

Dietary resistant potato starch alters immunological status and microbial populations in swine to limit *Salmonella*

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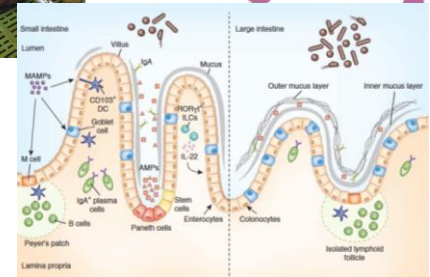
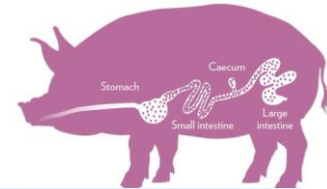
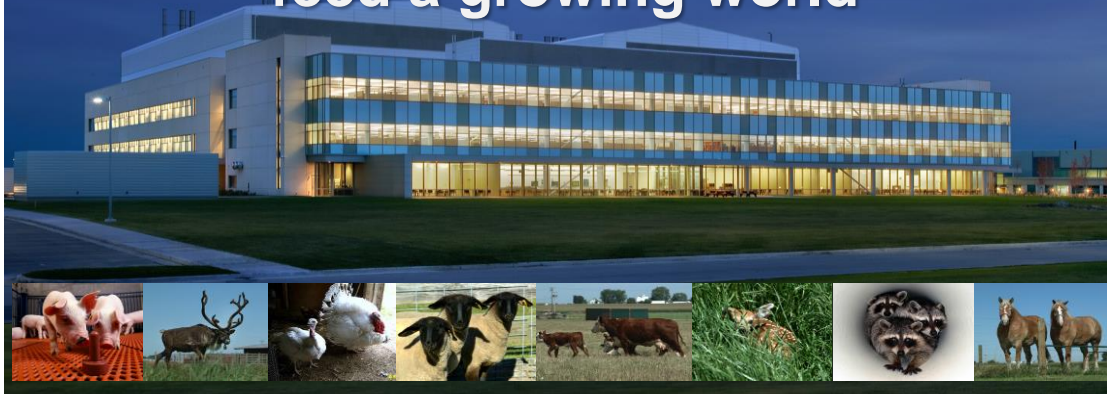
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Conduct basic and applied
research to improve livestock
health and enhance food safety to
feed a growing world



Nature Immunology, 14 (7) July 2013

What is purpose of alterbiotic administration?

*Alterbiotic – bio-approach to modulate intestinal status

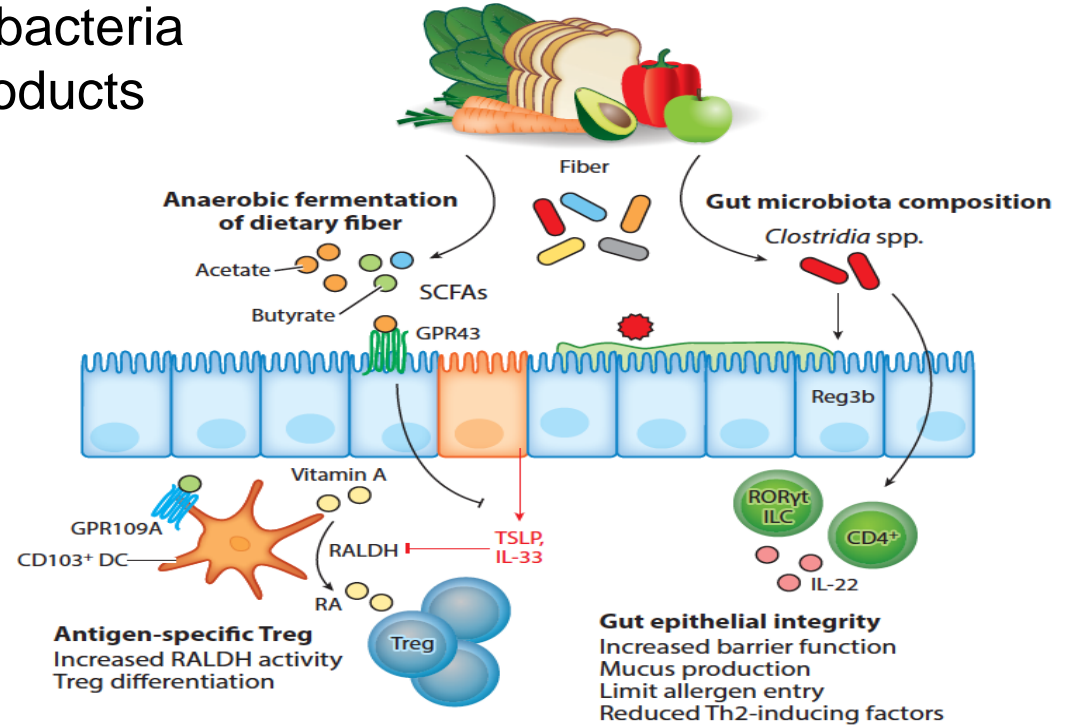
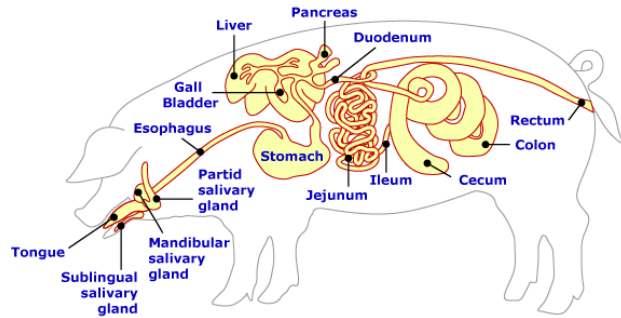
- Promote growth (nutrient uptake)?
 - Modulate immune status?
 - Enhance intestinal integrity?
 - Modulate microbial composition or function?
 - Prevent or limit infection/colonization?
-
- Determining the mode of action of various alterbiotics will allow for targeted applications in production.



Fermentation products can alter immune cellular responses and status

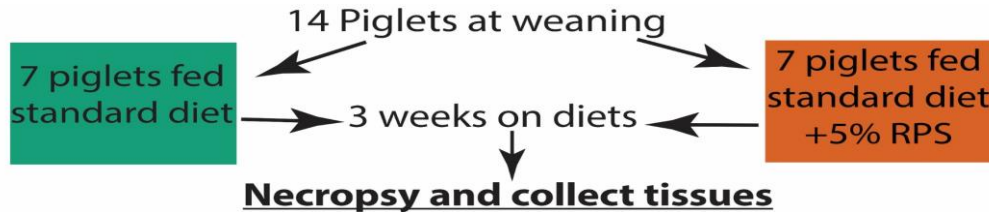
Enhanced growth of beneficial bacteria producing micronutrient end products (eg., short chain fatty acids):

- Butyrate, etc



Dietary resistant starch and immune status

- Does addition of 5% resistant starch (raw potato starch, RPS) to swine diet modulate:
 - Intestinal microbial populations & short-chain fatty acid levels?
 - Intestinal immune status?
 - T-regulatory cell populations? IgA-secreting cells?
 - Local defense proteins or cytokines?

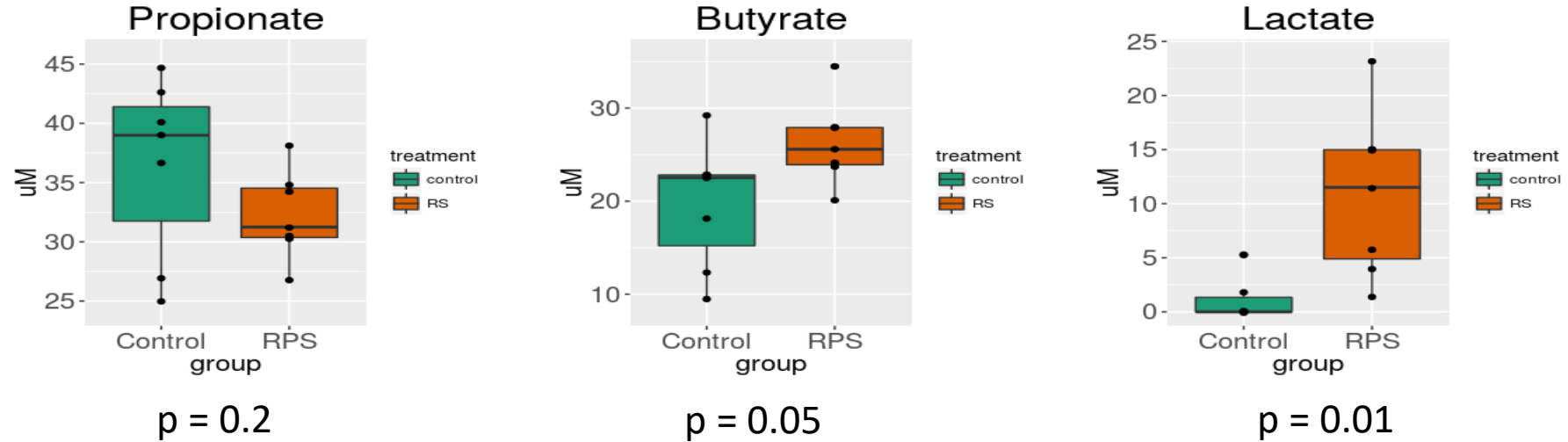


Dietary Resistant Potato Starch Alters Intestinal Microbial Communities and Their Metabolites, and Markers of Immune Regulation and Barrier Function in Swine

Julian Trachsel^{1,2}, Cassidy Briggs^{1,2}, Nicholas K. Gabler⁴, Heather K. Allen^{1*} and Crystal L. Loving^{1*}

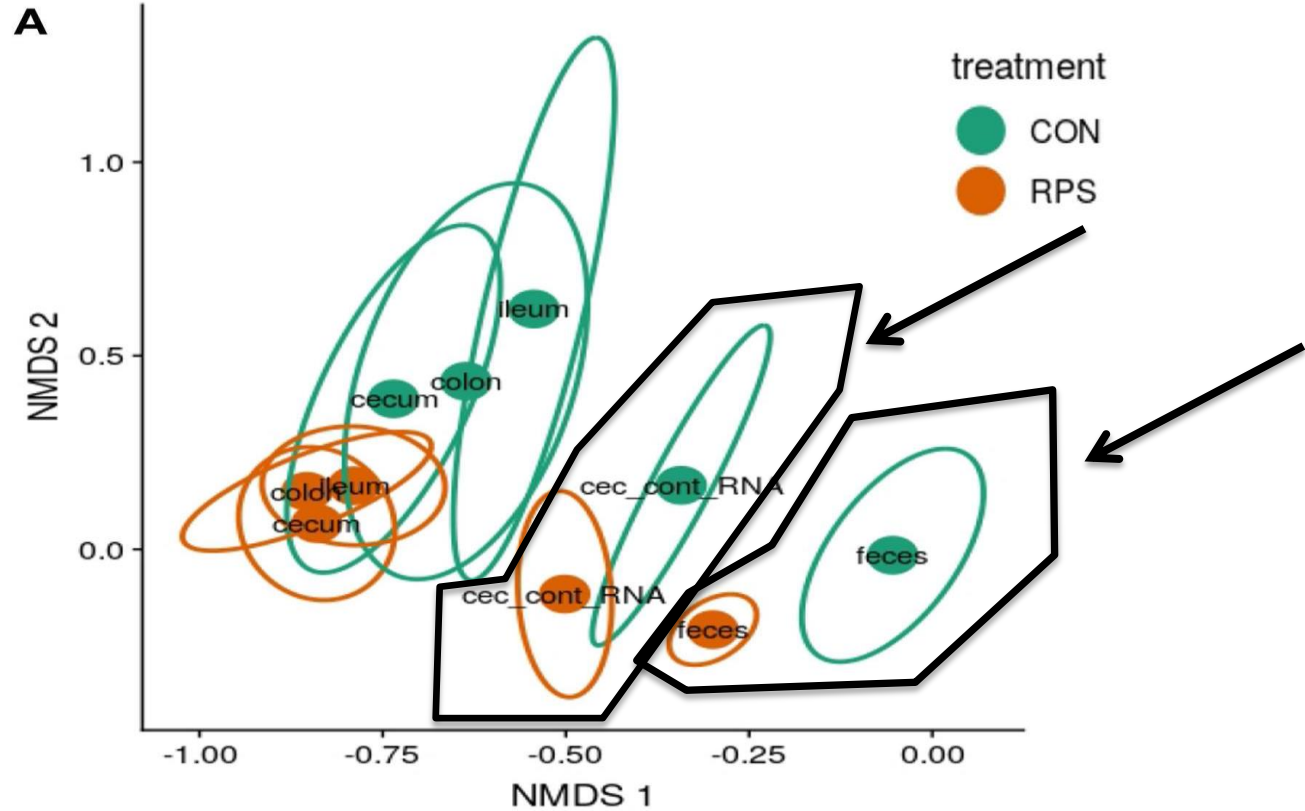
¹ Food Safety and Enteric Pathogens Research Unit, National Animal Disease Center, Agricultural Research Service, Ames, IA, United States, ² Interdepartmental Microbiology Graduate Program, Iowa State University, Ames, IA, United States, ³ Summer Scholar Research Program, College of Veterinary Medicine, Iowa State University, Ames, IA, United States, ⁴ Department of Animal Science, Iowa State University, Ames, IA, United States

Dietary resistant potato starch (RS) increased cecal SCFA levels

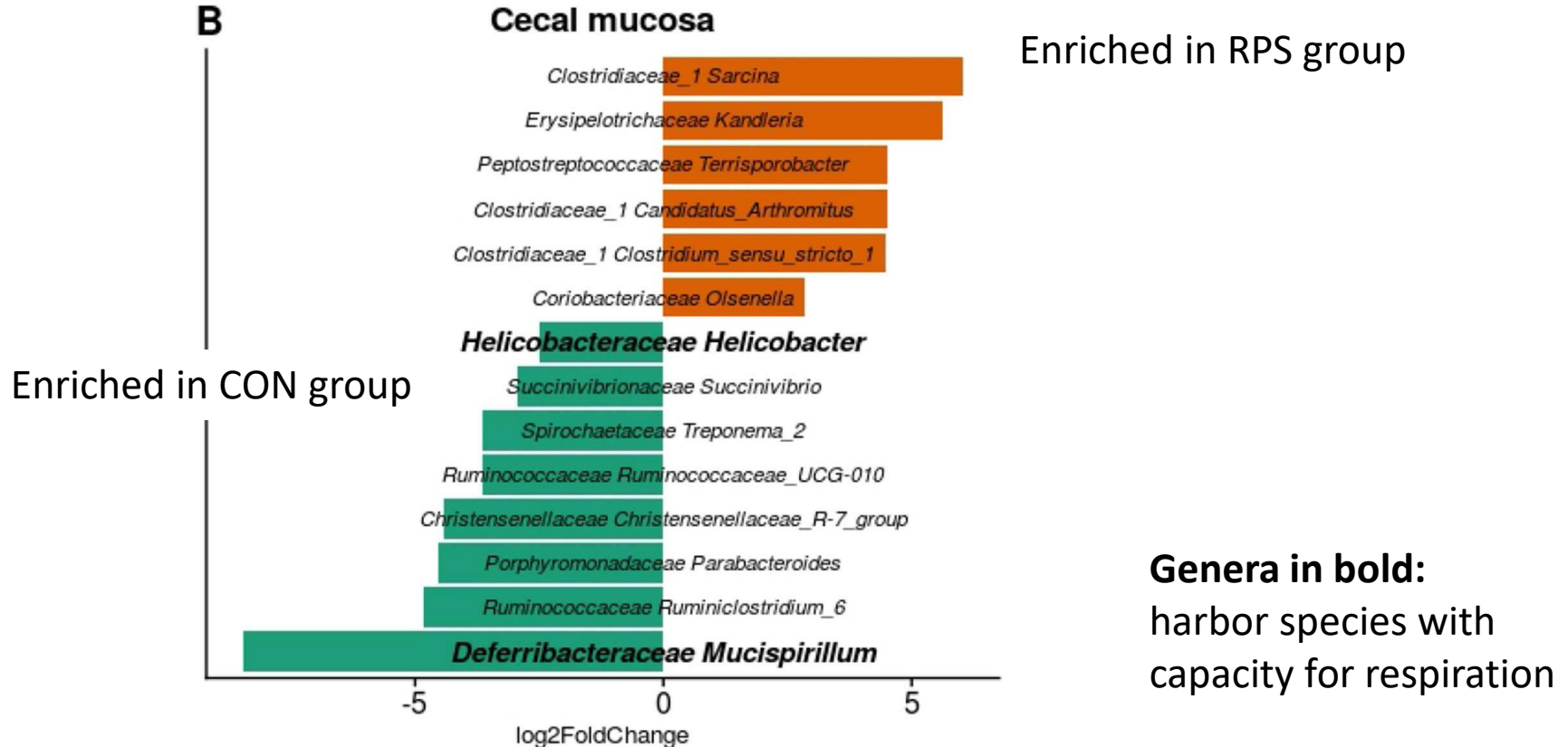


- Increased butyrate and lactate in cecal contents of pigs fed RPS (raw potato starch)

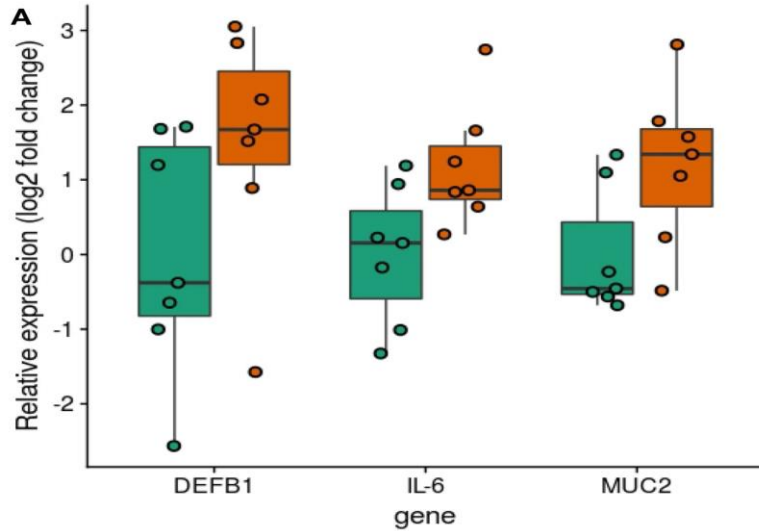
Dietary RPS modulated microbial community structure



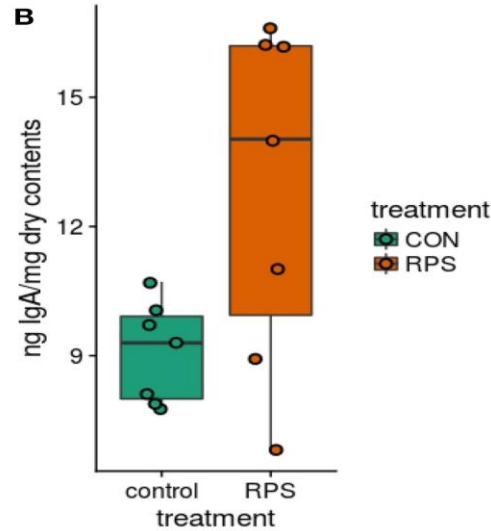
Microbial communities altered by dietary RS



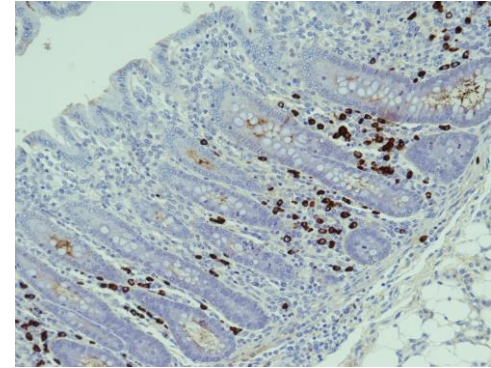
Dietary RPS altered mucosal immune status



Increased cecal expression of genes associated with epithelial integrity

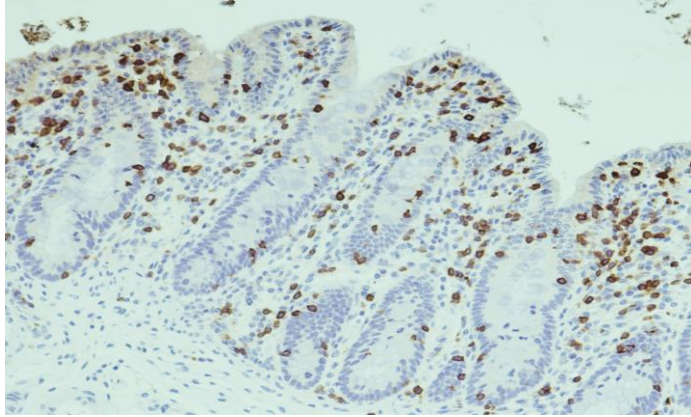


Increased levels of total IgA in feces, but no increase in IgA+ cells in cecum

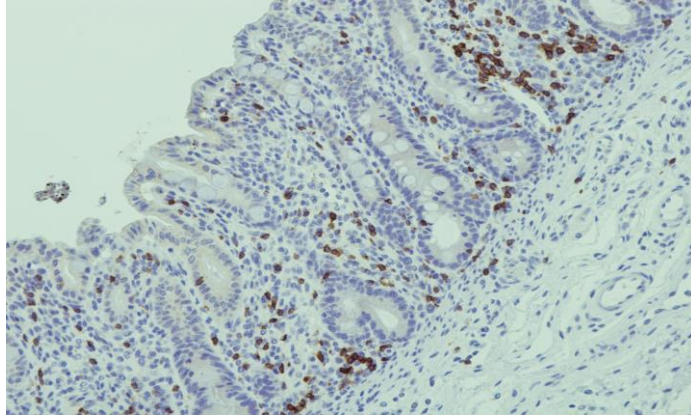


IgA staining - Cecum

RPS did not alter abundance of cecal T cells



Control Pig: CD3+ cells

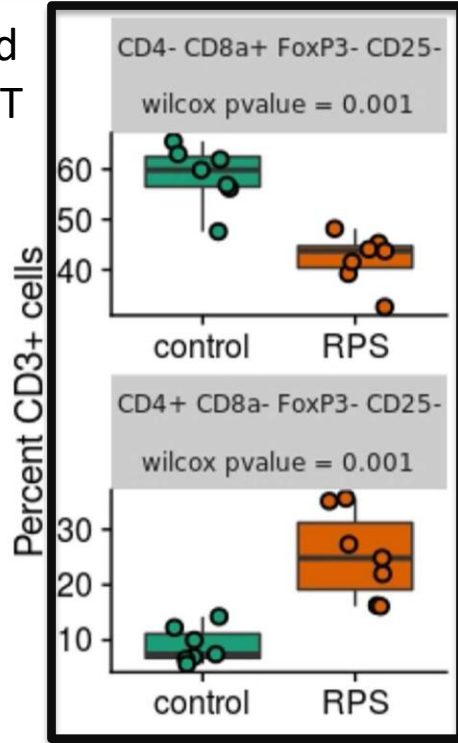


RS Pig: CD3+ cells

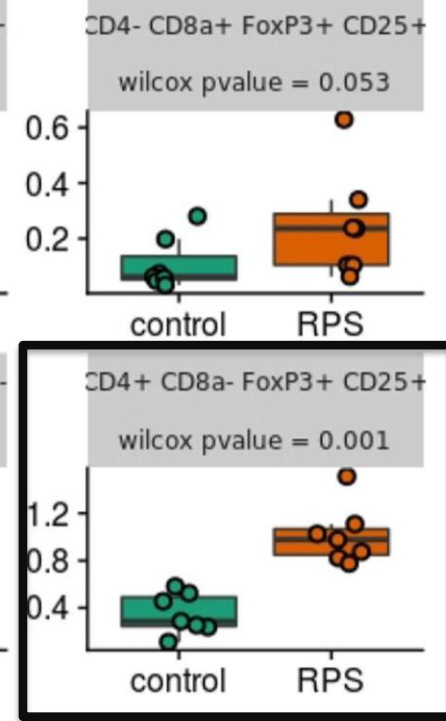
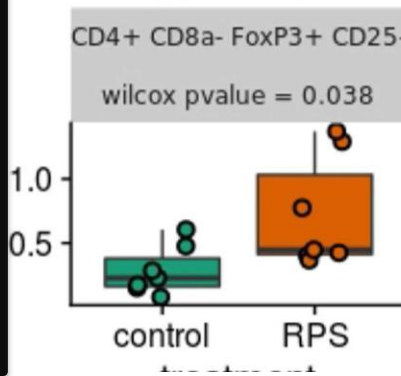
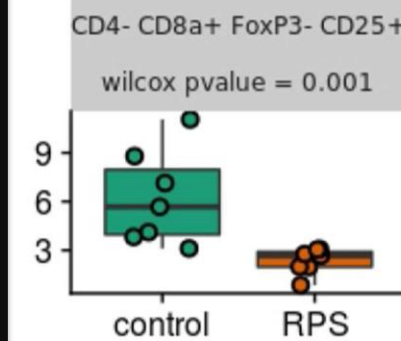
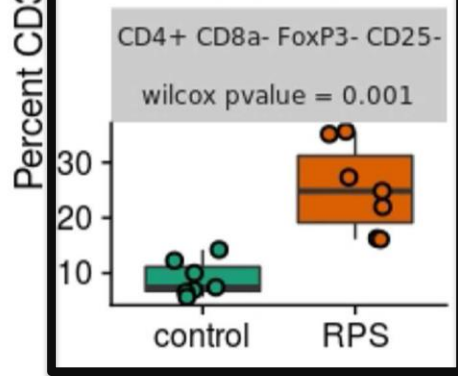
- No overall difference in #CD3+ T cells between groups

Dietary RPS altered cecal Treg cell frequency

Decreased
cytotoxic T
cells



Increased
CD4+



treatment
CON
RPS

Increased T regs

No differences in total CD3+ cells – differences in phenotype of T cells present

Dietary RPS as an alterbiotic

*Alterbiotic – bio-approach to modulate intestinal status

- Promote growth (nutrient uptake)?
- ✓ **Modulate immune status**
- ✓ **Enhance intestinal integrity**
- ✓ **Modulate microbial composition or function**
- Prevent or limit infection/colonization?
- Determining the mode of action of various alterbiotics will allow for targeted applications in production.



Dietary RPS as an alterbiotic

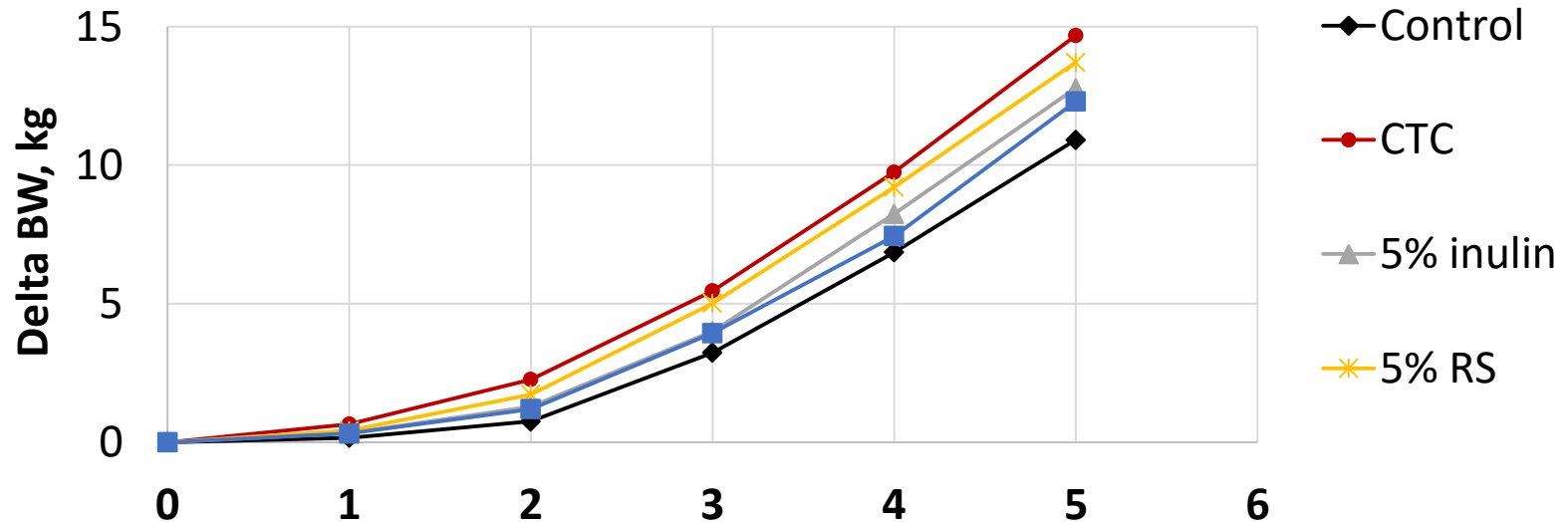
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Dietary RPS can alter production parameters

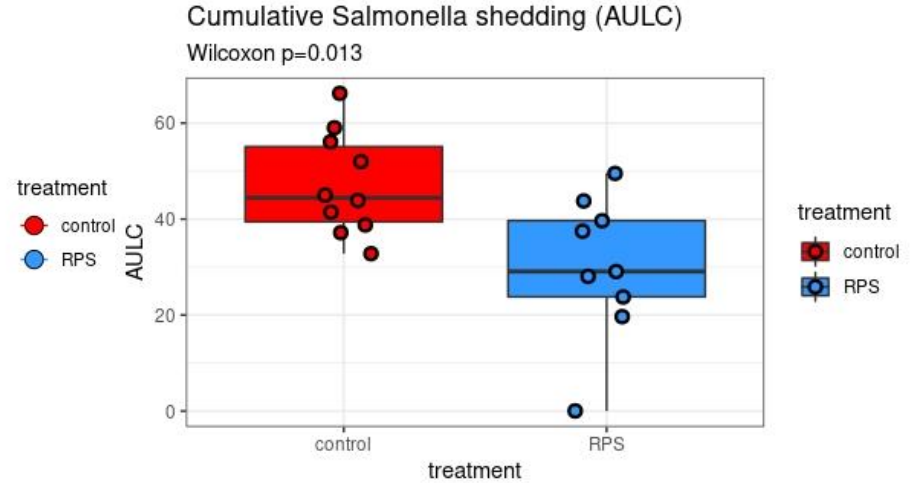
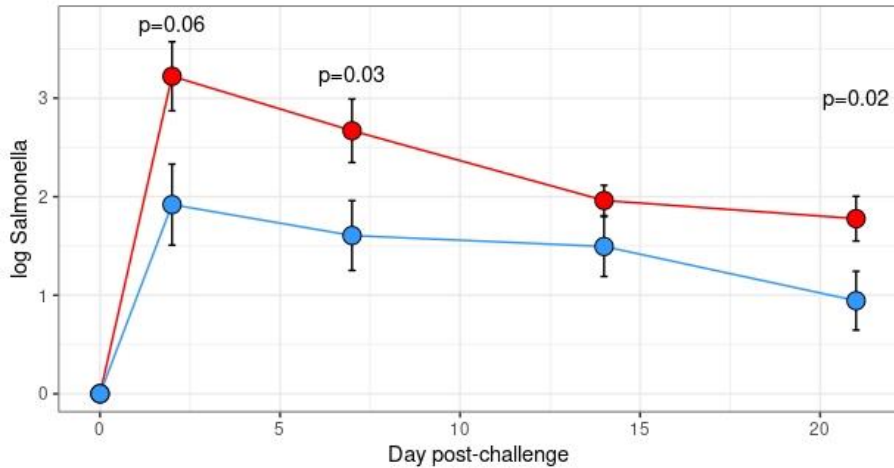
Diet	ADG, kg	ADFI, kg	G:F
Control	0.32 ^a	0.37 ^a	0.88
CTC	0.43 ^b	0.51 ^b	0.85
5% inulin	0.37 ^{ab}	0.44 ^{ab}	0.87
5% Resistant Starch (RS)	0.40 ^b	0.46 ^{ab}	0.87
0.25% Butyrate	0.36 ^{ab}	0.42 ^{ab}	0.88



Dietary RPS and *Salmonella* shedding

- Nursery pigs fed 5% RPS for 4 weeks
- Oral inoculation with monophasic *Salmonella*
 - Maintained on 5% RPS diet for additional 21 days
 - Fecal shedding overtime
 - SCFA analysis

Dietary RPS reduced *Salmonella* shedding



Proposed MOA – limit low level inflammation

Cell Host & Microbe

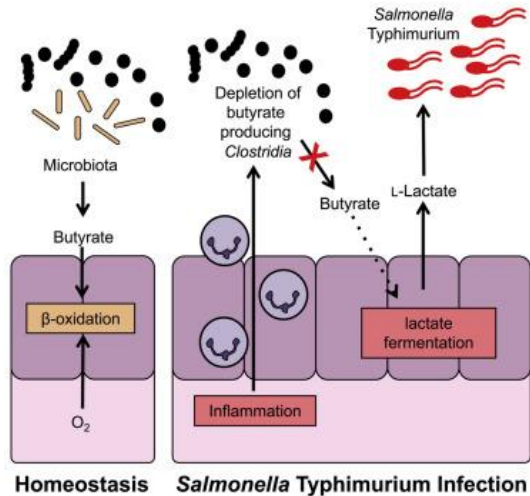
Volume 23, Issue 1, 10 January 2018, Pages 54-64.e6



Article

Dysbiosis-Associated Change in Host Metabolism Generates Lactate to Support *Salmonella* Growth

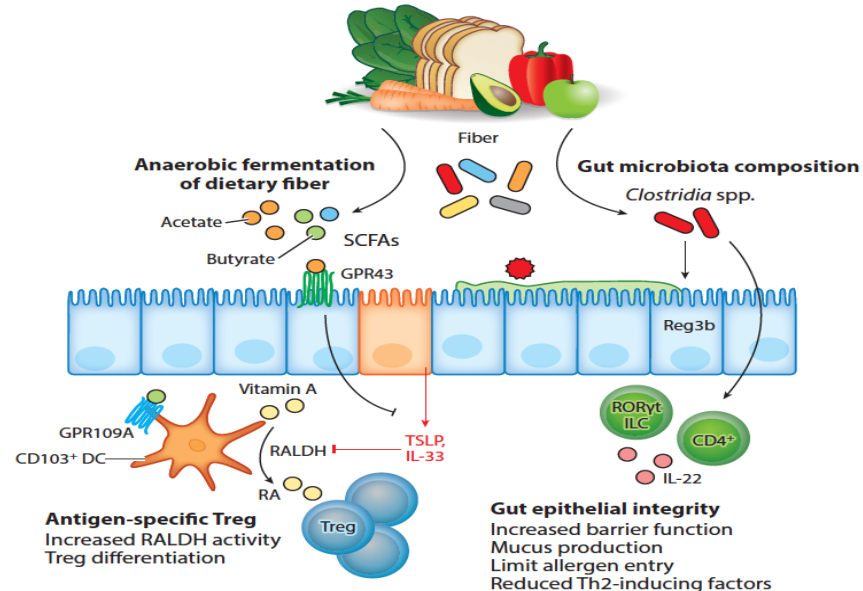
Caroline C. Gillis¹, Elizabeth R. Hughes¹, Luisella Spiga¹, Maria G. Winter¹, Wenhan Zhu¹, Tatiane Furtado de Carvalho², Rachael B. Chanin¹, Cassie L. Behrendt³, Lora V. Hooper^{3,4}, Renato L. Santos², Sebastian E. Winter^{1,5,*}



Reduced inflammation

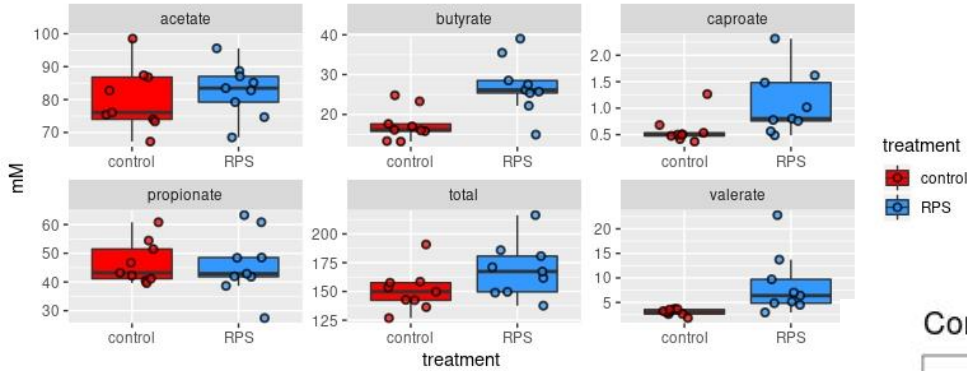
O₂ consumption by epithelial cells = anaerobic environment

Increased gut integrity



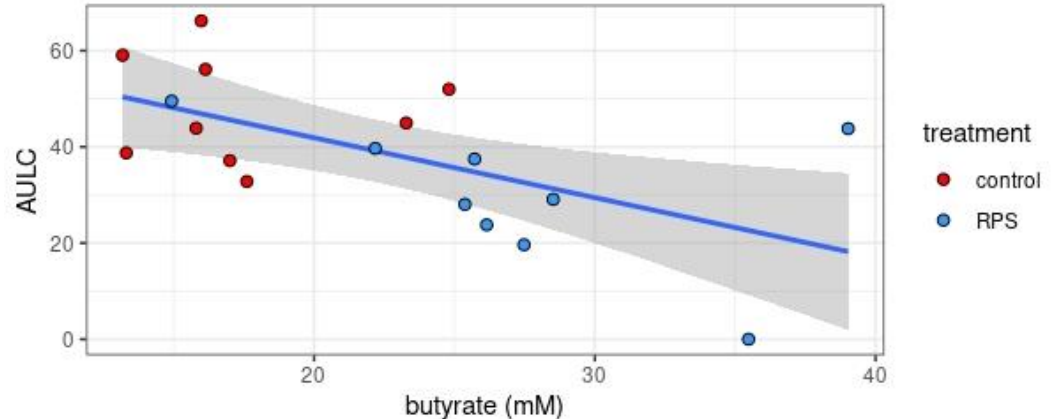
Negative correlation between butyrate and *Salmonella* shedding

Cecal SCFAs 21 days post challenge



Increased cecal butyrate even after *Salmonella* challenge
Reduced *Salmonella* shedding with increased butyrate levels

Correlation between AULC and cecal butyrate at D21



Dietary RPS can serve as alterbiotic

- Potential to promote growth
- Potential to modulate microbial communities
 - Modulate immune status
 - Enhance intestinal integrity
- Potential to limit *Salmonella* shedding

- Why potential? – dependent on food web of organisms that utilize RPS as substrate.
- Determining the mode of action of various alterbiotics will allow for targeted use in production.

Thanks to:



Dr. Julian Trachsel

Dr. Heather Allen

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Zahra Bond

Sam Humphrey

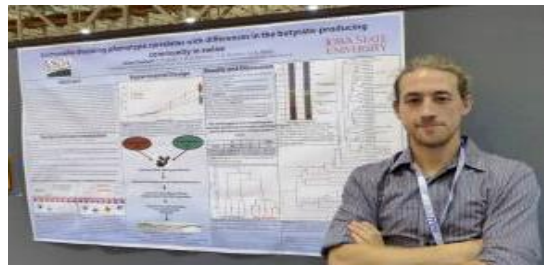
Jenn Jones

Elli Whalen

Dr. Shyamesh Kumar

Dr. David Alt

Animal Care Staff



Dr. Julian Trachsel



Dr. Kristen Byrne

Innate Training/immunomodulation
Poster MI11 & IM2



Dr. Shawn Bearson

Dr. Brad Bearson

Poster VA3

DIVA Salmonella vaccine



My peeps