

NOVUS[®]

SOLUTIONS SERVICE SUSTAINABILITY™

The commercial relevance of micro-mineral nutrition for monogastrics



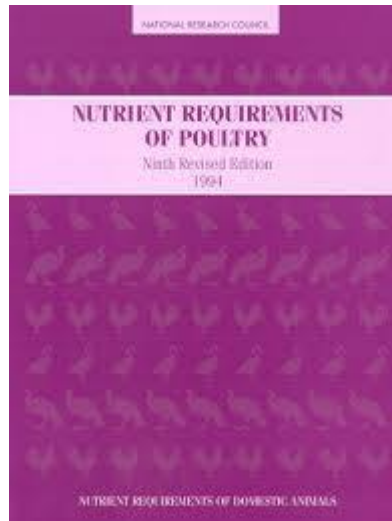
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Micro Mineral Nutrition World 2014

In spite of all the things we know and understand about nutrition we don't really know the actual mineral requirements of the animals we feed



“Thus there is a great deal of literature concerning the calcium and phosphorus requirements of the broiler and minimal research concerning requirements for trace elements” (NRC, 1994)

Many nutritionists don't worry much since they believe that, by using “safely high” levels of Zn, Cu and Mn with inorganic sources, they maximize performance

Micro Mineral Nutrition World 2014

Many nutritionists believe that using organic minerals is a luxury and can ONLY be used if the use reduces total feed cost, generally done by using lower mineral levels.

Many nutritionists don't think much about the differences between inorganic and organic mineral molecules and treat them as equivalents.

Mineral sources available for commercial animal nutrition

- Inorganic sources of micro minerals
 - Sulphates
 - Oxides
 - TBCC's
 - ETC
- Organic sources of micro minerals
 - True Chelates (Mintrex, Glycinates)
 - Non-Chelates (many)

Comparing Organic Trace Mineral Ligands

- Ligands are ions or neutral molecules that bond to a central metal atom or ion.
- *Zinc is Zinc, Copper is Copper, Manganese is Manganese*
- The only reason to feed an organic trace mineral (OTM) over a mineral salt is to deliver minerals to the blood
- The only point of **differentiation** among OTMs is their **ligand**.
- **Structure definition** is critical to determine physical and chemical characteristics as well as consistency from lot to lot
- Chemical and physical **stability** is critical to performance.
- If the molecule disassociates prior to reaching intestinal receptor sites, antagonists can bind the mineral, causing it to precipitate out of solution rendering it unavailable to the animal.
- Receptor site proteins will absorb the soluble mineral and “pass” it to specific “chaperone,” or transport, proteins or enzymes

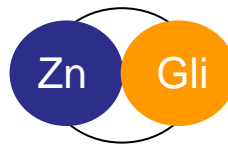
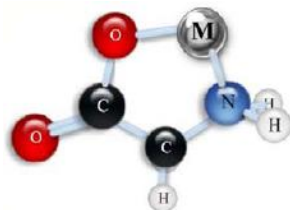
Organic Trace Minerals: Non chelates

AAFCO CLASS OF LIGAND*	MOLECULAR REPRESENTATION	SCHEMATIC MODEL	CHARACTERISTICS
<p>Metal (Amino Acid) Complex Product resulting from complexing a soluble metal salt with an amino acid(s)</p>			<ul style="list-style-type: none"> • Many of them do not have a defined structure ie, there can be changes in the ligand (different AAs, protein chains or carbohydrates) • Therefore it is not possible to determine the bond strength, the number of bonds, molecular size and the amount of ligands in the product. • The composition can vary and the amount of chelated mineral is uncertain. • Even when the ligand is well known, the protection to the metal is partial since the ligand covers only one side of it.
<p>Metal (Specific Amino Acid) Complex Product resulting from complexing a soluble metal salt with a specific amino acid</p>			
<p>Metal Proteinate Product resulting from the chelation of a soluble salt with amino acids and/or partially hydrolyzed protein</p>			
<p>Metal Polysaccharide Complex Product resulting from complexing a soluble salt with a polysaccharide solution</p>			

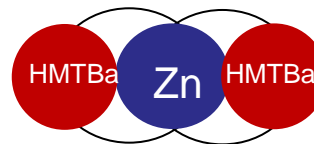
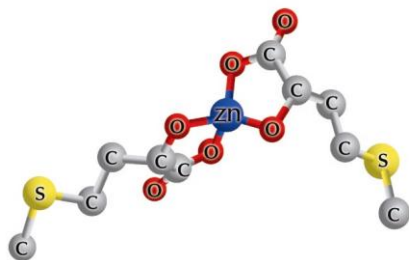
Organic Trace Minerals: True chelates

AAFCO CLASS OF LIGAND*	MOLECULAR REPRESENTATION	SCHEMATIC MODEL	CHARACTERISTICS
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Chelate Metal Amino acid



Chelate metal-HMTBA



- Are true chelates
- They have defined structure, composition and sizes

Minerals in the Gastric Environment (pH 2 -5)

Proventriculus (pH 2 - 5)

Small Intestine (pH 6 - 6,5)

CuO



CuSO4



Soluble
Complexes

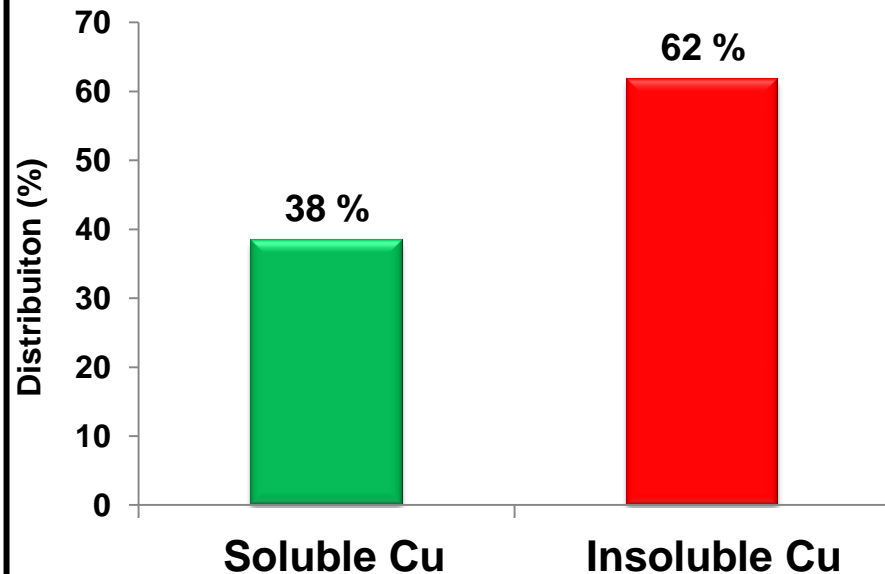
Insoluble
Complexes



Form of copper present in the duodenum and jejunum

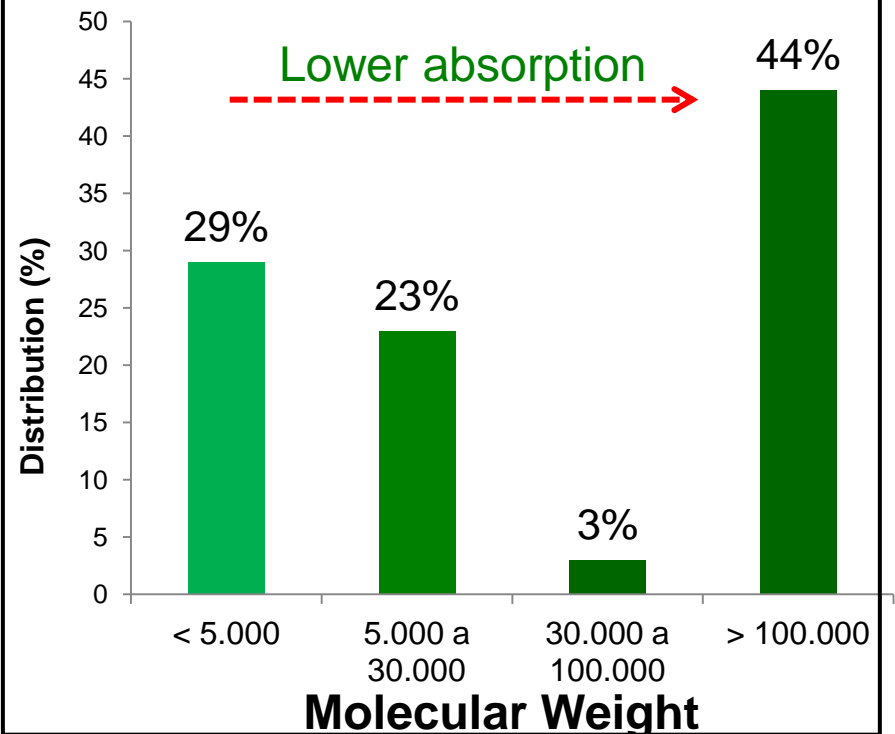
- Experiments done *in vivo*
- Copper Sulfate = 8 ppm Cu

Solubility of Cu in the small intestine



Soluble portion:

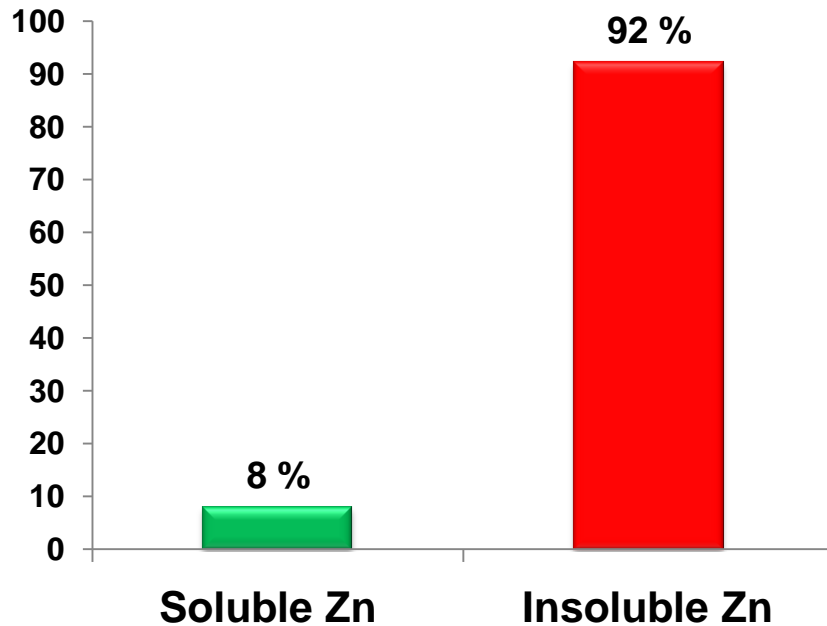
Size of the complex Cu intestine



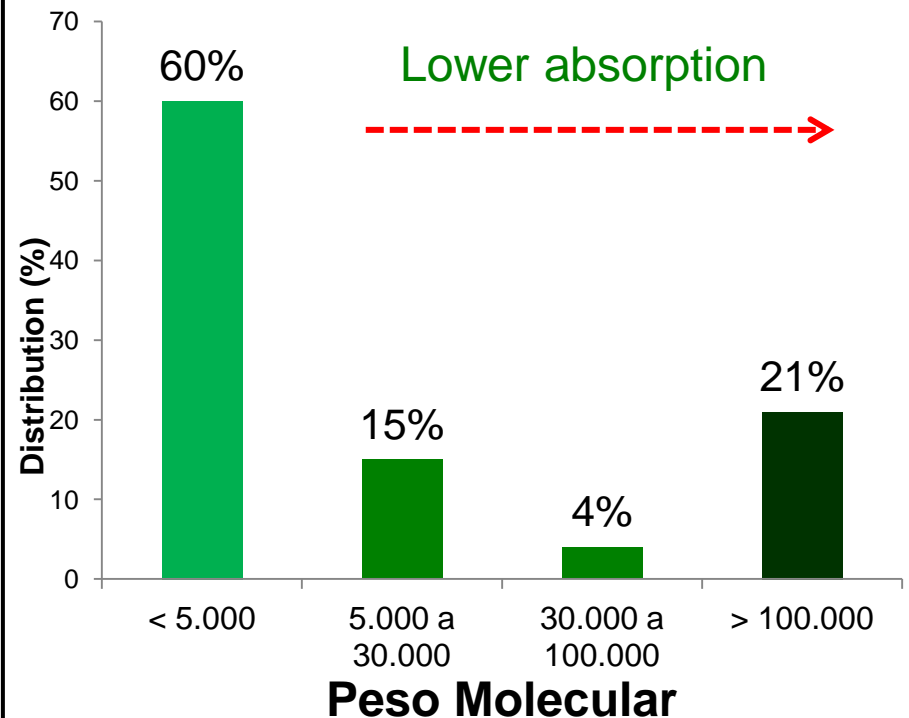
Form of Zinc present in the duodenum and jejunum

- Experiments conducted *in vivo*
- Zinc Oxide = 125 ppm Zn

Solubility of Zn in the small intestine



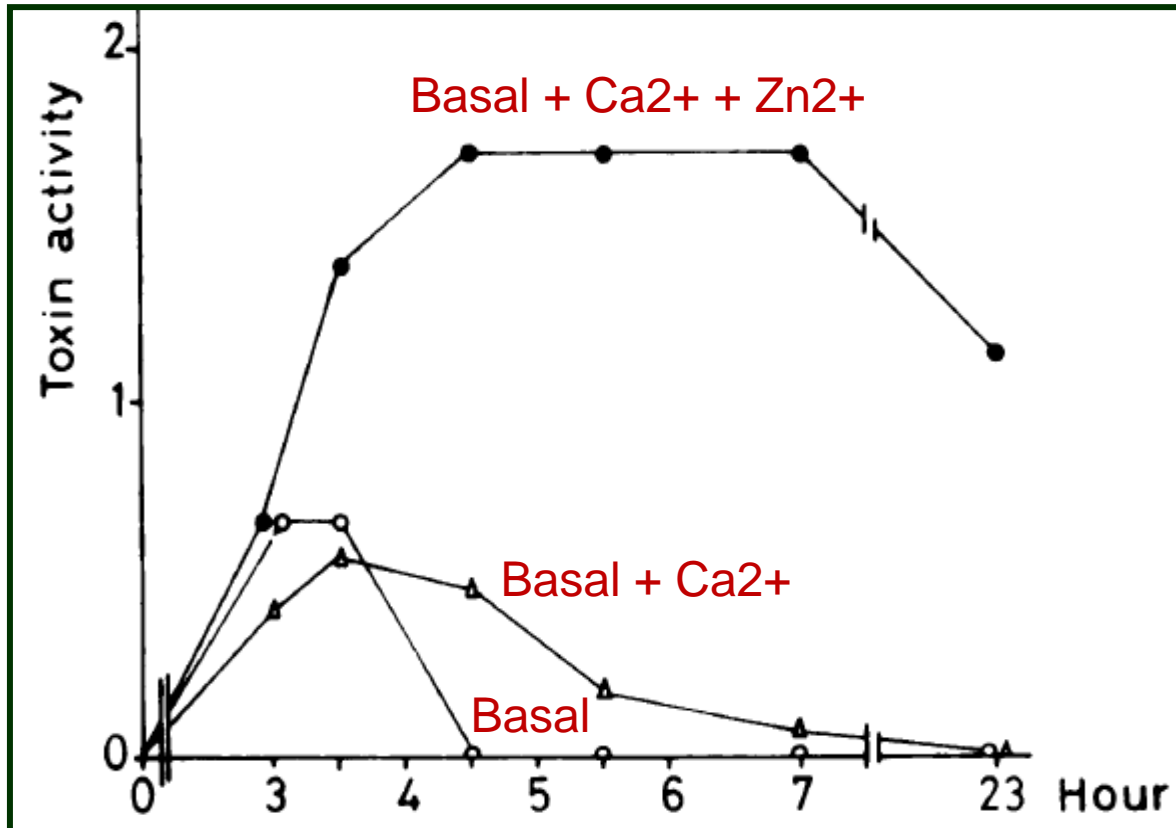
Soluble portion: Size of Zn complexes in the intestine



Ideas on Micro Mineral Nutrition World Few Years Ago

We don't understand what really happens to these minerals throughout the gut, the interactions, etc

Zn as activator of α -toxin of *Clostridium perfringens*

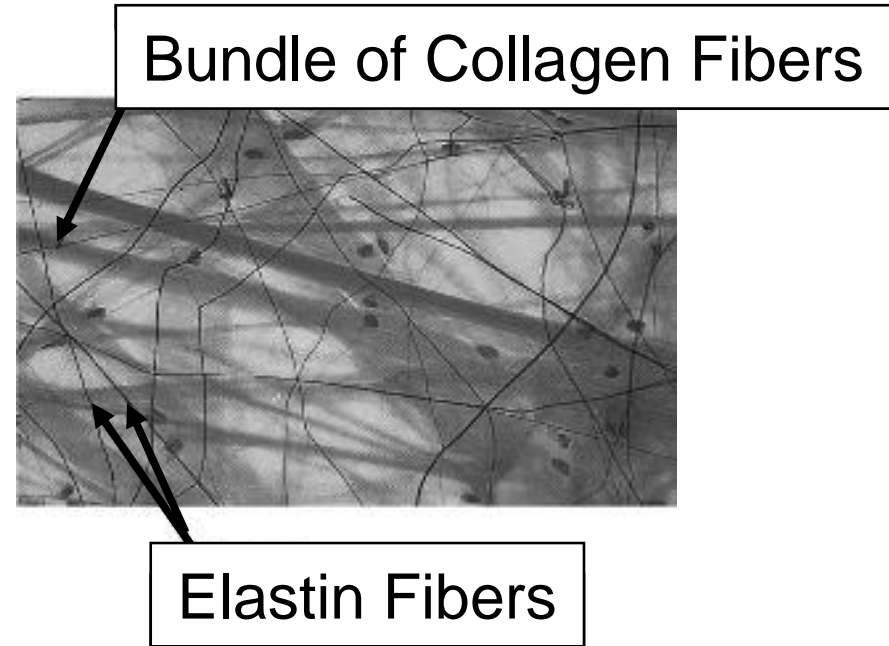


Ca induces denaturation of α -toxin while Zn activates and makes α -toxin stable

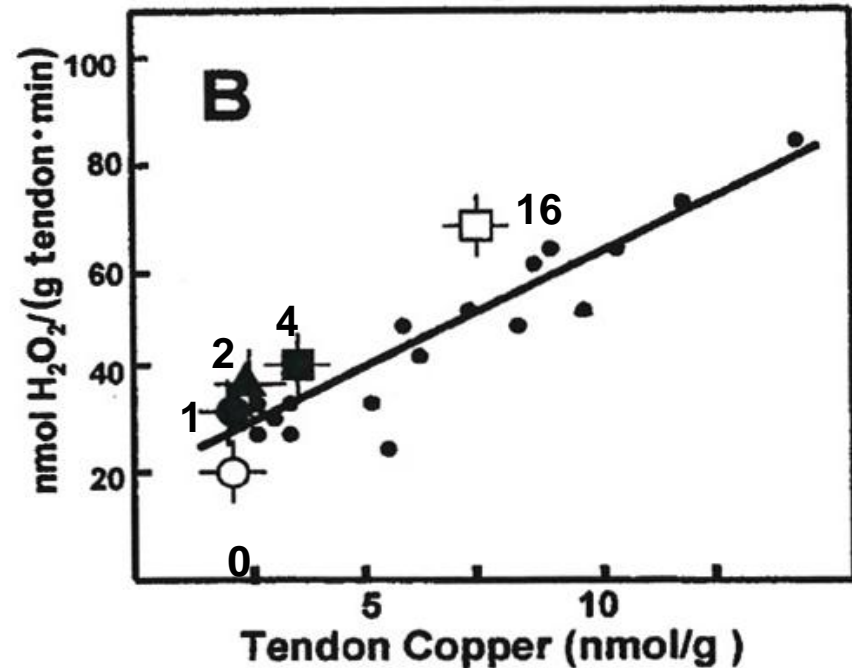
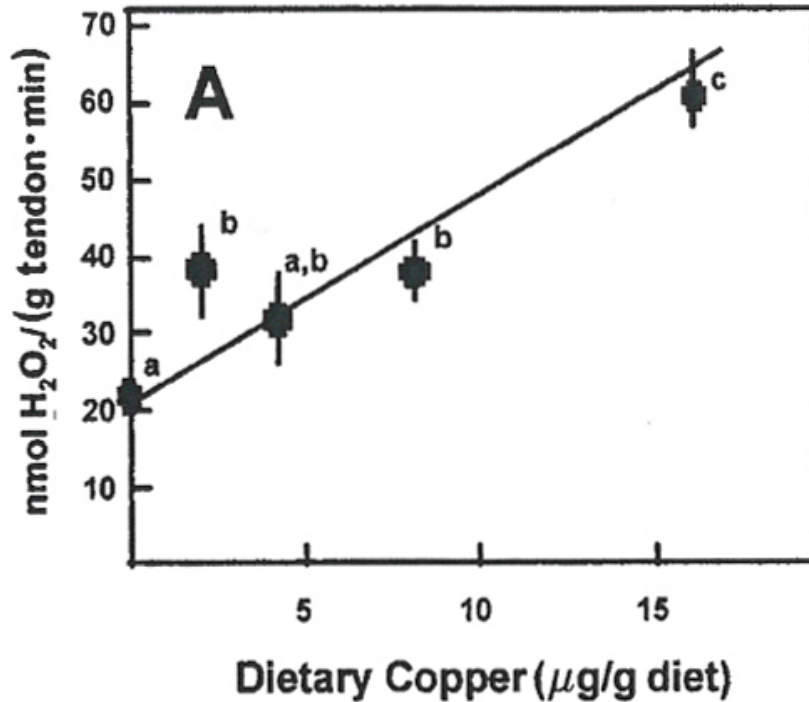
Some Biologic Functions of Zn, Cu and Mn



Connective Tissue: Essential for Function

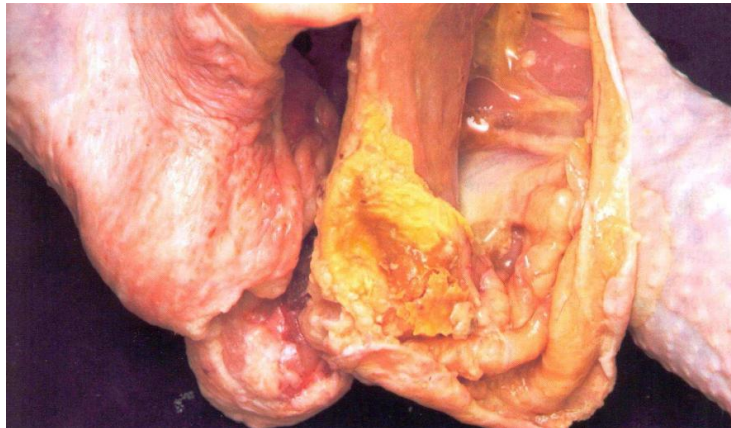


Collagen Crosslinking and Dietary Copper



Dietary Cu above that required for optimum performance was associated with increased chick tendon lysyl oxidase activity and tendon Cu was correlated with enzyme activity ($r^2 = 0.88$).

Partial condemnations due to skin problems

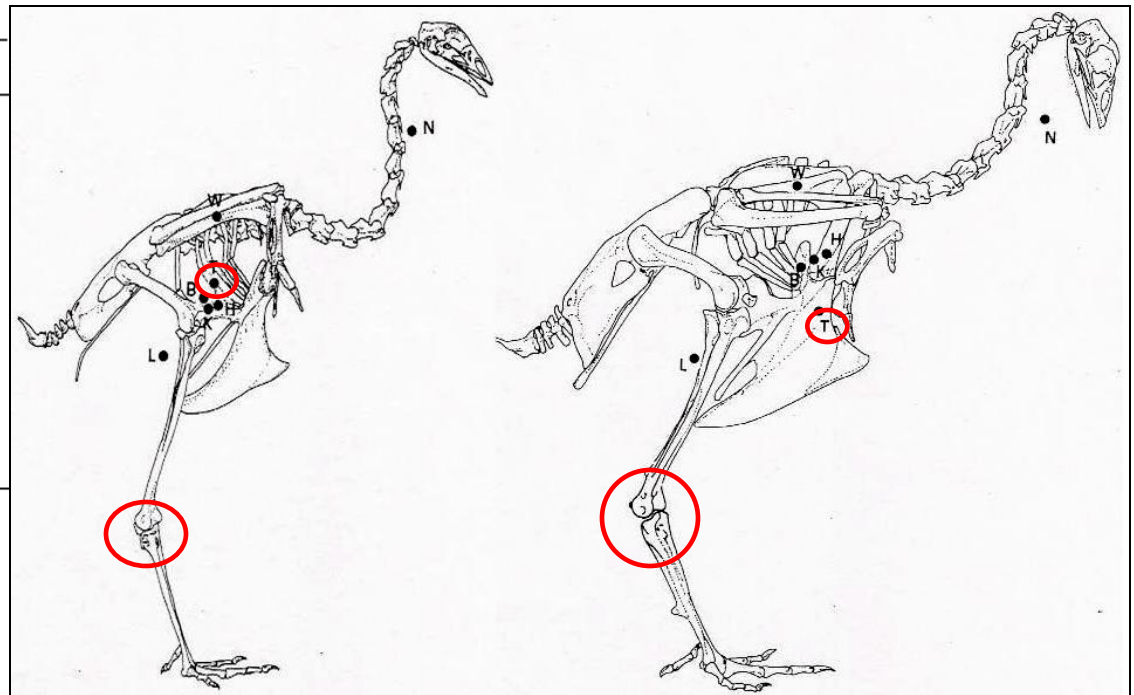


Broiler Breeder Hens: Carcass Effects

Effects of Body Weight and Feed allocation During Sexual Maturation in Broiler Breeder Hens. R.A. Renema

Breast Muscle Percentage At Sexual Maturity

Source	Breast muscle	
	Weight	Percentage ¹
	(g)	(%)
Feed ³		
AL	489.7 ^a	14.59 ^b
RF	446.7 ^b	16.20 ^a
SEM	7.7	0.23
Size ⁴		
LOW	442.2 ^b	15.47
STD	469.6 ^a	15.13
HIGH	492.8 ^a	15.59
SEM	9.5	0.28
Source of variation		
Feed	0.0002	0.0001
Size	0.002	0.48



Abourachid, 1993;
Emmerson et al, 1997;
Nestor et al, 2001.

Lameness (Bone Health) is currently the 4th most prevalent health issue in US broiler Industry



Financial loss estimated at \$120,000,000/year in USA (Cook, 2000)

Rank	Condition
1	Enteric
2	Respiratory
3	Skin
4	Lameness
5	Chicks

All top 4 issues associated with Barrier Health



Catalytic properties of enzymes involved in maintaining egg shell/membrane quality and supporting embryogenesis

- **Zn- Carbonic anhydrase**

- Activity of carbonic anhydrase, an essential enzyme for eggshell formation, has been directly related to zinc status of the hen
- Zn functions in collagen synthesis

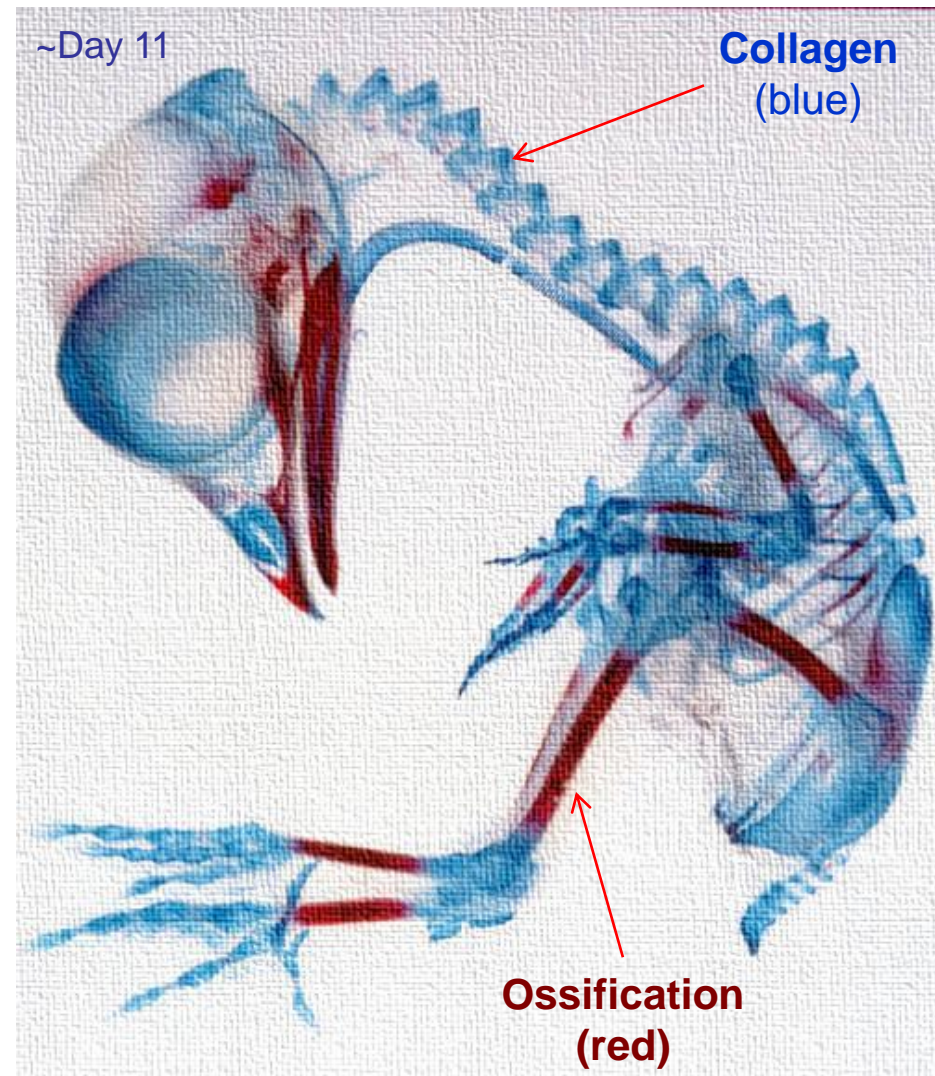
- **Cu- lysyl oxidase**

- Cu is involved in collagen cross linking
- Second most common metalloenzyme constituent (after zinc)

- **Mn - glycosyl transferases**

- Chondroitin sulfate is the structural component of cartilage and provides much of its resistance to compression

(Leeson and Summers, 2001; Hudson et al., 2004; Bellairs & Osmond, 2005; Dibner and Richards, 2006; Dibner et al., 2007; Ferket et al., 2009)

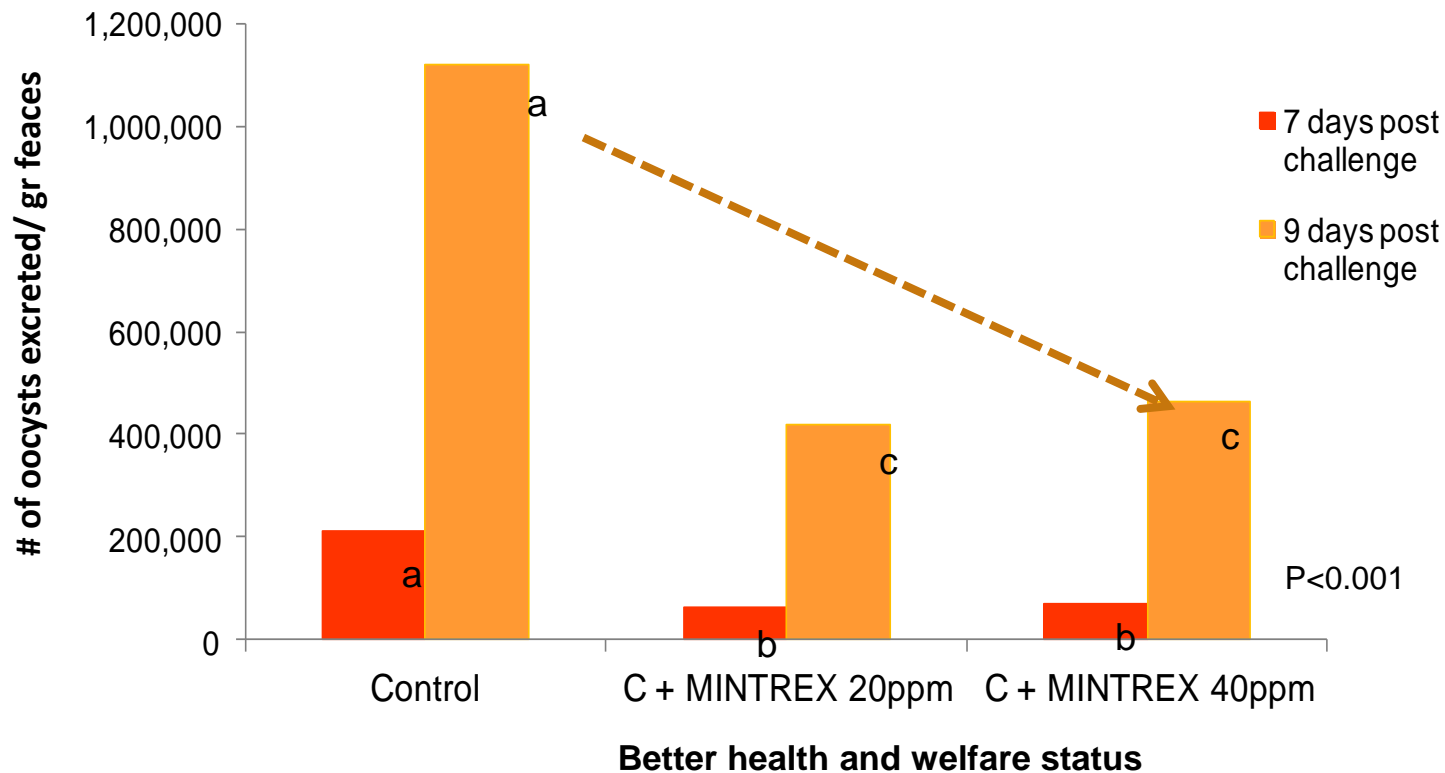


Immune function of trace minerals

- Zn:
 - Macrophages activity (Ferket et Qureshi, 1992)
 - Cofactor of DNA and RNA polymerase, it influences the number of leucocytes (Gupta et al., 1985)
- Cu:
 - Increases the immune response in an attack (Strain, 1994).
 - Ceruloplasmina: Cu is its cofactor. It protects the membranes from the free radicals of oxygen. The amount of this protein is increased during the attack of E.coli and Salmonella
- Mn:
 - It acts in the activity of neutrofiles and macrophages (Underwood y Suttle, 2001).

Trace mineral support in coccidiosis control

288 male broilers , supplemented with different levels of MINTREX[®] Zn and challenged with *E. tenella*.



Immunity and oxidative stress

**Maintaining
oxidative balance
=
stronger immunity**

Immunity

- Inflammatory process, stress hormones (adrenaline, cortisone):
➤ oxygen free radicals

Oxidative stress

- Susceptibility to disease
- Hyper activation of immune system
- Role in auto-immune diseases

Oxygen free radicals

- Damages to cell membranes

Some of the benefits attributed to Organic Trace Minerals

1. Catalyse Enzyme activity

- Hundreds of enzymes depend on Zn, Cu y Mn

2. Ensure tissue and structural health.

- Collagen, keratine, elastine and bone formation.
- Bone and feather structure, tendons, eggshell, gut and skin structure. wound healing. etc...

3. Optimize Immune function.

- Ig titters.
- B Cells, T Cells.

4. Reduce the impact of oxidative stress.

- Micro-minerals are cofactors of enzymes such as Superoxide dismutase and Gluthation peroxidase

5. Lower impact to the envirnment and safety of use

- Lower fecal excretion
- Dioxins, etc.

PARADIGM TO BREAK: MINTREX CAN HAVE A DIRECT IMPACT IN PRODUCTION PARAMETERS

- At practical levels of supplementation and by using commercially available sources of minerals, economically important parameters can be optimized:
 - Feed Conversion
 - Weight gain
 - Carcass yield
 - Chick quality
 - Egg production per housed hen,
 - Piglet weight at birth
 - ETC..
- AND, in spite of the economic importance of this topic, many nutritionists know relatively little about Zn, Cu, and Mn nutrition

Every Mineral Source Has a Unique Potential to optimize animal performance

- Inorganics

- Sulphates
- Oxides
- TBCC's
- ETC

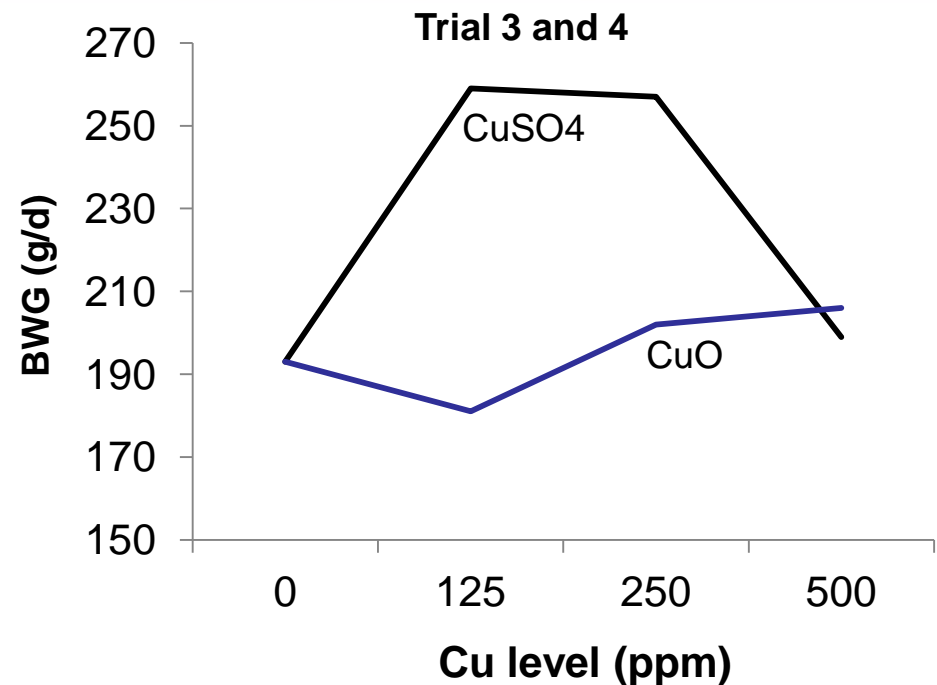
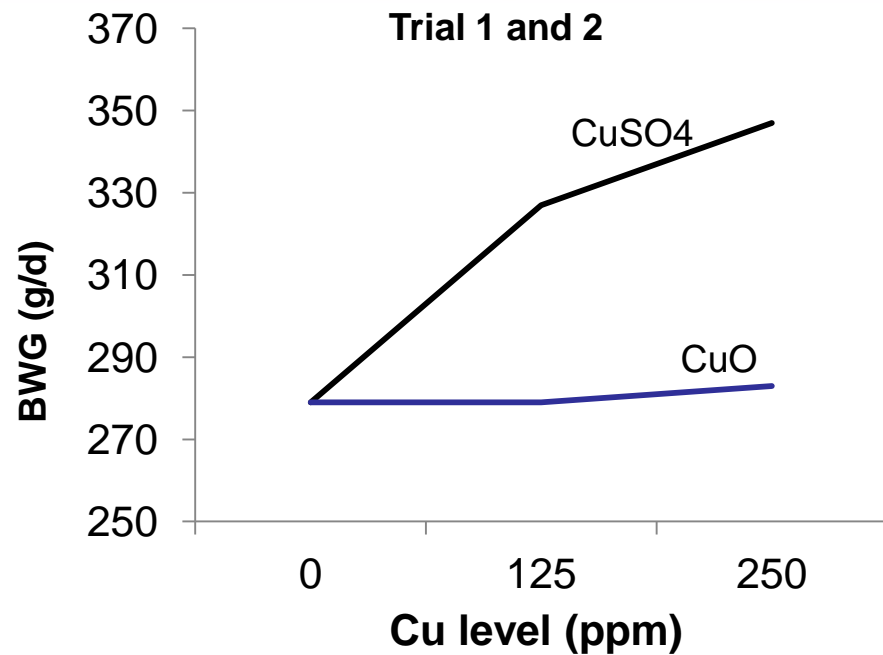
- Organics

- True Chelates (Mintrex, Glycinates)
- Non-Chelates (many)

Every source of mineral has its own unique characteristics and peculiarities

Their peculiarities can produce unique and distinctive animal responses

CuSO₄ is irreplaceable vs. CuO in improving growth rate and feed conversion ratio in swine



Copper improved piglet performance in its sulfate form. Copper when supplemented in its oxide form was ineffective, even at high concentrations.

This makes CuSO₄ an **IRREPLACEABLE** source of copper vs. CuO, to achieve optimal piglet performance

Important Questions to ask?

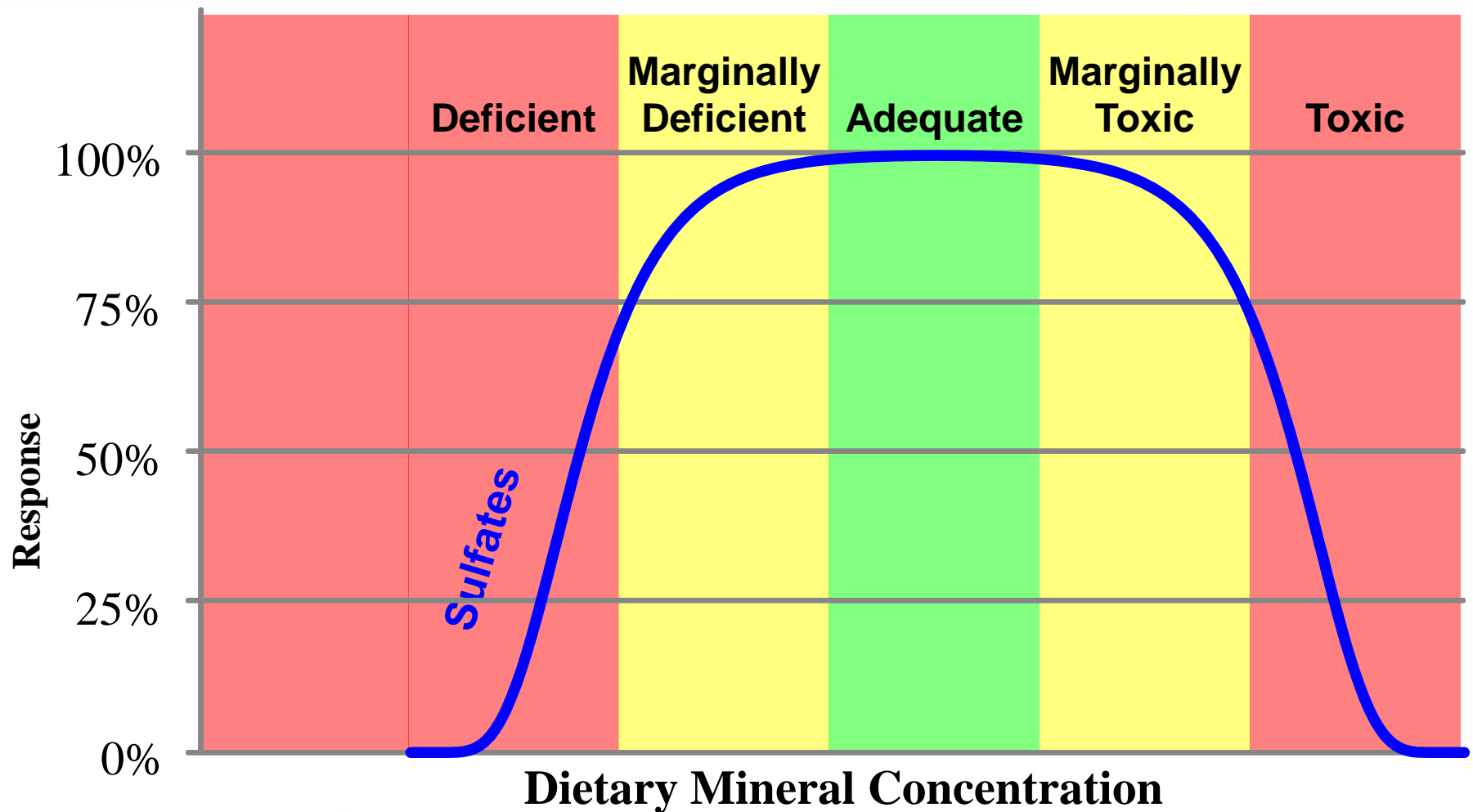
ARE ALL ORGANIC MINERALS IRREPLACEABLE
VS. INORGANIC MINERALS?

FOR PRODUCTION PARAMETERS?

Relevant Questions

DO ALL ORGANIC MINERALS SHOW THE SAME IMPROVEMENTS ON THE SAME PRODUCTION PARAMETERS?

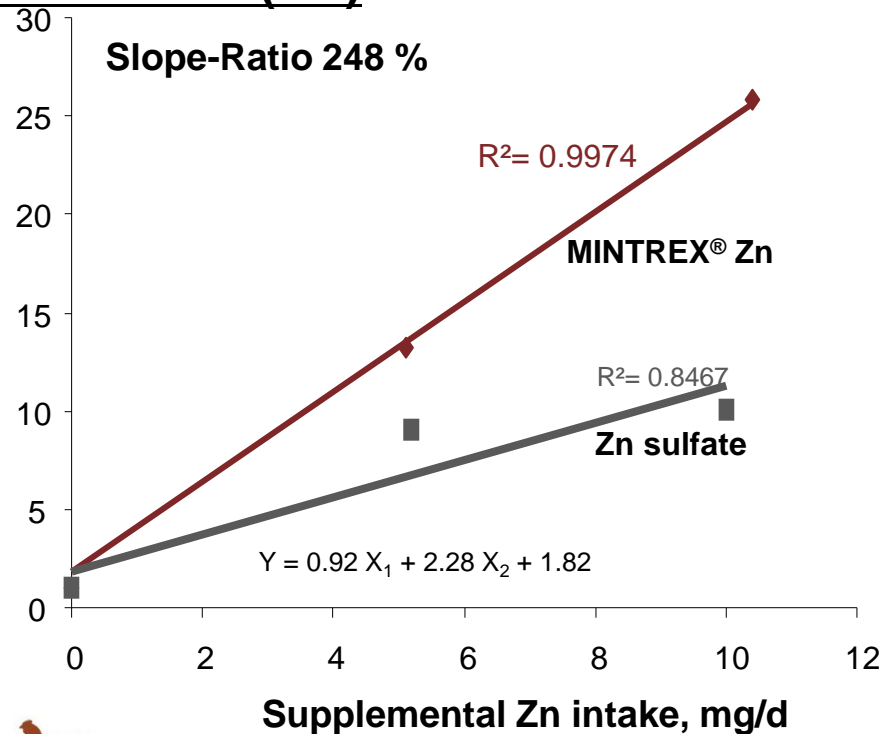
Dose-response of animals fed graded mineral concentrations



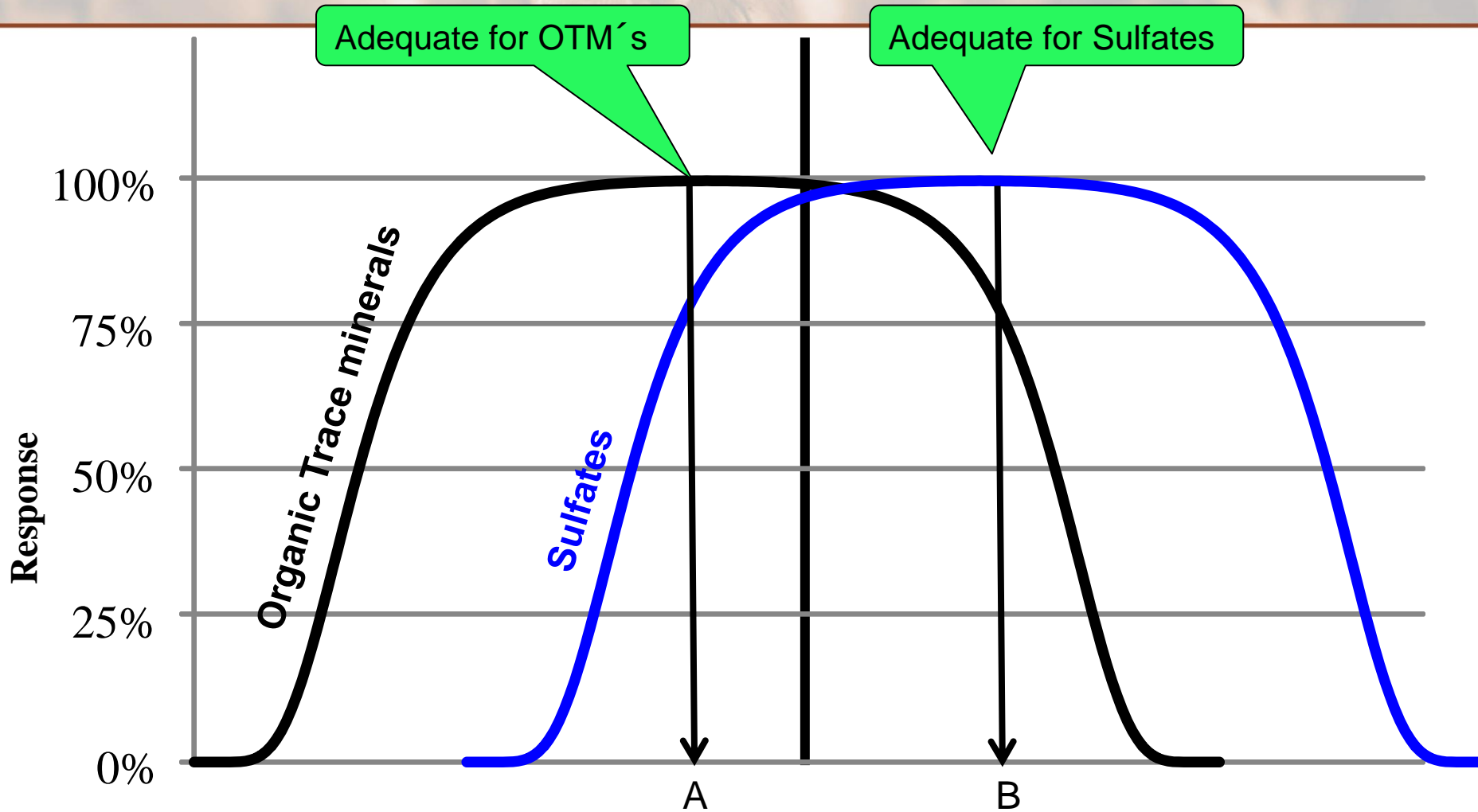
MINTREX[®] is more bioavailable than other mineral sources.. BUT, What does that really mean?

- Metallothionein (MT) mRNA expression is a marker for Zn absorption/status in multiple species
- Mintrex Zn is 248 % more bioavailable compared to ZnSO₄

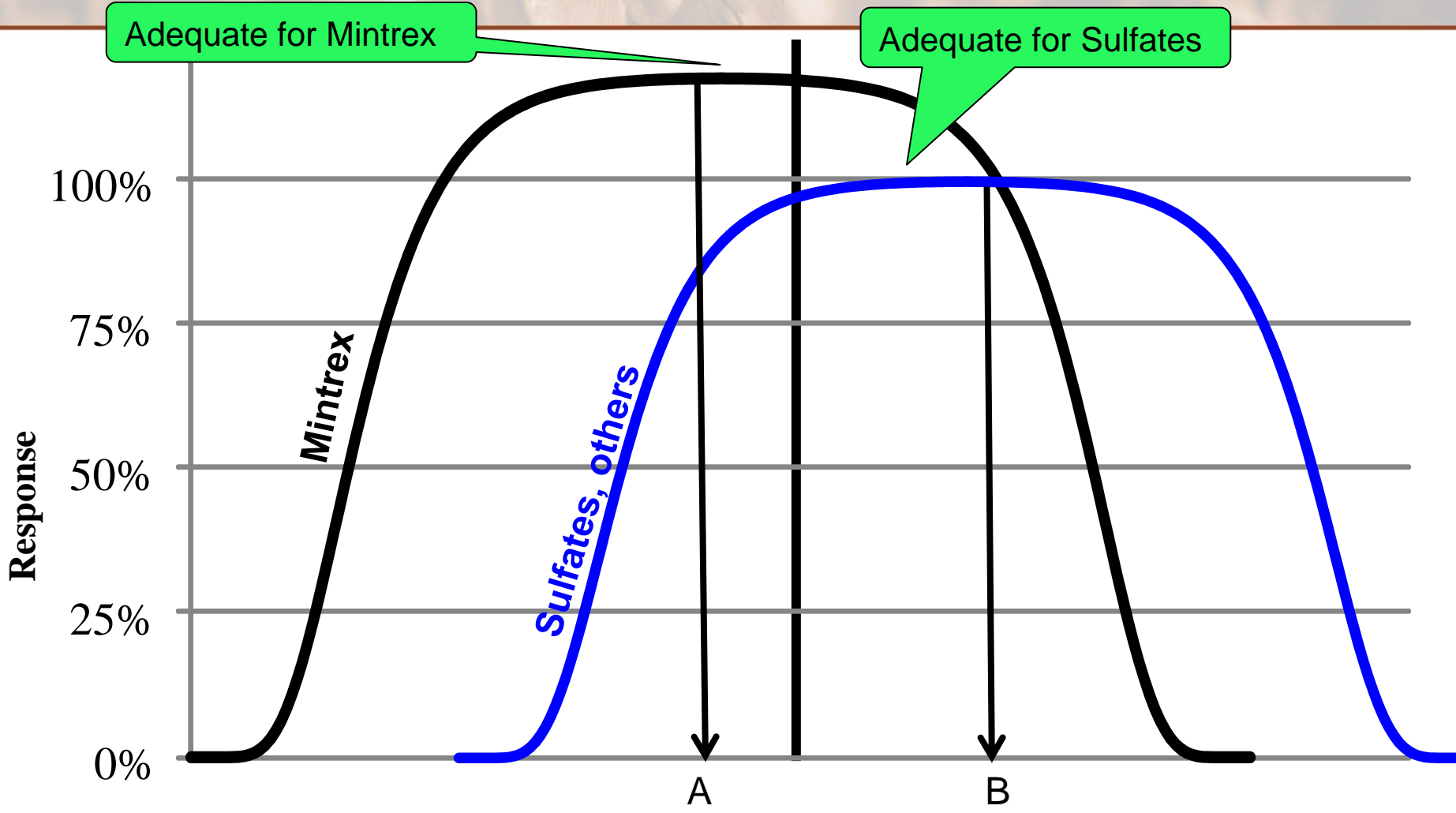
b) Metalothionein (MT)



Dose-response of OTM vs. ITM

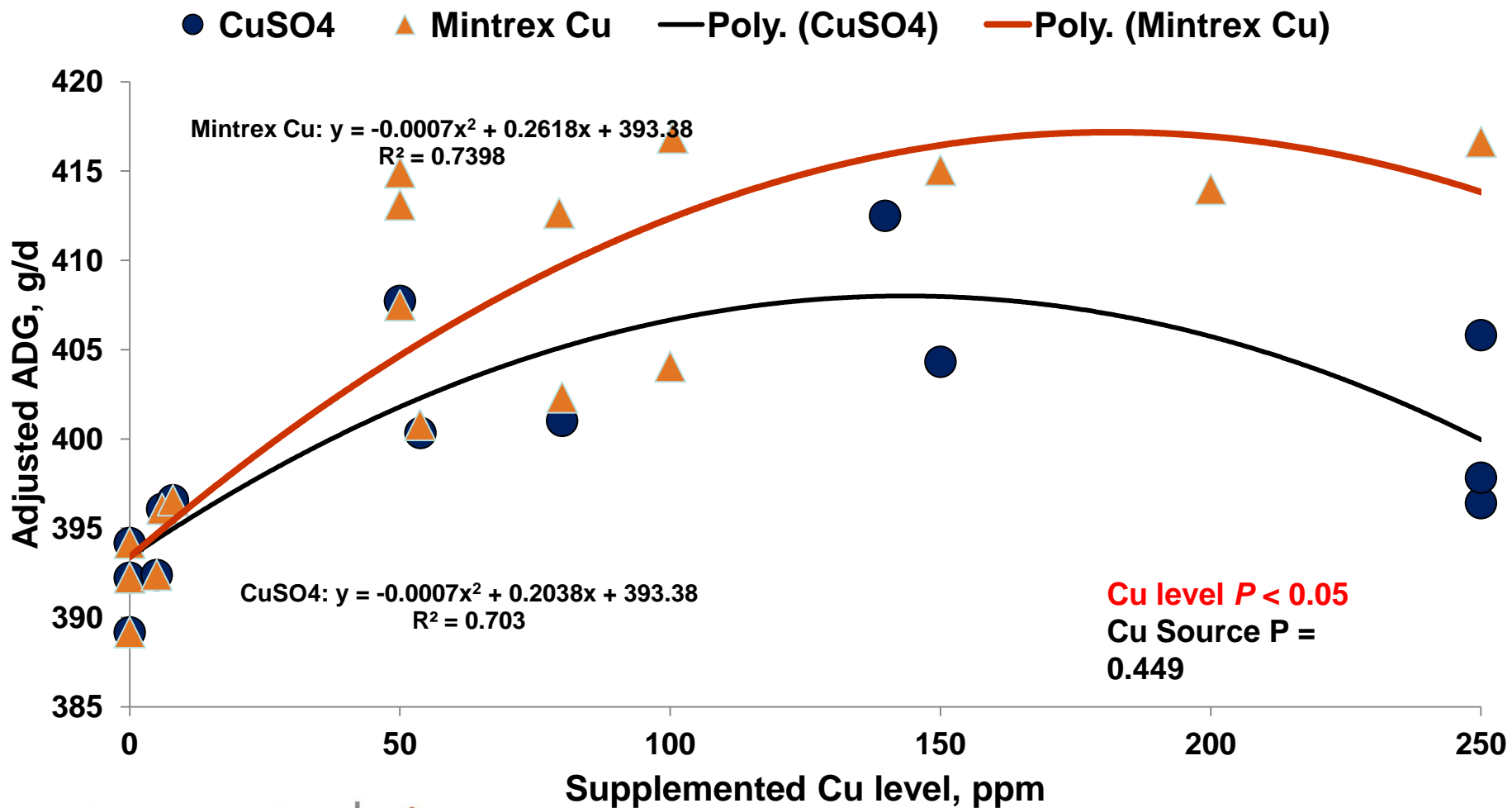


Mintrex vs Other Mineral Sources



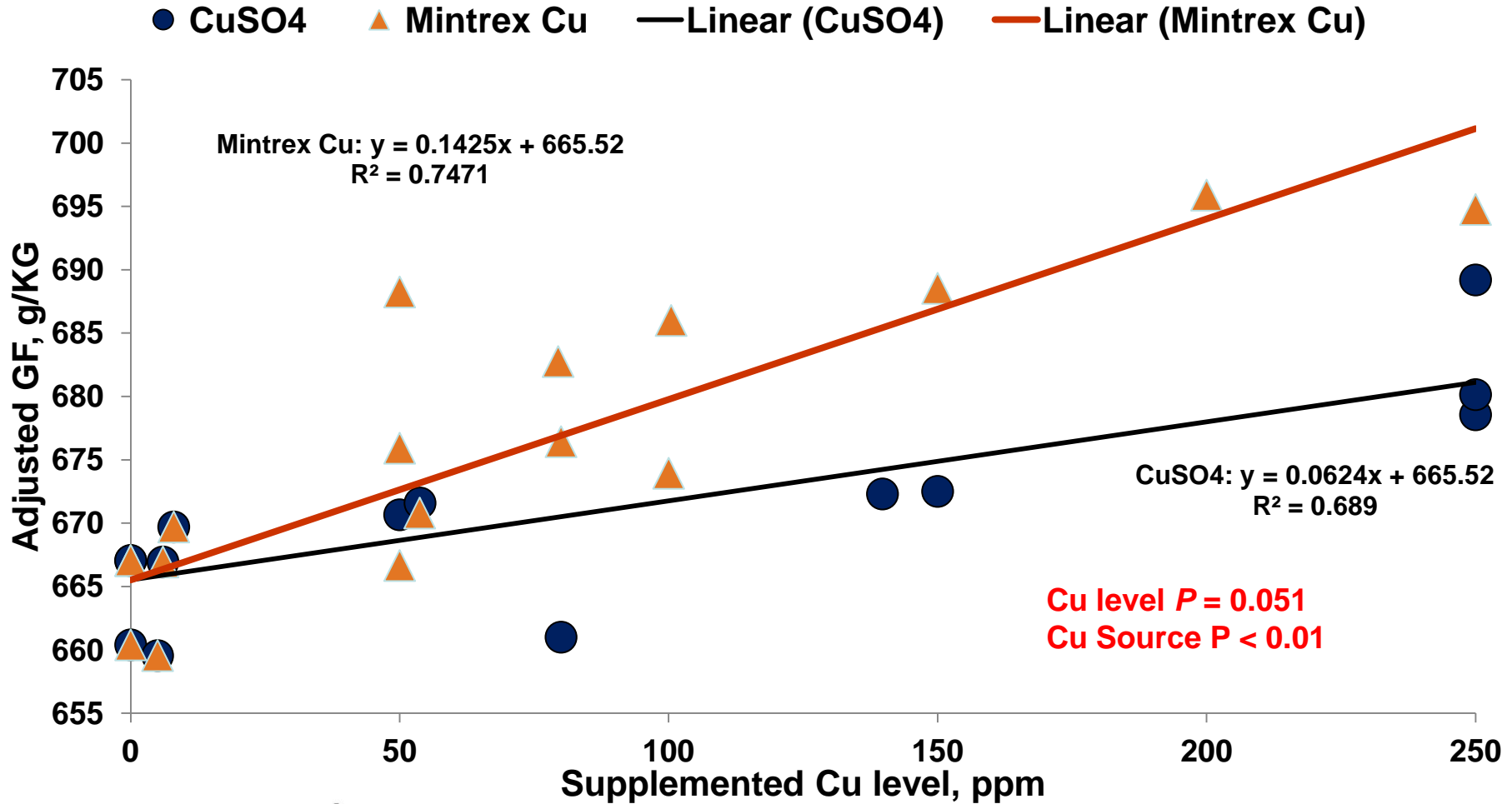
Effect of various levels of Mintrex Cu and CuSO₄ on overall nursery ADG (0~42 d)

(Data of 6 trials)



Effect of various levels of Mintrex Cu and CuSO₄ on overall nursery G:F (0~42 d)

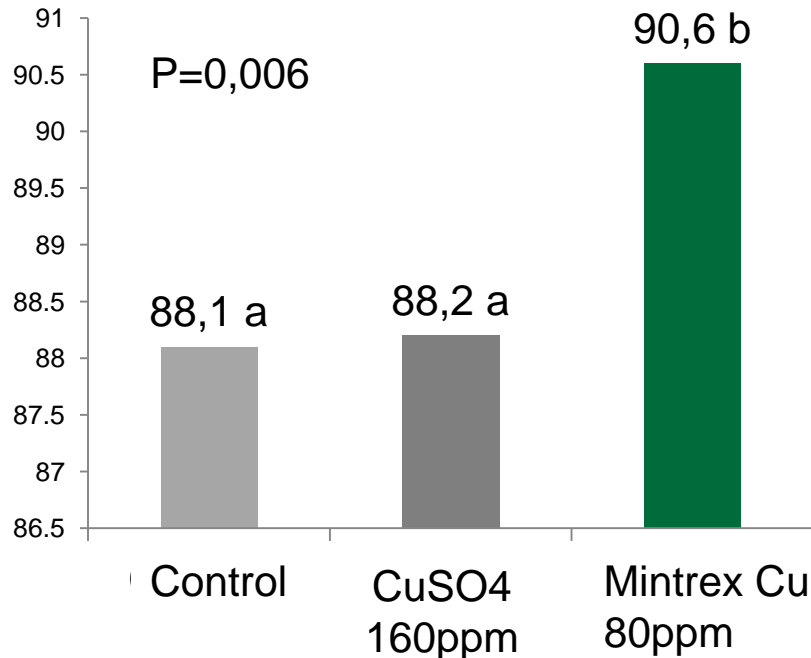
(Data of 6 trials)



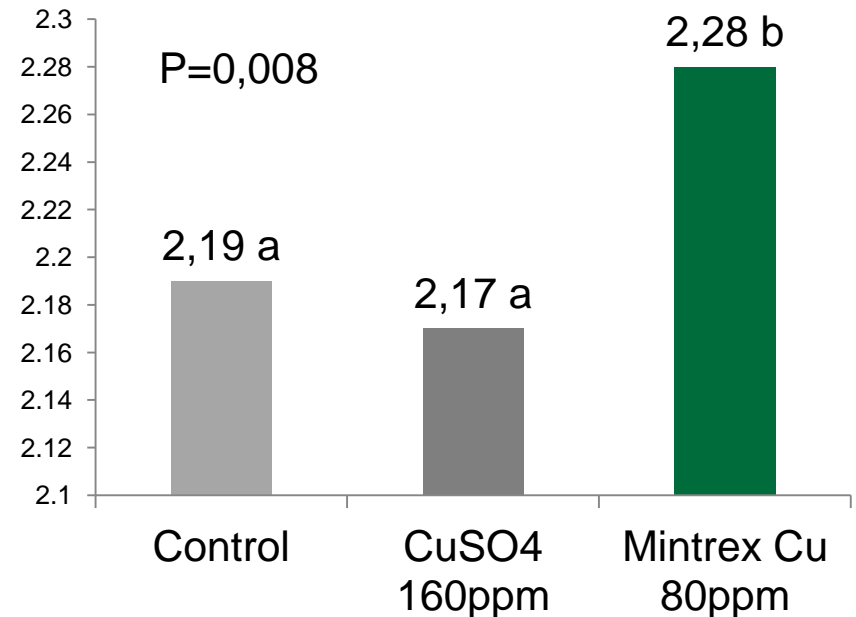
80 ppm of MINTREX Cu Optimizes Carcass Parameters in swine

Initial weight 32 kg / Final weight 120 kg

Carcass weight (kg)

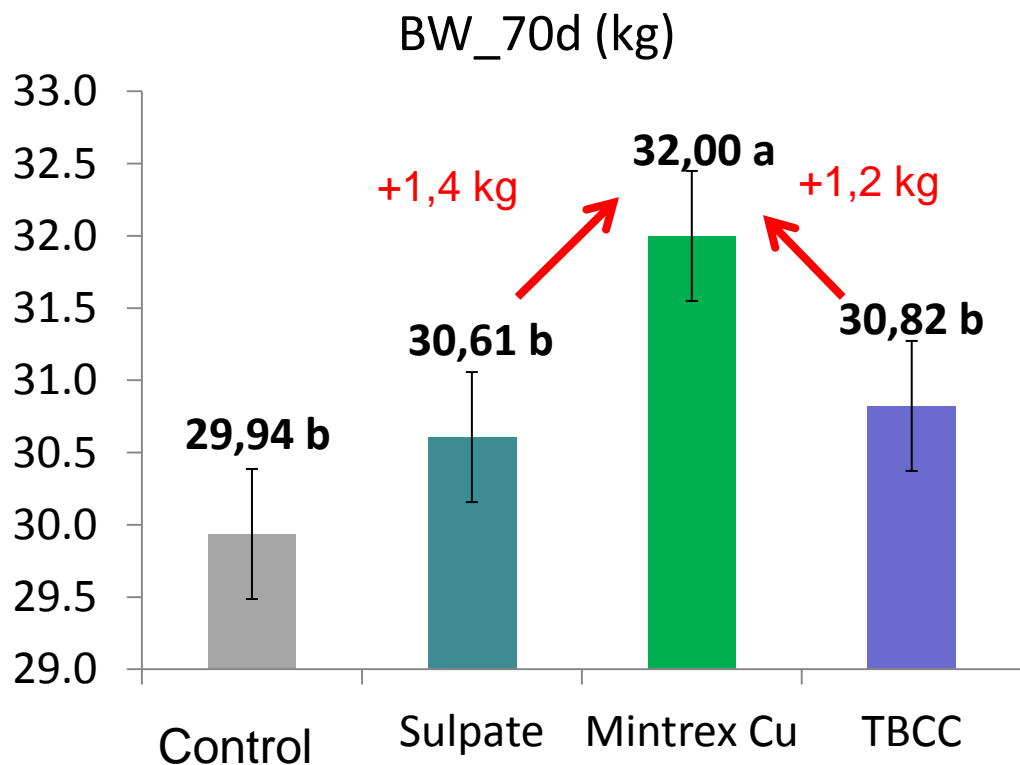


Loin depth (pol.)



	Control	CuSO4 160ppm	Mintrex Cu 80ppm
U\$/swine	118,7 a	119,4 a	122,8 b

A body weight comparison between 3 sources of Cu included in swine diets

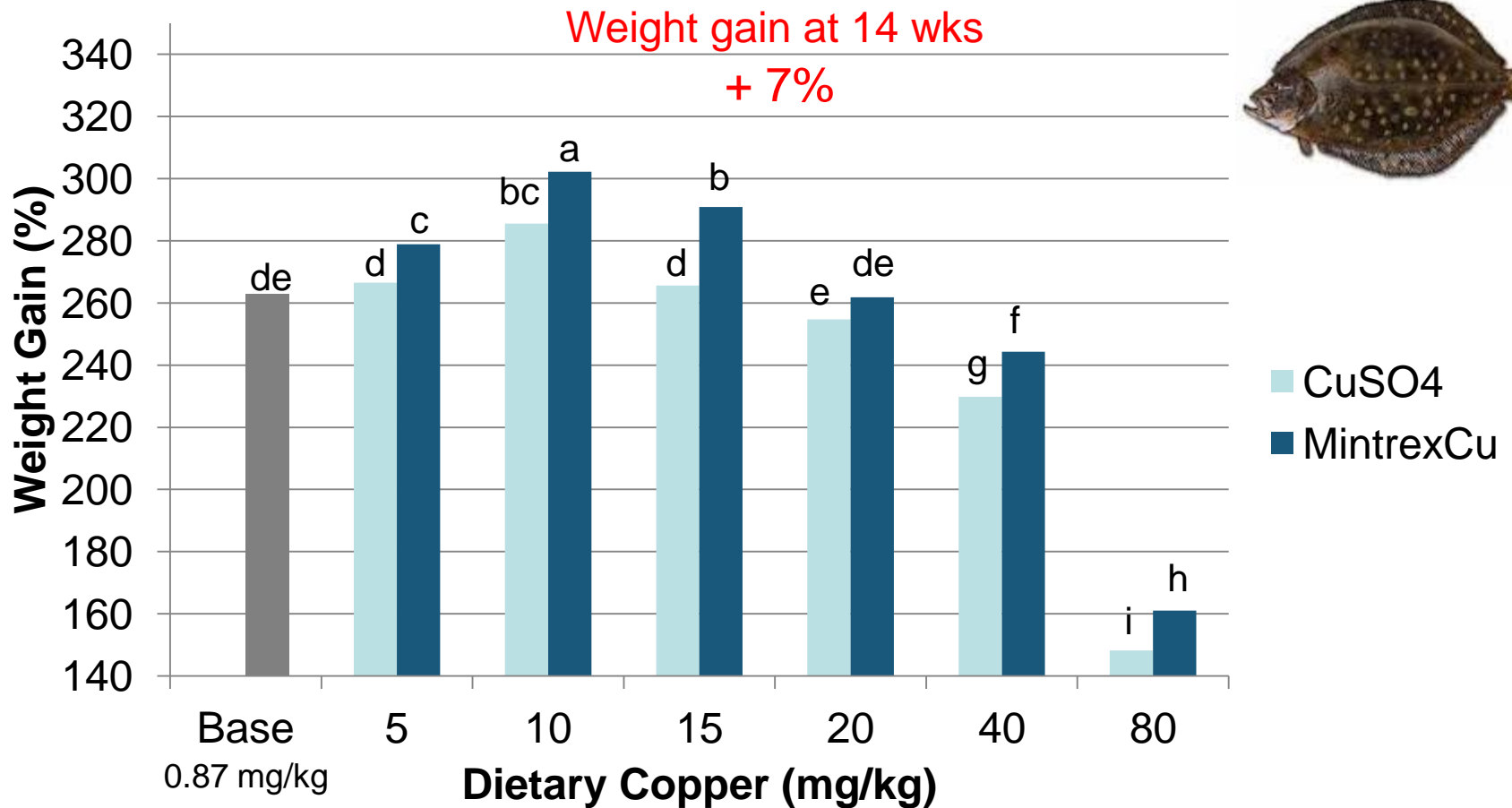


Orthogonal contrasts	
Control x All	0,0142
Mintrex x (Sulphate +TBCC)	0,0139
Sulphate x TBCC	0,7119

P= 0.0084
CV = 9.77%



In a trial conducted with Olive Flounder- Mintrex Cu improved weight gain when compared to CuSO₄



The distinctive dose-response curve of any mineral source may depend on a complex combination of unique characteristics such as

- Bio efficacy
- Site of absorption
- Solubility
- Interaction with other components in the diet
- Oxidative balance
- Antimicrobial effect
- ???
- Etc., etc.,

To illustrate how poorly we understand this interaction, try to explain why ZnO at 2,000 ppm benefits piglets that much more compared to ZnSO₄ or OTM.. It's certainly not due to higher bio efficacy...

Take home message

1. Minerals have unique and distinctive effects on animal performance
2. The performance potential of an animal may be determined by the source of mineral it is fed
3. Micro-mineral nutrition may impact economically important performance parameters and therefore deserve more attention