

*From sequence ...*

*...to consequence*

# ECO-FCE: Integrative genomic models for FCE in monogastrics

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# outline

*From sequence ...*

*...to consequence*

- **Selective Breeding**
- Models
- Simulation study
- Real data



The ECO-FCE project is funded by the European Union Seventh Frame Programme (FP7 2007/2013) under grant agreement No. 311794.



# Selective breeding

From

sequence

Phenotypic variation in the population



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Programme (2007/2013) under grant agreement No. 311794.



# Selective breeding

...to consequence



**Heritability**

Some of this variation is heritable:  
Related individuals have more similar  
phenotypes than unrelated individuals



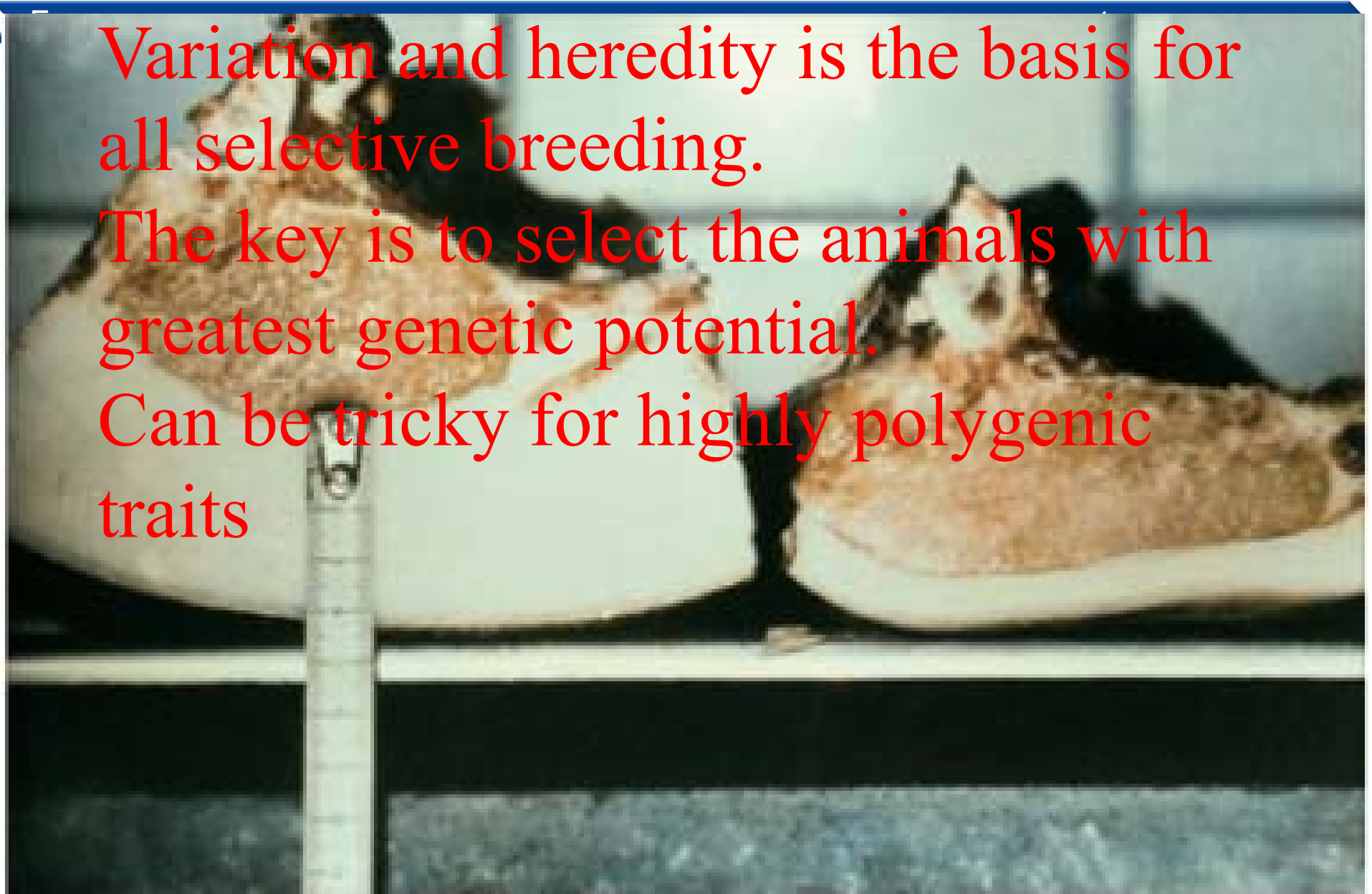
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Variation and heredity is the basis for all selective breeding.

The key is to select the animals with greatest genetic potential.

Can be tricky for highly polygenic traits



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# BLUP model

From sequence ...

...to consequence

$$\mathbf{y} = \mu + \mathbf{Z}_a \mathbf{a} + \mathbf{e}$$

Breeding values

Genetic relationship

$$\mathbf{a} \sim N(0, \mathbf{A}\sigma_a^2)$$

$$\mathbf{e} \sim N(0, \mathbf{I}\sigma_e^2)$$

Additive genetic variance



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# BLUP model

From sequence ...

...to consequence

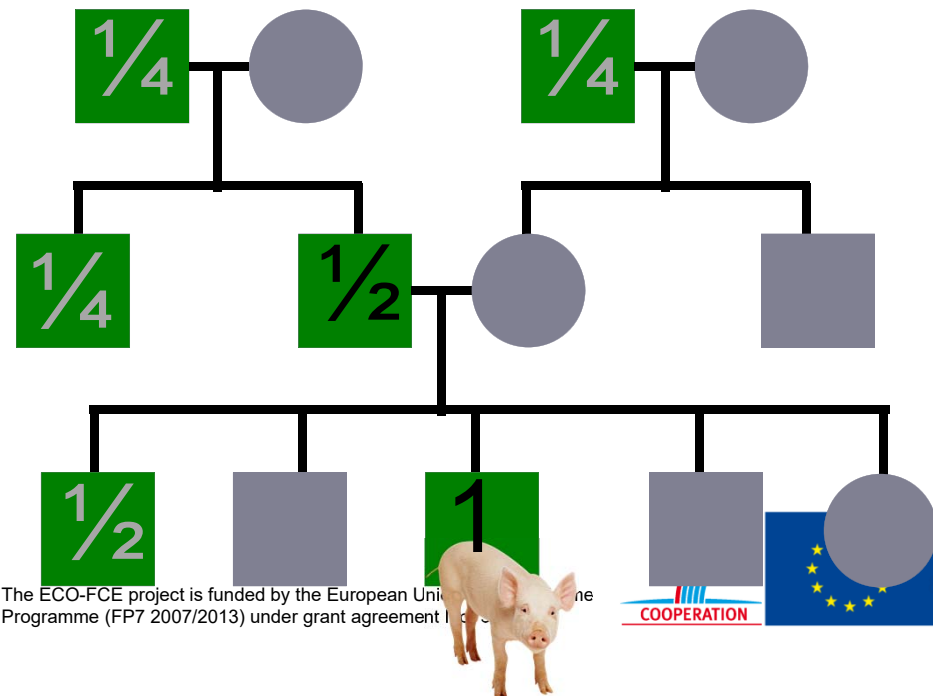
$$\mathbf{y} = \mu + \mathbf{Z}_a \mathbf{a} + \mathbf{e}$$

Breeding values

Genetic relationship

$$\mathbf{a} \sim N(0, \mathbf{A}\sigma_a^2)$$

$$\mathbf{e} \sim N(0, \mathbf{I}\sigma_e^2)$$



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# GBLUP model

From sequence ...

...to consequence

$$y = \mu + Z_g g + e$$

Genomic breeding value      Genomic relationship

$$g \sim N(0, G\sigma_g^2)$$

$$e \sim N(0, I\sigma_e^2)$$



# GBLUP model

From sequence ...

...to consequence

$$\mathbf{y} = \boldsymbol{\mu} + \mathbf{Z}_g \mathbf{g} + \mathbf{e}$$

$$\mathbf{g} \sim N(0, \mathbf{G}\sigma_g^2)$$

$$\mathbf{e} \sim N(0, \mathbf{I}\sigma_e^2)$$

All markers are from the same distribution - have same chance of being informative



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# GBLUP model

From sequence ...

...to consequence

$$y = \mu + Z_g g + e$$

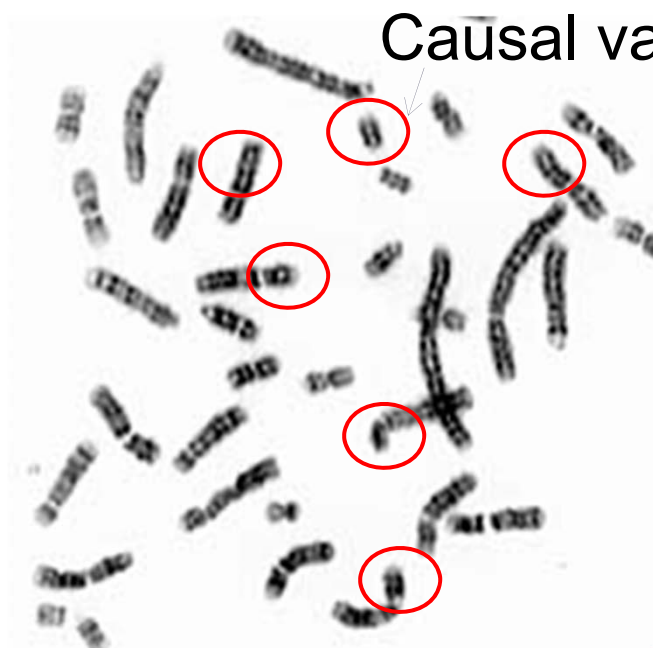
Genomic BV

Genomic relationship

$$g \sim N(0, G\sigma_g^2)$$

$$e \sim N(0, I\sigma_e^2)$$

Causal variation



# GFBLUP model

From sequence ...

...to consequence

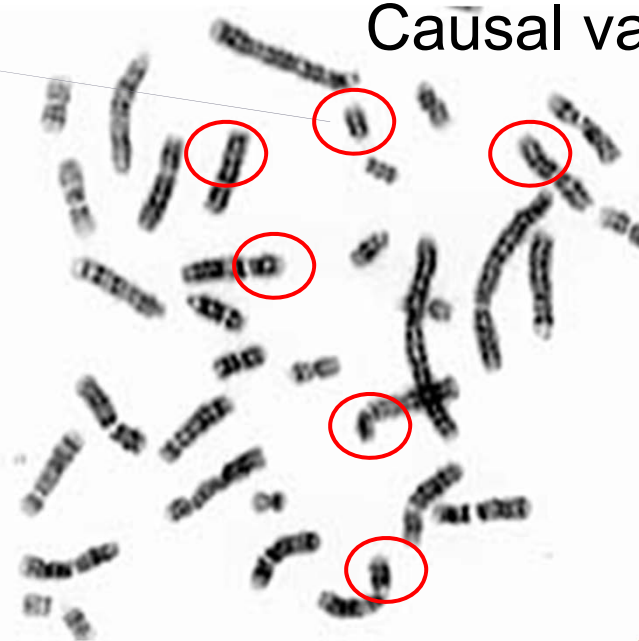
$$\mathbf{y} = \mu + \mathbf{Z}_f \mathbf{f} + \mathbf{Z}_r \mathbf{r} + \mathbf{e}$$

$$\mathbf{f} \sim N(0, \mathbf{G}_f \sigma_f^2)$$

$$\mathbf{r} \sim N(0, \mathbf{G}_r \sigma_r^2)$$

$$\mathbf{e} \sim N(0, \mathbf{I} \sigma_e^2)$$

Causal variation



Some markers are more equal than others



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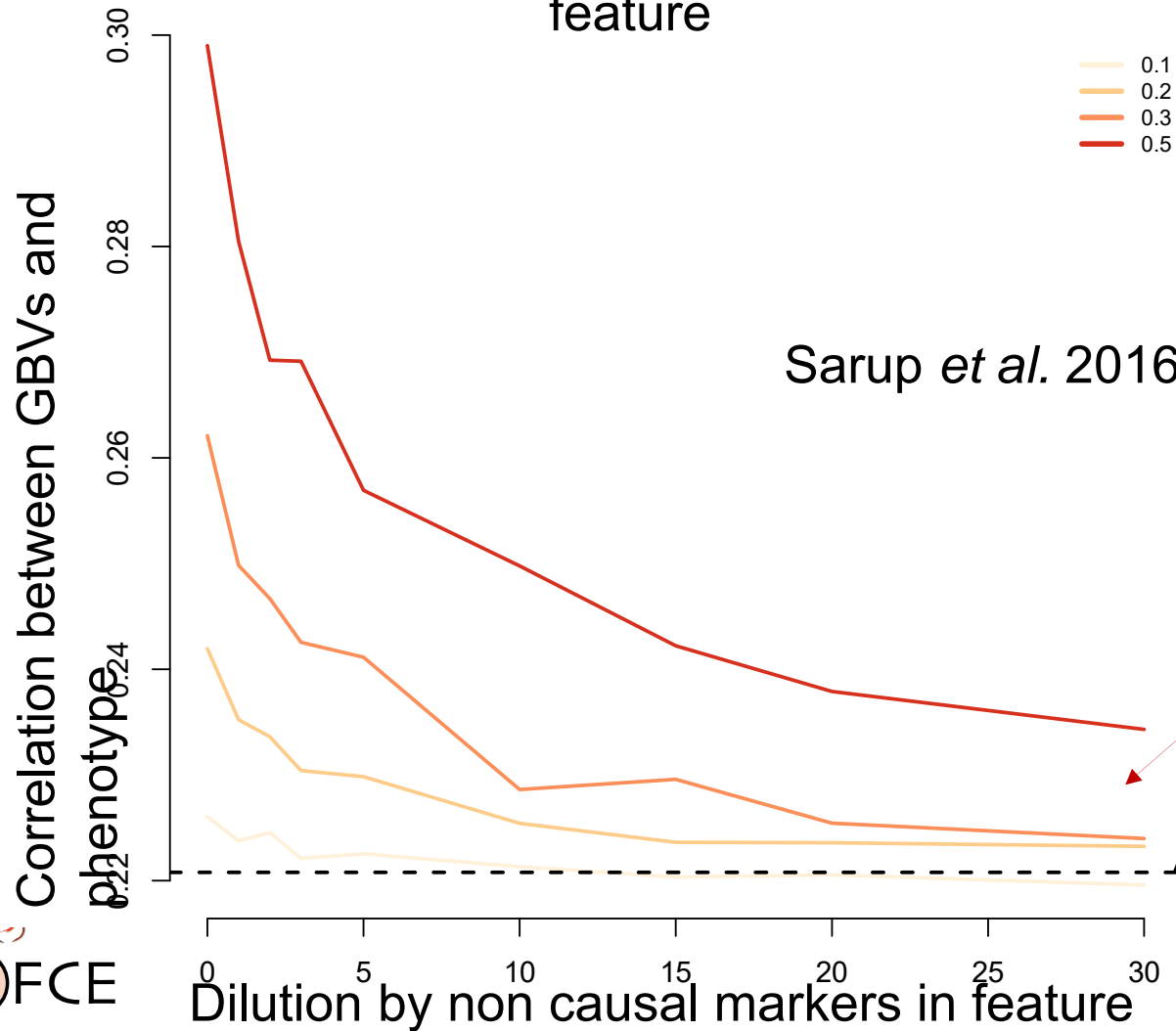


# Simulation study

From sequence ...

...to consequence

Proportion of genetic variation in genomic feature



Sarup *et al.* 2016, BMC genetics



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# Real data

From sequence ...

...to consequence

Genotypes for 2213 Hermitage MaxGro pigs  
from  
~55K SNPs.

Corrected phenotypes for genotyped individuals  
EBVs + residuals from BLUP on ~14K individuals

**Total Feed Intake TFI**  
**Residual Feed Intake RFI**

50 random 10 fold validation sets.



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# Defining genomic features

From sequence ...

...to consequence

Get information from

The study population

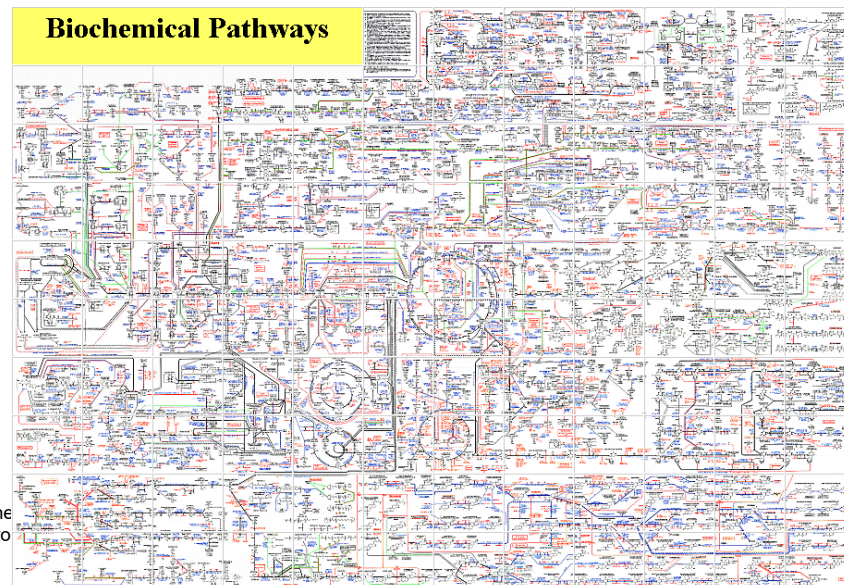
GWAS, **Gene expression**, proteomic, or metabolomic experiments

Literature/databases

QTL database

Gene Ontology

Kegg pathways



The Pro

# Defining genomic features

From sequence ...

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Some of the SNPs was linked to genes that differ in gene expression between high RFI and low RFI pigs



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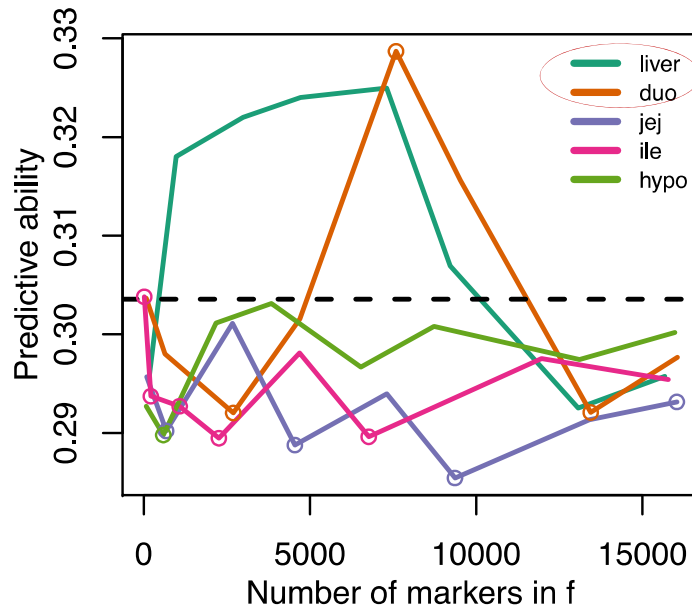


# results

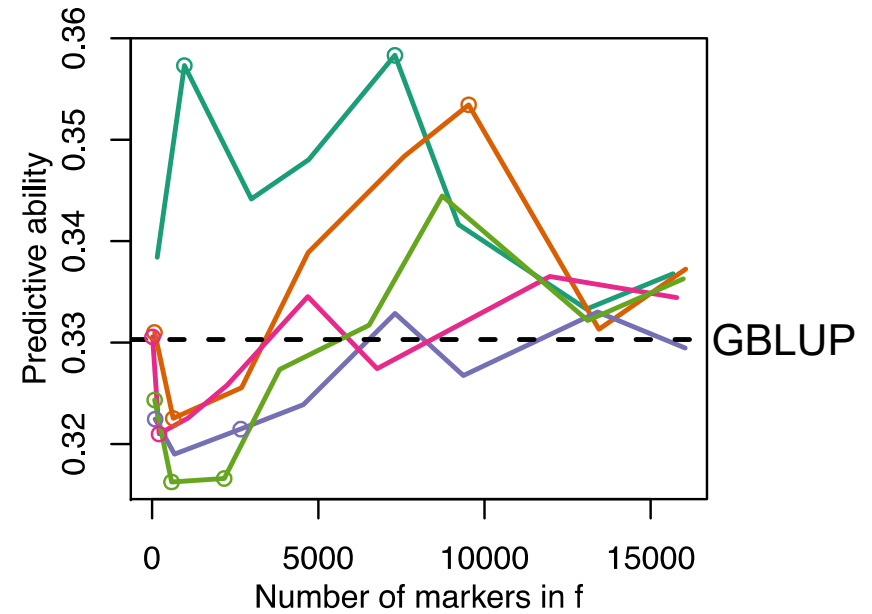
From sequence ...

...to consequence

### Residual Feed Intake



### Total Feed Intake



○ p < 0.001



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# Conclusions

From sequence ...

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Prediction accuracies improved by ~10%.

Genes that are differential expressed in liver and duodenum most informative.

**GFBLUP models can increase prediction ability and contribute knowledge on the biological background of complex traits**



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# Thank you for your attention

From sequence ... ..to consequence

Genotype and phenotype data provided by:



*P. Varley*



*T. Ostersen*

Gene expression experiments

*K. Wimmers, H. Reyer*



Model development and genomic prediction:

*P. Sarup, J. Jensen, M. Shirali, P. Sørensen*



More on models and analysis in  
session 55 Thursday 11.15 in

Auditorium  
ECO-FCE

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# Real data

From sequence ...

...to consequence

## Modeling of the phenotype

EBVs + residuals for genotyped individuals  
from BLUP on 13572 Hermitage tested  
animals

$$\text{TFI} = \text{CPG} + \text{SEX} + \text{LITTER} + A + \text{SBW} + \text{EBW} + e$$

$$\text{D110} = \text{CPG} + \text{SEX} + \text{LITTER} + A + e$$

$$\text{LMP} = \text{CPG} + \text{SEX} + \text{LITTER} + A + e$$

$$\text{RFI} = A_{\text{TFI}} - b_{\text{D110}} A_{\text{D110}} - b_{\text{LMP}} A_{\text{LMP}} + e_{\text{TFI}} - b_{\text{D110}} e_{\text{D110}} - b_{\text{LMP}} e_{\text{LMP}}$$



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