Impact of Interventions during the Neonatal development of Piglets

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Strategies to reduce antibiotic usages

"All Disease begins in the Gut"
Hippocrates ca. 460 – ca. 370 BC

Antibiotics are frequently used to treat intestinal problems.
Figure 3.1. Veterinary therapeutic autibiotic sales from 1999-2009 (FIDIX, 2010; vertical bars). The presents the trends in grams of active ingredients used per kg live weight.



Influence intestinal health without usage of antibiotics?





Gut is the gatekeeper of health



Earlier results intestinal development, immune and microbiome

- Critical transition moments for microbiota development
 - Prenatal
 - Birth
 - Weaning
- Interventions
 - Feed(-additives)
 - Management
- Effects
 - Intestinal immune development
 - Long lasting
 - Due to early colonisation





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Early-Life Environmental Variation Affects Intestinal Microbiota and Immune Development in New-Born Piglets

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Long-Lasting Effects of Early-Life Antibiotic Treatment and Routine Animal Handling on Gut Microbiota Composition and Immune System in Pigs

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Plasticity of intestinal gene expression profile signatures reflected by nutritional interventions in piglets

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Hypothesis

Long-lasting effects of intestinal health are due to different programming of the gut immune system early in life







Experimental set-up



- GOAL: To determine the effect of dietary intervention on intestinal development in piglets
- Two different feeding strategies to apply intervention:
 - Maternal administration
 - Neonatal administration by oral gavage







interventions



- Medium Chain Fatty acids: Probably affects microbiota composition proximal intestine
- Beta-glucan: Immunomodulatory effects
- <u>Galacto-oligosaccharides</u>: beneficial effects on humans → bifidogenic effect colon
- Control





Microbiota composition depends on administration route







- Small differences in microbiota
- Microbiota composition depends on administration route
- ~ 10 differences on family level between treatment and control
 - maternal_d.1_Jejunum_CONTROL maternal_d.1_Jejunum_MCFA maternal_d.31_Jejunum_CONTROL maternal_d.31_Jejunum_MCFA neonatal_d.1_Jejunum_CONTROL neonatal_d.31_Jejunum_CONTROL neonatal_d.31_Jejunum_MCFA

Effect on microbiota

Number of statistical changes in microbiota composition on family level

Intervention	Day	Number of statistically significant differences compared to control
1	1	45
	31	19
2	1	2
	31	0
3	1	5
	31	0

- MCFA induce changes in composition
 - Strongest effect on day 1, early life colonization
 - Suggestive for transmission effect via other routes than milk





Intestinal development depends on administration route



Maternal MCFA	%	Neonatal MCFA	
Lipid digestion, mobilization and transport	7/71 (10%)	Regulation of complement cascade	4/26 (15%)
Chylomicron-mediated lipid transport	4/17 (23%)	Immune system	37/1547 (2%)
Retinoid metabolism and transport	5/42 (12%)	Chemokine receptors bind chemokines	5/56 (9%)
Metabolism of fat soluble vitamins	5/51 (10%)	Interferon alpha/beta signalling	5/68 (7%)
Reversible hydration of carbon oxide	3/12 (25%)	Metabolism of AA and derivatives	12/335 (4%)
Lipoprotein metabolism	4/34	IRN3 mediated activation of type 1 IFN	2/6 (33%)
Chemokine receptors bind chemokines	4/56 (7%)	Metabolism	42/1908 (2%)
Digestion of dietary carbohydrate	2/8 (25%)	Complement cascade	5/80 (6%)
Erythrocytes take up $\rm O_2$ and release $\rm CO_2$	2/8 (25%)	ZBP1 mediated induction type I IFNs	3/26 (12%)
Visual phototransduction	5/97 (5%)	Innate immune system	21/807 (3%)
WAGENINGEN			



Conclusion



- Administration route determines the outcome of intestinal development and on microbiota composition
- Large differences in intestinal development of new-born piglets between the 3 maternal dietary interventions
- Neonatal administration of MCFAs induces immunological changes







Overall conclusions



Feed in early life has consequences for the programming of the host immune system
 Administration timing of importance for result
 Hereby it is possible to change gut health



And therefore this is an antibiotic reduction strategy.





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