

The history of EU43 and resistance to mandipropamid in Denmark

Jens G. Hansen & Isaac K. Abuley, Aarhus University

Outline

- EU43 resistance against mandipropamid sounding the alarm
- Key results on the Danish P. infestans population
- The evolution and spread of EU43 in Europe, 2017-2023 focus on DK with a reference to NL











Risiko for resistens mod Revus

I flere forsøg ses der nu en vigende effekt af Revus mod skimmel i kartofler. Derfor anbefaler landskonsulent, at midlet indtil videre ikke anvendes, hvor der er udbredt skimmel.



Forsøgsmæssigt er der anvendt ren Revus i forsøgmarken i Arnborg. Den vigende effekt giver mistanke om resistensudvikling. Arkivfoto

SEGES sounding the alarm

News from SEGES, 30 August 2022

Risk for resistance against Revus

(a.i. = mandipropamid). Obs from trials and commercial fields







Department of Agroecology

A new variant of the late blight pathogen Phytophthora infestans is threatening the potato production

The results of a study on late blight show 100% resistance to one of the most important fungicides in potato production. Researchers find the development of the new variant of late blight worrying in relation to future control in Danish fields.



among researchers from Aarhus University

https://agro.au.dk/en/current-news/news/show/artikel/kartoffelproduktionen-trues-afstigende-resistens-hos-kartoffelskimmel-mod-kemiske-bekaempelsesmidler

Resistance to mandipropamid in EU_43_A1 reported

Press release by Aarhus University, 6 Jan 2023

5 isolates tested – all resistant to mandipropamid





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Results of the EuroBlight potato late blight monitoring in 2022

EuroBlight now reports on the 2022 results. Approximately 1100 samples from 22 countries genotyped.

21. marts 2023 af Jens Grønbech Hansen

21 March 2023

Download the news story as pdf

Key findings:

- Late blight pressure in 2022 was lower than average across many parts of Europe. Disease outbreaks from 22 countries were sampled by 23 teams in 2022 resulting in 1098 genotyped samples
- The frequency of genotype EU_43_A1 (EU43) increased from 2 % in 2021 to 15.8 % of the population in 2022, mainly sampled in Denmark the Netherlands and Belgium. EU43 was also found in Norway, Sweden, Germany and Portugal
- Isolates of EU43 from several regions in Denmark were resistant to mandipropamid at doses of up to 100ppm. Other genotypes (EU_36_A2, EU_37_A2 & EU_41_A2) were all sensitive.
- Reduced sensitivity of EU37 to fluazinam has reduced its use, prevented management failures & driven a decline of this genotype to 2.6% of the sampled population
- Comprising 36% of the samples, EU36 was the most frequently sampled genotype which suggests it remains fitter than other clones but conclusive evidence of a specific fitness trait is challenging to demonstrate in vitro
- Primary inoculum is locally generated and spread. Better management of all inoculum sources is required
- The proportion of 'other' genotypes generated from sexual oospore inoculum remained stable between 20-30%



regions in Denmark were resistant to mandipropamid at doses of up to 100ppm. Other genotypes (EU_36_A2, EU_37_A2 & EU_41_A2) were all sensitive.

Plant Pathology An International Journal edited by the British Society for Plant Pathology



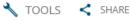


The EU43 genotype of *Phytophthora infestans* displays resistance to mandipropamid

Isaac K. Abuley 🔀 James S. Lynott, Jens G. Hansen, David E. L. Cooke, Alison K. Lees

First published: 28 April 2023 | https://doi.org/10.1111/ppa.13737

SECTIONS





Abstract

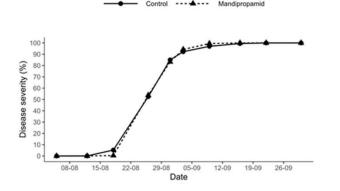
Mandipropamid is an active ingredient in the carboxylic acid amide group of fungicides and plays a key role in current potato late blight (Phytophthora infestans) management programmes. However, reports from Danish potato growers in 2022 suggested that mandipropamid had lost its efficacy. A study was therefore conducted to investigate the sensitivity of isolates collected from fields in which mandipropamid had been reported to be ineffective. Seventy-two isolates of *P. infestans* collected from potato fields in Denmark were genotyped using microsatellite markers, revealing a dominance of the clonal lineage EU43 and fewer isolates of EU41 and 'other' genetically distinct genotypes. Isolates belonging to the EU43 and EU41 lineages were selected, in addition to representative isolates of clones EU36 and EU37 from Scotland, and tested for sensitivity to mandipropamid at five concentrations ranging from 0.1 to 10 µg/mL on potato leaf discs (cultivar Maris Piper). The EU43 genotype infected leaf discs at all tested concentrations, and therefore no dose-response curve could be calculated. A dose response was observed for isolates of genotypes EU36, EU37 and EU41 with EC50 values ranging from 0.35 to 0.75 µg/mL. Field experiments confirmed resistance of tested isolates of genotype EU43 to mandipropamid, with no significant difference in the area under the disease curve between the untreated and mandipropamid treatments. Analysis of the Danish population of *P. infestans* showed that EU43 was widely distributed across the country. To our best knowledge, this is the first report of resistance to mandipropamid in P. infestans.

28 April 2023

Phytophthora infestans was isolated from late blight lesions



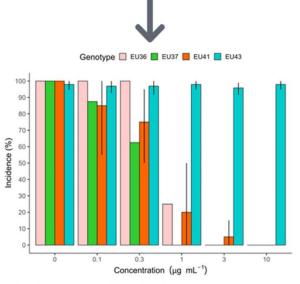
Field experiment with **EU43**



Disease development in the untreated control and mandipropamid treated plots

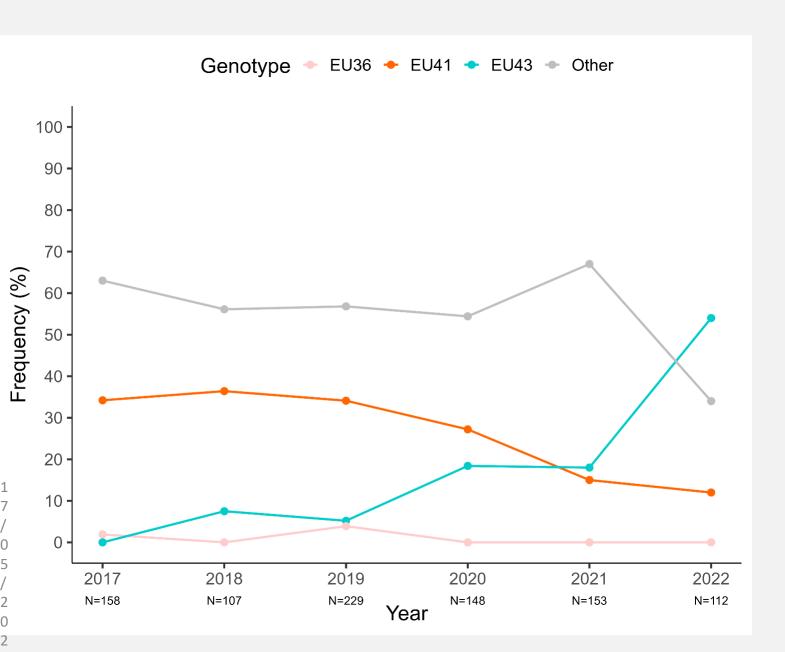
Isolates were tested for their sensitivity to mandipropamid



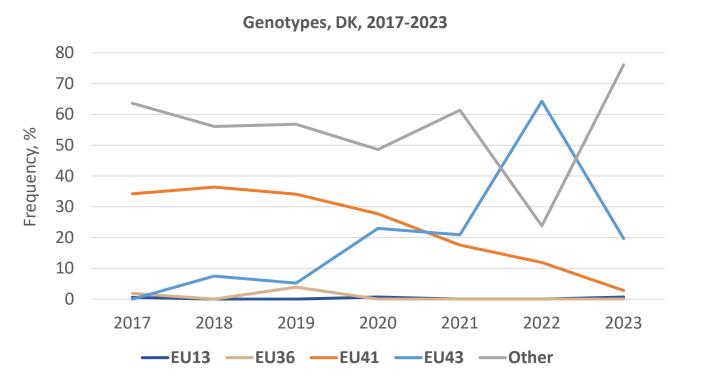


EU43 infected leaf discs at all concentrations of mandipropamid

The spatial and temporal distribution of EU43 in Denmark until 2022





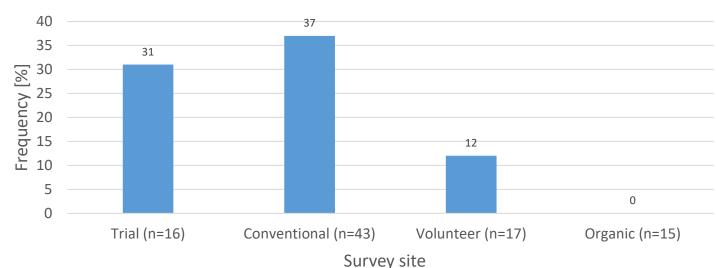


Status Denmark, 2023

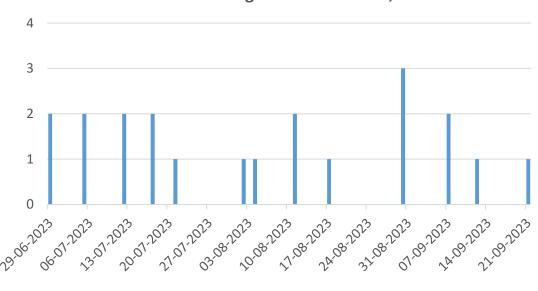
EU43: from 64% in 2022 to 19,7% in 2023.

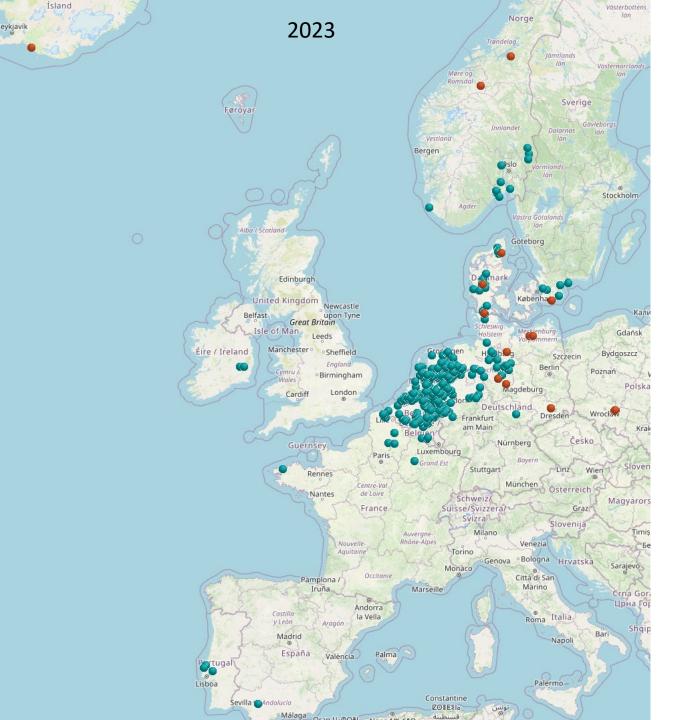
EU41: from 12 to 1% and Others: from 24% to 76%

Frequency of EU43 in different types of grown potato



Seasonal recordings of EU43 isolates, 2023





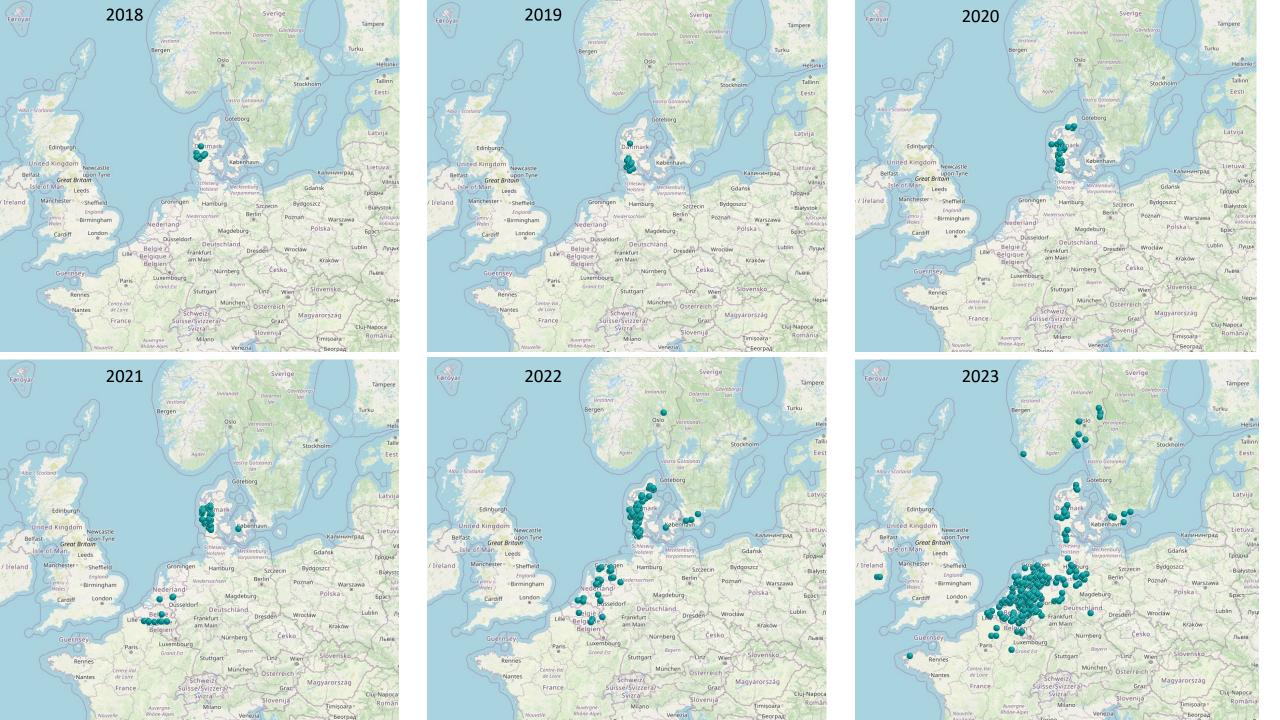
What is the history in time and space of two clones, initially found in Denmark

- EU41 in DK 2013
- EU43 in DK 2018





The evolution and spread of EU41 Go East and North



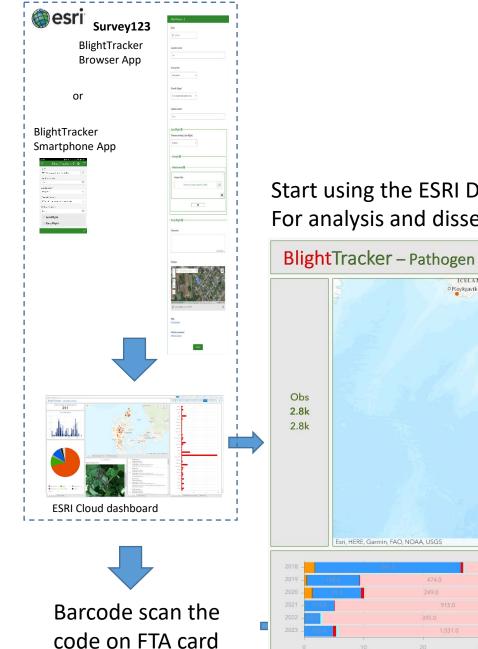


Ísland

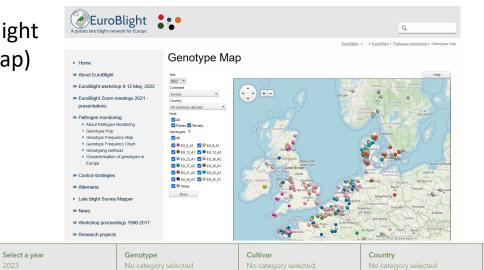
Dataflow – management – display and analysis:

- Late blight disease surveillance
- Isolate characterisation
- Genotype * host cultivar interactions

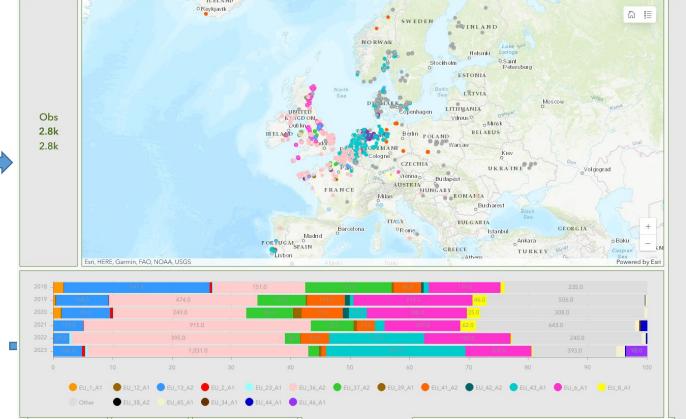
Building the Dashboard for analysis



Public in EuroBlight (Open Street map)

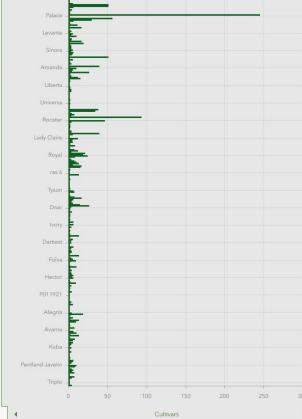


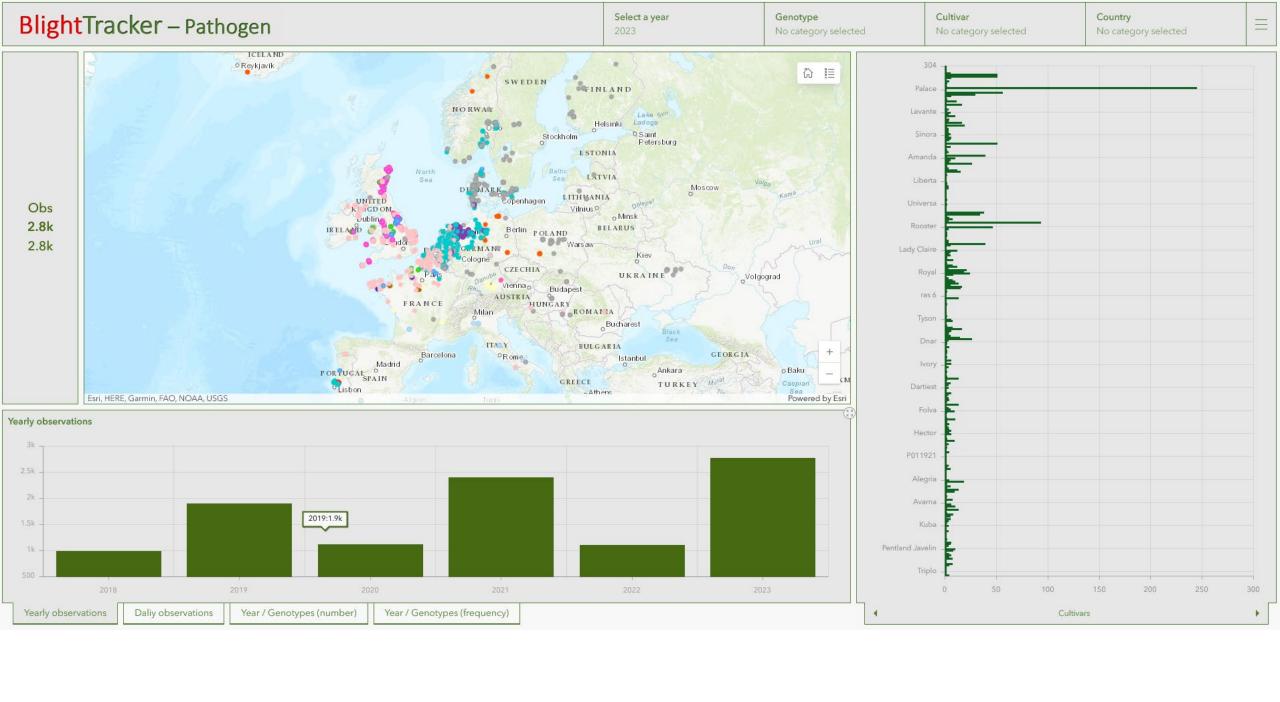
Start using the ESRI Dashboards For analysis and dissemination

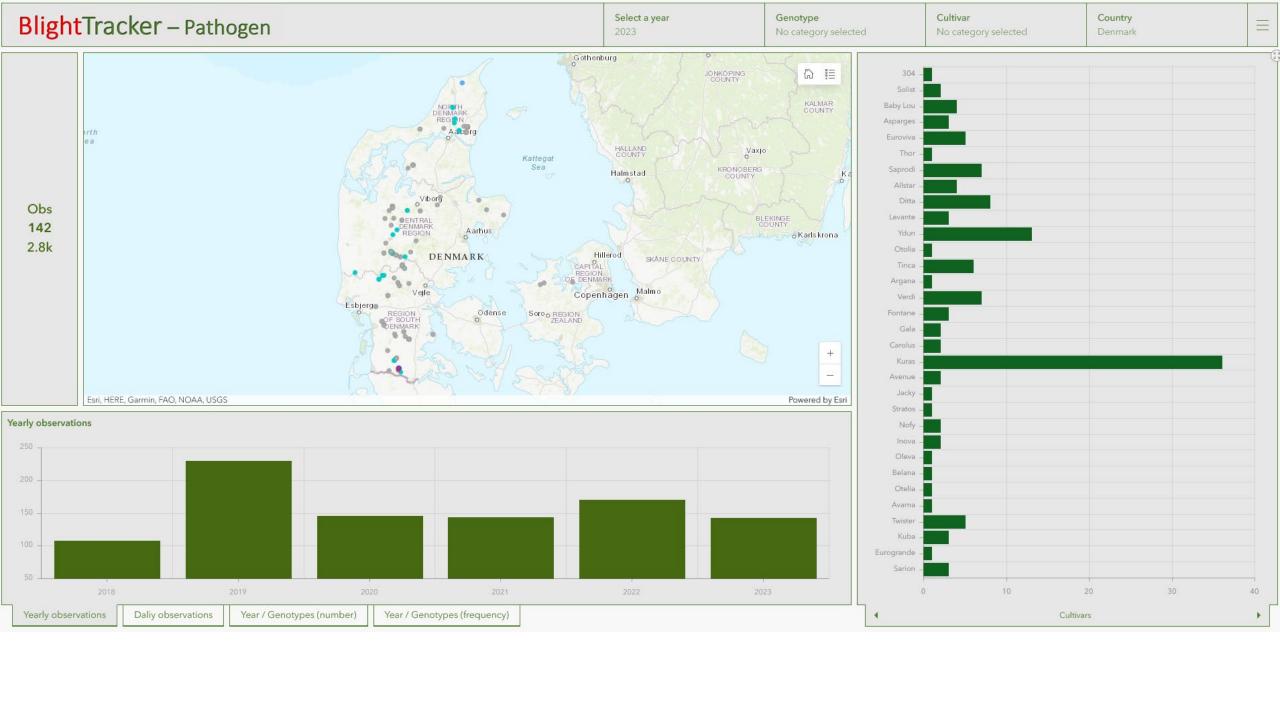


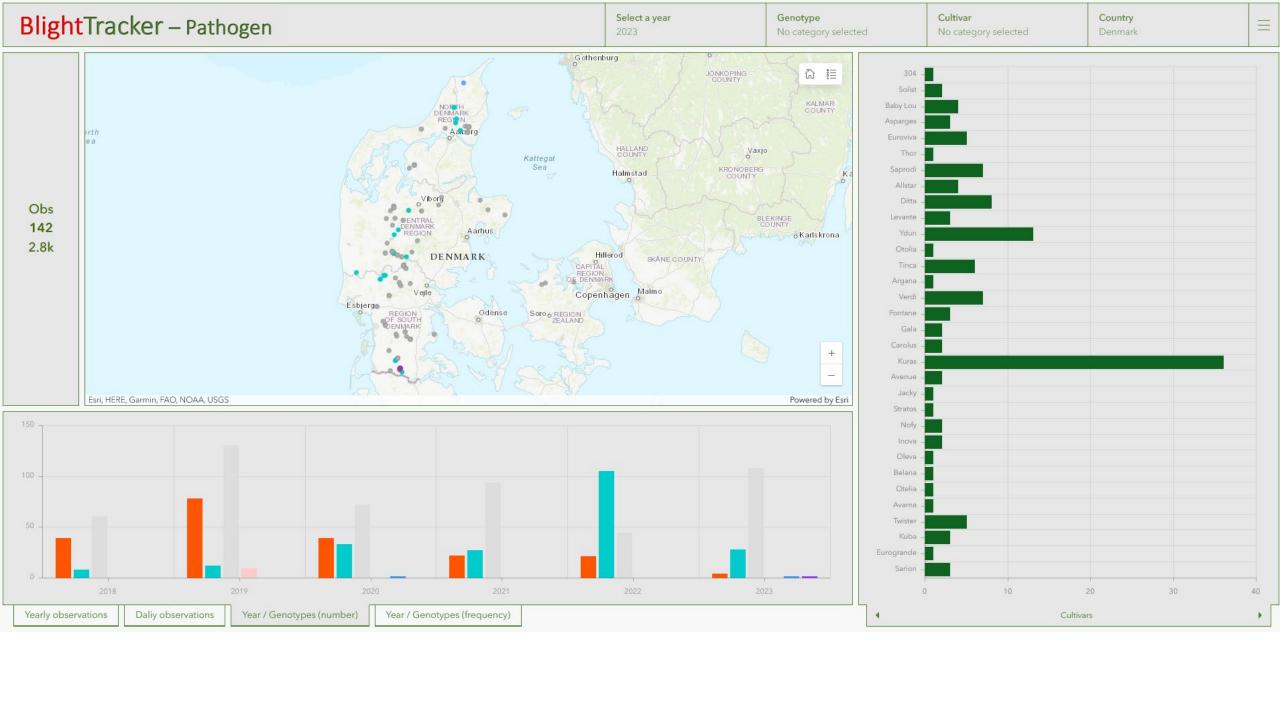
Year / Genotypes (frequency)

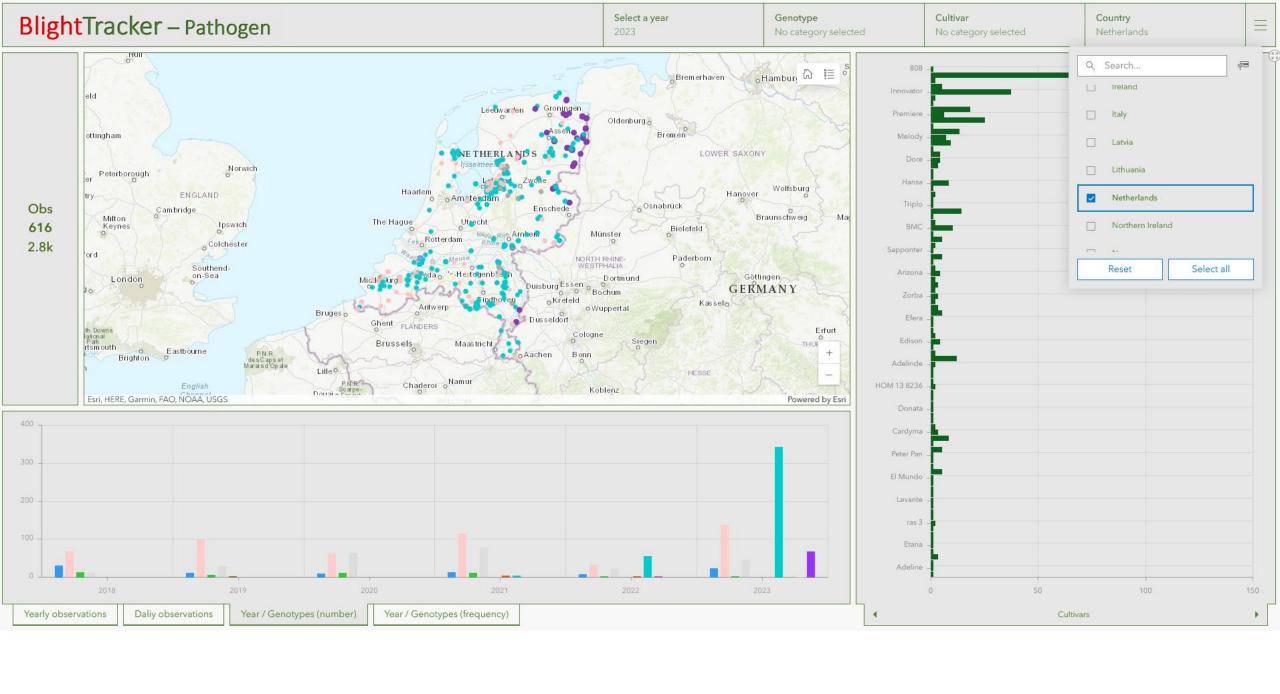
Year / Genotypes (number)

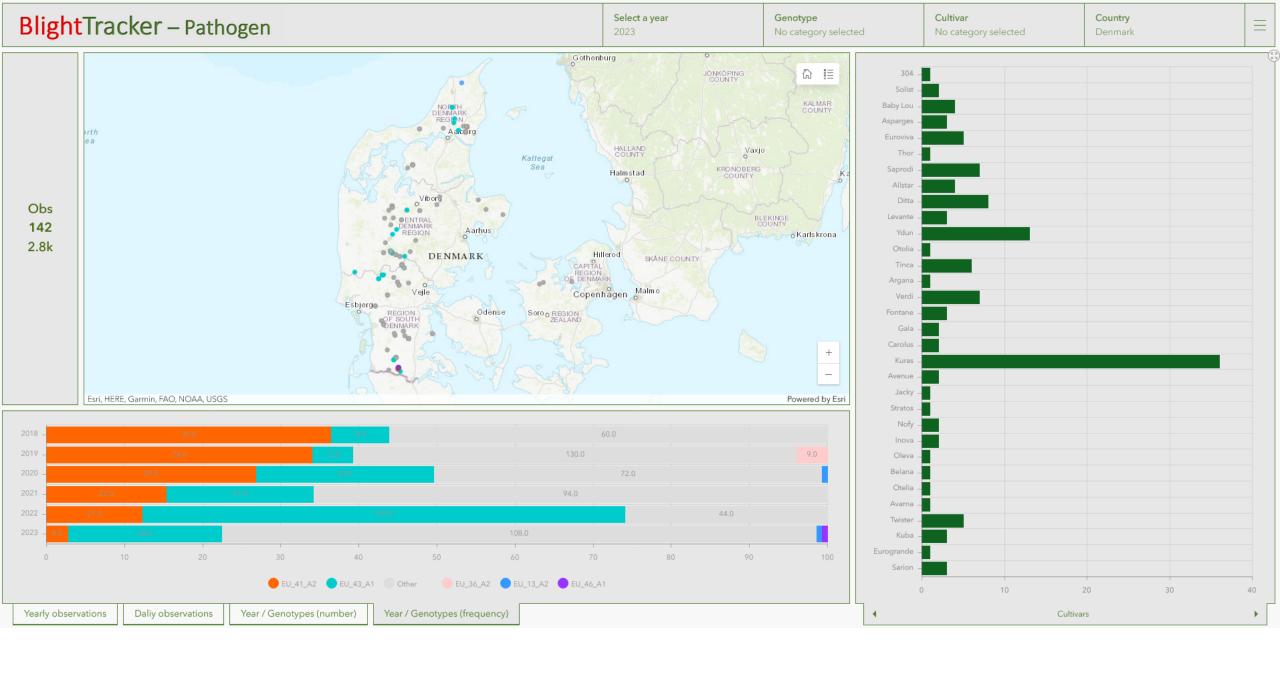


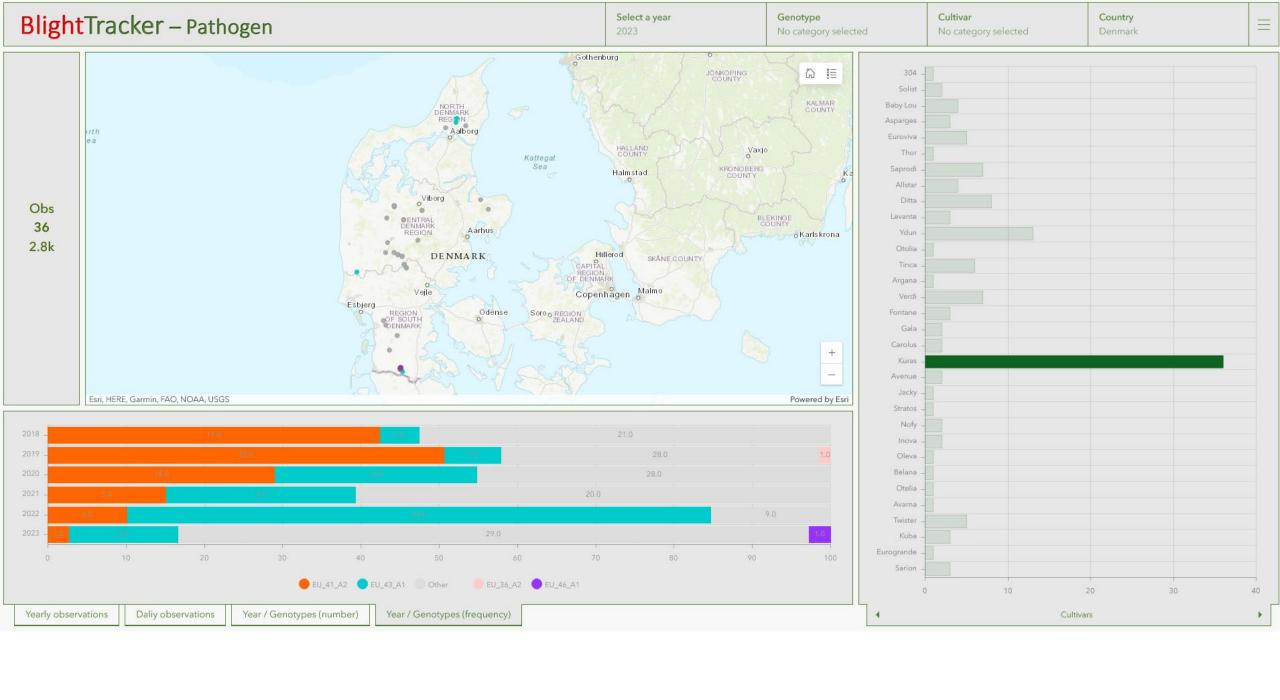


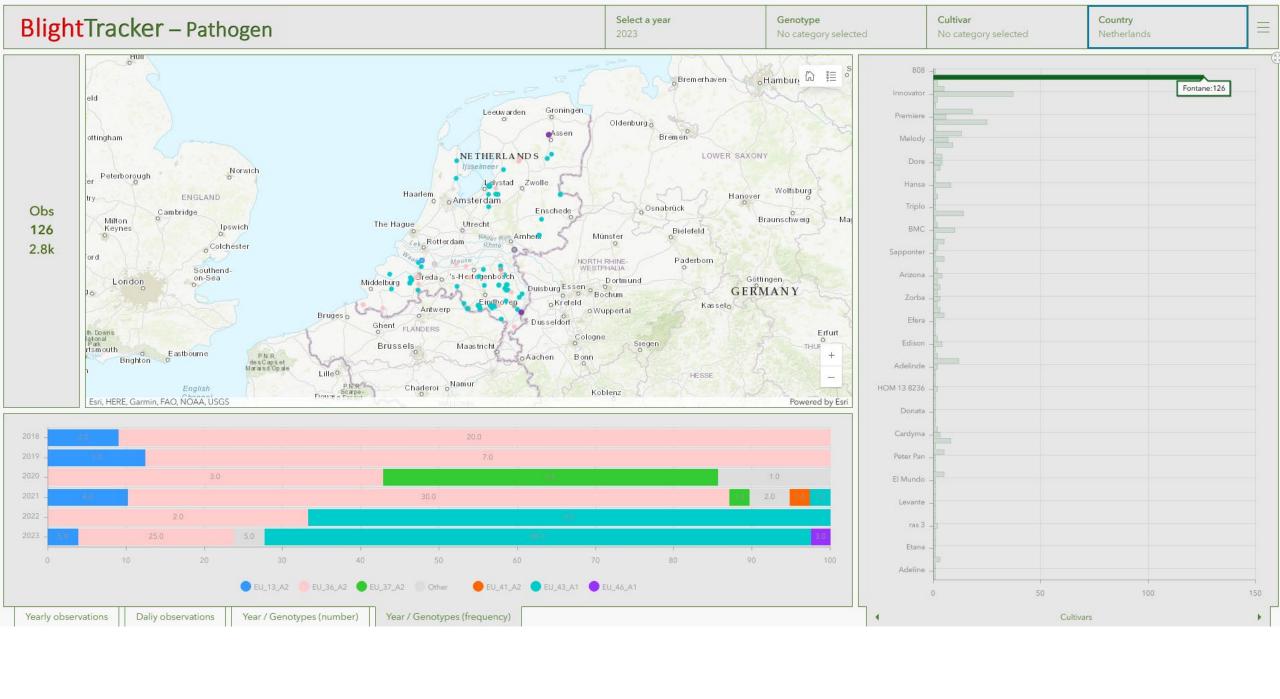












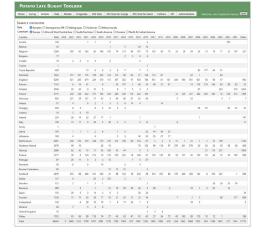
POTATO LATE BLIGHT TOOLBOX

Home Survey Samples Trials Models Fungicides DSS Tools DSS Tools for Sverige DSS Tools for Island Partners API Administration Welcome Jens Grønbech Hansen logout

Sample overview

Country	Total	2024 202	3 2022	2021	2020	2019	2018	2017	2016	2015	2014	2013	2012	2011	2010	2009	2008	2007	2006	2005	2004	2003	2002	2001	Olde
Austria	108			1	1		2					4											100		
Belarus	45						1		5		29	10													
Belgium	2369	30	5 43	242	42	148	103	74	173	98	187	75	193	58	73	35	24	54	26	13	16	17	32	107	23
Bulgaria	17								2	3	6	6													
Croatia	19		2 3	4	5		2				3														
Cyprus	1											1													
Czech Republic	409			15	6	8	3	6	7		7	5							80	177	64	31			
Denmark	1832	15	191	176	159	245	123	174	96	64	37	126	114	87				23	1			65			
England	8290	63	265	674	234	519	115	397	363	97	403	188	451	61	101	306	590	901	801	56	98	47			99
Estonia	1143	(5 16	41		9		24	107	1	68	54	49	51	41			14	118	119	144	84	89	83	2
Finland	3048	2	3 20	9	10	5		6	7	5	4	4							87			234		210	242
France	5715	23	128	342	137	103	244	260	230	116	222	185				12	125	331	382	303	444	72	136	213	149
Germany	926	22	1 29	121	71	45	3	94	86	87	62	40				3		53				3	1		
Greece	57		1	6	2	1	3	3	6	14	4			14											
Hungary	330		3 5		6	2	12	3	4										56	67			93	27	4
Iceland	18		2 6	10																					
Ireland	254	2	5 14	20	27	17	1				1								1						14
Italy	158	1-	1 17	9	26	5	49	4	2		2	13				6									1
Jersey	1												1												
Latvia	194		1 1	1	2	6		1	5			52	44	48	33										
Lithuania	180		1		4			5	5		40	53	25	27	17										
Netherlands	4001	63	5 156	259	148	179	129	252	198	80	154	87	5	3	6	15	1	21	1	1	25	109			153
Northern Ireland	2579	5	3 10			28	13				75	123	99	118	97	276	241	578	20	54	24	40	58	38	629
Norway	2686	6	2 42	10	11	93	100	42	44		5	5								21	115	331			180
Poland	3277	3) 8	124	215	10	133	259	262	14	204	61	126	150	89	59	121	45	129	93	46	22	30	149	89
Portugal	97	2	5 11	5	4	4	15			1	5	27													
Romania	36		3	5		14			5		4														
Russian Federation	60						15	15	25			5													
Scotland	4870	21	5 88	406	141	425	41	98	134	78	243	39	210	351	86	170	400	338	134	8	474	231		1	55
Serbia	137	(5		20	2	103		2		4														
Slovakia	121		2	1	1	1						2									26	26	26	36	
Slovenia	280		2		1		21	10	30	49	43	2	28		2			16	1	2	61		12		
Spain	143	2			14	3	9	2			20	24													3
Sweden	1558	7.			38	77	10	21	26	37	23	34				7		2	3			88		277	68
Switzerland	133		4		10	10	1	4	17	22	1	9						19	5						
Ukraine	26		7	5	4	8						2													
United Kingdom	10											3	3	4											
Wales	1535	6	5 86	99	118	74	27	45	63	47	91	43	57	34	73	45	160	89	110	10	12	1			18
Total	46663	0 284		2700	1457			1799	1904		1947	1282	1405	1006	618		1662		1955		1549	1401			11712

46663 samples from 38 countries



EuroBlight database



DK:

EU41

EU37

	2004	2010	2013	2018	2023
NL:	EU13	EU33	EU36		EU46

Why do NL and DK have the highest number of MLGs in Europe

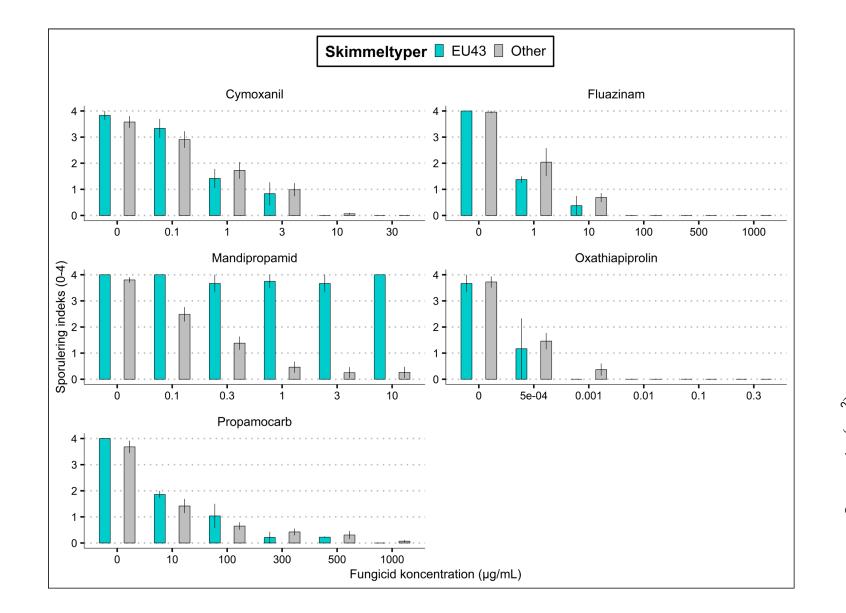
Where is the centre of diversity in Europe?

Country	No of MLGs in 2023	N
The Netherlands	114	633
Denmark	101	144
Eng, Sco & Wa	94	909
Belgium	81	305
France	69	235
Germany	67	219

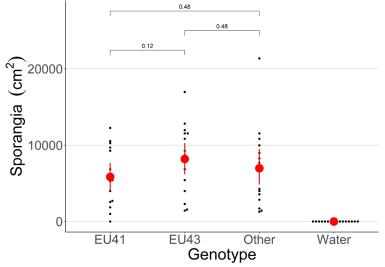
EU43

Except EU43 and mandipropamid, all tested isolates were sensitive to the fungicides tested

Agressiveness tests of EU43 See presentation by Isaac K Abulay later







This a game of Chess and <u>we won the first match against Phytophthora</u>. But the game is best out of 5 matches!





AF JENS GRØNBECH HANSEN & ISAAC KWESI ABULEY, AARHUS UNIVERSITET

SKIMMELTYPER 2023 - og hvad kan vi forvente i 2024?

Vi forventer, at EU43 kun i mindre grad udvikles fra inficerede knolde på grund af et sundt læggemateriale. Derfor skal der være fokus på de alternative smittekilder, bl.a. affaldsdynger, spildkartofler, spredning af smitstof med vinden fra Tyskland og indført inficeret læggemateriale fra områder i Holland, Belgien og Nordtyskland.

I 2022 kom der mere skimmel end forventet, fordi en Revus resistent variant, EU43, bredte sig markant i Danmark, fra 21 procent i 2021 til 64 procent i 2022 (Fig. 1).

Tilsvarende skete i Holland hvor EU43 steg fra en procent i 2021 til 42 procent i 2022. Det var et vink med en vognstang.

I 2023 ændrede anbefalingerne i Danmark sig til, at Revus ikke skulle bruges i starten, og at der skulle bruges blandinger af produkter med forskellig virkemekanisme. Målet var klart. Vi kan ikke undvære mandipropamid (aktivstof i Revus) i vores arsenal af aktivstoffer, dels fordi det ifølge EuroBlight er et af de bedste midler mod skimmel, dels fordi Ranman Top lige er blevet forbudt og endelig, fordi fraværet af de to midler vil lægge et alt for stort selektionspres på de resterende tilladte midler- ikke mindst fluazinam, som er aktivstoffet i bl.a. Shirlan.

Planen virkede, men kun fordi forsk-

ning, rådgivning, industri og avlerne arbejdede sammen om at forstå, hvad der skete og om at finde de rette løsninger. Aarhus Universitet har netop analyseret de sidste prøver for både genotypning (genetisk fingeraftryk) og fænotypning (resistens, sporuleringskapacitet, etc.). Vi kan nu lave en endelig opgørelse om genotyper i 2023. Kan man også forudsige, hvad der vil ske i 2024?

Måske, hvis man forstår fortiden.

Udviklingen af skimmeltyper i Danmark 2017-2023

I 2023 blev der fundet ét eksemplar af EU13, i en have i Nordjylland med aspargeskartofler. Der blev også fundet en ny skimmel type for første gang i Danmark, EU46, i en konventionel mark med Kuras tæt på den tyske grænse.

EU41 er faldet fra 30-40 procent i 2017-2019 til 2,8 procent (4 isolater) i 2023.

EU43 er faldet fra 64 procent i 2022 til 20 procent i 2023. Til gengæld er gruppen "andre" steget fra 24 procent til 76 procent (Fig. 1).

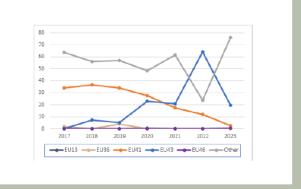
Forskel i hyppigheden af EU43 mellem konventionel og økologisk

I 2023 startede et nyt projekt om økologiske kartofler: "Sustainability and Resilience in Organic Potato Production - SROPP".

I dette projekt har vi indsamlet skimmel fra mange økologiske marker for at sammenligne populationen af skimmel i marker med forskellig dyrkning. Det viste sig, at der var stor forskel på hyppigheden af EU43 imellem i de forskellige dyrkningstyper.

Af skimmel, indsamlet i 69 konventionelle marker, var 25 procent EU43, tilsvarende var 27 procent EU43 i indsamlinger fra forsøg, 12 procent EU43 i indsamlin-

Figur 1. Udviklingen af skimmeltyper i Danmark, 2017-2023



Farmers' journal, May 2024

Variants of *P. infestans* in 2023 – What can we expect in 2024?

From Science

From Practice

to Science

to Practice



The 10 principles for control strategy 2024



AF LARS BØDKER LANDSKONSULENT SEGES INNOVATION

Der udvikles hurtigt resistens i kartoffelskimmel, når specifikke svampemidler anvendes i blokke med flere behandlinger efter hinanden uden blandingspartnere og hvis fx. Zorvec Enicade udsprøjtes på etableret skimmel i marken. Foto: Lars Bødker.



SKIMMELSTRATEGI 2024

Skimmelstrategien i 2024 er præget af meget få tilbageværende svampemidler til forebyggelse af kartoffelskimmel, resistens overfor Revus og en stærk stigende risiko for resistens overfor både Shirlan/ Banjo/Zignal og Zorvec Enicade. Det kan stadig lade sig gøre, men det kræver, at alle overholder nogle fælles principper.

Udvikling af resistens

Ligesom bakterier er i stand til at udvikle resistens overfor antibiotika, er kartoffelskimmel i stand til at udvikle resistens overfor både svampemidler med forskellige virkemekanisme men også bryde resistensgener i resistente kartoffelsorter. Jo flere gange, der behandles med det samme svampemiddel og jo oftere en sort dyrkes, jo nemmere er det for en mutation i en spore af kartoffelskimmel at blive opformeret og spredt ikke blot til Danmark men til hele Europa. Dette er sket med fx den resistente genotype af skimmel (EU43) overfor mandipropamid (Revus), som er muteret, opformeret og spredt fra Danmark helt til Portugal.

I de sidste 25 år har forskere, rådgivere

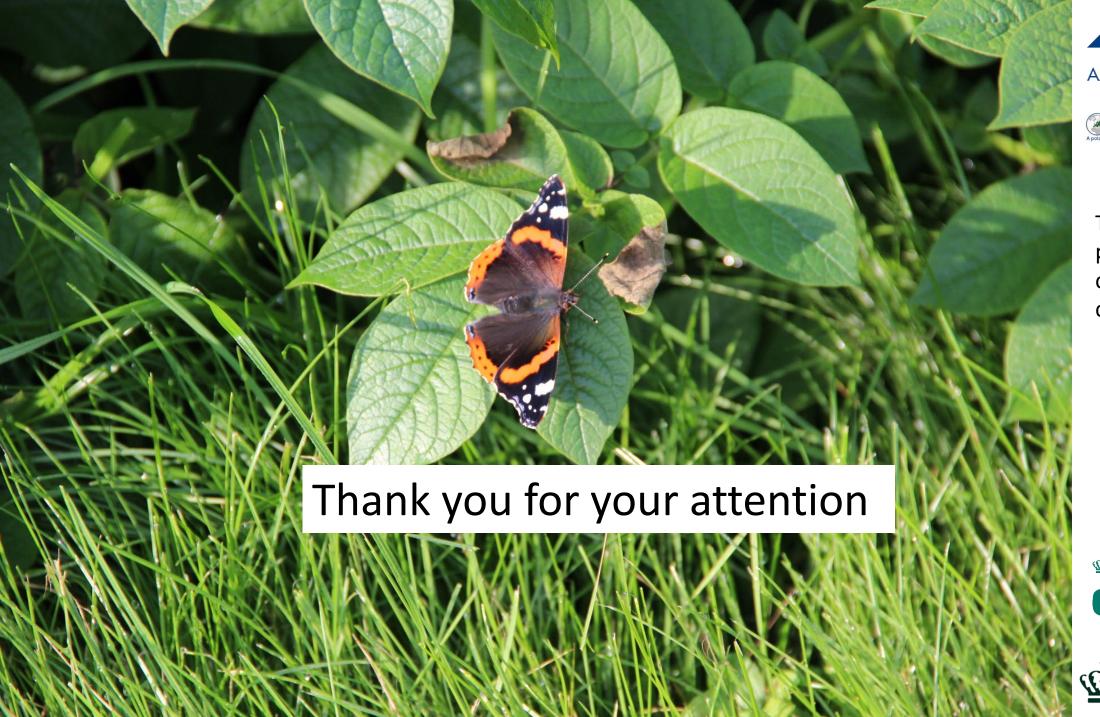
I 2023 forekom der udbredt resistens (EU43) overfor mandipropamid (Revus) i store dele af specielt Nordeuropa. Kilde: EuroBlight.



og avlere haft stor fokus på reduktion af forbruget af svampemidler ved bekæmpelse af kartoffelskimmel på grund af både et økonomisk og miljømæssigt fokus. Der er lavet et utal af forsøg med forskellige beslutningsstøttesystemer (NegFry, Skimmelstyring, BlightManager, PlantPlus, Akkerweb), som har gjort det muligt at reducere dosering og variere intervaller samt anvende de stærkeste midler på de mest skimmelfavorable tidspunkter. Dette har sikret, at der i Danmark over en 25-årig periode er brugt en langt mindre mængde svampemidler og ressourcer end i vores omgivende lande.

Har dette været en fejl, når vi nu ser udbredt resistens overfor svampemidler? Et bakspejl er nu bedst til at bakke efter, men det er OK at blive klogere set i lyset af historien og ny viden. Kartoffelskimmel kender ingen grænser, og der er skabt resistens overfor kartoffelskimmel i andre lande end Danmark med langt større forbrug af pesticider fx Ridomil Gold, Shirlan i flere omgange og nu også Zorvec Enicade efter blot få års brug. Men set i bakspejlet har det været forkert, specielt efter forbuddet mod det bredspektrede mancozeb (Dithane), at anvende specifikke soloprodukter udbragt i blokke med 2-3 behandlinger i træk på etableret skimmel. Dette har ført til et unødigt højt risiko for opformering og spredning af resistente mutationer, risiko for opformering og spredning af resistente mutationer. Dette er ikke kun sket i Danmark, men i hele Europa.

Nu står specielt dansk kartoffelavl i en situation, hvor der er udviklet resistens overfor Revus i (EU43) 2022, og hvor registreringen af Ranman Top er trukket tilbage i januar 2023. Der er kun Shirlan/ Banjo/Zignal, Zorvec Enicade, Sporax/ Raport og Cymbal WG/Option WG tilbage i Danmark, hvor aktivstofferne i de to sidstnævnte kendes fra Proxanil. I lande, som grænser tæt op til Danmark, er der nu resistensdannelse overfor både Shirlan/ Banjo/Zignal og Zorvec Enicade, men som endnu ikke er bekræftet fundet i Danmark. Dette betyder, at der skal lægges en ny strategi for skimmelbekæmpelse for 2024, men som grundlæggende bygger videre på samme principper som i 2023.









Thanks to many partners and colleagues for contributing





Miljøministeriet Miljøstyrelsen