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Differences in receptivity to gastrointestinal infections with nematodes in dairy ewes: Influence of age and of the level of milk production

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11 Abstract

Resistance to anthelmintics in populations of gastrointestinal nematodes is a major concern in small ruminants. One solution 12 to limit the spread of anthelmintic resistance is to apply treatments selectively by targeting the most susceptible animals within 13 a flock. In dairy goats, previous studies have shown that, within a flock, goats in first lactation and those with high level of milk 14 15 production were highly receptive to nematode infections. These results provided the rationale for targeted treatments. In dairy ewes, such epidemiological information on possible factors modulating the susceptibility to parasitism were still lacking. The 16 objective of the current study was therefore to examine differences in the level of parasite infection and in the pathophysiological 17 consequences in dairy ewes, depending on the age or on the level of milk production. In three farms, parasite egg excretion, and the 18 serum concentrations of pepsinogen and inorganic phosphate were compared on one hand between primiparous and multiparous 19 ewes; on the other hand, between ewes with the highest and the lowest level of milk production, within a cohort of 3-5-year 20 old animals. Overall, the results did not indicate significant differences for both either the parasitological or pathophysiological 21 measurements depending on the level of milk production. In contrast, significant differences were found according to age, 22 indicating higher levels of infections in the primiparous ewes than in the multiparous ones and suggesting that this category of 23 animals represents a particular parasitic risk within a flock. 24

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26 Keywords: Dairy ewes; Parasitic nematodes; Susceptibility; Epidemiology; Selective treatment

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1. Introduction

In small ruminants, the constant increasing development of anthelmintic resistances nowadays severely impairs the control of gastrointestinal trichostrongylo-31

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sis based on chemical drugs (Sangster, 1999; Jackson 32 and Coop, 2000). Resistances to the three main fami-33 lies of broad spectrum anthelmintics available for the 34 control of trichostrongyles have now been described 35 worldwide in most nematode species (Sangster, 1999; 36 Jackson and Coop, 2000; Kaplan, in press). Therefore, 37 there is an urgent need to seek alternative or comple-38 mentary solutions to anthelmintics as well as to im-39 prove the use of the drugs currently available in order 40 to preserve their efficiency in the future (Waller, 1999). 41 In many countries, recommendations have been emit-42 ted to reduce the selection pressure for anthelmintic 43 resistance in nematode populations. They rely mainly 44 on a reduction in frequency of treatments, the respect 45 of appropriate doses, the alternation of drug families 46 and the preferred use of narrow spectrum anthelmintic 47 when possible (Dash et al., 1985; Waller et al., 1995). 48 Another recommendation to slow down the develop-49 ment of anthelmintic resistance within worm popula-50 tions is to favour targeted anthelmintic treatments in-51 stead of their systematic use. The principle of targeted 52 treatments is to give anthelmintics only to the most 53 infected and/or the most susceptible animals within a 54 flock. By leaving some animals untreated, the method 55 allows to maintain a refugia of susceptible genes within 56 the worm population and this will contribute to slow 57 down the spread of anthelmintic resistance, by diluting 58 resistance genes (Barnes et al., 1995; Sangster, 1999; 59 Coles, 2002; Van Wyk, 2001). 60

One key point in any method of selective treatment 61 lays in the identification of the animals to be treated. 62 In tropical and subtropical areas where Haemonchus 63 contortus is the dominant species, a method based on 64 the individual evaluation of related clinical signs has 65 been developed. Both in sheep and goats, it has been 66 shown that this FAMACHA method lead to significant 67 reductions in the number of treatments applied per flock 68 although maintaining a good level of control of para-69 sites (Van Wyk and Bath, 2002). On the other hand, in 70 temperate countries where Haemonchus is not the most 71 prevalent genus, a method, based on epidemiological 72 data, has been proposed in dairy goats (Hoste et al., 73 2002a). Its rationale was provided by the assessment 74 of differences in receptivity to parasites between does 75 within a flock, depending on the age or on the level of 76 milk production, and therefore characterising the ani-77 mals to be treated preferentially. The same method ap-78 peared potentially applicable in milk producing sheep. 70

However, information on the influence of similar factors on the receptivity and/or susceptibility to nematode infections were not available for dairy ewes.

The present study was performed to determine within flocks of dairy ewes whether the age or the level of milk production might also modulate the response to gastrointestinal nematodes.

2. Materials and methods

The survey was conducted in 2002 in three dairy ewe 88 farms, from one of the main areas of ewe milk produc-89 tion in France, i.e. the Basque Country. In the three 90 farms, the ewes were grazing for the whole year with 91 lambing time occurring in November-December. Af-92 ter 40 days spent for lambs, the ewes were milked from 93 January to June/July. Three groups of 20 ewes were se-94 lected and surveyed within each farm. One group was 95 composed with 20 ewes in first lactation which were 96 randomly selected. The two other groups were com-97 posed with multiparous, adult ewes which were 3, 4 98 or 5 years old. Within each farm, within the pool of 99 these 3-5-year old animals, the ewes were classified 100 according to their level of milk production, based on 101 (i) the mean records of the previous year and (ii) the 102 mean yield of the first month of lactation, i.e. at a time 103 when nematode challenges were low. In each farm, af-104 ter classification of the ewes according to these two 105 criteria, the two additional groups of 20 multiparous 106 ewes were composed, one corresponding to ewes with 107 a relative high level of milk production and the second 108 one corresponding to the ewes with the low level of 109 production. The values of both criteria characterising 110 these two subgroups of selected multiparous ewes per 111 farm are described in Table 1. 112

During the grazing season, individual faecal and 113 blood samples were taken from the 60 ewes per farm, 114 four times per year, i.e. in February, at the end of March, 115 in May and at the end of September. In the three farms, 116 the ewes were treated with anthelmintics during sum-117 mer, i.e. respectively, with closantel (Seponver[®]) at the 118 beginning of August and at the start of September in 119 farms 1 and 2 and with fenbendazole (Panacur®) at the 120 beginning of August in farm 3. 121

Individual faecal egg counts (FECs) were performed using a modified Mc Master method (Raynaud, 1970). 123 These data were completed by larval cultures to as-

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Table 1

Mean values of the criteria used to discriminate between the 20 high producing and 20 low producing 3–5-year old ewes in the three farms of the survey

Size of the flock (% first lactation)	Farm 1 340 (20.6%)		Farm 2 440 (25%)		Farm 3 180 (27.8%)	
Low producers	161	0.80	212	0.81	133.4	0.692
High producers	228	1.347	282	1.43	203.4	1.220

sess the generic composition of nematode populations 125 on each farm (Gevrey et al., 1963). The blood sam-126 ples were collected by venipuncture into vacuum tubes 127 and were used to measure the serum concentrations of 128 pepsinogen and inorganic phosphate. Pepsinogen con-129 centrations were measured according to the method de-130 scribed by Kerboeuf (1975). Inorganic phosphate con-131 centrations were determined according to the method 132 described by Robinson et al. (1971). 133

For all the measurements (faecal egg counts and pathophysiological parameters), the comparisons were performed using a repeated measure analysis of variance (SYSTAT 9.0 software for Windows 1998, SPSS Inc., Chicago, USA). Values of eggs per gram were log (x + 1) transformed before being compared.

140 3. Results

141 3.1. Parasitological data

142 3.1.1. Egg excretion (Fig. 1)

Overall, the levels of mean egg excretion in the three
farms, in the different groups of ewes, remained under
500 eggs per gram with the only exception being the
values measured in September for the group composed
of the first lactating ewes in farm 3.

The statistical comparisons of the mean egg excretions between the high and the low producing ewes in
each farm did not indicate any significant differences
(Fig. 1a).

In contrast, the values of egg excretion were usually found to be higher in the first lactating ewes than in the multiparous ones in the three farms. This observation was confirmed through the statistical comparison of the mean egg excretions between the two groups which confirmed significant differences in farms 2 and 3 (P < 0.01) (Fig. 1b).

3.1.2. Generic composition of nematode populations (Fig. 2)

When pooling the data obtained on the four different sampling dates, the infective larvae of *Teladorsagia* sp. were found to represent, respectively, 78.0, 72.9 and 69.9% of the nematode larvae recovered from the larval cultures in the three farms. *Haemonchus* larvae were present on the three farms at a low level, ranging from 0 to 16.4% depending on the farm and the time of the year.

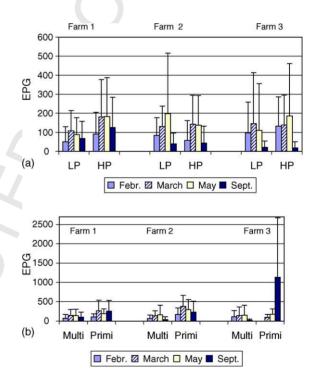


Fig. 1. Mean egg excretions in the three farms of the survey. (a) Comparison between high producing (HP) and low producing (LP) ewes; (b) comparison between multiparous vs. primiparous ewes. Significant statistical differences (P < 0.01) were assessed between the primiparous and multiparous ewes in farms 2 and 3 after values of egg excretion have been log (x + 1) transformed.

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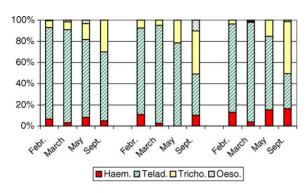


Fig. 2. Generic composition of the population of third stage infective larvae collected after larval culture in the three farms on the different dates of the survey.

- Moreover, the same pattern of the generic compositionwas found with time showing an increase in the number
- 169 of *Trichostrongylus* larvae in autumn.

3.1.3. Pathophysiological parameters (Figs. 3 and 4)

The statistical comparisons of the mean values for the two pathophysiological measurements between the

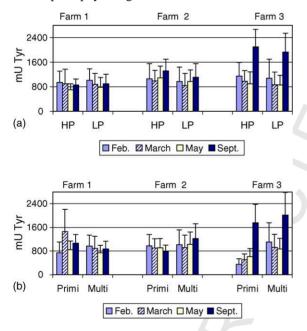


Fig. 3. Mean pepsinogen concentrations in the three farms of the survey. (a) Comparison between high producing (HP) and low producing (LP) ewes; (b) comparison between multiparous vs primiparous ewes. Significant statistical differences (P < 0.05) were assessed between the primiparous and multiparous ewes in farm 2.

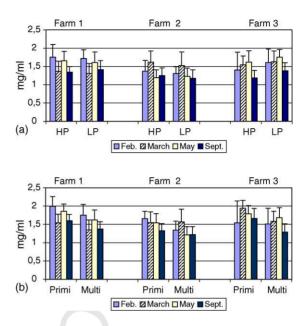


Fig. 4. Mean concentrations of serum inorganic phosphate in the three farms of the survey. (a) Comparison between high producing (HP) and low producing (LP) ewes; (b) comparison between multiparous vs. primiparous ewes. Significant statistical differences (P < 0.05) were assessed between the primiparous and multiparous ewes in farms 1 and 2.

high and the low producing adult ewes in the three farms did not indicate any differences between these two groups, with the exception of the phosphate values in farm 3, which were significantly lower in the HP than in the LP group (P < 0.04) (Figs. 3a and 4a). 178

In contrast, on the three farms, differences were re-179 peatedly observed between the first lactating and the 180 multiparous ewes both for the pepsinogen and the phos-181 phate values (Figs. 3b and 4b). In general, these statisti-182 cal differences corresponded to higher pepsinogen val-183 ues and lower inorganic phosphate values found in the 184 multiparous ewes compared to the primiparous ones. 185 Significant differences between these two groups were 186 found for the two measurements in farm 2 (pepsinogen: 187 P < 0.05; phosphate P < 0.01), and for the phosphate 188 values in farm 1 (P < 0.01). 189

4. Discussion

Overall, in the three farms, no statistical differences ¹⁹¹ were found between the high and the low produc-¹⁹²

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ing ewes in the egg excretion at any date of the sur-193 vey. This result suggests that a similar level of infec-194 tion with gastrointestinal nematodes was present in the 195 ewes regardless of the level of milk production. This 196 conclusion is also supported by the lack of statistical 197 differences in the two pathophysiological parameters 198 which were measured. Because the blood concentra-190 tions of pepsinogen and inorganic phosphate are usu-200 ally considered to reflect mucosal damage and thus, 201 indirectly, the size of the worm populations occurring 202 in the abomasum and in small intestine, respectively. 203 This absence of difference tends to confirm a similar 204 level of parasitism in the two subgroups of multiparous 205 ewes. 206

207 In dairy goats, several studies have indicated that the high producing animals within a flock were less resis-208 tant and resilient to nematode infections than does from 209 the same flock with a low level of milk production. Evi-210 dence supporting these conclusions have been acquired 211 both in experimental infections (Hoste and Chartier, 212 1993; Chartier and Hoste, 1997) and through epidemio-213 logical surveys (Hoste et al., 1999, 2001, 2002b). It was 214 postulated that such differences in host response de-215 pending on the level of production could be explained 216 by the excessive nutritional demands related to high 217 milk vield according to the nutritional framework pro-218 posed by Coop and Kyriazakis (1999). 219

Our results in dairy ewes are thus in contrast with 220 data acquired in goats. Because the mean quantity of 221 milk exported by dairy goats is nearly twice those ex-222 creted by ewes, it is suspected that the nutritional de-223 mands in ewes are less important than in goats, and this 224 could explain the discrepancy. Moreover, in the cur-225 rent survey, half of the samples were taken at the end 226 of lactation (May) or when ewes were out of produc-227 tion (September). On the other hand, it is also usually 228 admitted that sheep are more likely than goats to ex-229 hibit an immune response against gastrointestinal ne-230 matodes (Pomroy et al., 1986; Huntley et al., 1995). 231 Therefore, it can be postulated that the differences in 232 response to parasites depending on the level of milk 233 production might reflect some differences in the priori-234 tisation of functions between the two small ruminant 235 species. 236

Whereas only minor differences in parasitism were
found between the ewes depending on the level of production, differences appeared to occur depending on
the age, by the comparing primiparous and multiparous

animals. This conclusion is supported by consistent re-241 sults obtained for all three measurements. In the three 242 farms, the egg excretions were generally higher in the 243 first lactating ewes than in the multiparous ones and 244 these differences were statistically significant on two 245 farms. This result suggests the presence of higher worm 246 populations in the youngest animals and is similar to 247 previous findings in goats (Hoste et al., 1999, 2001, 248 2002b). As has been suggested for goats, it can be hy-249 pothesized that the higher receptivity of the first lacta-250 tion ewes is due to the lack of or to the low intensity 251 of previous contacts with trichostrongyles and the ab-252 sence of an immune response able to regulate the worm 253 populations as in older ewes. 254

Overall, when examining data from the three farms, 255 the multiparous ewes were usually found to present 256 higher pepsinogen and lower phosphate values. These 257 results suggest that more severe pathophysiological 258 changes occurred in the multiparous than in the primi-259 parous ewes. Somehow, this appeared paradoxical with 260 the lower levels of egg output found in the adult ani-261 mals, suggesting lower levels of infections. However, 262 this apparent contradiction might be explained if one 263 postulates that the immune response developed by the 264 adult ewes, which regulates the worm biology, also neg-265 atively affects the digestive tissues and functions of 266 the host. Previous circumstancial evidence have been 267 acquired supporting such a hypothesis of the involve-268 ment of some immunopathological processes in the ori-269 gin of the structural and/or functional damages associ-270 ated with the presence of gastrointestinal nematodes 271 (Pullman et al., 1991; Larsen et al., 1999; Meeusen, 272 1999; Balic et al., 2000). 273

In dairy goats, the differences observed in recep-274 tivity and/or susceptibility between does within a flock 275 depending on the age or on the level of production were 276 used to provide the rationale for a selective application 277 of anthelmintics (Hoste et al., 2002a). Our current re-278 sults confirm that ewes in first lactation should repre-279 sent the target of specific surveillance and possible se-280 lective treatments within dairy flocks as it has also been 281 suggested in meat producing sheep (Leathwick et al., 282 1995). However, in multiparous ewes, it appears from 283 our study that the identification of animals to be treated 284 could not rely only on epidemiological information on 285 differences in receptivity to nematode infection. Con-286 sequently, innovative methods of diagnosis should be 287 developed and evaluated for this category of animals.

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288 Uncited reference

289 Sangster (2001).

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