# UltraSorb Overview

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## Mycotoxin occurrence



#### Factors affecting Mycotoxin occurrence in the food chain



CAST mycotoxin report 2003



## Mycotoxin occurrence







CLIMATE PREDICTION CENTER, NOAA Computer generated contours Based on preliminary data



## Characteristics of mould growth



Moulds can grow over a wide range of temperatures



# **Europe Summary**



- Still early days but a lot of factors have been present to show increased levels of mycotoxins
- DON and T2 still represent the major risk in feed ingredients particularly in Cereals and Co-Products
- Warmer temperatures have favoured a higher incidence of Aflatoxins, especially in the southern of Europe

Risk maps for aflatoxin contamination in maize at harvest in 3 different climate scenarios



## Mycotoxin impact on animal health



Mycotoxin	Toxic Effect	Species
AFB1	Damage to liver, immune disorders	All animal species
DON	Reduced feed efficiency Feed refusal / anorexia, Emesis, immune disorders	All animal species, pigs being the most sensitive
ZEA	Reproductive disorders immunosuppression	All animal species, pigs being the most sensitive
FB	Damage to organs (liver, lungs and heart), nerve damage, Reduced immune response	All animal species, highly toxic to horses
T-2 Toxin	Cutaneous lesions, Reproductive disorders, Immunosuppression	All animal species, highly toxic to cats





# Mycotoxin-detoxifying agents

"substances for reduction of the contamination of feed by mycotoxins: substances that can suppress or reduce the adsorption, promote the excretion of mycotoxins or modify their mode of action."

Definition regulation (EC) N°386/2009 12/05/2009

## Options

# Mycotoxin modifiers / Bio-transforming agents:

- Degradation to non toxic metabolites
- Bio transforming agents include bacteria / yeasts / fungi and enzymes

#### Adsorbing agents (binders)

**Clays** - Bentonite, montmorillonite, aluminosilicate, activated charcoal

Yeast cell walls - Binding Ochratoxin & Zearalenone



# What is Mycotoxin Remediation?

## Controlling mycotoxin risks by:

- Specially selected biologically active ingredients to bind, transform and degrade mycotoxins within the animal
- Unique species-specific formulations, targeted to the types of mycotoxin commonly seen in their feeds
- Essential oils to counteract adverse effects from toxins







Because animal feeds contain differing amounts of forage, grains and corn, the range of mycotoxins they are exposed to are different.

Volac tailor the formulation of each product in its range to support the industry and exposure level of mycotoxin **Different Relative Toxicity** 

Toxin	Poultry	Swine	Ruminant
Aflatoxins	+++	+	+
Ochratoxins	+++	+	+
Fumonisins	+	+++	+
Zearalenone	+	+++	+++
Deoxynivalenol	+	++	++
T2 Toxin	+++	++	+++



## UltraSorb is Species Specific



### Different Gut pH and Transit Time



Ruminant	Time, h
Esophagus/Mouth	0
Rumen	<24
Omasum	<5
Obamasum	<9



Pig	Time, h
Esophagus/Mouth	0
Stomach	0-6
Small intestine	2-6
Large intestine	20-43



Poultry	Time, h
Esophagus/Crop	50
Proventriculus/gizzard	90
Duodenum	5 -8
Jejunum	20-30
lleum	50-70
Caecum	24-48 h
Colon/rectum	24



## Mode of Action



#### Binds

A blend of highly adsorbent minerals to bind a wide range of mycotoxins and their derivatives to render them harmless to the animal.

#### Degrades

Degrades mycotoxins into smaller less toxic molecules making them easier to bind

#### Transforms

Biologically active YEAST DERIVATIVES transform the molecular structure of mycotoxins to improve binding efficacy

#### Protects

A unique blend of essential oils protects against adverse effects of toxin-forming moulds which may be present in the feed.



## **UltraSorb** Composition



Inorganic mineral binders binders are selected to adsorb species specific polar mycotoxins at pH 3 and pH 7. (70%)

Organic binders selected to adsorb species specific non-polar mycotoxins. (20%)



The core is based on a blend of yeast components derived from **Saccharomyces cerevisae** to drive transformation and degradation processes. (10%)



## In vitro Aflatoxin binding at pH3 and pH7







## Degradation of ZON





**Time Minutes** 



## Degradation of ZON Using R404 HPLC





# Why Use UltraSorb over Others?



- UltraSorb uses the optimum mineral binders, proven and tested against others
- UltraSorb uses a range of yeast components
- UltraSorb has a range of products which are species specific
- Tailored to have the optimum combinations of amounts for the mycotoxins you are likely to encounter in feed
- Bespoke UltraSorb Core based on lab analysis of local clays available in the market

### FEW COMPANIES OFFER THIS UNIQUE COMBINATION



## Negative effects in ruminants











## Dose and application in ruminant species





Product	Species	Usage	Mycotoxin Risk	Cattle Dosage (inclusion) rate g/head/day	Calves, Sheep & Goats: Dosage rate g/head/day	
UltraSorb R	All	Compounders & pre-mixers		Low to med. risk 10g Med. to high risk 20g	Low to med. risk 3-5g Med. to high risk 6-10g	
UltraSorb R Farm Pack		On farm - TMR		Low to med. risk 20g Med. to high risk 40g	Low to med. risk 6-10g Med. to high risk 12-20g	
UltraSorb R Dairy	ELL registered for doiny	Compounders & pre-mixers	Any	Low to med. risk 10g Med. to high risk 20g	Low to med. risk 3-5g Med. to high risk 6-10g	
UltraSorb R Dairy Farm Pack	cattle	On farm - TMR		Low to med. risk 20g Med. to high risk 40g	Low to med. risk 6-10g Med. to high risk 12-20g	
UltraSorb R Extra		Compounders and pre- mixers, also on farm - TMR	<i>Fusarium sp</i> . (Fumonisin, T2, DON, HT2, ZON). Damp & cool.	Low to med. risk 10g Med. to high risk 20g	Low to med. risk 3-5g Med. to high risk 6-10g	
UltraSorb R Plus	All	at advised dose rates.	<i>Aspergillus sp.</i> (Aflatoxins). Warm & dry climates.	Low to med. risk 25g Med. to high risk 50g	Low to med. risk 7-12g Med. to high risk 15-25g	
UltraSorb R Core Compounders & pre- mixers. Mix with a binder. Mix 650-850kg of selected my add 150-350kg of UltraSorb F at a dose of 10-20g/h/d in fee capacity.				cotoxin binder with suitable de- Core and mix until blend is uni d depending on the mycotoxin c	dusting agent. Afterwards form. Apply complete mixture contamination level or binding	



## Negative effects in swine











## Dose and application in swine





Product	Usage	Mycotoxin	Sows: dry & gestating (kg/tonne)	Boars: Developing & mature(kg/tonne)	Piglets: Pre-starter, starter (kg/tonne)	Growers & finishers (kg/tonne)
UltraSorb S		All	Low to med. risk 1kg Med. to high risk 2kg	Low to med. risk 0.5kg Med. to high risk 1kg	Low to med. risk 1kg Med. to high risk 2kg	Low to med. risk 0.5kg Med. to high risk 1kg
UltraSorb S Extra	Compounders and pre- mixers, or on farm at appropriate dose rates.	Increased risk of Fusarium sp. (Fumonisin, T2, DON, HT2, ZON). Cool & damp.	Low to med. risk 1kg Med. to high risk 2kg	Low to med. risk 0.5kg Med. to high risk 1kg	Low to med. risk 1kg Med. to high risk 2kg	Low to med. risk 0.5kg Med. to high risk 1kg
UltraSorb S Plus		Increased risk of Aspergillus sp. (Aflatoxins). Warm & dry climates.	Low to med. risk 2.5kg Med. to high risk 5kg	Low to med. risk 1.25kg Med. to high risk 2.5kg	Low to med. risk 2.5kg Med. to high risk 5kg	Low to med. risk 1.25kg Med. to high risk 2.5kg
UltraSorb S Core	Compounders and pre- mixers. To be mixed with a binder.	All	Mix 650-850kg of selected mycotoxin binder with suitable de-dusting agent. Then add 150-3 UltraSorb S Core and mix until blend is uniform. For best mixing properties avoid the use of oils as a de-dusting agent. Apply complete mixture at a dose of 0.5 – 2kg/t feed depending c mycotoxin contamination level or binding capacity.			



## Negative effects in poultry









## Dose and application in avian species





Product	Usage	Mycotoxin	Broiler chickens (kg/t complete feed)		Breeders & layers (kg/t complete feed)		Ducks, turkeys, geese (kg/t complete feed)				
				Low to med. risk	Med. to high risk		Low to med. risk	Med. to high risk		Low to med. risk	Med. to high risk
			Starter	1	2	Pullet	1	2	All	1	2
UltraSorb P		All	Grower	1	2	Pre-layer	1	2			
		Finisher	0.5	1	Layer Breeder	0.5	1			-	
UltraSorb P Extra UltraSorb P Extra UltraSorb P Extra Compounde rs, premixers and on farm T2, HT2	High risk of Fusarium species (Fumonisin, T2, DON, HT2, ZON)	Starter	1	2	Pullet	1	2	All	1 2		
		Grower	1	2	Pre-layer	1	2			2	
		Finisher	0.5	1	Layer Breeder	0.5	1				
		High riskof	Starter	2.5	5	Pullet	2.5	5			
UltraSorb P Plus	Aspergillus	Grower	2.5	5	Pre-layer	2.5	5	All	2.5	5	
		species (Aflatoxins)	Finisher	1.25	2.5	Layer Breeder	1.5	2.5			
UltraSorb P Core	Mix with binder	Mix 650 - 850kg selected mycotoxin binder with suitable de-dusting agent. Afterwards add 150 - 350kg UltraSorb P Core and mix until the blend is uniform. For the best mixing properties avoid the use of vegetable oil as a de-dusting agent. Apply complete mixture at an inclusion rate of 0.5 - 2 kg/t of feed depending on mycotoxin contamination level.									



# Next Generation R&D: Endotoxin Challenge



## Improving host immunity



# Endotoxins – what are they and how do they stimulate inflammation?





#### What are Endotoxins?

- Vital component of the outer cell wall of bacteria – cell structure, adhesion and protection
- Found on GNB (lipopolysaccharide, LPS) and GPB (lipoteichoic acid, LTA)
- Pathogen associated molecular pattern (PAMPs)

#### How do they stimulate inflammation?

- Detected by cell membrane bound receptors (Toll-like receptors TLRs)
- Cause a signalling cascade and production of proinflammatory cytokines



Alexandrov et al. 2020; Eckel and Amtaj, 2016; Kawai and Akira, 2010

## Endotoxin Related Challenges in Animal Production

- The Obvious:
  - Mastitis
  - Metritis

#### The Less Obvious:

- Acidosis
- Transition period
- Heat stress
- Receiving phase
- Immunocompromised animals
- Environment



#### 70% of the immune system is located in the gut



# **Example Pathogenesis: Acidosis**

- Over 20 billion bacteria in 1 teaspoon of rumen fluid
- Many of which are gram negative and have LPS
- During acidosis, lysis of GNB is the biggest contributor to free LPS in rumen fluid
- Acidotic diet also increases GNB growth, contributing further to free LPS
- Increased gut permeability tight gut junctions
- TLR4 signalling pathway leading to proinflammatory response
- Endotoxocis



Table 6. The mRNA expression of tight junction proteins of the colonic epithelium of control (CON) dairy goats and dairy goats with SARA (n = 6 goats/group)

	Trea	tment			
Item	CON	SARA	SEM	<i>P</i> -value	
CLDN1 CLDN4 CLDN7 OCLN ZO1	$\begin{array}{c} 0.98 \\ 1.01 \\ 1.03 \\ 0.93 \\ 0.91 \end{array}$	1.65 1.21 0.88 1.54 0.83	$\begin{array}{c} 0.128 \\ 0.073 \\ 0.046 \\ 0.055 \\ 0.047 \end{array}$	$\begin{array}{c} 0.030 \\ 0.826 \\ 0.583 \\ 0.016 \\ 0.590 \end{array}$	



# Additive challenge of mycotoxins and endotoxins?

- T-2 and LPS simultaneous exposure lead to:
  - Increased mortality
  - Hypothermia
  - TNF $\alpha$  production
  - Thymic atrophy



Indirect signalling through TLR4 pathway – proinflammatory cytokine production



## **UltraSorb Next Generation**

### LPS binding

- Reducing endotoxin load at times of stress
- Gut integrity
  - Supporting gut function



### Liver function

• Supporting other biological functions



## Questions?



