

# Sustainable control of worms in sheep



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SCOPS is an industry-wide initiative including representation from

NSA (Chair) SNFU Defra SAC NOAH AHDA RUMA CSL RVC SVS



#### The SCOPS terms of reference are :

- To advise and disseminate new recommendations on sustainable parasite control to the sheep industry, initially concentrating on internal parasites
- To provide a forum for feedback from the sheep and animal health industries, veterinary profession and allied groups
- To consider new developments, feedback and information and revise the recommendations accordingly
- To facilitate mechanisms to inform all stakeholders in the sheep industry. Ensure that the messages have consistency and clarity.



### Introduction

Over the past 20-30 years :

- 1. dependence on anthelmintic use has increased
- 2. anthelmintic resistance has emerged as a problem worldwide and, latterly, in the UK
- 3. parasite epidemiology has changed
- 4. there is new understanding of AR and its control
- 5. some of the strategies which have been recommended for worm control select for AR



# The increase in ewe numbers by sector 1971 to 2000





Data from Moredun Research Institute

## The parasites

The major worm parasites of sheep in the UK include

- Gastrointestinal nematodes
- The trematode Fasciola hepatica
- The lung worms



### The parasites

 This presentation is concerned with the gastrointestinal nematodes, and not the lungworms,

 and brief mention is made of Fasciola hepatica.



### **Anthelmintics**

Broad-spectrum anthelmintics fall into three main classes:

- BZ
  benzimidazoles
- LM
   levamisole and morantel
- ML
  - macrocyclic lactones



# What is anthelmintic resistance (AR)?

### AR exists if

- the parasite can tolerate anthelmintic doses which are normally lethal
- the ability to do so is heritable



### How is resistance measured?

- faecal egg count reduction trials (FECRT)
  - resistance is declared if dosing does not reduce FEC by at least 95%
  - anthelmintics may 'appear' to be still working even if reduction in FEC is only 60% to 80%
- resistance is also measured in laboratorybased larval development assays



## How does resistance appear?

- resistance alleles pre-exist in most worm populations even before anthelmintics are ever used
- then, when the anthelmintic is used, the very few worms with resistance alleles are favoured
- resistance develops slowly at first, then more rapidly as allele frequency increases



Frequency of resistance alleles and homozygous resistant worms in a worm population developing anthelmintic resistance



Frequency of resistance alleles and homozygous resistant worms in a worm population developing anthelmintic resistance



Frequency of resistance alleles and homozygous resistant worms in a worm population developing anthelmintic resistance

![](_page_16_Figure_1.jpeg)

Will resistance go away if the farmer stops using the anthelmintic?

- the short answer is 'No!'
  - once resistance to an anthelmintic emerges, reversion to susceptibility is unlikely to occur

![](_page_17_Picture_3.jpeg)

### **Reversion to susceptibility**

- Resistant alleles make worms less fit to survive in the absence of anthelmintic
- So, in theory, reversion to susceptibility should occur when the anthelmintic is not used
- Possibly, this happens in zone A
- It appears, however, that once AR is in zone B, co-adaptation to survival means that resistant parasites are equally fit for survival as susceptible ones.

![](_page_18_Figure_5.jpeg)

## What factors influence the rate of AR development?

- 1. The relative size of the *in-refugia* population.
- 2. Frequency of treatment
- 3. Rate of re-infection after dosing
- 4. Dose rates

![](_page_19_Picture_5.jpeg)

#### Exposed population

![](_page_20_Figure_1.jpeg)

#### **Exposed** population

![](_page_21_Figure_1.jpeg)

#### In-refugia population

Factors influencing the rate of AR development 1. The relative size of the *in-refugia* population

- The larger the *in-refugia* population, relative to the exposed population, the slower AR will develop.
- When an entire group of sheep is treated prior to a move to a low-contamination pasture, the in-refugia population is relatively <u>small</u>.

![](_page_22_Picture_3.jpeg)

## What factors influence the rate of AR development?

- 1. The relative size of the *in-refugia* population.
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![](_page_23_Picture_5.jpeg)

#### Factors influencing the rate of AR development

2. Frequency of treatment

- More frequent treatment selects faster for AR
- Treatment is particularly selective when frequency approaches the pre-patent period
- Treatment gives the resistant worms a reproductive advantage over susceptible worms

![](_page_24_Picture_5.jpeg)

## What factors influence the rate of AR development?

- 1. The relative size of the *in-refugia* population.
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- 3. Rate of re-infection after dosing
- 4. Dose rates

![](_page_25_Picture_5.jpeg)

#### Factors influencing the rate of AR development

#### 3. Rate of re-infection after dosing

- After dosing, resistant parasites have a period of reproductive advantage
- The period is shorter if the sheep become quickly reinfected.
- If re-infection is delayed, resistant survivors have the advantage for longer.

![](_page_26_Picture_5.jpeg)

#### Rapid re-infection after dosing

- The factors which influence re-infection rates after dosing are
  - the infectivity of the pasture
  - the susceptibility of the sheep
    - lambs >> ewes
- Dosing of immune ewes may be a significant factor selecting for AR

![](_page_27_Picture_6.jpeg)

## What factors influence the rate of AR development?

- 1. The relative size of the *in-refugia* population.
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![](_page_28_Picture_5.jpeg)

#### Factors influencing the rate of AR development

#### 4. Dose rates

- Under-dosing encouraged the rapid appearance of AR to the BZ and LM anthelmintics
- Under-dosing allows heterozygous parasites to survive
- Full doses should kill all but homozygous-resistant parasites

![](_page_29_Picture_5.jpeg)

# What can be done to delay AR?

- 1. Rotations of anthelmintics
- 2. Combinations of anthelmintics
- 3. Prevent the entry of resistant worms onto farms from other farms.

![](_page_30_Picture_4.jpeg)

# What can be done to delay AR?

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![](_page_31_Picture_4.jpeg)

# What can be done to delay AR?

- 1. Rotations of anthelmintics
- 2. Combinations of anthelmintics
- 3. Prevent the entry of resistant worms onto farms from other farms.

![](_page_32_Picture_4.jpeg)

# The new guidelines

- An 8 step strategy
- Many of the recommended steps are unchanged from previous guidelines
- There are some key new recommendations, as a result of research and experience in UK and other countries
- Importance of involving expert advice is emphasised

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#### The new guidelines for anthelmintic use and worm control

- 1. Work out a control strategy with your veterinarian or advisor.
- 2. Use effective quarantine strategies to prevent the importation of resistant worms in introduced sheep and goats
- 3. Test for AR on your farm
- 4. Administer anthelmintics effectively
- 5. Use anthelmintics only when necessary
- 6. Select the appropriate anthelmintic for the task
- 7. Adopt strategies to preserve susceptible worms on the farm
- 8. Reduce dependence on anthelmintics

1. Work out a control strategy with your veterinarian or advisor.

- The need for specialist consultation is greater now than before.
- Decisions about judicious use of anthelmintics in worm control programs are complex, and will require on-going consultations

![](_page_35_Picture_3.jpeg)

- Introduction of resistance alleles is considered a major cause of AR in UK flocks.
- The recommended strategy involves three steps:

![](_page_36_Picture_3.jpeg)

#### Step 1

- Treat all introduced sheep and goats with levamisole plus an ML
- Do not mix, dose sequentially
- Give full doses of each drug

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![](_page_37_Picture_6.jpeg)

- Step 2
  - After treatment, hold animals off pasture for 24-48 hours, to empty out any worm eggs
  - Supply feed and water during that time
  - Collect faeces passed during that time
    - do not apply to pastures
    - consider incineration, for example

![](_page_38_Picture_7.jpeg)

- Step 3
  - Then place sheep on contaminated pastures
    - to allow dilution of eggs from any surviving worm parasites
    - to encourage rapid re-infection with worms endemic to the farm.

![](_page_39_Picture_5.jpeg)

#### 3. Test for AR on your farm

- Sheep farmers must be strongly encouraged to test for AR
- A knowledge of each drug's efficacy is fundamental
- Without this knowledge
  - adequate worm control may not occur
  - sensible drug rotations cannot be planned

![](_page_40_Picture_6.jpeg)

# 4. Administer anthelmintics effectively

- Dose for the heaviest in the group
- Check the gun is
   working satisfactorily
- Administer the drug correctly

![](_page_41_Picture_4.jpeg)

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![](_page_42_Picture_4.jpeg)

![](_page_42_Picture_5.jpeg)

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![](_page_43_Picture_4.jpeg)

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# 5. Use anthelmintics only when necessary

 Carefully evaluate the need to dose ewes at tupping

![](_page_44_Picture_2.jpeg)

# 5. Use anthelmintics only when necessary

- Carefully evaluate the need to dose ewes at tupping
- If dosing ewes at turnout
  - use highly efficacious treatments
  - leave some ewes untreated
  - treat well before the end of PPRI

![](_page_45_Picture_6.jpeg)

![](_page_45_Picture_7.jpeg)

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- Carefully evaluate the need to dose ewes at tupping
- If dosing ewes at turnout
  - use highly efficacious treatments
  - leave some ewes untreated
  - treat well before the end of PPRI
- Use FEC monitoring to assist decision-making

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#### 6. Select the appropriate anthelmintic

- Use narrow-spectrum drugs when possible
   *eg*, closantel for *Haemonchus*
- Avoid off-target use
  - particularly in fluke-nematode combinations
- Rotate anthelmintics when appropriate
  - do not let rotation choice over-rule decisions about quarantine treatment, or narrow-spectrum drugs
- Consider risks & advantages of persistency of some anthelmintics

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# 7. Preserve susceptible worms on the farm

- The dose-and-move strategy has been identified as potentially selective for AR
  - part-flock treatment is expected to reduce selection
    - leave 10% untreated (5% to 20%)
    - use highly efficacious treatments (>99% efficacy)
  - delay the 'move' after the 'dose'

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# 8. Reduce dependence on anthelmintics

 Use grazing management, rather than anthelmintics, to provide 'safe' grazing

![](_page_49_Picture_2.jpeg)

![](_page_49_Picture_3.jpeg)

# 8. Reduce dependence on anthelmintics

FEC EBV = -50 EPG

EC EBV +50 EPC

- Use grazing management, rather than anthelmintics, to provide 'safe' grazing
- Use rams selected for low FEC to breed ewe replacements

- Faecal egg counts (FECs) can give a useful guide to the level of parasitism in a flock of sheep
- But, there are important limitations to their use as a monitoring tool

![](_page_51_Picture_3.jpeg)

#### <u>Sample size</u>

- At least 10 animals should be sampled to estimate a group mean FEC
- A 'group' is a flock of animals of the same sex, age, reproductive status and treatment history, running in the same field
- The faeces from 10 sheep may be pooled at the laboratory it should not be mixed before then.

![](_page_52_Picture_5.jpeg)

- What is a suitable group?
  - Animals that are fully-fed and in satisfactory health
    - Results are reported as eggs per gram of faeces
    - If feed intake is impaired, faecal volume is reduced, and results are impossible to interpret

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#### <u>Collection of faeces</u>

- Gather the group, hold quietly in one area, then gather faeces from the pasture

![](_page_54_Picture_3.jpeg)

![](_page_54_Picture_4.jpeg)

#### <u>Collection of faeces</u>

- Gather the group, hold quietly in one area, then gather faeces from the pasture
- Place faeces in airtight container and cool
- Deliver to laboratory within 48 hours

![](_page_55_Picture_5.jpeg)

![](_page_55_Picture_6.jpeg)

#### **Collecting faecal samples**

- Gather the group into one place in the field.
- Remove the dog, and let them stand quietly.
- For a group of 200 ewes, 3-4 minutes is sufficient. Smaller groups require more time.

![](_page_56_Picture_4.jpeg)

![](_page_56_Picture_5.jpeg)

#### **Collecting faecal samples**

- Let the sheep move quietly away.
- Pick up faeces from the pasture and place in a container or small plastic bag.
- Select only warm, freshly-dropped specimens.
- Keep each specimen in a separate bag or container.

![](_page_57_Picture_5.jpeg)

#### Interpretation of results

- Interpret with local knowledge
- Remember: FECs cannot 'detect' burdens of immature worms
- Consider the different relationships between worm numbers and egg numbers in
  - different worm species
  - sheep of different age and reproductive status

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- Price and availability
  - A FEC test is available from a number of laboratories and veterinary practices
    - VLA labs offer a pooled test (10 samples) for £15.60 + VAT

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# Faecal egg count reduction test (FECRT)

- FECs can be used to detect the presence of AR
  - Simple tests
    - 7 to 14 days post-treatment
    - A quick and easy test for the presence of AR
  - Formal tests
    - Set up with randomised groups, and controls
    - Calculate a percent reduction
    - < 95% reduction implies resistance</li>

![](_page_60_Picture_9.jpeg)

# The liver fluke -*Fasciola hepatica*

![](_page_61_Picture_1.jpeg)

![](_page_61_Picture_2.jpeg)

## Liver fluke

- Liver fluke control is based on a number of drugs (fasciolicides) with different activities
- Resistance to some fasciolicides has developed in the UK and other countries
- Control programmes should consider the need to reduce selection pressure for resistance to these drugs
- Quarantine strategies should aim to reduce the risk of importing resistant fluke.

![](_page_62_Picture_5.jpeg)

![](_page_63_Figure_0.jpeg)

Multiplication up to 500 times or more in snail

#### Efficacy of fasciolicides available for use in sheep in the UK against susceptible fluke populations

(adapted from Fairweather and Boray, 1999).

	Age of fluke (weeks)													
Flukicide	1	2	3	4	5	6	7	8	9	10	11	12	13	14
Albendazole										50 -	70%	80 - 99%		
Oxyclozanide														
Nitroxynil						50 - 90%			91 - 99%					
Closantel														
Triclabendazole (T	CB)	90 -	99%	99 - 99.9%										

Preventing the development of resistance to flukicides

- Rotational use of TCB, closantel or nitroxynil, where appropriate
- Consider the use of drugs other than TCB when fluke burdens are expected to be entirely or mostly of adult fluke

![](_page_65_Picture_3.jpeg)

# Quarantine treatments for liver fluke

#### Performed for one of three reasons

- 1. Farm has no snail habitat
  - treat to improve the health of the sheep
- 2. Farm has snail habitat, but no fluketreat to prevent entry of all fluke
- 3. Farm has endemic fluke
  - treat to prevent entry of resistant fluke

![](_page_66_Picture_7.jpeg)

# Quarantine treatments for liver fluke

#### Develop a strategy after considering:

- 1. Resistance to TCB is still relatively uncommon in the UK
- 2. Treatment of TCB alone will not remove TCB-resistant fluke
- 3. Treatment with closantel or nitroxynil is expected to prevent the output of fluke eggs for at least 8 weeks
- 4. Resistance to closantel and to nitroxynil can occur.
- Treatment with more than one product will reduce the risk of introducing fluke with resistance to any one product.
  - but the use of two products at the same time may be injurious to health

![](_page_67_Picture_8.jpeg)

### The end

This presentation was brought to you by SCOPS.

Sustainable Control of Parasites in Sheep.

See also www.nationalsheep.org.uk

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![](_page_68_Picture_5.jpeg)