, or313 both, could lead to granuloma. This requires further study.

314

315 SH data were available for 3 years, 2004 and 2013 and 2014; respondents from 2014 316 were a subset of those who replied in 2013. In all 3 years, routine footbathing was associated 317 with increased risk of SH, providing a consistent association. In 2013, we identified increased 318 risk associated with formalin footbaths. Formalin is carcinogenic, causing nasopharyngeal 319 cancer in animals and humans (Swenberg et al., 1980; Hauptmann et al., 2004) and at 320 concentrations >5% in footbaths it causes skin inflammation in cattle (Cornelisse et al., 1982) 321 and sheep (Ross, 1983), and hard keratinous-like material in the interdigital skin (Pyror, 322 1959). Formalin is also used post mortem to fix tissues. It acts by dehydrating cells which 323 harden and form cross-links to create an insoluble meshwork (Thavarajah et al., 2012). The 324 combination of inflammation of living skin and tough dead cells in the outer layers of skin 325 and horn could exacerbate the micro- and macro-disintegration of the white line observed in 326 SH (Conington et al., 2010a). The association between SH and formalin could possibly be 327 reverse causality, with farmers with SH in their flock using formalin footbaths. However, 328 footbaths are not a recommended treatment for SH and farmers rarely change their behaviour 329 (Wassink et al., 2005; Wassink et al., 2010a), so it is likely that these farmers had been using 330 formalin footbaths for many years and are still reporting SH.

331

The associations between foot trimming and SH were inconsistent; therapeutic foot trimming was a risk in 2004 but not in other years and all routine foot trimming was not associated with SH in any study, although careful routine foot trimming with no bleeding, was only associated with reduced risk in 2013, but this association was confounded by the presence of severe footrot in the flock. Many observational and interventional studies have

reported that foot trimming is detrimental. It increases the flock prevalence of all lameness
(Kaler and Green, 2009; Winter et al., 2015), severe footrot (Wassink et al., 2003) and
CODD (Dickins et al., 2016), delays healing (Kaler et al., 2010) and increases the
reoccurrence of footrot (Kaler et al., 2010). We consider that the inconsistent and weak
association and confounding associated with the presence of severe footrot indicates that foot
trimming is not a potential cause of SH and that the recommendation to avoid foot trimming
sheep is still robust.

344

345 In our study, SH was more likely to occur in flocks at stocking densities of fewer than 346 four ewes per acre compared with more than eight ewes per acre. This is consistent with SH 347 being more common in flocks on poor quality pasture. Winter and Arsenos (2009) suggested that physical damage was a risk factor for SH. Features of poor quality pasture include 348 349 uneven surfaces, stones, or very dry or wet pasture, which might increase the risk of physical 350 damage to the white line and predispose to formation of SH. The presence of farm tracks are 351 a risk factor for white line disease in dairy cattle (Barker et al., 2009) and so give some 352 credence to this hypothesis by analogy (Bradford Hill, 1965). Additionally, if the grass was 353 poor quality, sheep could be malnourished and develop poor quality horn, as hypothesised by 354 Conington et al. (2010a). Sheep kept at low stocking densities are often different breeds from 355 those farmed at higher stocking densities; this might indicate breed differences in 356 susceptibility to SH, as well as the within-breed differences reported by Conington et al. 357 (2010b). However, there are a plethora of other management factors associated with low vs. 358 high stocking densities that could act as risk factors for SH.

359

360 There were no significant associations and varying patterns of positive and negative
 361 coefficients between biosecurity variables and granulomas and SH between the years of study

362 (Supplementary Tables 1 and 2). This indicates that there were no strong or consistent risks 363 with biosecurity variables and we hypothesise that granulomas and SH are unlikely to be of infectious origin, introduced into flocks by incoming sheep. Additionally, the distribution of 364 365 granulomas and SH within flocks provides further clues, as many flocks did not have lesions, 366 most affected flocks had very low prevalence and only a few had very high prevalence (Kaler 367 and Green, 2008a). This is not typical of infectious disease and suggests that granulomas and 368 SH are non-infectious diseases and that where there is a high prevalence there is a farm-369 specific risk.

370

371 In studies by Conington et al. (2010a; 2010b), the prevalence of SH in 27 flocks 372 ranged from 0-76%. In the current study, the range of SH was 0% to 37-95%, indicating there 373 were some flocks with a similar prevalence of SH to those reported by Conington et al. 374 (2010a; 2010b), but many hundreds with low prevalence. The within-flock prevalence of 375 lesions was a farmer estimate from recollection of foot examinations. It is therefore likely to 376 be an underestimate of the true prevalence of SH, since the farmers are unlikely to have 377 examined all feet simultaneously. Even if the prevalences reported here and elsewhere are not 378 representative of the within-flock prevalence of granulomas and SH across the population of 379 English sheep flocks, our data are robust enough to estimate the risk of granulomas and SH in 380 flocks, as long as the presence of a lesion did not bias farmers' responses. This seems 381 unlikely but cannot be determined from our data. When findings were consistent across the 382 questionnaires, the robustness of the results is greater. However, because farmers from 2013-383 2015 were the same, the robustness is less than for independent studies. 384

385 Conclusions

386	This is the first study to associate footbathing, formalin and foot trimming with
387	granulomas and SH in sheep flocks. These management protocols were associated with
388	reduced risk of SH, but less consistently. We conclude that the current recommended
389	management protocols of avoiding footbathing and foot trimming to minimise lameness and
390	footrot and CODD also contribute to reduced risk of granulomas. Further work is required to
391	understand whether granulomas are more likely to occur when formalin is used and to
392	investigate the role of formalin footbaths and low stocking density on the occurrence of SH.
393	
394	Conflict of interest statement
395	None of the authors has any other financial or personal relationships that could
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403	partner.
404	
405	Appendix A: Supplementary material
406	Supplementary data associated with this article can be found, in the online version at doi:
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540 Median (interquartile range) flock prevalence of lameness, granuloma and shelly hoof (SH)
541 identified by farmers from questionnaires completed by sheep farmers in England in 2004,

542 2013, 2014 and 2015.

	Granuloma	Granuloma		SH	SH				
Year	correctly	incorrectly	$P^{a}$	correctly	incorrectly	$P^{a}$			
	identified	identified		identified	identified				
Lameness %									
2004	8 (5-10)	8 (5-10)	0.88	10 (5-15)	10 (5-15)	0.25			
2013	-	-		4 (2-5)	4 (2-6)	0.88			
2014	-	-		3 (2-5)	3 (2-5)	0.53			
2015	5 (2-7)	15 (1.5-15)	0.28 -		-				
		(	Granuloma	%					
2004	0.25 (0.0-1.0)	0.5 (0.0-1.1)	< 0.01	0.6 (0.3-1.1)	0.8 (0.3-1.5)	0.72			
2013	-	-		-	-				
2014	-	-		-	-				
2015	0.9 (0.0-1.0)	5 (0.0-10.0)	0.68	-	-				
		S	Shelly hoof	%					
2004	0.6 (0.0-2.0)	0.5 (0.0-2.1)	0.40	10 (5-40)	20 (10-50)	0.03			
2013	-	-		2.0 (1.0-5.0)	3.0 (1.0-5.0)	0.78			
2014	-	-		2.0 (1.0 -5.0)	3.0 (1.4-5.0)	0.11			
2015	-	-		-	-				

544 - Data not collected

543

<sup>a</sup> *P* calculated from Mann-Whitney tests comparing prevalence of SH and granulomas by
 correct and incorrect farmer identification of lesions.

548 Binomial regression models of factors associated with the presence of granulomas in 2004549 and 2015

Model term	Category	Number of	%	OR	95% CI	
		flocks				
2004						
Footbathing <sup>a</sup>	No	265	34.42	1.00		
	Yes	500	64.94	1.71	1.26	2.32
Trim to treat sheep	No	35	4.55	1.00		
lame with footrot	Yes	723	93.90	2.07	1.03	4.32
2015						
Routine footbathing <sup>a</sup>	No	68	43.04	1.00		
	Yes	90	56.96	2.38	1.19	4.83
Routine foot trim	No	131	82.91	1.00		
	Yes	27	17.09	3.17	1.11	11.47

550

551 OR, odds ratio; 95% CI, 95% confidence interval

<sup>a</sup> Footbathing product data not collected

554 Binomial regression models of factors associated with the presence of shelly hoof (SH) in

# 555 sheep flocks in 2004, 2013 and 2014

Model term	Category	Flocks,	%	OR	95% CI	
		n				
2004						
Footbathing <sup>a</sup>	No	261	34.71	1.00		
	Yes	486	64.63	1.81	1.33	2.46
Trim to treat sheep lame	No	35	4.65	1.00		
with footrot	Yes	705	93.75	2.24	1.12	4.68
2013						
Ewe stocking rate	<4/acre	385	42.73	1.00		
	4-8/acre	451	50.06	0.75	0.55	1.02
	>8/acre	36	4.00	0.34	0.17	0.68
Footbathing <sup>a</sup>	No	341	37.85	1.00		
	Yes: formalin used	411	45.62	1.65	1.19	2.30
	Yes: formalin not	145	16.09	1.38	0.90	2.16
	used					
2014						
Routine foot trim	No trim	295	48.92	1.00		
	Trim: no bleeding	53	8.79	0.55	0.30	1.00
	Trim: bleeding	244	40.46	1.13	0.78	1.64

556

557 OR, odds ratio; 95% CI, 95% confidence interval

- <sup>a</sup> Footbathing with any product was associated with significantly higher OR for shelly hoof in
- 559 2013; footbathing product data not collected in 2004 and 2014

561 The number and percentage of flocks by management protocols for lameness in England in

Variable	2004 ( <i>n</i> =7:	2004 ( <i>n</i> =752)		2013 ( <i>n</i> =901)		2014 ( <i>n</i> =603)		2015 (n=158)	
	n	%	п	%	n	%	n	%	
Trimmed feet as footro	t treatment								
No	35	5					62	39	
Yes	705	94					88	56	
Never			38	4	28	5			
Always			345	38	110	18			
Sometimes			161	18	257	43			
Usually			292	32	174	29			
Routine foot trimming	practised								
No	173	23	395	44	295	49	131	83	
Yes	572	76	487	54	303	50	27	17	
Never									
Bled during routine									
trimming									
No trimming			395	44	295	49			
No bleeding			60	7	53	9			
Bleeding			408	45	244	40			
Used footbaths									
No	261	35	341	38	248	41	68	43	
Yes	486	65			355	59	90	57	
Yes: Used formalin			411	46					

562 2004, 2013, 2014 and  $2015^{a}$ .

Yes: Did not use formalin			145	16				
Used Footvax								
No	649	86	693	77	470	78	98	62
Yes	97	13	194	22	132	22	55	35
Checked feet of sheep pre-pur	chase							
Always			254	28	130	22		
Usually			182	20	126	21		
Sometimes			125	14	75	12		
Never			135	15	97	16		
Did not purchase			192	21	167	27		
Isolated new sheep								
Never			102	11	50	8		
Sometimes			73	8	39	6		
Usually			133	15	93	15		
Always			393	44	266	44		
No new arrivals			176	20	138	23		
Sheep left the flock then return	ned							
No			548	61				
Yes			339	38				
Flock mixed with other flocks								
No			838	93				
Yes			35	4				
Don't know			5	1				
Farm type								
Lowland			766	85			140	87

		25	3				
		94	10				
						16	10
		849	94				
		41	5				
361	48	385	43	266	44	69	44
241	32	451	50	296	49	84	53
102	14	36	4	28	5		
	241	241 32	94 849 41 361 48 385 241 32 451	$\begin{array}{cccccccccccccccccccccccccccccccccccc$	$\begin{array}{cccccccccccccccccccccccccccccccccccc$	$\begin{array}{cccccccccccccccccccccccccccccccccccc$	$\begin{array}{cccccccccccccccccccccccccccccccccccc$

<sup>a</sup> Blank cells,

# 565 Figure legend

- 567 Fig. 1. Example photograph and description of granuloma (A), and shelly hoof (B), also
- 568 known as white line separation or degeneration, provided to farmers in questionnaires to
- 569 investigate their ability to identify common foot lesions.