Anticoagulant Resistance Management Strategy For Pest Management Professionals, Central And Local Government And Other Competent Users Of Rodenticides
Technical Monograph 2003

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AIM

This document seeks to provide guidance to advisors, authorities, professionals and others on both the application of rodenticide use strategies that will avoid the development of resistance, and the recognition of anticoagulant resistance and how to manage it.

INTRODUCTION

It has now been shown over some fifty years that the use of anticoagulant rodenticides forms the most effective method of controlling commensal rodent populations.

The continued use of these anticoagulant rodenticides has however led to the development of resistance in some of the commensal rodent species. Such resistance is restricted geographically.

Remember, resistance is characterised by the ability of individuals within a rodent population in the field to continue feeding on the anticoagulant bait over many weeks, without being killed. It is not characterised by the reluctance of the rodents to feed on the baits.

Continuous feeding from anticoagulant baits may not only be due to resistance, but may also be caused by under-baiting or immigration. However, once these alternatives have been eliminated the probability that the cause of the continued feeding activity is anticoagulant resistance is high.

From the point of view of those undertaking field rodent control the term Practical Resistance is used to identify resistance that has led to the difficulty to control rodents in field situations.

"Anticoagulant resistance is a major loss of efficacy in practical conditions where the anticoagulant has been applied correctly, the loss of efficacy being due to the presence of a strain of rodent with a heritable and commensurately reduced sensitivity to the anticoagulant". Greaves, 1994*.

1 GENERAL INSTRUCTIONS ON THE CORRECT USE OF RODENTICIDES

BACKGROUND
The three most cosmopolitan rodent pest species are

- Ship (black, roof, fruit, house) rat - Rattus rattus
- Norway (brown, sewer, common) rat - Rattus norvegicus
- House mouse - Mus musculus spp

Here lies the key issue in any effective rodent control programme, “know the species”!

Each species has distinct behavioural and ecological requirements that must be clearly understood if effective management is to be achieved. The control techniques available must be clearly targeted to the characteristics of the species concerned. Mis-identification or failure to appreciate these characteristics will reduce the chances of effective control significantly.

The problems caused by rodents should not be seen as the problems of individuals, but as community problems that need to be addressed by the community as a whole. Attempts to solve rodent infestation across wider areas needs to be addressed on a community basis if the efforts are to be cost effective.

The effective control of rodent infestations should not be treated simply as a question of killing the rodents. The problem must be seen more broadly, as an infestation within a vulnerable environment.

The control strategy should be developed both in terms of reducing rodent numbers but also in terms of managing the environment. This will not only prevent the development of the rodent population but also prevent rodents within that environment coming into conflict with man and his interests.

The name given to this “total environment” approach to rodent control is termed Integrated Pest Management (IPM) and incorporates three main components

- Habitat management
- Control of rodent movement through proofing
- Control of the rodent population using appropriate chemical and physical control measures.
Before any IPM is undertaken it is first essential to understand the nature of the problem through the completion of a comprehensive survey and inspection of the area/s concerned. The objective of this survey is to identify

- The species involved.
- The extent of the infestation three dimensionally (horizontally and vertically).
- The severity of the infestation (roughly how many rodents there are).
- Where the rodents have come from.
- The harbourage being used.
- The food source(s) being used by the rodents.
- Non-target hazards and associated risks.

Only once these components have been identified can an effective IPM strategy be developed.

**HABITAT MANAGEMENT**

For a rodent infestation to thrive there must first be an environment that provides the three main features that rodents require. These are

- Food
- Harbourage
- Water (free water not necessarily required by house mice)

Therefore removing these components of the habitat as far as is practicable, or protecting them from access by rodents will make it less likely to be infested by commensal rodents.

**EXCLUSION AND PROOFING**

In situations where the rodent problem is caused by rodents moving from one area to another, it may be possible to prevent the rodents reaching the vulnerable crop, store, processing unit or building by the simple expedient of ensuring that physical barriers are placed in their way.

*Figure 1: Potential rodent access routes into a building.*
Mice are so small they can also be ‘delivered’ into premises in infested goods, e.g. inside a pallet containing sacks of grain or processed food products. In such cases, the warehouse or food store should have adequate ‘quarantine’ procedures to inspect incoming goods for evidence of mice.

CONTROL
One of the key characteristics that make rodents such successful mammals and pests is their ability to breed very rapidly. Whilst the reproductive rates of individual species will vary, a typical female commensal rodent can produce an average of eight young every month! Thus, to achieve effective control high levels of mortality are required, otherwise populations will return rapidly to their pre-control levels and the control operations will not be cost effective, see fig. 2.

Figure 2: Rates of recovery.

The most cost effective technique for controlling commensal rodent populations is to use chemical rodenticides. It is essential however that these are used safely so that non-target species are not harmed either through primary or secondary poisoning.

Chemical Control
The development of the chronic anticoagulant rodenticides in the early 1950s and subsequently has revolutionised rodent control. With the chronic rodenticides 100% mortality became the expectation rather than the exception and these chronic rodenticides are now the most widely used group of rodenticides.
They achieve their effect by interfering with the blood coagulation in the rodents that feed on them, resulting in death by haemorrhage. The particular advantage of these anticoagulants is that they have a delayed action and the minimum time to death is 2 days, with most deaths occurring between days 4 and 7. This delayed action prevents the rodent associating its illness with the rodenticide until it has consumed a dose of anticoagulant large enough to ensure it will die. The objective when using these chronic rodenticides is therefore to ensure that all rodents are able to feed at the bait points.

**Anticoagulant rodenticides available worldwide**

<table>
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<tr>
<th>First Generation Anticoagulants</th>
<th>Second Generation Anticoagulants</th>
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<tr>
<td>Warfarin</td>
<td>Difenacoum</td>
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<tr>
<td>Chloropacinone</td>
<td>Bromadiolone</td>
</tr>
<tr>
<td>Coumachlor</td>
<td>Difethialone</td>
</tr>
<tr>
<td>Diphacinone</td>
<td>Brodifacoum</td>
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<tr>
<td>Coumatetralyl</td>
<td>Flocoumafen</td>
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It is essential that the anticoagulant rodenticide is placed in such a way as to ensure that the target rodents are able to feed on the baits, but that children and non-target species do not have access to the bait:

- by using suitable materials available on site to cover and secure bait points,
- by using commercially available tamper-resistant bait stations,
- by using self made bait stations,
- by placement only in secure premises,
- by placing baits in areas inaccessible to non-target species,
- by undertaking frequent visits to the site.

**Remember** that before using an anticoagulant rodenticide
- only use a product that has been approved in the country of use,
- read the label and follow the directions for use,
- undertake risk assessments relating to both the safety of the user, the environment and non-target species.
Remember when using an anticoagulant rodenticide

- to place the bait in areas of rodent activity identified in the survey,
- ensure that revisits are made to the baiting points at weekly intervals or more frequently where necessary,
- prevent non-target access to the baits,
- replenish eaten bait at the revisits as instructed on the label,
- search for dead and dying rodents at the revisits and dispose of these safely,
- when dead and dying rodents are being found frequently, increase the frequency of the re-visits, to reduce potential impact on the environment,
- ensure that no spillage of bait is taking place and remove any spillage that does occur,
- wear the protective clothing recommended on the label,
- inform those living and working in the area of the rodent treatment,
- keep records of bait placement and monitor bait consumption at revisits.

Remember after completion of the anticoagulant treatment

- remove all surplus baits
- the baiting positions may be left in place as monitoring points or to reduce neophobic response during subsequent treatments
- non toxic or placebo baits may be left in position to assist with the monitoring for rodent activity apply environmental, proofing and hygiene requirements as a part of the Integrated Pest Management strategy
- store all anticoagulant rodenticides safely
- complete the records to assess the success of the treatment.

Where the anticoagulant rodenticide is used correctly 100% control of the rodents should be achieved. However if the infestation is proving persistent follow the Checklist for Anticoagulant Rodenticide Users (Appendix 1).
Record Keeping
It is essential to maintain effective records throughout the treatment, these should include the following elements
- positions of all bait placements,
- record details of bait used,
- quantities of bait laid,
- details of visits to check and replenish baits,
- bait consumption determined at revisits,
- details of IPM strategy implementation.

The maintenance of an effective record keeping system enables the more reliable determination of the progress of the control operations and the determination of the causes of treatment failure, including resistance.

If necessary the records may be plotted in a form similar to Figure 3 to determine progress with the treatment.

Figure 3: Progress of treatment.
2 RECOMMENDATIONS FOR THE TREATMENT OF POPULATIONS CONTAINING RESISTANT INDIVIDUALS

PRACTICAL RESISTANCE

Early evidence of resistance in house mouse and Norway rats involved first generation anticoagulants warfarin and diphacinone. In such cases, it was not possible to control the rodents with these rodenticides.

Subsequently, laboratory studies have produced a series of tests to identify resistance against a range of active ingredients. However, many of the tests are very sensitive, and the level of resistance they identify may not be sufficient to affect practical control. This is particularly the case for the second generation anticoagulants. The term “Technical Resistance” is used to describe such resistance.

Recent work funded by RRAC of CropLife International and published on the web site (www.croplife.org, at library, monographs) is the first practical attempt to address this issue. Susceptibility baselines and tests have been developed for nine active ingredients against Norway rat, and five active ingredients against house mice, which allow the determination of a Resistance Factor as a measure of the level of resistance.

Resistance Factor is simply the multiple of the baseline dose required by the non-resistant, susceptible, animal that achieves the same response in a resistant animal. These tests can be used to give a clear indication of the likely practical impact of the resistance on field efficacy.

With Practical Resistance, Resistance Factors are high and treatment outcome is adversely affected; with Technical Resistance, Resistance Factors may be low, and there is not necessarily a discernible affect on treatment outcome.

For the Field Technician, the first evidence of Practical Resistance will be high bait consumption records on the third and subsequent weeks of treatment (Fig 3). The assessment of the number of bait points with take on each bait application (plotted as indicated on Fig 3), is a simple method requiring minimal additional effort, that can be used to provide evidence of Practical Resistance, although it should be remembered that under-baiting and immigration can have a similar effect on overall bait take; and that immigration may be difficult to assess in the field.
STRATEGIC USE OF RODENTICIDES

The important issues here are firstly to identify strategies for avoiding the development of resistance in susceptible rodent populations and secondly to identify strategies for managing resistance to the anticoagulants when it is suspected or identified.

Remember that the normal strategy used for managing resistance in populations of insects, weeds or other pests is to rotate the control between different groups of pesticide, targeting as they do, different control mechanisms.

Unfortunately, the anticoagulant rodenticides all work in much the same way and the nature of the resistance to the different anticoagulants is so similar that simply rotating between the anticoagulants is not a reliable means of managing anticoagulant resistance. However, using anticoagulants of higher toxicity plays a major part in resistance management. In case of confirmed practical resistance, an anticoagulant rodenticide of higher toxicity compared to that, which is hit by resistance, should be used to eradicate the infestation. In some cases, especially with mice, alternations with non-anticoagulants can be part of the strategy.
SUMMARY

TO AVOID THE DEVELOPMENT OF RESISTANCE IN SUSCEPTIBLE RODENT POPULATIONS

- Use anticoagulant rodenticides. Ensure that all baiting points are inspected weekly and old bait replaced where necessary.
- Undertake treatment according to the label until the infestation is completely cleared.
- On completion of the treatment remove all unused baits.
- Do not use anticoagulant rodenticides as permanent baits routinely. Use permanent baits only where there is a clear and identified risk of immigration or introduction or where protection is afforded to high-risk areas.
- Monitoring of rodent activity should be undertaken using visual survey, through the use of non-toxic placebo monitors or by other effective means.
- Record details of treatment.
- Where rodent activity persists due to problems other than resistance, use alternative baits or baiting strategy, extend the baiting programme or apply alternative control techniques to eliminate the residual infestation (acute or sub-acute rodenticides, gassing or trapping).
- Ensure that complete elimination of the infestation is achieved.
- As appropriate during the rodenticide treatment apply effective Integrated Pest Management measures (remove alternative food sources, remove water sources, remove harbourage and proof susceptible areas against rodent access).

TREATMENT OF RODENT INFESTATIONS CONTAINING RESISTANT INDIVIDUALS

- Where rodent infestations containing resistant individuals are identified, immediately use an alternative anticoagulant of higher potency. If in doubt, seek expert advice on the local circumstances.
- Alternatively use an acute or sub-acute but non anticoagulant rodenticide.
- In both cases it is essential that complete elimination of the rodent population is achieved. Where residual activity is identified apply intensive trapping to eliminate remaining rodents. Gassing or fumigation may be useful in specific situations.
- Apply thorough Integrated Pest Management procedures (environmental hygiene, proofing and exclusion).
- Do not use anticoagulant rodenticides as permanent baits as routine. Use permanent baits only where there is a clear and identified risk of immigration or introduction or where protection is afforded to high risk areas.
- Record details of treatment.
APPLICATION OF AREA OR BLOCK RODENT CONTROL TO ELIMINATE RESISTANCE

- Where individual infestations are found to be resistant or contain resistant individuals it is possible that the resistance extends further to neighbouring properties.
- Where there are indications that resistance may be more extensive than a single infestation, apply area or block control rodent programmes.
- The area under such management should extend at least to the boundaries of the area of known resistance and ideally beyond.
- These programmes must be effectively coordinated and should encompass the procedures identified above.
APPENDIX

RRAC Checklist for Rodenticide Users Experiencing Difficulties.

Issued by RRAC. RRAC is a group formed by senior technical specialists within CropLife International. Members are BASF, Bayer CropScience, Liphatech S.A., Rentokil, Sorex and Syngenta. The committee would like to thank Adrian Meyer for facilitating the development of this advice.
The information contained in this monograph is accurate to the best of the knowledge of CropLife International, but no liability can be accepted whatsoever in respect of the use of this information nor in respect of any advice contained herein.
The Rodenticide Resistance Action Committee (RRAC) is a working group within the framework of CropLife International. Participating companies include: Bayer CropScience, LiphaTech S.A., BASF, Pelgar, Rentokil, Syngenta and ZAPI. Senior technical specialists, with specific expertise in rodenticides, represent their companies on this committee.

The objective of RRAC is to advise international agencies, government bodies, regulatory bodies and rodenticide users on technical matters relating to rodenticide resistance by: producing guidance leaflets for rodenticide users on the safe and effective use of rodenticide products; organizing seminars and conferences whereby industry members can meet and exchange ideas with experts from universities, governments and international organizations; participating in trade exhibitions and other similar events; sponsoring research projects on rodenticide resistance; developing and advocating the use of effective resistance management strategies.

Rodenticides are extremely valuable products in the prevention of human and animal disease and the protection of property, agriculture and the environment. New molecules are difficult and expensive to bring on to the market. Through its activities, the overriding intention of RRAC is to develop its understanding of resistance and resistance management in order to maintain the effectiveness of currently available rodenticides.

WHAT IS RRAC?

Rodenticide Resistance Action Committee (RRAC)
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1. Not all rodenticides are labelled for use against all rodent species because rodents differ in susceptibility to certain active ingredients. When using concentrates ensure that mixing is carried out exactly according to label instructions.

2. The positioning of baits is often critical. They should be appropriately placed, where there are signs of rodent activity, for example in runs between the rodent harbourage and normal feeding points, in areas where droppings and other signs of activity are seen. A bait point placed even a meter away from a well used run may not be discovered. Thorough site exploration is essential see 5.

In sites where the rodents' natural food is highly attractive some may not be sufficiently palatable. A change of bait base or a change to another approved product can often solve this problem. When possible alternative food should be removed or sealed.

3. All anticoagulant rodenticides are slow acting, several days are required to exhibit a lethal effect. In addition, even in a moderate infestation it may take some individuals several days to take the bait. Complete eradication may take some weeks.

4. It is important not to underestimate the size of the infestation. In these cases, completely consumed baits are a sure sign that inadequate quantities and/or number of baiting points are being used.

5. It is important to bait not only areas where activity is obvious but to discover harbourages which are hidden or away from the main site. If these are neglected they will act as a reservoir of population. Thorough investigation is absolutely essential.

6. When an infestation has been eliminated by the effective use of a rodenticide, neighboring rodents may rapidly invade the de-populated territory and give the impression that the product has failed. Check surrounding properties for signs of infestation and bait if possible and/or consider perimeter baiting and proofing.

Product labels contain precautionary information and advice on usage. As with all pesticides it is essential that labels are read and understood before attempting to use a rodenticide.