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1 **Qualitative assessment of the entry of capripoxviruses into Great Britain from the**
2 **European Union through importation of ruminant hides, skins and wool**

3

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17

18 **Keywords:** Capripoxviruses, sheep pox and goat pox virus, lumpy skin disease virus, risk
19 assessment, skins, hides

20

21 **Abstract**

22 Sheep pox and goat pox (SPGP) virus and lumpy skin disease (LSD) virus belong to the
23 genus Capripoxvirus and cause disease with economic impacts in sheep/goats and cattle
24 respectively. In 2013/14, outbreaks of SPGP were reported in sheep in Greece and Bulgaria
25 and LSD outbreaks were reported in cattle in Turkey, Egypt and some countries in the Middle
26 East. Clinical signs for both diseases include pox lesions, papules and scabs on the skin
27 which may contain virus. This, together with the fact that Great Britain (GB) currently
28 imports cattle hides, sheep skins and wool from European Union (EU) countries without the
29 requirement for treatment prior to export, raises concern that capripoxviruses could be
30 introduced into GB. A qualitative assessment presented here concluded that the current risk
31 of entry of SPGP virus into GB through the importation of one untreated sheep skin, hide or
32 wool bale from an EU Member State (MS) with similar flock prevalence to that in sheep in
33 Greece in 2013/14 is low. In terms of SPGP virus levels, those infected sheep skins/hides
34 entering GB are more likely to be from infected animals with normal skin (i.e., not showing
35 lesions) and hence carrying lower levels of virus than those from animals showing papules
36 and scabs which contain very high virus levels and are easier to detect. The predicted risk of
37 importation of LSD virus per cattle hide/skin is also low (assuming LSD were to emerge in an
38 EU MS with similar herd prevalence to that reported for SPGP in Greece in 2013/14). The
39 levels of LSD virus on an infected cow's hide, if imported, may be very low. It is
40 recommended that the risks for entry of capripoxviruses are recalculated if outbreaks occur
41 elsewhere within the EU.

42 **Introduction**

43 Lumpy skin disease (LSD), sheep pox and goat pox are pox diseases of cattle, sheep and
44 goats, respectively. They are characterised by fever, nodules on the skin, internal lesions,
45 enlarged lymph nodes and sometimes death [1,2,3,4]. The diseases are of economic
46 importance as they cause damage to hides and can result in death due to secondary bacterial
47 infections [2] together with resulting disruption of trade in livestock and livestock products
48 [1]. LSD can cause a temporary reduction in milk production in cattle and sterility in bulls [2]
49 with subsequent production impacts. The World Organisation for Animal Health (OIE) has
50 categorised LSD and sheep pox and goat pox as notifiable diseases [4,5].

51

52 OIE consider sheep pox and goat pox to be a single disease entity [5], referred to here as
53 sheep pox and goat pox (SPGP). The viruses causing these diseases are members of the
54 Capripoxvirus genus of pox viruses (family Poxviridae) and are clinically indistinguishable.
55 Strains of sheep pox virus (SPPV), goat pox virus (GTPV) and lumpy skin disease virus
56 (LSDV) cannot be differentiated serologically [5]. There is close genetic relatedness of
57 capripoxvirus isolates, which average no less than 96% nucleotide identity between strains of
58 SPPV, GTPV and LSDV [1].

59 LSD is currently present throughout most of the continent of Africa, with only Libya,
60 Algeria, Morocco and Tunisia in the north still considered free [4]. It has spread out of the
61 African continent into the Middle East with the first cases in Israel in 1989 after the disease
62 appeared in Egypt the previous year [4]. LSD outbreaks have been reported in the Middle
63 Eastern region since 1990 including Kuwait, Lebanon, UAE, Israel and Oman. Tuppurainen
64 and Oura [4] write that there are no geographical or epidemiological reasons why LSD cannot
65 spread further north into Turkey and Europe, or further east into Asia and they cite the impact
66 of climate change on the abundance and distribution of mechanical vector populations as

67 possible reasons for this. Indeed, outbreaks of LSD occurred in south eastern Turkey in
68 2013/14 [6].

69 SPGP is found in Africa north of the equator, the Middle East and Asia including India,
70 Nepal and parts of China [5]. It has spread into Europe on several occasions [5], with
71 outbreaks reported in sheep in Bulgaria and Greece in 2013/14 [7]. Distinct host preferences
72 exist with most strains of SPPV and GTPV causing more severe disease in the homologous
73 host [1] and new introductions are generally only identified in one of the two animal species
74 concerned (i.e. goats or sheep) depending on the strain introduced [5]. For example, goat pox
75 was introduced into Bangladesh in 1984 from India, and sheep pox has caused occasional
76 outbreaks in Italy (1983), Greece (1988, 1995, 1996, 1997, 1998 and 2000) and Bulgaria
77 (1995 and 1996) having spread from Turkey, probably in illegally imported animals [5].

78 Spread of capripoxviruses can occur through trade of infected animals and their products
79 such as wool and hides [8]. Such products may be treated or untreated. Untreated hides and
80 skins are defined in Regulation (EU) No 142/2011 [9] as cutaneous and subcutaneous tissues
81 that have not undergone any treatment, other than cutting, chilling or freezing. There is
82 currently no requirement for treatment of these products imported to GB from within the EU.
83 Under Regulation (EU) No 142/2011 [9] fresh hides and skins must, however, comply with
84 the animal health conditions for fresh meat laid down under Council Directive 2002/99/EC
85 [10]. Thus, skins and hides must not come from slaughterhouses in which animals infected
86 with sheep pox and goat pox virus (SPGPV) or LSDV were present during the slaughtering
87 or production process. This is important because it means that if a positive animal is detected
88 at the farm or slaughterhouse in the EU then the whole batch (including other infected
89 animals which may have been missed) is condemned.

90 Given the ongoing outbreaks of SPGP in south-eastern Europe and LSD in Turkey, there is
91 potential for further spread of these capripoxviruses to and/or within Europe. This, together
92 with the fact that GB currently imports cattle hides, sheep skins and wool from European
93 countries without the requirement for treatment prior to export, raises concern that
94 capripoxviruses could be introduced into GB. This paper describes a qualitative assessment
95 of the risk of importation of one infected product (i.e. skin/hide or bale of wool) through legal
96 trade into GB.

97

98 **Methods**

99 **Risk question and scope**

100 The specific risk question was: What is the probability that an individual whole skin/hide or
101 bale of wool **legally** imported from an EU Member State (MS) with an ongoing outbreak is
102 infected with capripoxvirus at the point of entry into GB? Thus, following the OIE Terrestrial
103 Animal Health Code definition [11], an entry assessment was undertaken. SPGPV and LSDV
104 are very similar and each step of the risk assessment considers both viruses together, while
105 highlighting any subtle differences that warrant a separate consideration in terms of risk. The
106 products (skins/hide/wool) are considered collectively. Trade levels to GB and
107 transmission/spread, once within GB, were not considered. An infected product was defined
108 as one that contains one or more infectious virus particles.

109 **Risk pathway**

110 The risk pathway has four component steps: (i) the herd/flock from which an animal comes is
111 infected (with probability P_1), (ii) an individual animal is infected with the virus (with
112 probability P_2), given the herd/flock is infected, (iii) the infected skin/hide/wool bale enters

113 the export chain (with probability P_3), and (iv) the virus survives packaging and transport of
114 the skin/hide/wool to GB (with probability P_4). The probabilities P_2 , P_3 and P_4 are
115 conditional probabilities and the overall probability of virus entry (R) is given by:

$$116 \quad R = P_1 P_2 P_3 P_4 \quad (1)$$

117 Although the level of virus is not explicitly considered as an output from the assessment
118 (virus entry is defined as one or more infectious virus particles), it is important for the
119 estimation of some of the pathway probabilities. In particular P_3 and P_4 are dependent on the
120 levels of the virus in the skin of infected animals. For this reason, virus level was considered.

121

122 **Levels of virus on skins/hides of infected animals**

123 A distinction was made between skins/hides from infected animals showing clinical signs
124 (i.e. pox lesions, papules or scabs) and those from infected animals with normal skin (i.e.,
125 skin with no apparent gross pathology) and no clinical signs. This distinction was made
126 because most virus is found in the skin papules about six days after their first appearance [5].
127 Bowden et al [1] estimated that the normal skin of goats experimentally infected with GTPV
128 has $10^{3.0}$ to $10^{4.4}$ tissue culture infectious dose 50% (TCID₅₀) per gram between 8 and 13 days
129 post inoculation (dpi) while the papules have loading ranges over 100-fold higher than the
130 normal skin at $10^{5.2}$ to $>10^{7.2}$ TCID₅₀ per gram over the same time scale. Similarly, genomic
131 copies of SPPV in normal sheep skin were 4-log₁₀ (per 100 ng total DNA) at 8 dpi compared
132 to 6.5-log₁₀ for the secondary skin nodules [1] with a >5 log₁₀ difference at 13 dpi. In
133 experimentally infected sheep, SPPV titres of 10^7 TCID₅₀ per gram of skin (at sites where
134 virus was inoculated) were detected by day 7 to 8 [12]. In cattle experimentally infected with
135 LSDV through the jugular vein, skin nodules contained high levels of virus [13] with 5.1 and

136 5.3 log₁₀ plaque forming units (pfu) per gram at 12 and 15 dpi, respectively [13]. In contrast
137 to sheep and goats in the study of Bowden et al [1], infectious virus was absent from normal
138 skin of LSDV-infected cattle [13]. Furthermore, while levels of viral DNA in the skin
139 nodules of cattle were very high between 4.6 and 8.6 log₁₀ copies per µg tissue, levels of viral
140 DNA in normal skin of LSDV-infected cattle were in general undetectable [13]. Based on
141 these data, it was assumed that the titre of virus on a hide/skin is directly proportional to the
142 number of lesions or papules on that hide/skin and the time since infection. The papules and
143 scabs are likely to contain very high levels of virus, while normal skin from SPGPV-infected
144 goats and sheep is likely to contain medium levels of virus. Normal skin from LSDV-infected
145 cattle contains very low levels of virus.

146

147 **Levels of virus in wool from infected animals**

148 There is little information on levels of SPPV or GPPV in wool. Following experimental
149 intradermal inoculation, the virus replicates in the cells of the dermis and glandular hair cells
150 at the base of the hair follicles [15]. Unlike skin, the virus will not be able to replicate within
151 the wool itself, and therefore any virus present will be due to contamination of the wool with
152 skin fragments, including fragments of scab material. In this respect the wool could contain
153 fragments of lesion with high loadings of virus. The papules may cover the whole body or be
154 restricted to the more hairless or woolless parts of the skin [1,5]. In lambs and kids naturally
155 infected in the Duhok area of Iraq, the presence of pox lesions occurs in areas of the hide
156 with less wool and hair [14]. Similarly in sheep in Iran, the gross lesions in adults occurred in
157 woolless or sparsely woolled areas of skin [15]. However, the gross lesions were all over the
158 skin and in some internal organs in lambs [15]. It was assumed that wool from infected adult
159 sheep contains low levels of infectivity while wool from lambs contains medium levels of

160 infectivity. This reflects the fact that in some lambs lesions occur all over the skin rather than
161 in the woolless areas observed in adults.

162

163 **Qualitative probabilities**

164 The entry assessment describes the probability of entry of the virus into GB through the
165 importation of one product item from other regions of the EU. Following the European Food
166 Safety Authority (EFSA) definitions, the probabilities in Equation (1) are expressed
167 qualitatively as negligible, very low, low, medium, high or very high [16,17]. The definitions
168 of these terms were taken from [16] namely, negligible: so rare that it does not merit to be
169 considered; very low: very rare but cannot be excluded; low: event is rare but does occur;
170 medium: event occurs regularly; high: event occurs very often; and very high: event occurs
171 almost certainly. To estimate the risk of release, R, the qualitative probabilities were
172 combined as in Equation (1) using the reasoning described previously [18]. In summary, as
173 each qualitative probability P_1 to P_4 can be considered quantitatively as taking a value
174 between 0 and 1, it follows that the product R will be at most, the minimum of P_1 to P_4 . The
175 qualitative value of R is thus set as the minimum of the qualitative values of P_1 to P_4 . **The**
176 **probability definitions given above apply to all the qualitative probabilities within the risk**
177 **assessment, i.e. R, P_1 , P_2 , P_3 and P_4 .**

178

179 **Estimation of P_1 : Probability that a herd/flock is infected**

180 Data on the recent outbreaks in Greece were used to estimate P_1 . Hadjigeorgiou et al [19]
181 reported that there are around 9,200,000 sheep and 5,600,000 goats in Greece on about
182 300,000 farm units. Counting the units with more than 10 adult female animals, this equates

183 to about 155,000 farms. OIE [7] give data on the number of farm units in which outbreaks
184 occur. Between Aug 2013 and January 2014 (six months), outbreaks of SPGPV were
185 reported in sheep in Greece in 82 farm units [7]. Over a period of one year, therefore, double
186 that number of outbreaks, i.e. 164, might be expected. This would represent about one in a
187 thousand of the 155,000 goat and sheep farms in Greece. LSDV has never been reported in
188 Europe [4], and it is assumed here P_1 would be similar to that for SPPV in Greece.

189 **Estimation of P_2 : Probability that an individual animal within a positive herd/flock is**
190 **infected**

191 Data from the recent outbreaks of SPGPV in sheep in Greece and Bulgaria were used to
192 estimate this probability. Between Aug 2013 and January 2014, a total of 1,472 cases (250
193 deaths) of SPGPV were reported in Greece in 17,735 susceptible sheep in 82 infected flocks
194 (Figure 1) [7]. The OIE definition [23] of susceptible animals is, “Animals present in the
195 outbreaks at the start of the period in question”. Thus the number of susceptible animals
196 recorded by OIE [6, 7] includes all animals on the farm which are susceptible to the virus
197 whether infected or not. Therefore the average within-flock prevalence in sheep may be
198 calculated as $1,472/17,735 = 0.083$. The range of within-flock prevalences was from 0.0035
199 (1 case in 284 susceptible sheep) to 1.0 (13 cases in 13 susceptible sheep) with 5th and 95th
200 percentiles of 0.007 and 0.552 respectively. Linear regression analysis of the data for 82
201 infected sheep flocks in Greece [7] showed that there is a statistically significant relationship
202 between decreasing within-herd prevalence and increasing herd size, ($P < 0.001$) (Figure 1).
203 There is uncertainty associated with why this relationship could occur, but one possibility is
204 that this represents an under-reporting in larger flocks because once a single case is detected
205 the whole flock is condemned, and there is little point in looking for every last case in a large
206 flock.

207 In Bulgaria there were three outbreaks in 2013 [7], with a total of 37 cases in 558 susceptible
208 sheep giving an average within-flock prevalence of 0.066. The reported outbreaks from
209 Greece and Bulgaria give estimates of the mean within-flock prevalence (P_2) to be between
210 0.066 and 0.083. Pox lesions, however, may be missed due to their restricted distribution on
211 some sheep [8]. Thus, the true within-flock prevalence may be higher than these values. In
212 terms of disease symptoms between individual animals, SPGPV typically exhibits a uniform
213 range of responses in the respective host species [8] such that infected animals typically show
214 symptoms. However in the case of cattle, not all animals infected with LSDV exhibit clinical
215 signs thus potentially hindering detection of cases on farm. In 21 outbreaks in Turkey
216 between Nov 2013 and Feb 2014 [6], 837 LSD cases were reported in 21,829 susceptible
217 cattle giving an average within-herd prevalence of 0.038 and roughly half that of SPGP
218 reported in sheep in Greece and Bulgaria. According to Tuppurainen and Oura [4], only 50%
219 of LSDV-infected cows are likely to show clinical signs, even though the majority of
220 experimentally infected cows become viraemic. The observed within-herd prevalence of
221 LSDV was therefore multiplied by factor of two for the purpose of this risk assessment. Thus
222 the estimated within-herd prevalence for a LSDV-positive herd in Turkey is around 0.076 and
223 comparable to that reported for SPGPV-positive sheep flocks in Greece and Bulgaria. The
224 range of within-flock prevalences for LSDV-positive herds in Turkey was from 0.0007 (1
225 case in 1,372 susceptible cattle) to 0.67 (2 cases in 3 susceptible cattle). Linear regression
226 analysis of the data for 21 LSDV-infected cattle herds in Turkey between Nov 2013 and Feb
227 2014 [6] showed that there is a statistically significant relationship between decreasing
228 within-herd prevalence and increasing herd size, ($P=0.002$) (data not shown).

229

230 **Estimation of P_3 : Probability that infected skin/hide/wool bale enters the export chain**

231 The probability P_3 relates to the detection of an infected animal and therefore whether the infected
232 hide is prevented from being exported, rather than whether the animal was slaughtered for domestic
233 consumption or export. With reference to the risk question (see above) it is given that the hide has
234 been legally imported into GB and therefore comes from an EU farm registered to export where EU
235 142/2011 is enforced. The probability P_3 depends on the probability that an individual infected
236 animal/skin is not detected either on the farm or at the approved slaughter house. In the 82
237 reported outbreaks of SPGPV in sheep in Greece between Aug 2013 and Jan 2014, flocks
238 with as few as one case in 284 susceptible sheep were reported [7]. This suggests that some
239 farmers/slaughter house operators are good at spotting low frequency occurrences of clinical
240 cases in a large number of animals although it is not known how many cases were present in
241 that flock of 284 sheep and were thus missed. It would seem unlikely that the remaining 283
242 susceptible sheep in that flock were tested to confirm they were negative. However, at the
243 other extreme, 270 cases were reported in a flock of 390 susceptible sheep, suggesting that
244 the probability of detecting an infected animal with clinical symptoms is relatively high. To
245 estimate P_3 it was assumed that the probability of detection of an infected animal at an
246 approved slaughter house or on a farm is directly proportional to the number of lesions on
247 that animal (i.e. the more papules the greater the chance that the farmer or slaughterhouse
248 worker will see one). Therefore, those animals with high titre hides/skin/wool have a high
249 probability of being detected, while the lower titre animal hides from infected animals
250 without lesions are more likely to be missed. For SPGPV, the distribution of pox lesions in
251 the skin can be widespread with over 50% of the skin surface affected [8] facilitating
252 detection of cases. However, more commonly in enzootic areas, the lesions in sheep and
253 goats are restricted to a few nodules under the tail and are thus only detected on close
254 examination [8], increasing the probability of not detecting a case. Furthermore some animals
255 in the slaughterhouse may be at a stage where infection has taken place in the skin but
256 clinical symptoms have not yet developed. Thus, virus was detected in normal skin of sheep

257 at 4 dpi [1] while macules did not develop before 5 dpi. Moreover the number of SPGP viral
258 genomic copies was $\sim 4\text{-log}_{10}$ in normal skin in sheep at 6 dpi when macules faded on
259 exsanguination prior to necropsy [1]. Thus macule detection efficiency could be reduced at
260 slaughterhouses, while significant levels of virus are present in normal skin and in the
261 macules themselves. The macules enlarge and develop into papules within 1 to 2 days and
262 then to scabs within the following week [1]. Papules and scabs are less likely to be missed.
263 For LSDV-infected cattle, only half of those infected show symptoms (discussed above).

264 The probability P_3 is also related to the number of clinical cases in the flock/herd on the farm
265 or in the slaughterhouse batch (which could include more than one flock or herd). Thus the
266 more clinical cases in an infected flock/batch, the greater the chance that at least one is
267 detected and that all animals in that flock/batch and thus their products are condemned
268 according to EU 142/2011. Analysis of the outbreak data for SPGPV in sheep in Greece [7]
269 showed that the statistical distribution for the number of cases per infected flock is skewed
270 with a significant proportion of infected flocks having a few cases and a small proportion
271 having a large number of cases (Figure 2). Thus, although the average was 18.0 cases per
272 infected flock (1,472 cases in 82 flocks), some 33% of infected flocks had just 1, 2 or 3 cases.

273 The statistical distribution of the number of LSDV cases in infected cattle herds (Figure 2) is
274 even more skewed than for SPGPV based on the data for 21 LSDV outbreaks in Turkey
275 between Nov 2013 and Feb 2014 [6]. Thus of 21 infected herds, 14 (66%) has just one or two
276 cases while three herds had >200 cases.

277 About 110 fleeces may go into a bale of wool [20]. The shearing process on the farm may
278 expose skin lesions and allow detection of infected animals at an earlier stage. However,
279 although many animals contribute wool to a bale, many infected sheep flocks have few cases
280 of SPGPV (Figure 2).

281 **Estimation of P₄: Virus on wool/hide/skin survives transport to GB.**

282 Although SPGPV is very susceptible to direct sunlight, it can persist for months in dark
283 conditions, such as contaminated animal sheds [5] and has been shown to remain infectious
284 for periods of at least 3 months in scab material obtained from animals which have recovered
285 from the infection [21] LSDV is stable between pH 6.6 and 8.6 and showed no significant
286 reduction in titre after 5 days at 37°C over this pH range [22]. For LSDV in the skin lesions
287 of infected animals, the virus can persist for at least 33 days even though the necrotic portions
288 of skin have completely dried out [22]. Skin/hides and wool are likely to be transported to
289 GB via trucks and ships. Various travel blogs report that the drive from Greece to England
290 requires 4 - 7 days. With temperatures below 37°C and in the dark, it was assumed that little
291 or no inactivation of the virus would occur during this time for transport.

292

293 **Results**

294 On the basis of 2013/14 data for Greece and taking into account the number of sheep/goat
295 farms in Greece, it was concluded that the probability, P₁, that a herd or flock is infected, is
296 low.

297 Assuming that the data for SPGP in Greece and Bulgaria [7] and LSD in Turkey [6] give a
298 reasonable description of within-flock prevalence for capripoxviruses in any EU country
299 which could potentially have outbreaks or undisclosed infection, it was concluded that P₂ is
300 mostly medium. It is noted that smaller flocks appear to have higher within-flock prevalences
301 (Figure 1) (1.0 in two flocks). Thus on a flock to flock bases, P₂ may vary between medium
302 and very high.

303 As described previously a SPGPV-infected sheep could be missed because the macules may
304 be localised to areas where they are hidden such as under the tail and a LSDV-infected cow
305 could be missed because it is not showing clinical symptoms. Taking this into account
306 together with the evidence above that virus could be present in skin before macules have
307 developed, the probability of not detecting an SPGPV/LSDV-infected animal is judged to be
308 medium. This is at the level of the individual animal. For those flocks/herds with a large
309 number of cases, P_3 was considered to be negligible (because at least one case would be
310 detected resulting in condemnation of the whole flock/batch before it could enter the export
311 chain) while for those flocks/herds with lower numbers of cases, P_3 was considered medium
312 (reflecting the chance of missing a single case). Since a significant proportion of SPGPV-
313 infected sheep flocks and the major proportion of LSDV-infected cattle herds have only 1, 2
314 or 3 cases (Figure 2), it was concluded that overall, P_3 is medium. This represents a worst
315 case scenario. Although a batch at a slaughterhouse may include more than one flock, it is
316 very unlikely that more than one flock would be positive in a given daily batch because the
317 between-flock prevalence (P_1) is low. Thus even those batches at slaughterhouses comprising
318 multiple flocks/herds are still likely to have only a few cases based on Figure 2.

319 Given that the SPGP/LSD virus in a hide/skin is unlikely to undergo significant decay within
320 the travel time to GB together with the medium initial titres of virus in normal skin of
321 infected sheep and goats, it was concluded that the probability of virus survival, P_4 , is high.

322 The individual probabilities and the overall probability R are given in Table 1. The lowest
323 probability is for P_1 (low). Thus by combining the qualitative probabilities in Table 1, using
324 the method described by Gale et al [18], it was estimated that the overall probability, R , is
325 low. This represents the probability that an individual raw hide/skin or bale of wool is
326 infected with SPGP/LSD virus on legal import to GB from an EU MS with ongoing

327 outbreaks with similar between-flock (P_1) and within-flock (P_2) prevalences to those reported
328 for sheep in Greece in 2013/14.

329 Descriptions of uncertainty and key assumptions are also presented in Table 1 for each
330 probability term. This process identifies that the risk assessment is highly dependent on the
331 data available from the SPGP outbreak in Greece and there is therefore uncertainty associated
332 with the estimate of risk of another EU country having an outbreak. It is therefore
333 recommended that this risk assessment is revisited if outbreaks occur elsewhere so that the
334 estimates for P_1 , P_2 and P_3 and hence the overall estimate of risk can be verified.

335

336 **Discussion**

337 Outbreaks of SPGP have been reported in sheep in Greece and Bulgaria in 2013/14. LSD
338 outbreaks have occurred in cattle in south-eastern Turkey and there is no reason to assume
339 that LSDV will not spread into Europe at some stage [4]. Given this situation, there is
340 concern within GB that capripoxviruses could be imported via the legal trade of skins/hides
341 and wool from the EU. Using data from the 2013/14 outbreaks of SPGP in Greece and
342 Bulgaria and LSD in Turkey, together with microbiological data from the literature, a
343 qualitative entry assessment was undertaken. It was estimated that the probability of entry of
344 SPGPV/LSDV in a single hide/skin/wool bale imported from a MS with ongoing outbreaks is
345 currently low. Entry of infection was defined in terms of importation of one infected product
346 (skin, hide or bale of wool), contaminated with one or more infectious virus particles.

347 Although there are quantitative data available for many of the parameters including P_1 , P_2 and P_4 , a
348 qualitative approach was adopted here because of the lack of any quantitative data for estimation of
349 P_3 .

350 Viral load was considered to some extent within the assessment although it was not explicitly
351 stated in the probability of entry. While normal skin of SPGPV-infected sheep and goats does
352 contain infectivity, the titres are very much lower than for those in papules and nodules. In
353 view of the inspection processes at approved slaughterhouses, it is considered here that any
354 imported product infected with the virus would most likely have come from an infected
355 animal not yet displaying clinical signs. Alternatively the hide/skin material imported may
356 exclude those regions of the skin (woolless areas or under the tail) where nodules more
357 commonly occur [8]. The viral load on the hide/skin of such an animal is likely to be at a
358 medium level rather than the very high levels found in skins with lesions and papules. The
359 viral levels in wool from an infected bale would also be medium although there may be some
360 variation depending on whether the wool is from lambs or adult sheep. Whether or not this
361 medium level would be important for transmission within GB would depend on several
362 factors including the dose-response relationship and the potential routes of exposure for GB
363 cattle, sheep and goats. SPGPV is spread through aerosols and/or close contact and by
364 indirect means such as contamination of cuts and abrasions (Babiuk et al, 2008a). The high
365 concentration of virus in the skin may also contribute to spread via insect vectors [1] although
366 it is not clear whether this could happen from hides/wool in GB. Normal skin of LSDV-
367 infected cows has very low levels of virus [8]. Thus the capripoxvirus levels on an LSDV-
368 infected cattle skin/hide given it has entered GB from the EU may be much lower than that
369 for SPGPV-infected hides.

370 The assessment did not consider the volume of trade in skins/hides/wool from the EU. Thus
371 the probabilities of entry per year or per batch were not estimated. There are currently no data
372 available to determine the volume of trade. Should these data become available in the future,
373 the assessment could be extended to include such estimates.

374 The method used to combine the qualitative probabilities associated with the risk of entry of
375 virus makes use of the fact that these probabilities are conditional; they correspond to a
376 sequential set of events, all of which are necessary for the importation of an infected product.
377 In a comparable quantitative assessment, the rules of probability mean that the conditional
378 probabilities are multiplied to give the joint probability which represents the estimate of risk.
379 The absolute maximum of this joint probability will be the minimum of the conditional
380 probabilities. It is intuitive to consider the same multiplicative process when dealing with
381 conditional probabilities that are qualitative. However, in this case, risk may be over-
382 estimated because no account is taken of where on the qualitative category an individual
383 probability will lie. Furthermore, if all four probabilities P_1 to P_4 were low, for example, then
384 R would still be low as it would if three were high and just one were low. Thus, the low
385 estimate of virus entry may very well be an over-estimate in this case. As it currently stands,
386 the value of P_1 is the determining probability for R as it is the only probability with a value of
387 low. Thus, based on the current data and assessment, the risk of entry of virus depends on the
388 herd/flock prevalence in the countries in which there have been recent outbreaks i.e. Greece
389 and Bulgaria. Should the situation in Greece, Bulgaria or any other EU country change, the
390 estimate of risk would need to be updated.

391 Although there is considerable variation in the within-herd prevalence for SPGPV-positive
392 sheep flocks in Greece (Figure 1) (and for LSDV-positive herds in Turkey (not shown)) this
393 range could reflect natural variation, for example due to differences in exposure resulting
394 from the intensity of the sheep/sheep contacts (sheep density), differences in environmental
395 factors between flocks and differences in the susceptibility of individuals/breeds within a
396 given flock (i.e. dose-response). There is a statistically significant relationship between
397 decreasing within-herd prevalence and increasing herd size both for SPGP in sheep in Greece
398 (Figure 1) and for LSD in cattle in Turkey (not shown). As discussed previously, this may

399 relate to some bias within the data due to failure to detect all of the infected animals within a
400 positive herd, particularly in the larger herds. However, using the approaches described above
401 for the combining of probabilities, only a significant decrease in the magnitude of P_2 (such
402 that P_2 is less than P_1) would affect the predicted value of R in this assessment. Under EU
403 142/2011 all animals in the flock/slaughterhouse batch are condemned if at least one case is
404 detected. Therefore the probability of an infected skin/hide/wool bale entering the export
405 chain (P_3) is dependent on the statistical distribution of the number of cases within an
406 infected flock/herd/batch. Thus the more cases in a flock, the greater the chance that at least
407 one is detected and that the whole flock/slaughter house batch is condemned (under EU
408 142/2011). An increase in within-herd prevalence or emergence of a more virulent strain
409 which meant fewer infected flocks had just one or two cases, would greatly decrease P_3 .
410 Indeed should P_3 decrease in magnitude below low, so too would R . While some
411 capripoxvirus-infected animals do not show symptoms (see above) and would not be detected
412 on an individual basis it is unlikely that multiple infected animals in a given flock/herd would
413 all be symptomless at time of inspection. Thus a high within-herd prevalence not only
414 increases the probability of at least one case with symptoms being detected, but also increases
415 the probability of at least some cases displaying detectable symptoms. The statistical
416 distribution of the number of cases within those infected herds/flocks is therefore central to
417 understanding the uncertainty in P_3 .

418 In conclusion, based on the 2013/14 outbreak data for countries in south-east Europe, the
419 probability of entry of SPGPV into GB from the importation of a single hide/skin/wool bale
420 from an EU MS with ongoing outbreaks has been assessed as low. The predicted risk is also
421 low for LSDV in a single cattle skin/hide should this virus emerge in an EU MS at some
422 stage. These estimates are sensitive to the herd/flock prevalence during an outbreak in the
423 EU.

424

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Table 1: Estimated qualitative probabilities for SPGP and LSD.

Probability	Qualitative probability	Key assumptions and uncertainties
Herd/flock infected (P_1)	Low	Data only available for SPGP in Greece, therefore uncertain of probability of a herd/flock being infected elsewhere in the EU. No cases of LSD in EU so assumes P_1 is similar to that for SPGP in sheep in Greece and Bulgaria.
Animal infected, given herd is positive (P_2)	Medium to Very High	Within-flock prevalence data only available for SPGPV in sheep in Greece and Bulgaria (and LSDV in Turkey), therefore uncertain of the value of P_2 for outbreaks in other EU countries.
Infected skin enters export chain (P_3)	Medium	Infected animals with fewer lesions or earlier stages of infection may be missed. P_3 tends to negligible for herds with many infected animals.
Virus survival (P_4)	High	None
Risk of release for one product item (R)	Low	Limited or no data available for likely prevalence of SPGP and LSD within or between flocks/herds in EU countries other than Greece and Bulgaria. Therefore considerable uncertainty associated with the risk of release if an outbreak is reported in another country.

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499

500 Figure legends

501

502 Figure 1: The reported within-flock prevalence for cases of SPGPV-infected sheep decreases
503 with the size of the flock, $P = 2.3 \times 10^{-7}$ (slope $-0.217/\log_{10}$ flock size, 95% c.i. -0.14 to $-$
504 0.29). Data for 82 SPGPV-infected sheep flocks in Greece between Aug 2013 and Jan 2014
505 (OIE, 2014b).

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508 Figure 2: Distribution of number of detected cases per infected herd for SPGPV outbreaks in
509 sheep in Greece (OIE, 2014a) and LSDV outbreaks in cattle in Turkey (OIE, 2014b).

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