

# Fungus Species

- More than 50 species of Fungus produce more than 400 types of Mycotoxins:
- Fusarium sp
- Aspergillus sp
- Penicilium sp
- Claviceps sp

# Important Points

- Generally, Mycotoxins in feed are mixed and rarely occurs singular (**Solution should take care of Mixed toxins**)
- Mycotoxins are synergistic in action- Mixed mycotoxins are more Dangerous
- Aflatoxins entry from feed to milk average 6%
- Mycotoxins can enter Milk within 6 hours after animal has taken feed
- As Calves have Rumen at developing stage, mycotoxins are more toxic for Calves.
- Toxins also interact with Amino acids and Minerals absorption present in the feed.

## Source of mycotoxins in ruminant feed

Ingredient	Mycotoxin occurrence	Inclusion rate in diet	Contribution in total mycotoxin risk
Maize silage	All mycotoxins, high occurrence	High	High
DDGS	All mycotoxins, very high occurrence	Medium	High
Maize	All mycotoxins, high occurrence	High	High
Grass (pasture)	Ergot alkaloids, occurrence depends on year	High	Medium
Wheat bran	Trichotecenes, ZEA, ergot alkaloids, occurrence depends on year	High	Medium
Grass silage	Low occurrence	High	Low
Oilseeds	OTA, ZEA	High	Low
Wheat	Trichotecenes, ZEA, ergot alkaloids, occurrence depends on year	Medium	Low

# Mycotoxins in ruminants

## Reproductive system

### ZEA

- Irregular heats
- Low conception rates
- Ovarian cysts
- Embryonic loss

## Central nervous system

### ERG

- Neck paralysis

## Gastro-intestinal tract

### FUM, T2, DON

- Gastroenteritis
- Intestinal hemorrhages
- Impaired rumen function
- Diarrhea
- Ketosis



## Udder

### AFL, T2, DON,

- Milk contamination
- Decreased milk production
- Mastitis

## Hoofs

### FUM, DON, ERG

- Laminitis

## Performance

### T2, DON

- Decreased feed intake
- Lower milk production
- Decreased feed efficiency

# Negative effects of mycotoxins in ruminants

- Calves
- Heifers

## Common symptoms:

- Decreased feed intake
- Reduced weight gain
- Inhomogenous growth
- Higher mortality
- Diarrhoea
- Gastroenteritis
- Enlarged and reddened teats in heifers
- Poor response to vaccination
- Lesions on the muzzle, lips, tongue, and pharynx

- Dairy cattle
- Beef cattle

## Common symptoms:

- Decreased feed intake, increased feed refusal, anorexia
- Reduced weight gain, weight loss
- Decreased milk production, agalactia
- Depressed butterfat in milk
- Elevated somatic cell count, mastitis
- Diarrhoea
- Gastroenteritis
- Higher incidence of ketosis (fatty liver), retained placenta, metritis, fertility disorders, displaced abomasum
- Lameness, hoof necrosis
- Ovarian cysts
- Vaginitis, vaginal secretions
- Increased body temperature/heat stress
- Staggering gait
- Poor response to vaccination
- Aflatoxin M1 in milk





# Effect of mycotoxins on milk production and quality

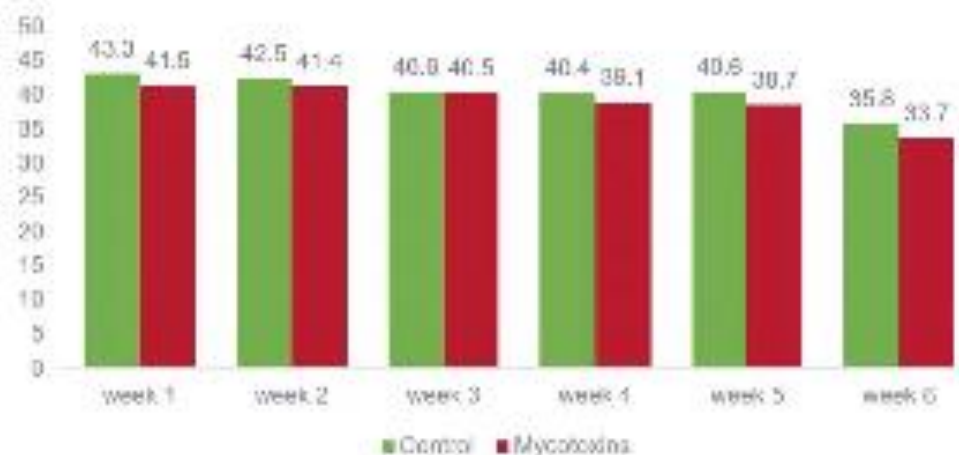
↓ milk production

↓ milk fat

↑ SCC

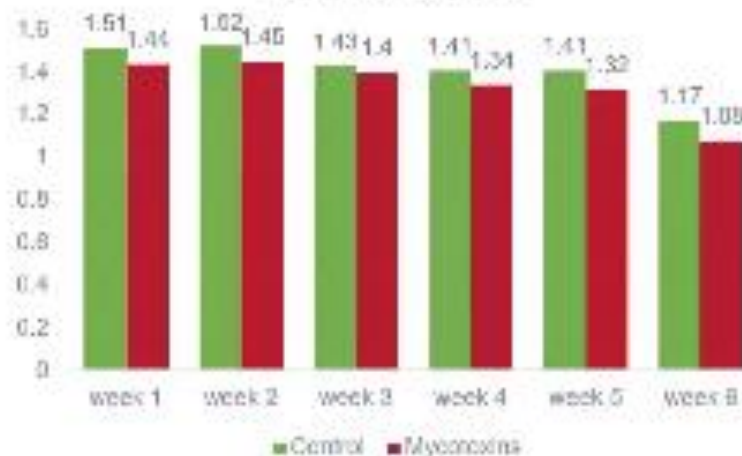
↑ Milk contamination by Aflatoxin M1

Energy correlated milk, kg/day

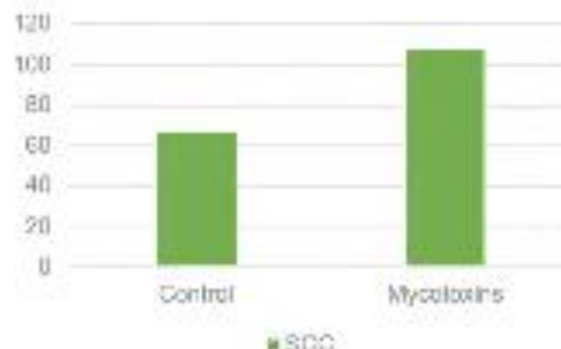


FUM 1054 ppb, ZEA 628 ppb, DON 4771ppb

Fat yield, kg/day



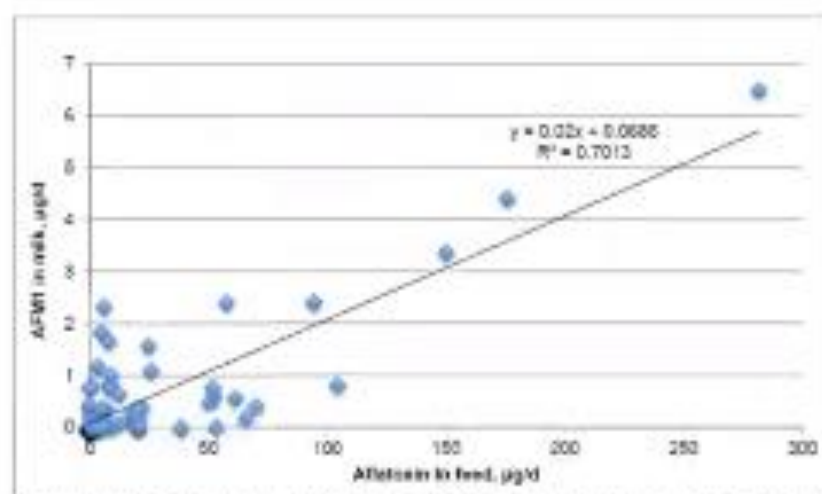
SCC



natural contamination -  
fusarium mycotoxins DON,  
15-acetyl DON and ZEA  
(Korosteleva et al. 2009)

# Typical symptoms of mycotoxicosis

- Aflatoxin B1 – hepatotoxic
  - 0.05 ppb
- Aflatoxin B1 – food quality concern
  - 0.002 ppb (0.02 ppb):



# Life phase – different metabolism

medium



heifer

Lower feed/mycotoxins turnover in rumen – better ruminal deactivation (e.g. DON)

Consider health of cow-fetus due to stable mycotoxins (e.g. ZEA, ERG, AFL)

Mycotoxins stable in rumen can cause abortion (e.g. ZEA, ERG)

high



dry cow

Lower feed/mycotoxins turnover in rumen – better ruminal deactivation (e.g. DON)

Consider health of cow-fetus due to stable mycotoxins (e.g. ZEA, ERG, AFL)

Mycotoxins stable in rumen can cause abortion (e.g. ZEA, ERG)

Mycotoxins can cause oxidative stress (AFL)

high



1/3 lactation

Higher feed/mycotoxins turnover in rumen – worse ruminal deactivation (e.g. DON)

Mycotoxins will affect performance (DON – yield and fat) and milk quality (AFL)

ZEA can cause false oestrus and ineffective insemination

medium



2-3/3 lactation

High feed/mycotoxins turnover in rumen – worse ruminal deactivation (e.g. DON)

Mycotoxins will affect performance (DON – yield and fat) and milk quality (AFL)

Mycotoxins stable in rumen can cause abortion (e.g. ZEA, ERG)



## Maximum legal limits in complete feed, ppb

Mycotoxin	Dairy cows	Calves, lambs	Others
Aflatoxin*	5	10	50
DON**	5.000	2.000	5.000
Ochratoxin	200	100	200
Zearalenone**	500	500	500
T+HT-2 toxins	100	100	100
Fumonisin B1+B2**	50.000	20.000	50.000

\*DIRECTIVE 2002/32/EC

\*\*COMMISSION RECOMMENDATION 2006/576/EC



# strategy in mycotoxin management → **Protect**

## Multiple Toxin Binder- Toxynil

Detoxify



TOXY-NIL®

UNIKE® PLUS



### Adsorption

High adsorbent modified clay minerals

Bentonite → AFB1  
Sepiolite → ZEA, OTA, FUM, ERN  
Yeast wall → ZEA  
(*In vitro* evaluation)



### Bio-inactivation

Yeast and fermentation extracts

inactivation of mycotoxins by the intestinal microflora or by host cells, stimulated by the yeast

Repair



### Immune system revival

Fermentation extracts

➤ Innate immunity (macrophages, cytokines)  
➤ Adaptive immunity (antibodies, ➤ vaccination efficacy)



### Oxidative stress inhibition

Antioxidant and preservative mixture

Strong oxidant: ZEA  
Moderate oxidant: Fum, AFB1,  
Low oxidant: DON, OTA  
→ ROS production and then, oxidative damage



### Aid to affected organs

Botanicals



# R&D results to support the mode of action of **TOXY-NIL<sup>®</sup>**



University of Missouri

## Protocol

- ❑ 24 mid-lactation Holstein cows (183 ± 70 DIM)
- ❑ Randomized block design, blocks formed on days in milk, milk yield, and parity (n=8/treatment)
- ❑ Trial duration : 21 days (7 days adaptation / 7 days experiment / 7 days recovery (NC))
- ❑ 3 treatments :
  - ❑ **NC**: no aflatoxin B<sub>1</sub> and no mycotoxin deactivator
  - ❑ **PC**: 2.8 mg of aflatoxin B<sub>1</sub>/cow/day\*
  - ❑ **TN**: 2.8 mg of aflatoxin B<sub>1</sub>/cow/day\* + 100 g of **TOXY-NIL<sup>®</sup>**/cow/day
- ❑ Measured parameters :
  - ❑ Aflatoxin M<sub>1</sub> concentrations in milk and urine
  - ❑ Gene expression (RNA-sequencing) in blood leucocytes and milk somatic cells

*\*aflatoxin B<sub>1</sub> (650 mg/kg) culture material was incorporated into the diets. 5X FDA limit*

**ADISSEO**  
A BUNGE COMPANY





# R&D results to support the mode of action of TOXY-NIL®



University of Missouri

## Results: aflatoxin intake, excretion, clearance



	NC	PC	TN	SEM
<b>Intake of aflatoxin, <math>\mu\text{g}/\text{kg}</math> of diet</b>	0.0 <sup>a</sup>	106.5 <sup>b</sup>	107.6 <sup>b</sup>	2.9
<b>Intake of adsorbent, % of diet</b>	0.0 <sup>a</sup>	0.0 <sup>a</sup>	0.4 <sup>b</sup>	0.0
<b>Milk</b>				
<b>AFM<sub>1</sub> concentration, <math>\mu\text{g}/\text{kg}</math></b>	0.0 <sup>a</sup>	0.6 <sup>b</sup>	0.2 <sup>c</sup>	0.1
<b>AFM<sub>1</sub> excretion, <math>\mu\text{g}/\text{d}</math></b>	0.0 <sup>a</sup>	20.5 <sup>b</sup>	8.1 <sup>c</sup>	1.7
<b>AF transfer, %</b>	0.0 <sup>a</sup>	2.7 <sup>b</sup>	1.0 <sup>c</sup>	0.2
<b>Clearance of AFM<sub>1</sub>, %/d</b>	0.0 <sup>b</sup>	46.1 <sup>a</sup>	66.5 <sup>a</sup>	6.7
<b>Urine</b>				
<b>AFM<sub>1</sub> concentration, <math>\mu\text{g}/\text{l}</math></b>	0.5 <sup>c</sup>	14.7 <sup>a</sup>	6.9 <sup>b</sup>	1.5
<b>AFM<sub>1</sub> excretion, <math>\mu\text{g}/\text{d}</math></b>	15.4 <sup>c</sup>	521.6 <sup>a</sup>	225.8 <sup>b</sup>	53.1
<b>AF transfer, %</b>	0.0 <sup>c</sup>	18.6 <sup>a</sup>	8.0 <sup>b</sup>	1.9





# R&D results to support the mode of action of TOXY-NIL®

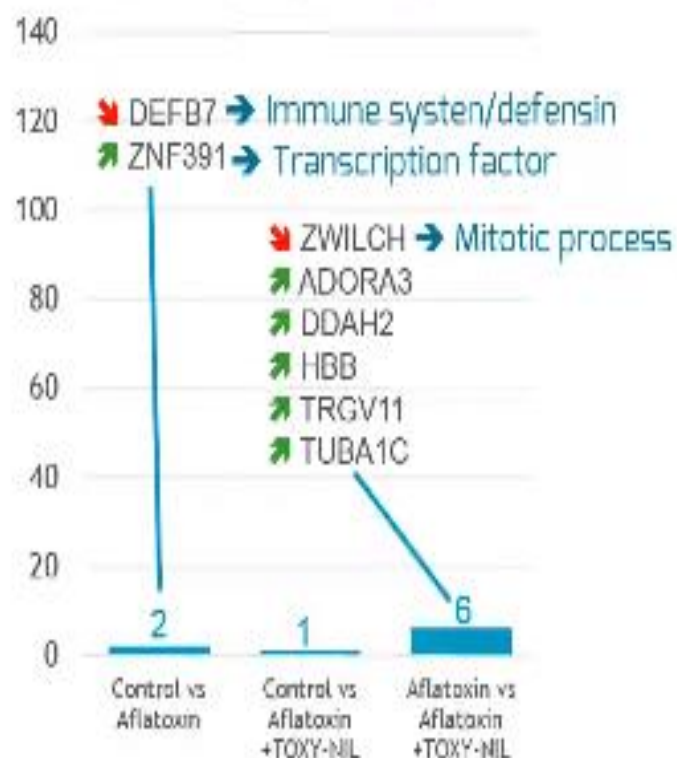


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## Results: gene expression

### Blood leucocytes

Differently expressed genes



Downregulation  
Upregulation

### Milk leucocytes

Differently expressed genes

