

EFFECT OF GUT MICROBIOTA ON PRODUCTION TRAITS, INTERACTION WITH GENETICS

Hélène Gilbert

S Lagarrigue, L. Verschuren, O Zemb, M Velasco, JL Gourdine, R. Bergsma, D Renaudeau, JP Sanchez, H Garreau

INRA, France; Topigs Norsvin Research Center, The Netherlands; IRTA, Spain



Between feed and meat

Digestion = building a microbiota

Maternal transmission

Genetics

Age

Sex



Physiological status

Contact with (others') feces

MANAGEMENT

use of anti-microbials

feed

water

animal density

temperature and humidity

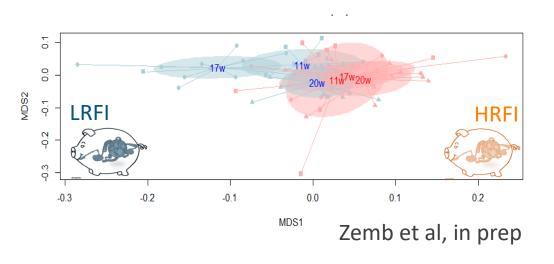
• • •

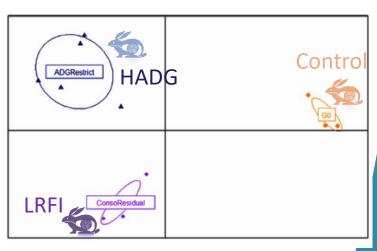
- → What should be accounted for to improve performances thanks to microbiota?
- → How to disentangle and test the effects of these factors?



Genetic determinism of microbiota composition

Based on selected <u>lines</u>





Drouilhet et al, JAS, 2016

→ Separate lines selected for different feed efficiency based on microbiota composition (DA-PC applied to OTU abundancy matrix)

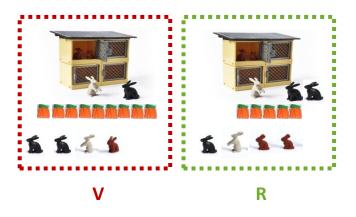
Feed-a-Gene



Adapting the feed, the animal and the feeding techniques to improve the efficiency and sustainability of monogastric livestock production systems

Genetic determinism

Within population



Bayesian estimation of variance components

Model **na**: no σ^2 a

Model **a**: σ^2 a

→ Deviance Information Criterion and BayesFactor computations

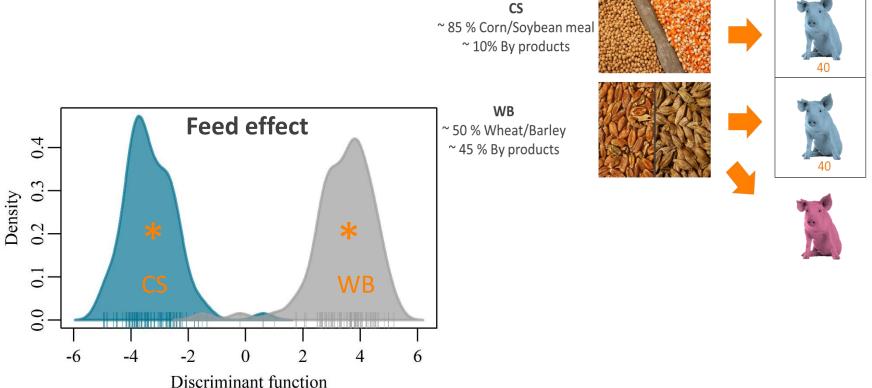
| Trait | h² |
|---------------------------|------|
| Phylum relative abundance | |
| Euryarcheota | 0.13 |
| Actinobacteria | 0.10 |
| Bacteroidetes | 0.09 |
| Cyanobacteria | 0.11 |
| Firmicutes | 0.09 |
| Proteobacteria | 0.11 |
| Tenericutes | 0.08 |
| Verrucomicrobia | 0.08 |
| Alpha index diversity | |
| #OTUs | 0.17 |
| Chao1 | 0.18 |
| Shannon | 0.11 |
| Inverse Simpson | 0.11 |
| Principal components (PC) | |
| PC1 | 0.09 |
| PC2 | 0.11 |

Feed-a-Gene



Adapting the feed, the animal and the feeding techniques to improve the efficiency and sustainability of monogastric livestock production systems

Feed effect



→ Contrasted feed compositions highly affect fecal microbiota

Day before slaughter





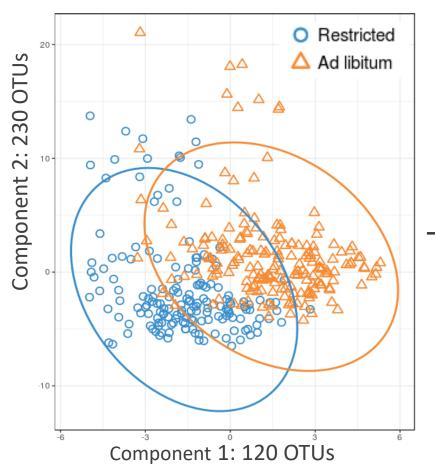


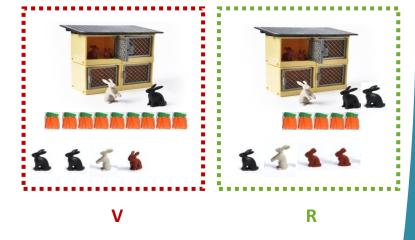


Verschuren et al, JAS, 2018

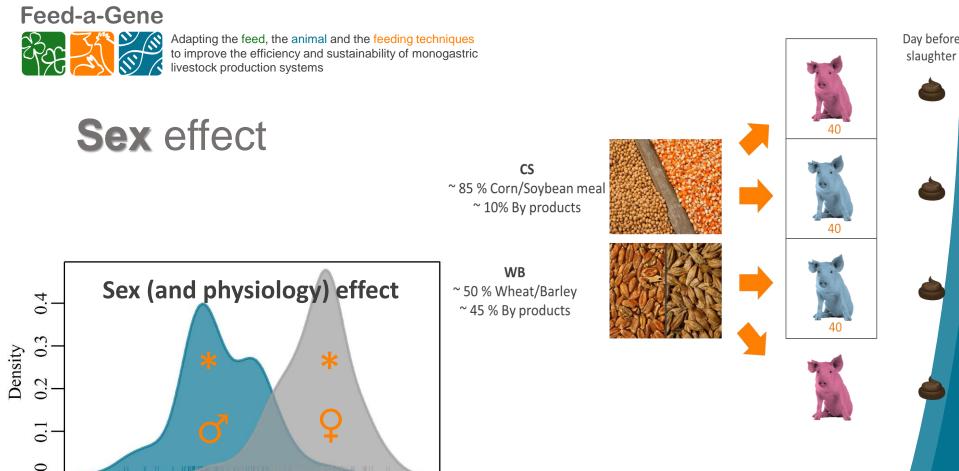


Feeding level effect





→ Feeding restricted or ad libitum affect fecal microbiota composition



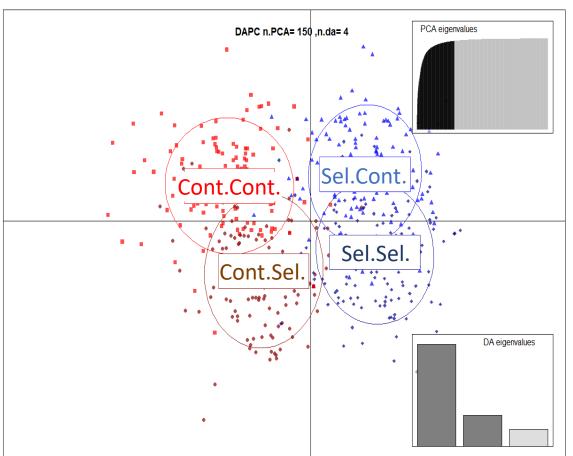
→ Sex, potentially via sexual maturity, affects gut microbiota composition

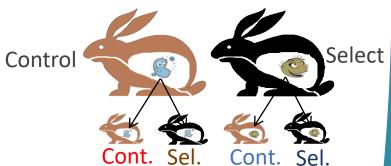
Discriminant function



Maternal transmission

Cross-fostering experiment





→ The kit line is the primary effect, but the dam line also nicely separates the individuals

Production traits: accounting for genetics

1. Model ignoring the effect of microbial composition

$$y = X\beta + a + e$$

FIXED EFFECTS Xβ RANDOM EFFECTS

Additive direct genetic: a ~ NMV (0, Aσ²a)

2. Model accounting for the effect of microbial composition

$$y = X\beta + a + b + e$$

RANDOM EFFECTS

- Additive direct genetic: a ~ NMV (0, Aσ²_a)
- Effect of the microbiome: $\mathbf{b} \sim \text{NMV}(0, \mathbf{B}\sigma_B^2)$

B =?
microbiability
$$\mathbf{b^2} = \sigma_B^2 / \sigma_P^2$$

13/09/2018



Production traits models: accounting for all confusion factors

Model accounting for the effect of microbial composition

$$y = X\beta + a + c + m + j + b + e$$

RANDOM EFFECTS

• Additive direct genetic $\mathbf{a} \sim N(0) \begin{bmatrix} \sigma_a^2 & \sigma_{am} \\ \sigma_{am} & \sigma_m^2 \end{bmatrix} \otimes \mathbf{A}$ • Maternal genetic $\mathbf{m} \sim N(0) \begin{bmatrix} \sigma_a^2 & \sigma_{am} \\ \sigma_{am} & \sigma_m^2 \end{bmatrix}$

- Litter: $\mathbf{c} \sim \text{NMV} (0, \mathbf{I}\sigma^2)$
- Pen: $\mathbf{j} \sim \text{NMV}(0, \mathbf{I}\sigma^2_i)$
- Effect of the microbiome: $\mathbf{b} \sim \text{NMV} (0, \mathbf{B}\sigma_B^2)$

axb?

13/09/2018



Production traits models





| | w/o b | with b |
|----------------|---------------|-------------|
| h² | 0.15 (0.04) > | 0.06 (0.03) |
| c ² | 0.26 (0.06) > | 0.14 (0.05) |
| b ² | - | 0.56 (0.11) |
| j² | 0.12 (0.03) > | 0.05 (0.02) |

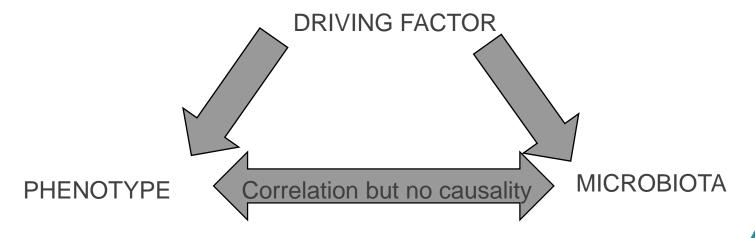
b ~ NMV (0, $\mathbf{B}\sigma^2_B$) **B**=f(wUnifrac dm)

→ An important % of the total phenotypic variance is explained by microbial composition



Summary

- ▶ Plenty of factors significantly affect gut microbiota composition → need to disentangle
- Some microbiota are related to production traits (statistical associations)
 - Causality? Joint effect of other factors?



13/09/2018

Discussion

- Ignored the diversity of OTU tables (content and rules to obtain them)
- Ignore the diversity of B
- Plenty is missing, keep digging
- What can be used for livestock at the moment?
 - For management?
 - For breeding?

Genetics
Maternal transmission
Age
Contact with (others') faeces
Sex
Physiological status

use of anti-microbioals

feed

water

animal density

temperature and humidity (see Le Sciellour et al,

Thursday morning)

13/09/2018