



Mancozeb: Banana agriculture & Resistance management

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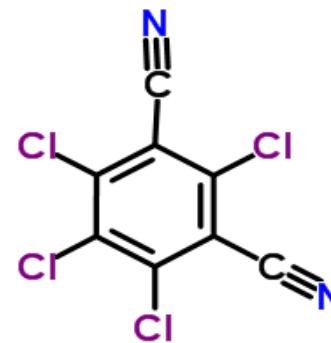
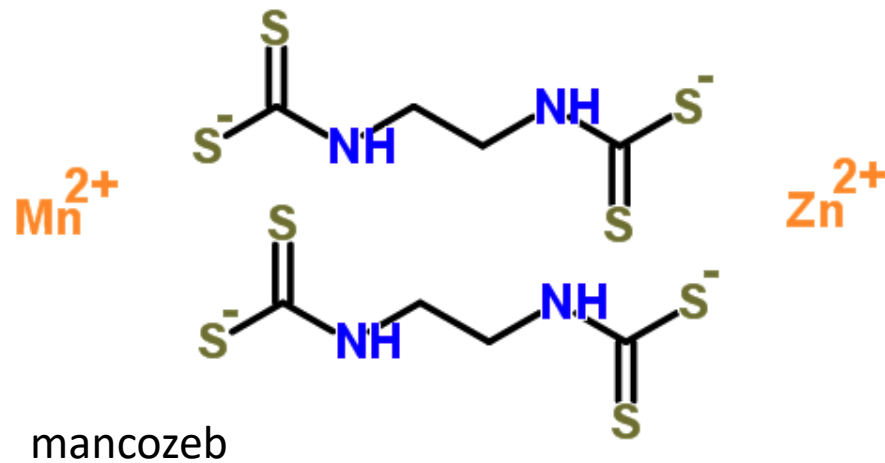
Mancozeb – mode of action

Protectant contact multi-site fungicides (e.g. mancozeb) will prevent spores from germinating and infecting the plant if applied prior to spore release. Once the infection has occurred and the fungus has penetrated the leaf, these fungicides will no longer control the disease.

Updated 12/03/15



Mancozeb – molecule

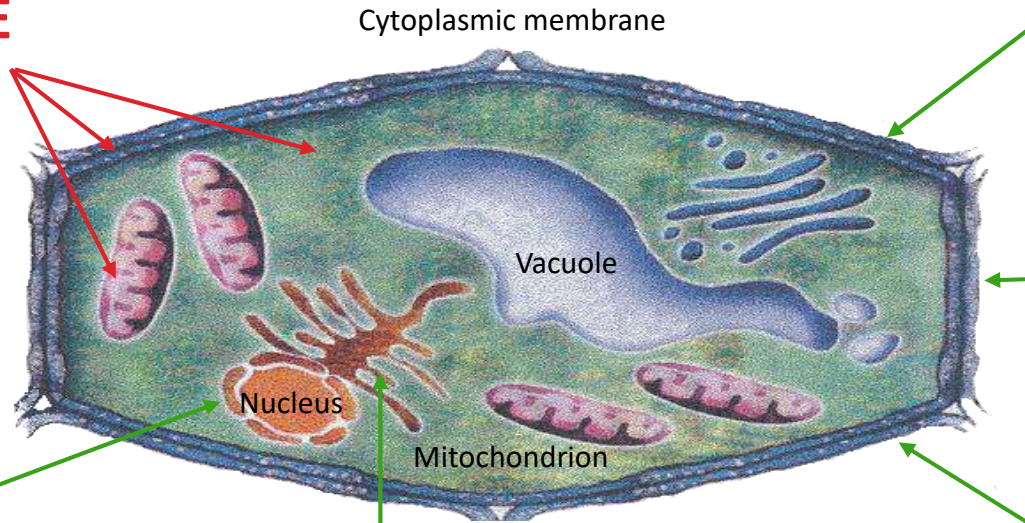


chlorothalonil

Mancozeb – mode of action

MULTI-SITE
mancozeb

Fungal cell



B: Mitosis and cell division
B3 zoxamide
B5 fluopicolide

A: Nucleic acid synthesis
A1 metalaxyl-M
benalaxyl-M

H: cell wall biosynthesis
H5 dimethomorph
mandipropamid
benthiavalicarb
valifenalate

F: lipid synthesis and membrane integrity
F4 propamocarb

C: Respiration, mitochondria
C3 famoxadone
fenamidone
C4 cyazofamid
amisulbrom
C5 fluazinam
C8 ametoctradin

Ⓜ Mancozeb is an ethylene bisdithiocarbamate (EBDC) fungicide, belonging to the group of dithiocarbamates

Fungal disease in bananas

- Black Sigatoka, black leaf streak/spot caused by ascomycete *mycosphaerella fijiensis*
- Pathogen produces conidia and ascospores, both are infective affecting photosynthesis indirectly
- High moisture conditions, dispersed via wind, for conidia, also by rain and irrigation water, yellow in cooler and black in warmer environment
- Untreated, yield can be affected by 35-50%, also shortening period between harvest & ripening - Most commercially important disease for export crop to control

Symptoms for Black Sigatoka



Fungicide control in bananas

- Large plantations reliant on chemical controls by aerial application
- Predominantly mancozeb due to solo or mixture use
- Mancozeb is often combined or rotated with morpholine, demethylation inhibitors (DMI), or strobilurin (QoI) fungicides to prolong the life other chemistries
- Resistance to other fungicides is widespread in many production areas eg: Strobilurins

Mancozeb – Importance for Bananas

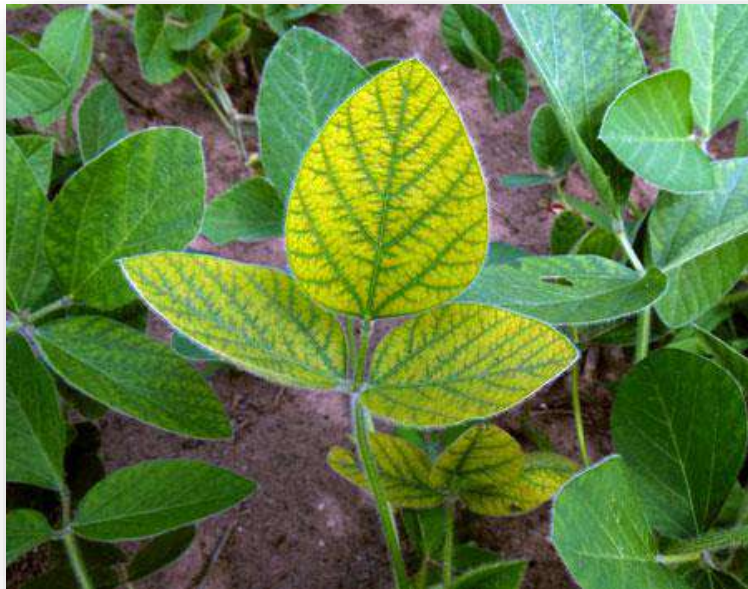
- Mancozeb products control the broadest number of diseases in the greatest number of crops of any fungicide
 - Registered on >70 crops for >400 diseases globally
 - Including a large variety of minor uses in most Fruit &Vegetables, flowers, forest trees
 - New areas of development
- Fungus (Oomycetes primarily but not only) and some bacteria
- A cost effective key multi-site fungicide which can be utilized to protect key single site actives
- Mancozeb, as a dithiocarbamate, belongs to the “Low Risk” group under
- FRAC mixtures recommendations due to a proven history:
 - FRAC Recommendations for fungicide mixtures, 2010 & FRAC List 2017
 - FRAC codes Mode of Action – Mancozeb (M3)

Mancozeb Products

- Vondozeb 75 WG (mancozeb 750g/kg)
- Vondozeb 62 OD (mancozeb 620g/kg)
- High compatibility with other products
- Registered for control of black & yellow sigatoka
- Economic solution to the farmer
- Effective at low doses (1.5kg/ha solo/0.75 mix)
- Nutrients like Zinc and Manganese enrich crop
- Non volatile, short half life – quick degradation
- Refined production process ensures good rainfastness

Mancozeb – micronutrients

Mancozeb is a complex of zinc and maneb containing 20% Mn and 2.5% Zn (ref. FAO Spec.)



Manganese deficiency in potato plant

- Ⓢ Boosts bulking
- Ⓢ Increases the yield of tubers
- Ⓢ Improves disease resistance
- Ⓢ Improves skin finish
- Ⓢ Increases tuber dry matter content
- Ⓢ Increases starch levels

FRAC Banana Recommendations

“Multi-site fungicides (**Mancozeb**...and other fungicides of low resistance risk) can be applied for control of black sigatoka in the following way:

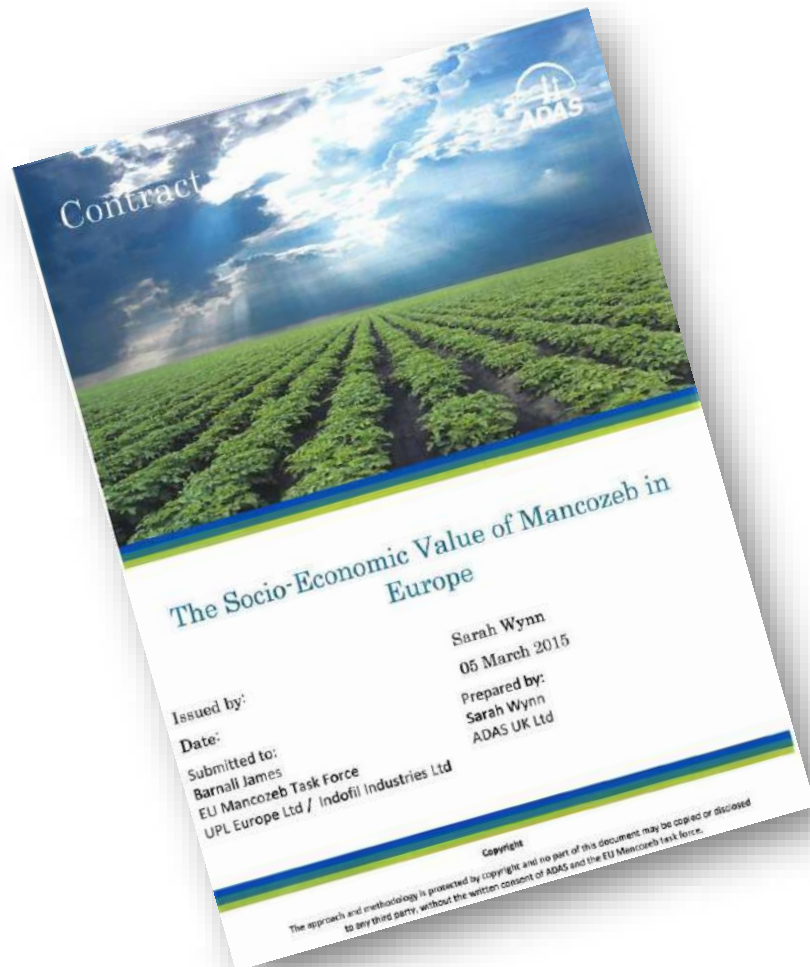
Multi site fungicides can be used solo or in mixtures with partners at manufacturers' recommended effective rates. There are no limitations or restrictions concerning the number of applications, the timing or the sequence as long as it is within the limits of the manufacturers' labels.”

FRAC Banana Group, 2014

FRAC Banana - Research

- Research at Wageningen University, The Netherlands - Home of blight research
 - PromoBanana project looking at Panama Disease and Black Sigatoka for earlier detection to assist growers.
 - UPL collaborated with Wageningen to explore the role of mancozeb in resistance management for potato early and late blight
 - Potential for new collaboration for banana research with a view to helping sustainability

Mancozeb – socio economic value of mancozeb



UK
France
Germany
Ireland
Italy
Netherlands
Spain
Greece

Mancozeb — socio economic value of mancozeb

- Ⓢ 90% of advisers were applying mancozeb to 100% of their crop at least once in the season
- Ⓢ Average number of applications of mancozeb was 7 per season
- Ⓢ Cost of production estimated to increase between €48 and €160 per ha, reducing the financial viability of potato enterprises
- Ⓢ Yield losses from uncontrolled blight are reported at up to 40% which is equivalent to 2.2 M tonnes of potatoes
- Ⓢ Loss of mancozeb will impact on resistant management strategies and increase risk of resistance developing to other active substances

Mancozeb – socio economic value of mancozeb



Mancozeb - research

Potatoes

 Wageningen University (Netherlands)

Cereals

 ADAS (UK)

 Rothamsted Research (UK)

Various crops

 University of Bologna (Italy)

Mancozeb - working with Wageningen

- Ⓢ Testing the efficacy of fungicides to control different strains of potato blight.
 - Ⓢ August 2013
 - Ⓢ May 2014
- Ⓢ Testing the efficacy of fungicides to control early blight
- Ⓢ Project conducted through Hub Schepers - Head of Euroblight
- Ⓢ Results presented at Euroblight 2015 by Serge Devauchelle (Retired French ministry potato advisor)

Mancozeb – resistance (potatoes)

Mode of Action	Target site	Group name	Example of common names	Predicted FDR (yrs)	Resistance reported* & strain colour	FRAC code
U	U	cyanoacetamide-oxime	cymoxanil	6.8	No (L to M)	27
A	A1	PA-fungicides	metalaxyl-M benalaxyl-M	5.9	Yes (H)	4
C	C4	Qil fungicides	amisulbrom	3.5	No (M to H)	21
C	C3	Qol-fungicides	famoxadone fenamidone	4.1	No (H)	11
C	C8	QoSI-fungicides	ametoctradin	7.8	No (M to H)	45
B	B3	benzamides	zoxamide	6.1	No (L to M)	22
	B5		fluopicolide	5.5	No (?)	43
H	H5	CAA - fungicides	dimethomorph benthiavalicarb mandipropamid	4.2	No (L to M)	40

SOURCE: FRAC LIST

* Reported in *P.infestans* (late blight)

Resistance risk

FDR = First detection of resistance

? = unknown

H = High, M to H = Medium to High, L to M = Low to Medium

Mancozeb - working with Wageningen

Strains	Background	Trial detached leaf	Trial pot in greenhouse
Blue 13 (13_A2)	Detected in NL and DE in 2004, in UK and FR in 2005, in Ireland in 2007, also present in many other EU countries since 2012. Strain more aggressive at low temperatures (8°C), resistant to phenylamides	Tested	Tested
Green 33 (33_A2)	Detected in NL in 2011, also present in BE and PL. Fluazinam is less efficacious on this strain. Expresses a weak fitness, therefore regresses when fluazinam solo is not any more used in a systematic way. Represented more than 20% of the strains in 2010 and 2011 but only 6% in 2012	Tested	Tested
Pink 6 (6_A1)	Detected in NL in 2002, in UK and FR in 2004. Seems very aggressive on leaf at temperatures around 10°C. Dominant in UK in 2011 and 2012, tends to regress during the last years	Tested	Tested
Orange	Old strain of A1 type, still present today but in a minority proportion	Not tested	Tested
Population VK98014	Old strain stored in the lab, supposed to be fully susceptible to fungicides	Tested	Not tested

Mancozeb - working with Wageningen

Type of strain	Untreated	Mancozeb 1500 g/ha
	% necrotic surface 7 days after inoculation	% necrotic surface 7 days after inoculation (efficacy)
Blue 13	99.8	17.3* (82.7)
Green 33	96.8	2.5* (97.4)
Pink 6	95.5	5.0* (94.8)
VK 98074	91.2	2.5* (97.3)

* values statistically different from untreated at 5% threshold in the 2 trials

Comparatively - Table 4 from August 2013 report:

Metalaxyl gave poor control of Blue 13 proving that it is metalaxyl resistant

Mancozeb - working with Wageningen

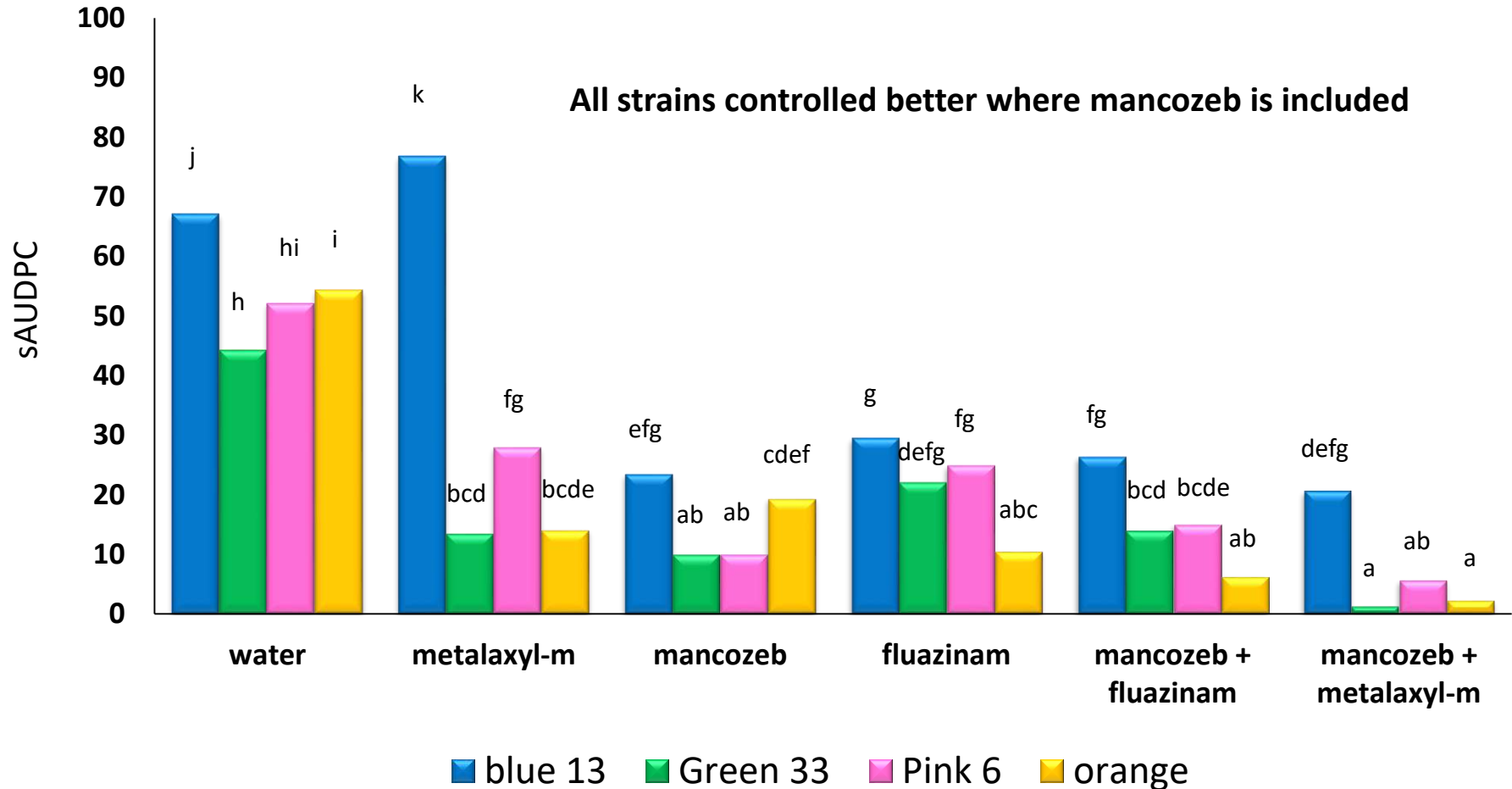
Greenhouse trial: protocol

Code	Fungicide	Dose rate g ai / ha
A	water (untreated control)	-
B	mancozeb 75% WG	1500
C	mancozeb 64% + metalaxyl-M 8% WG	1600 + 200
D	mancozeb 75% WG + fluazinam 500g/L SC	1500 + 200
E	metalaxyl-M 25% WG	200
F	fluazinam 500g/L SC	200

- Ⓢ Percentage necrotic foliage assessed 7, 9, 10, 13 and 15 days after inoculation. A Standard Area Under Disease Progress Curve (stAUDPC) was calculated.
- Ⓢ 4 replicates (1 replicate = 1 potato plant)
- Ⓢ Analysis of variance based on the percentage of necrotic foliage
- Ⓢ Rates comparable to those used for Bananas

Mancozeb - working with Wageningen

Greenhouse trial: results (2/2)



Mancozeb - working with Wageningen

- Ⓢ Mancozeb is effective against each type of strain
- Ⓢ Mancozeb strengthens significantly the efficacy of fluazinam and metalaxyl-m, even on strains sensitive to these fungicides
 - Stability of performance ensured whatever the type of strains met in the field
 - Essential for resistance management program

Mancozeb - working with Wageningen



- Ⓢ Early blight (*Alternaria solani*)
- Ⓢ Global disease
- Ⓢ Yield penalties of 30% plus

- Ⓢ UPL commissioned Wageningen University to study the efficacy of mancozeb against *Alternaria* (early blight), especially against strains resistant to strobilurins – ongoing
- Ⓢ mancozeb, azoxystrobin and boscalid + pyraclostrobin
- Ⓢ Similar results found – mancozeb controlled all strains where included

Mancozeb – working with Rothamsted



Dr Bart Fraaije (FRAG UK)

Ⓢ *In vitro* sensitivity testing of *Zymoseptoria tritici* field strains and mutants with resistant phenotypes to site-specific fungicides used for Septoria leaf blotch control



ROTHAMSTED
RESEARCH

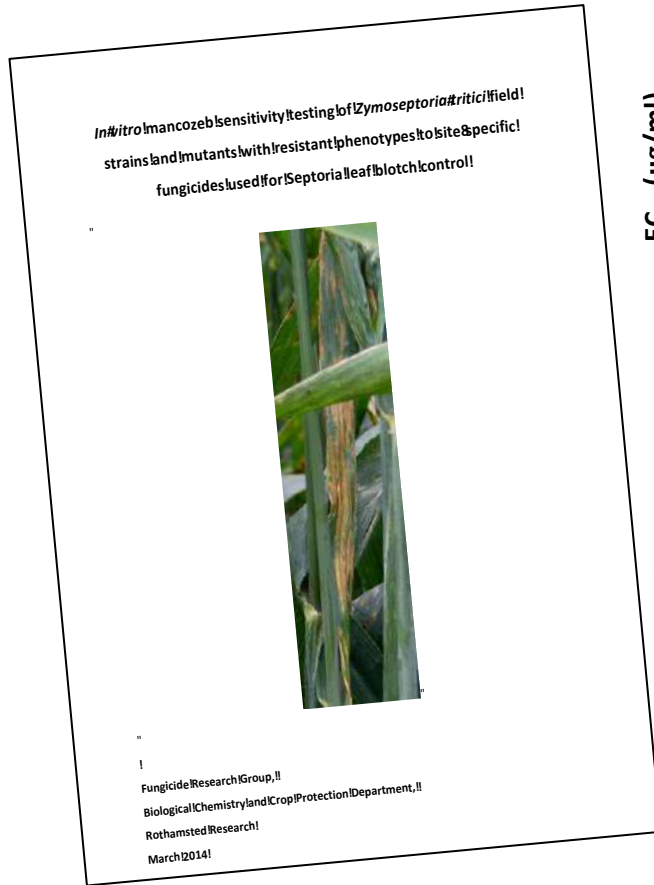
Mancozeb – working with Rothamsted

🔌 Objective

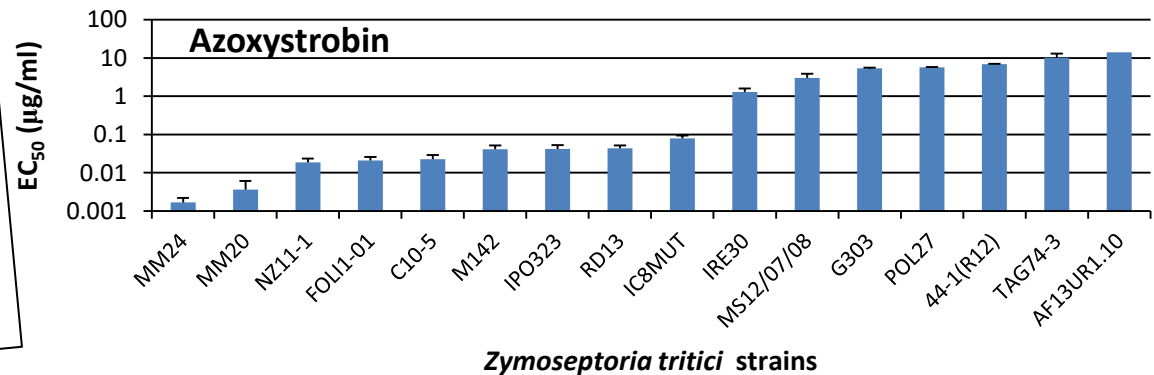
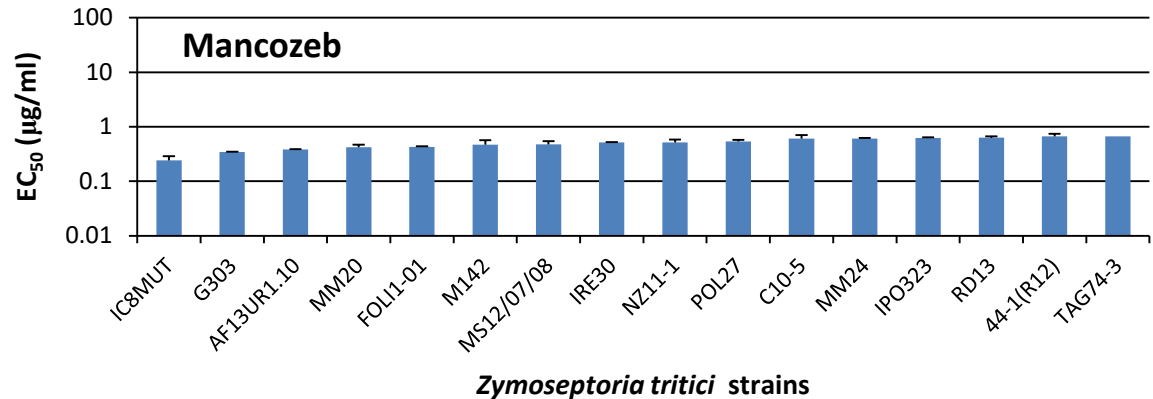
- 🔌 To test the efficacy and cross resistance patterns between a selection of fungicides on *Z.tritici* field strains (13) and lab strains (3)
- 🔌 Fungicides tested were:-
 - 🔌 Epoxiconazole, azoxystrobin, bixafen and multi-sites
- 🔌 Approximately 2500 spores of *Z.tritici* added to each well which was filled with appropriate fungicide. Plates were incubated for four days and then measured
- 🔌 Growth measured and reported as EC₅₀



Mancozeb – working with Rothamsted



Completed March 2014



Consistent level of control where mancozeb is used

Mancozeb – working with Rothamsted

- Ⓢ Resistant mutants (SDHI fungicides) and field strains (MBC, Qol and DMI fungicides) were all well controlled by mancozeb
- Ⓢ *In vitro* susceptibility testing of *Z.tritici* field strains showed no cross resistance between mancozeb and the single-site fungicides epoxiconazole, azoxystrobin, carbendazim and bixafen
- Ⓢ EC₅₀ values were narrow for the multi-sites
- Ⓢ EC₅₀ values were large for the other fungicides
- Ⓢ All multi-site fungicides were able to control strains with high levels of insensitivity to one or more classes of site-specific fungicides

Mancozeb – working with Rothamsted

- Ⓢ The sensitivity of European field strains of *Zymoseptoria tritici* (Septoria tritici) to mancozeb
- Ⓢ Sensitivity of epoxiconazole (triazole) was included in the project
- Ⓢ 11 concentrations of the test products were used to determine EC₅₀ values

Mancozeb – working with Rothamsted

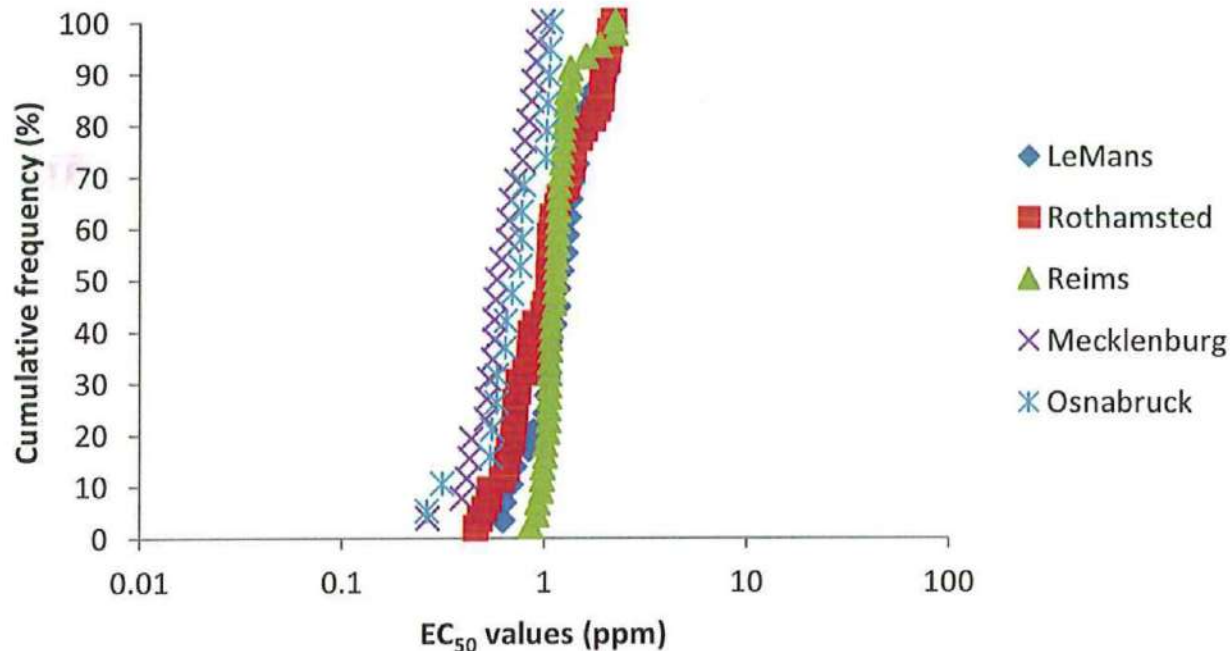
	Site name	Number of strains tested
1	Mecklenburg	26
2	Osnabruck	19
3	Reims	44
4	Le Mans	29
5	Rothamsted	53



5 sites across Europe to give a representative spread

Mancozeb – working with Rothamsted

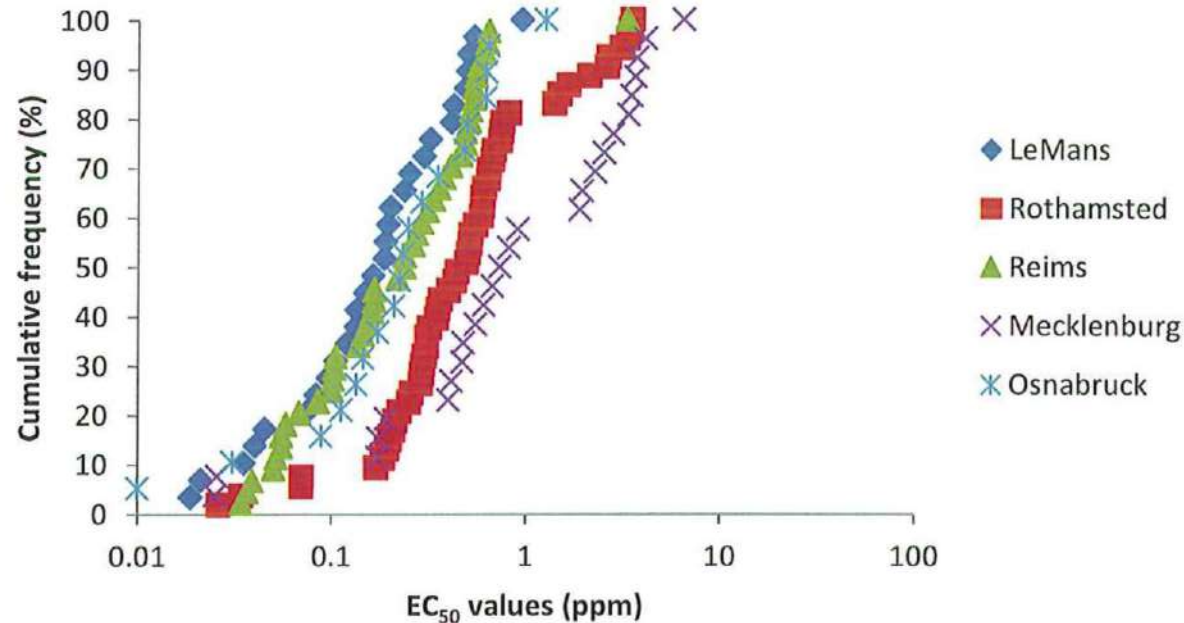
Mancozeb sensitivity test results (EC₅₀ values, ppm)



- Ⓢ Limited variation in sensitivity within sites, between 0.26 and 2.3 ppm for all strains
- Ⓢ Similar sensitivity distributions between sites
- Ⓢ No evidence for emergence of insensitive strains

Mancozeb – working with Rothamsted

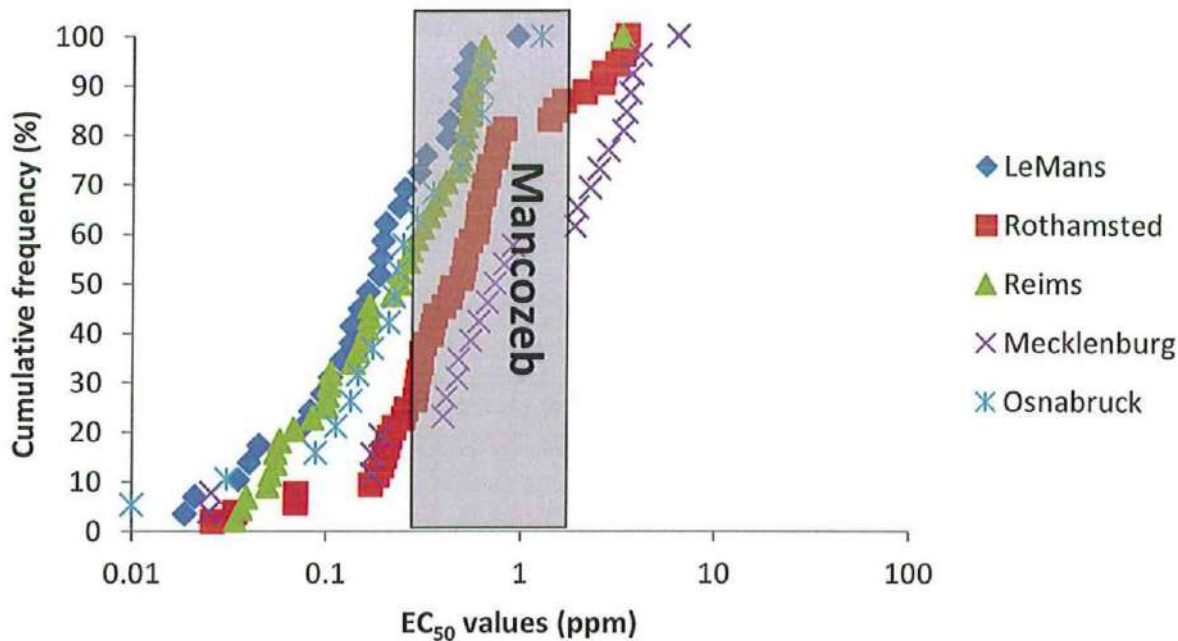
Epoxiconazole sensitivity test results (EC₅₀ values, ppm)



- ① Wide range of sensitivity within sites, between 0.01 and 6.4 ppm for all strains
- ① Similar sensitivity distributions between sites
- ① Evidence of resistance shifts and emergence of specific genotypes (e.g. CYP51 variants and strains over-expressing CYP51 and/or efflux pumps)

Mancozeb – working with Rothamsted

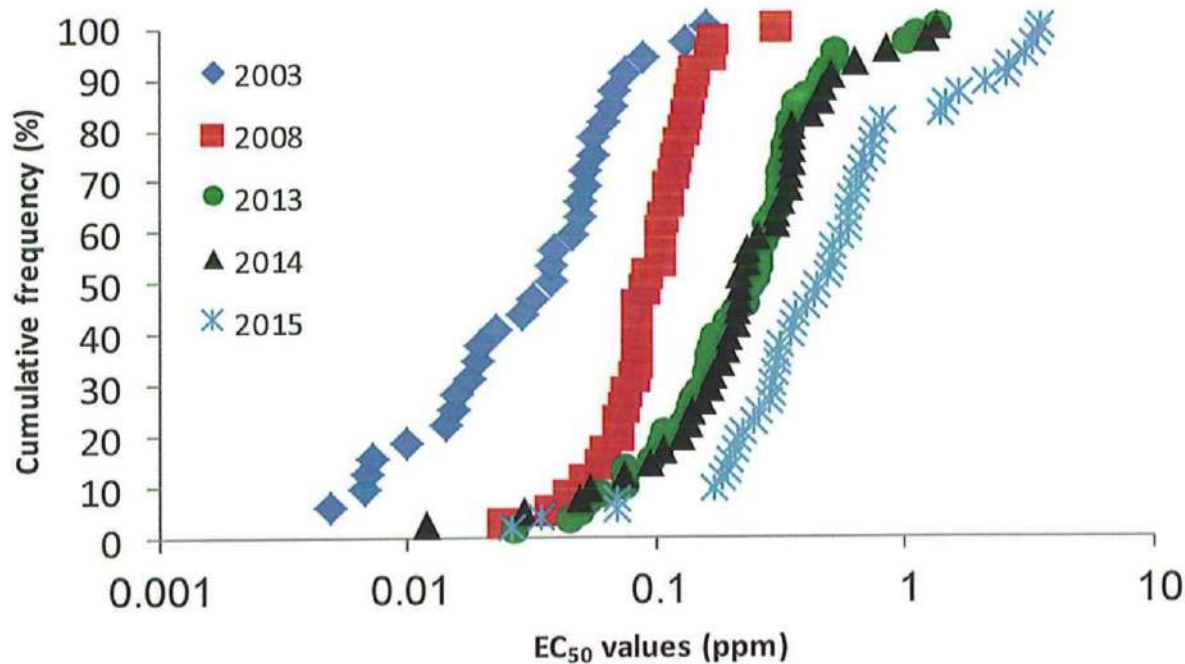
Combined sensitivity profiles – a resistant management tool in action



- ➊ Adding a multi-site inhibitor (mancozeb), with a low resistance risk, as mixing partner to a fungicide that is at-risk of resistance (epoxiconazole) reduces the rate of selection for fungicide resistance

Mancozeb – working with Rothamsted

Shifts in epoxiconazole sensitivity measure for *Zymoseptoria tritici* populations sampled at Rothamsted



- Ⓢ Long-term exposure to a fungicide that is at-risk of resistance leads to an increased rate of selection for fungicide resistance

Mancozeb – Working with University of Bologna

🌀 Apples

- 🌀 Apple scab (*Venturia inaequalis*)
- 🌀 Alternaria (*Alternaria spp.*)

🌀 Pears

- 🌀 Pear brown spot
(*Stemphylium vesicarium*)

🌀 Grapes/vines

- 🌀 Grape Downy mildew
(*Plasmopara viticola*)
- 🌀 Phomopsis Cane and Leaf spot
(*Phomopsis viticola*)
- 🌀 Black rot
(*Guignardia bidwellii*)



DIPARTIMENTO DI SCIENZE AGRARIE

Mancozeb – Working with University of Bologna

Apple scab (*Venturia inaequalis*) – conidial germination

- Ⓢ mancozeb was highly active against two tested isolates analysed, one was sensitive and the other resistant to strobilurins

Pear brown spot (*Stemphylium vesicarium*) and *Alternaria spp.* from apple – conidial germination

- Ⓢ Mancozeb was highly selective against the tested isolates

Grape Downy mildew (*Plasmopara viticola*), Phomopsis Cane and Leaf spot (*Phomopsis viticola*) and Black rot (*Guignardia bidwellii*)

- Ⓢ Mancozeb was highly selective against the tested isolates



DIPARTIMENTO DI SCIENZE AGRARIE

Petri dishes Assay in order to assess activity of mancozeb @ 6 concentrations (0-10-20-50-100 mg/L) on mycelial growth of *Phomopsis viticola* and *Guignardia bidwellii*.

Efficacy following Abbott's Formula in term of reduction of mycelial growth vs untreated from 6 to 15 days

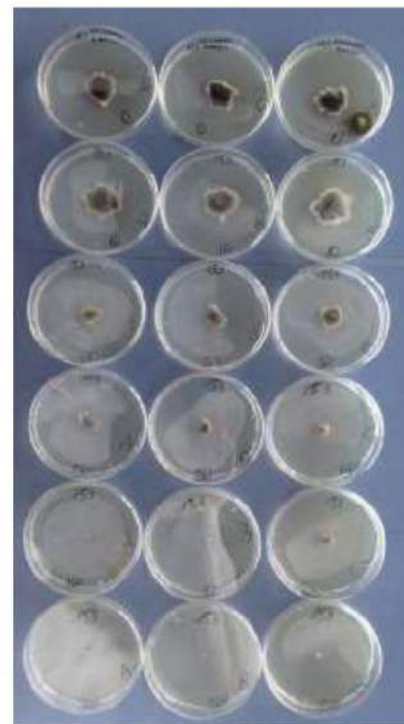
Active	Pathogen Isolate	Concentrations of mancozeb in mg/L				
		1	2,5	5	10	20
After 6 days	<i>Phomopsis viticola</i>	51,4	100	100	100	100
	<i>Guignardia bidwellii</i>	18,5	45,9	51,8	83,7	100
After 8 days	<i>Phomopsis viticola</i>	44,0	91,6	100	100	100
	<i>Guignardia bidwellii</i>	4,1	35,3	52,9	86,4	100
After 11 days	<i>Phomopsis viticola</i>	38,3	87,1	100	100	100
	<i>Guignardia bidwellii</i>	1,5	35,9	54,1	83,7	100
After 15 days	<i>Phomopsis viticola</i>	21,2	85,8	100	100	100
	<i>Guignardia bidwellii</i>	1,3	42,3	59,9	87,9	100

Petri dishes Assay in order to assess activity of mancozeb @ 6 concentrations (0-10-20-50-100 mg/L) on mycelial growth of *Phomopsis viticola* and *Guignardia bidwellii*.

Efficacy following Abbott's Formula in terms of reduction of mycelial growth vs untreated from 6 to 15 days



Phomopsis viticola – after 15 days



Guignardia bidwellii – after 15 days

Mancozeb 0 g/L

Mancozeb 200 g/L

Conclusions and final summary

- Effective and economic option for the farmers toolbox
- No resistance after over 60 yrs
- Highly compatible with other products
- UPL products show good rainfastness
- Quick degradation, safer to environment
- Key in resistance management
- Multisite





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