



Rodenticide Resistance: Introduction and the RRAC

Dr. Stefan Endepols
Envu and RRAC

SKABRA
Trondheim, September 10th 2024



The Rodenticide Resistance Action Committee (RRAC) is a working group within the framework of CropLife International. Participating companies are:

BASF

Envu

LiphaTech S. A.

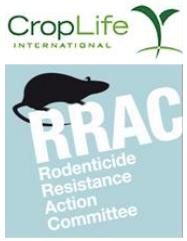
Pelgar

Rentokil Initial

Syngenta

ZAPI

Aktiva



The RRAC Resistance guide Co-Authors

Provided un-published data to resistance maps:

**Colin Prescott, Clare Jones & colleagues, D Rymer, K Baert, Tanja Blazic,
H-M Kohn, Freise & Runge, J Mooney, Y Motro, A Iacucci, T Montalvo,
RRAC, St Endepols, and others**

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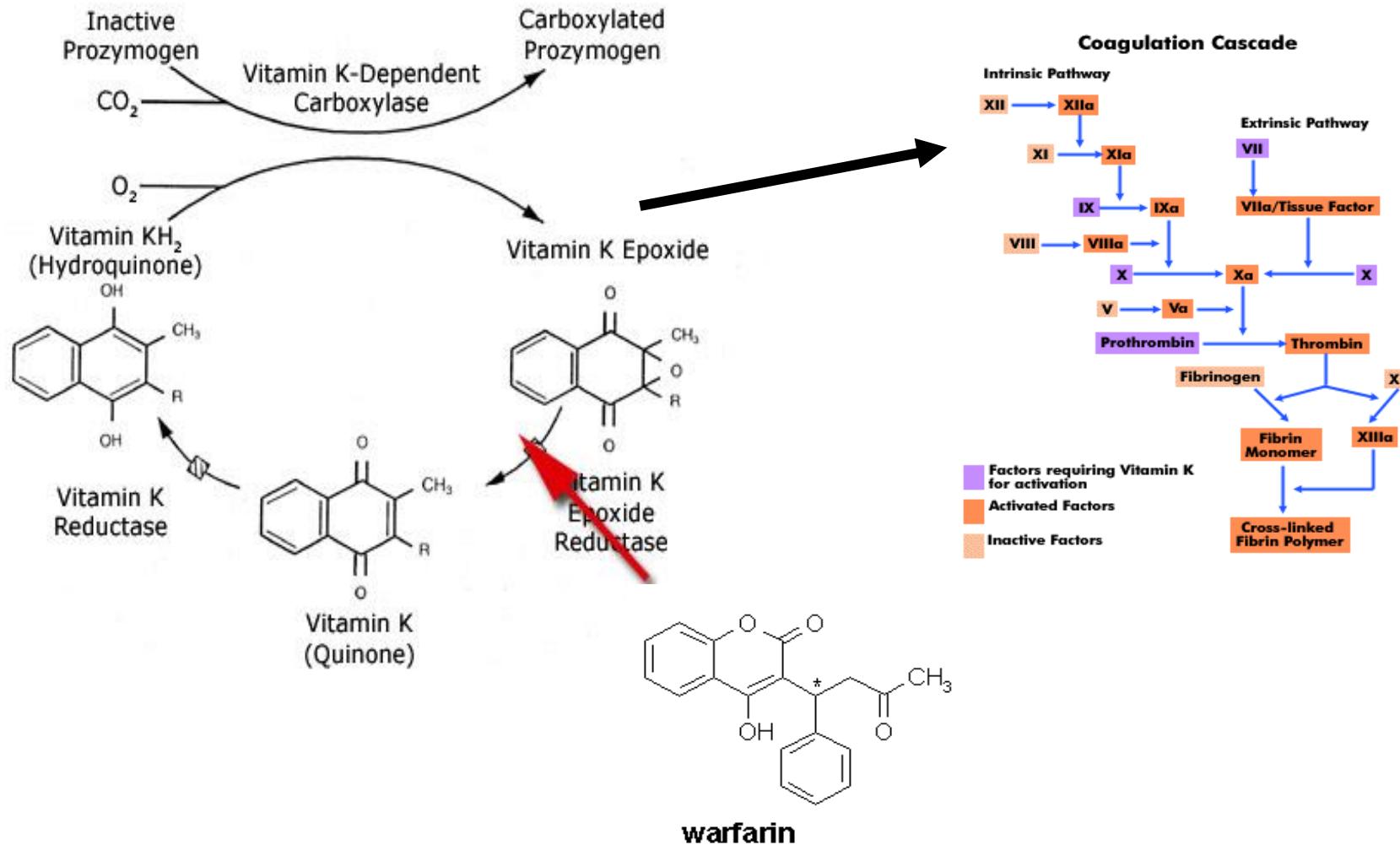
Webdesign:

Heiko Heeren, Kölner Medienwerk GmbH, Cologne



Principle of Anticoagulant Resistance and Resistant Strains

Anticoagulants inhibit the *Vkorc1* gene that drives the vitamin K cycle and blood coagulation

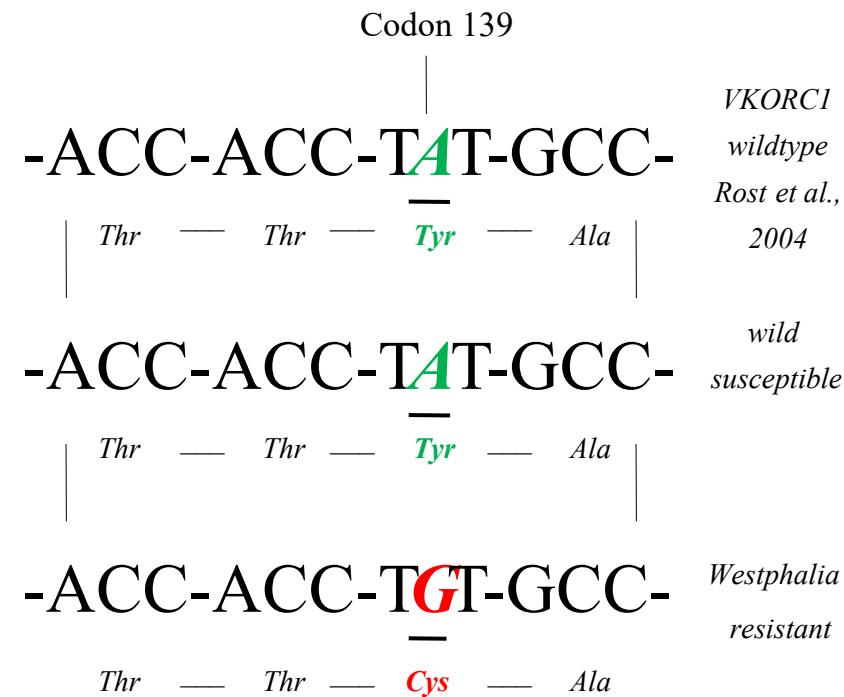


Rat: VKOR Variant Strain Westphalia resistant

Gene variant **Tyr139Cys (= Y139C)**:

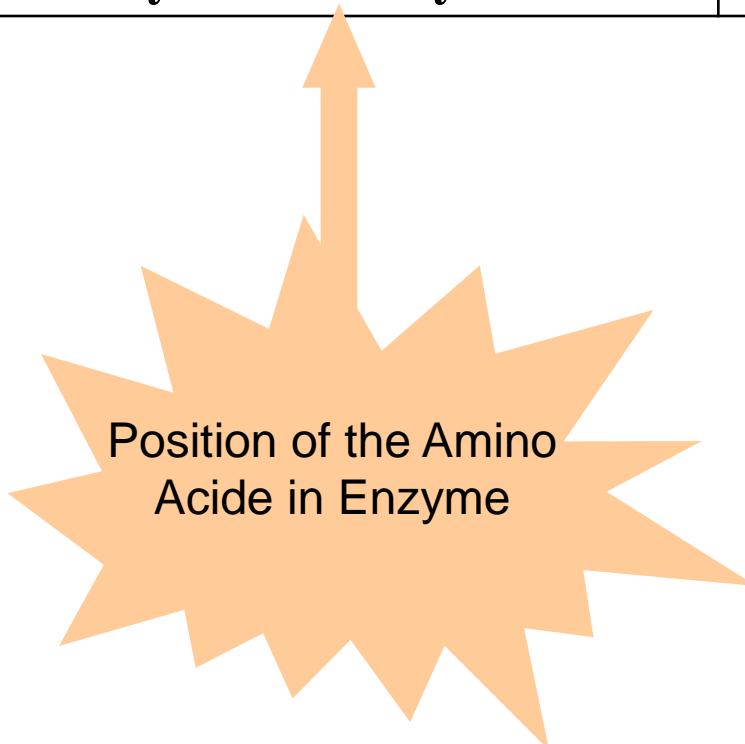
At Position 139 in the VKOR-gene: exchange of amino acid *Tyrosin* by *Cystein*. Due to the exchange on the gene of one nucleotide (*Adenine/Guanine*)

Marker of strain Westphalia-resistant, also present in NL, Be, UK, F, DK.



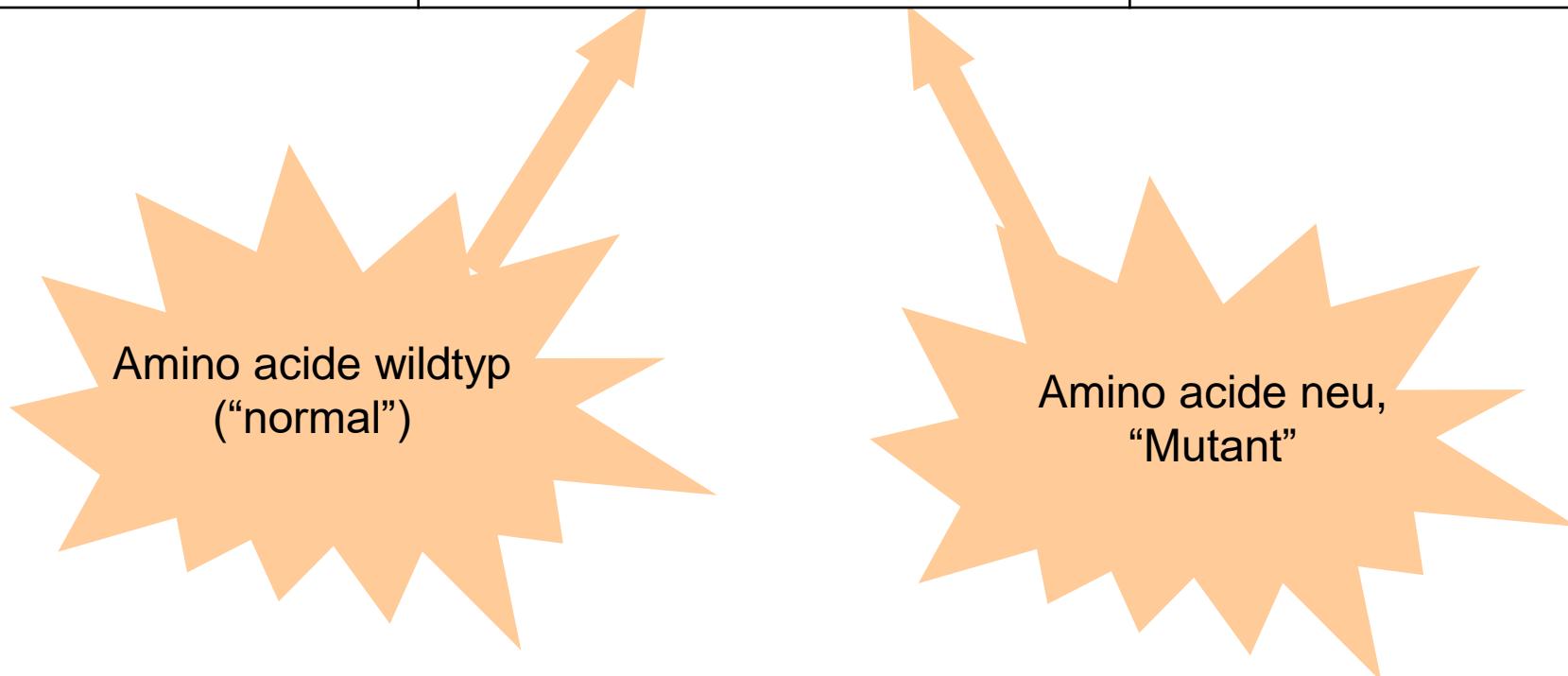
Marking the Genotypes of VKOR

| Resistance area | VKOR Polymorphism | |
|--------------------------------------|----------------------------|-------------------------|
| Westphalia, DK, NL, F, UK | Tyrosin 139 Cystein | Tyr139Cys, Y139C |



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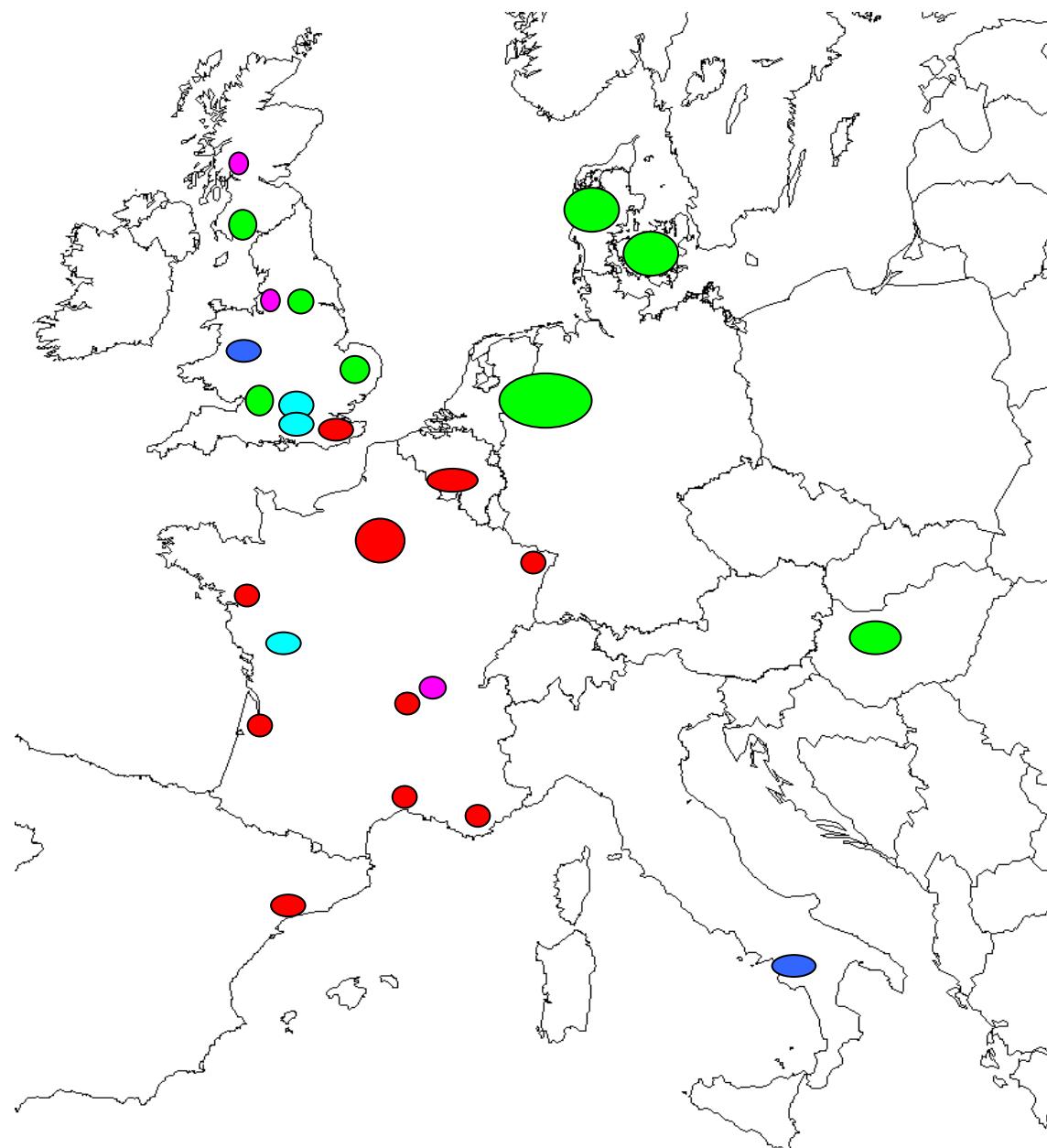
RRAC, study at JKI: Rat **Y139C**

Determination of effective doses in BCR test:
ED₅₀Baseline (mg/kg) and Resistance Factor (RF).

| | ED₅₀ male | ED₅₀ female | RF m | RF f | |
|---------------|---------------------------------|-----------------------------------|-----------------|-----------------|----------------------|
| Coumatetralyl | 0.36 | 0.44 | 42.2 | 94.1 | Endepols et al. 2007 |
| Bromadiolone | 0.47 | 0.62 | 17.0 | 15.5 | Endepols et al. 2007 |
| Difenacoum | 0.65 | 0.79 | 1.6 | 2.9 | JKI |
| Brodifacoum | 0.22 | 0.23 | 1.2 | 1.8 | JKI |
| Flocoumafen | 0.29 | 0.34 | 0.8 | 1.0 | JKI |
| Difethialone | 0.43 | 0.49 | 0.5 | 0.8 | JKI |

Norway rat: Resistant Strains in Europe

Y139C ●
Y139F ●
Y139S ●
L120Q ●
L128Q ●



SNPs confer practical resistance to AVKs of Brown Rat (*R. norvegicus*) and House Mouse (*M. musculus*)

AVKs Active Ingredients are

- Either active in practice and recommended to control these strains 
- Or partially or totally inactive and not recommended for control of them 

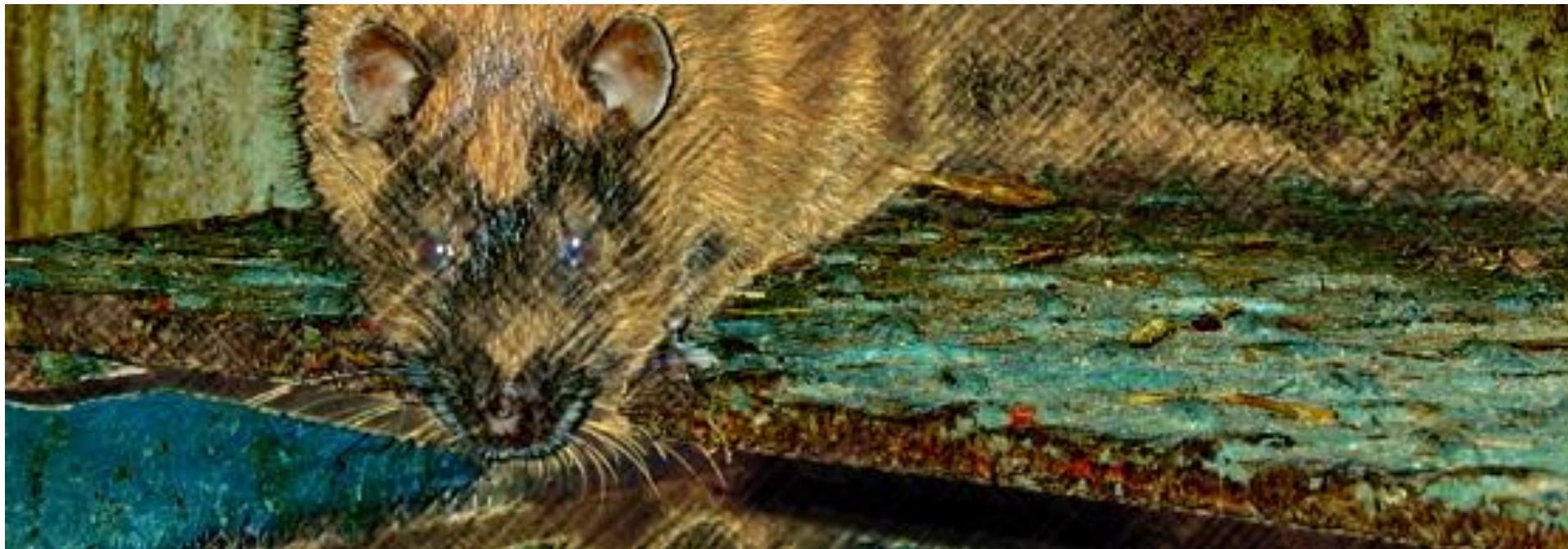
Brown Rats

| VKOR SNPs | FGARS | Bromadiolone | Difenacoum | Brodifacoum | Flocoumafen | Difethialone | Different MoA: Cholecalciferol |
|-----------|-------|--------------|------------|-------------|-------------|--------------|--------------------------------|
| L120Q | | | | | | | |
| L128Q | | | | | | | |
| Y139C | | | | | | | |
| Y139F | | | | | | | |
| Y139S | | | | | | | |

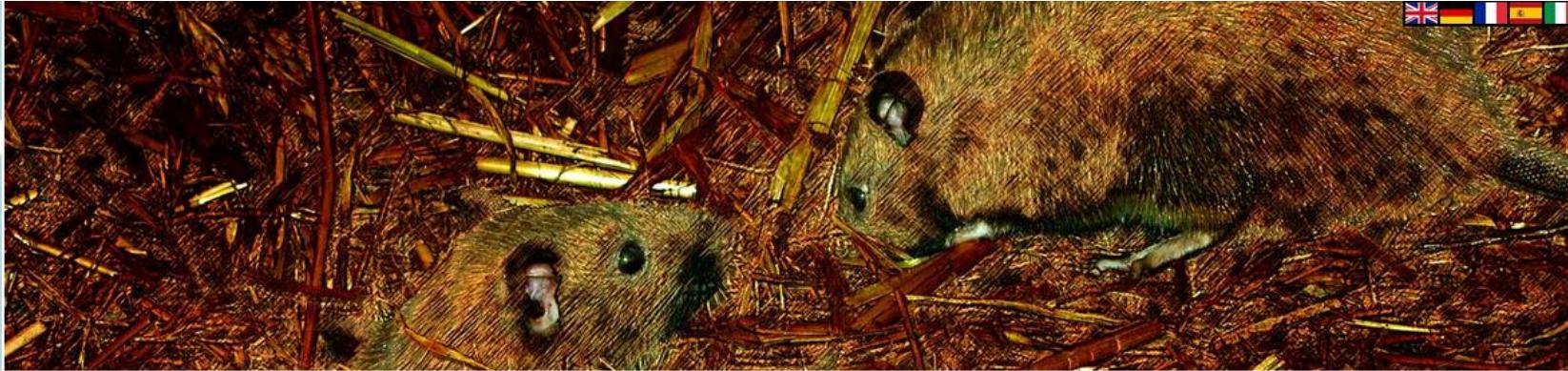
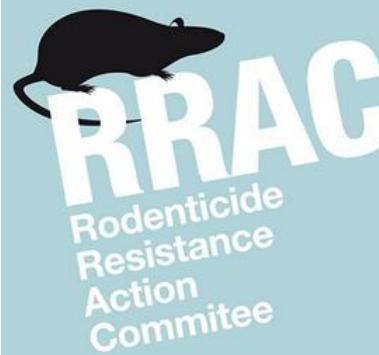
House Mice

| VKOR SNPs | FGARS | Bromadiolone | Difenacoum | Brodifacoum | Flocoumafen | Difethialone | Different MoA: Cholecalciferol |
|--------------------------|-------|-------------------|------------|-------------|-------------|--------------|--------------------------------|
| L120Q | | no data available | | | | | |
| L128S | | | | | | | |
| Y139C | | | | | | | |
| Y139S | | no data available | | | | | |
| spretus introgression | | | | | | | |

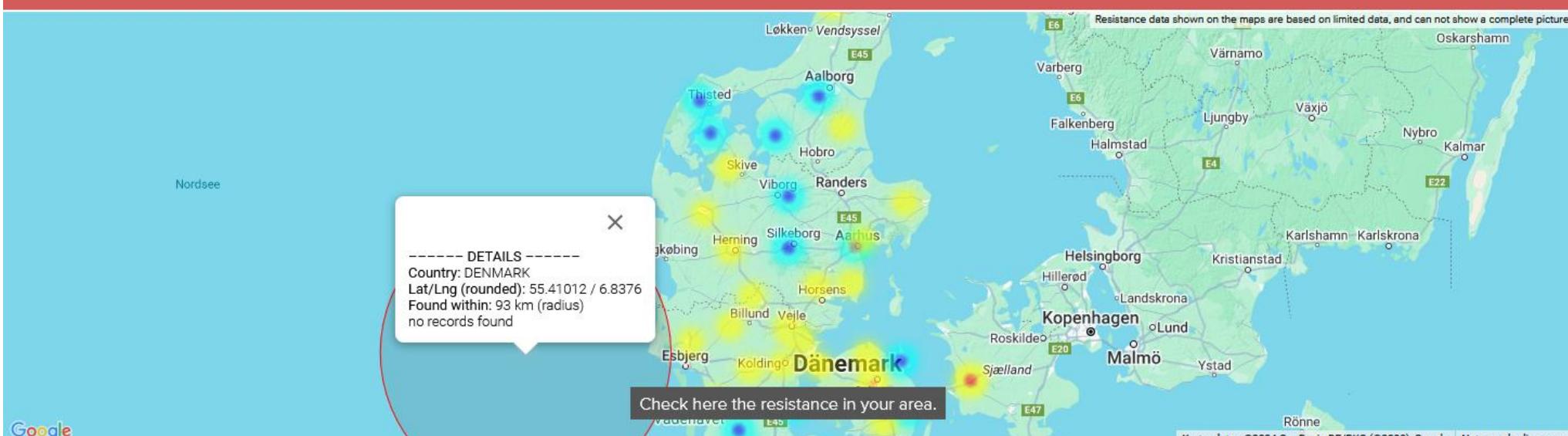
RRAC – www.rrac.info



Website Content



Now included in resistance maps: **4,550** Datapoints



NEW

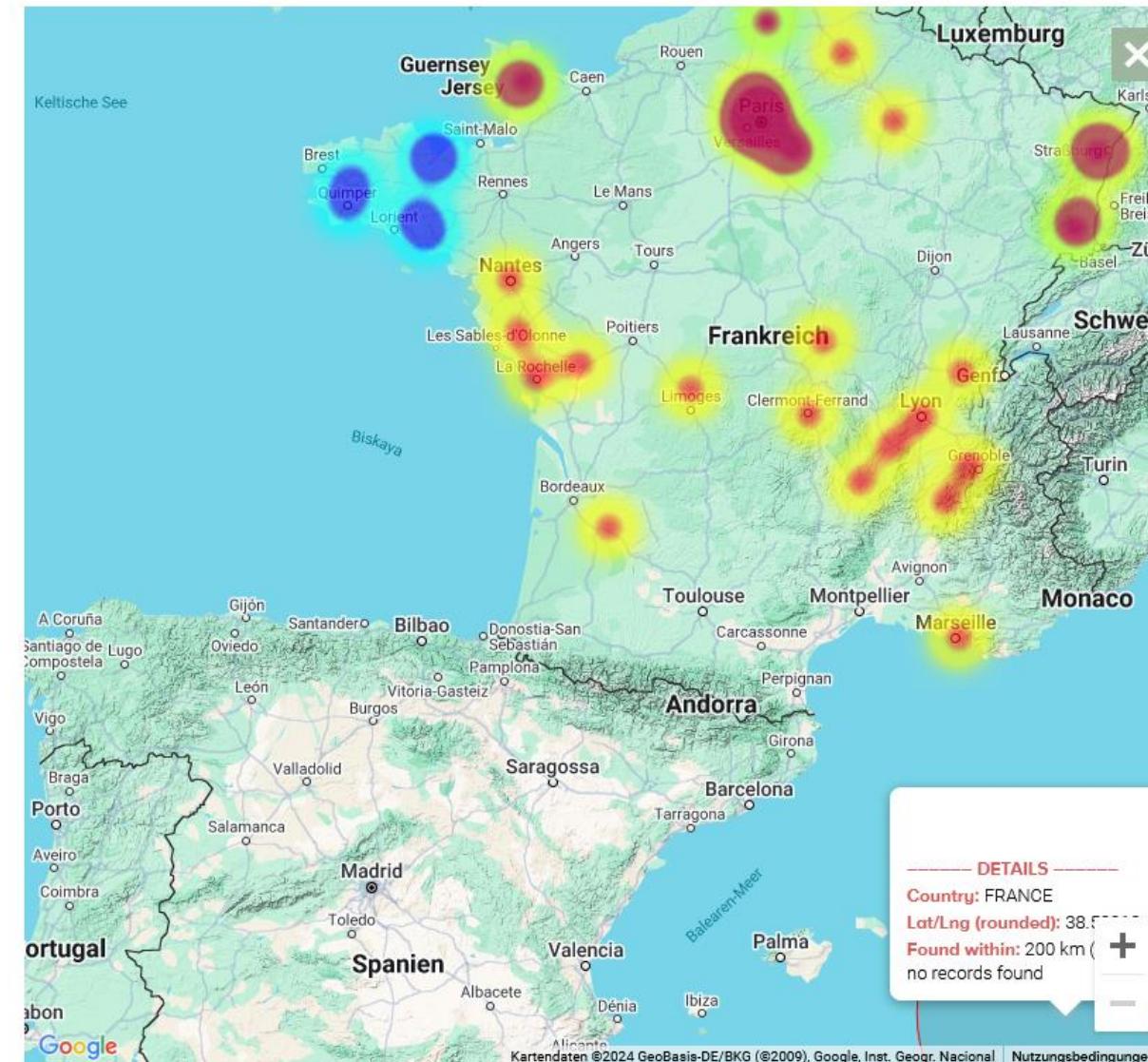
Hybrid-resistant mice strains

Proofs of the recently discovered hybrid-resistant mice strains are now visible on resistance maps of France, Germany, and the United Kingdom.

Resistance maps

[Norway Rat](#)[House Mouse](#)[Africa](#)[America](#)[Asia](#)[Europe](#)[Austria](#)[Azores](#)[France](#)[Germany](#)[Greece](#)[Ireland](#)[Italy](#)[Netherlands](#)[Serbia](#)[Spain](#)[Switzerland](#)[United Kingdom](#)

France



Resistance

Susceptible

Leave "RRAC guidelines", go back to "RRAC home".

With the following buttons you enable ("on") or disable ("off") points on the map.

Type

 Resistance Susceptible

Strains

 L128S Y139C spretus spretus + L128S

More functions

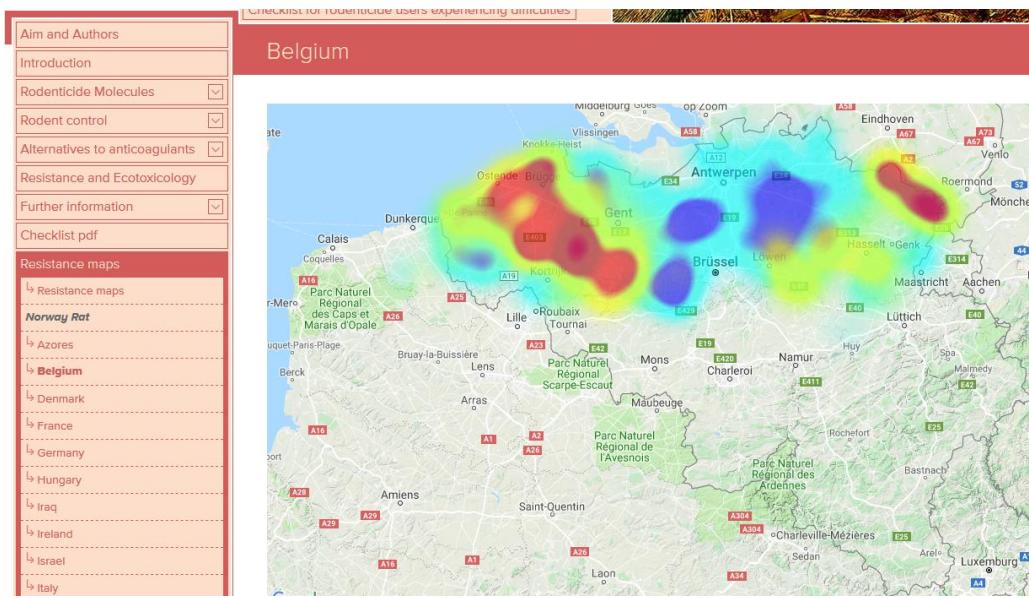
 Reset map position to France Disable surrounding info on map. Show map fullscreen.

Time range: 2016 - 2017

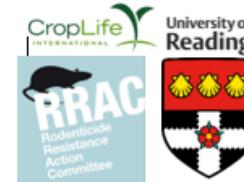
Heatmap intensity: 0.6

Heatmap opacity: 0.6

RRAC – www.rrac.info



Need for samples from white areas:
Contact me for details.



RRAC Anticoagulant Resistance Mapping Project SAMPLE COLLECTION PROTOCOL

With this initiative, the RRAC aims to further complete the picture on the distribution of anticoagulant resistance in Norway rats and in house mouse. We ask persons, companies and institutions to support this project by sending in samples to the University of Reading, our research partner for genetic analysis. The RRAC funds a certain number of analysis per year, in particular samples coming from areas with no data on resistance and susceptibility, respectively. This research is not aimed for quick operational support, however, as soon as data are generated, these will be visualised on resistance maps on our website www.rrac.info. Prior to collecting and sending samples, the site can also be visited to check if data on resistance already exist for the sample point of concern.

How to collect tails

- Please collect 1-3 tails per site.
- Collect tails from dead bodies or preferably trapped rodents (fresh, clean and intact bodies are needed for tests to work. If you suspect bodies are more than 3 days old and are not of good quality, do not use it)

How to process tails

- A tail tip (3-5 cm) is required to provide DNA from each rodent. Each tail tip must be removed using a clean blade or sturdy scissors and stored in a sealable plastic bag (e.g. Zip-Lok). Please put each tail in a separate bag.

How to store and send tails

- Once the tail sample has been collected and placed in a bag, it should either be frozen (within 12 hrs. of collection) or sent to the University of Reading for DNA testing.
- An exact location must be provided with a sample (GPS co-ordinates OR a post code OR Zip code), see form attached, or it cannot be processed.
- The samples must be labelled correctly and packed in a way that samples cannot be touched by un-authorized people

PLEASE NOTE: the quicker a fresh tail can be posted off OR stored in a freezer, the better the chances of successful results. To preserve samples, we recommend adding salt (NaCl) into the collection bags, in excess.

RRAC website

Thank you for attention!
www.rrac.info

A close-up photograph of a small brown mouse, likely a vole or similar rodent, standing on a dark, textured surface. The mouse has a light-colored belly and a darker, mottled back. It is looking directly at the camera with its large, dark eyes. Its front paws are visible, and it appears to be in a natural, outdoor setting.

Stefan.Endepols@envu.com



| VKOR SNPs | FGARS | Bromadiolone | Difenacoum | Brodifacoum | Flocoumafen | Difethialone |
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