

Modulation of the redox state in pigs differing in feed efficiency as revealed by a proteomic analysis

F. Gondret¹, S. Tacher¹ & H. Gilbert²

¹UMR Pegase, 35590 Saint Gilles, France

²UMR GenPhySe, 31326 Castanet-Tolosan, France

Florence.gondret@rennes.inra.fr

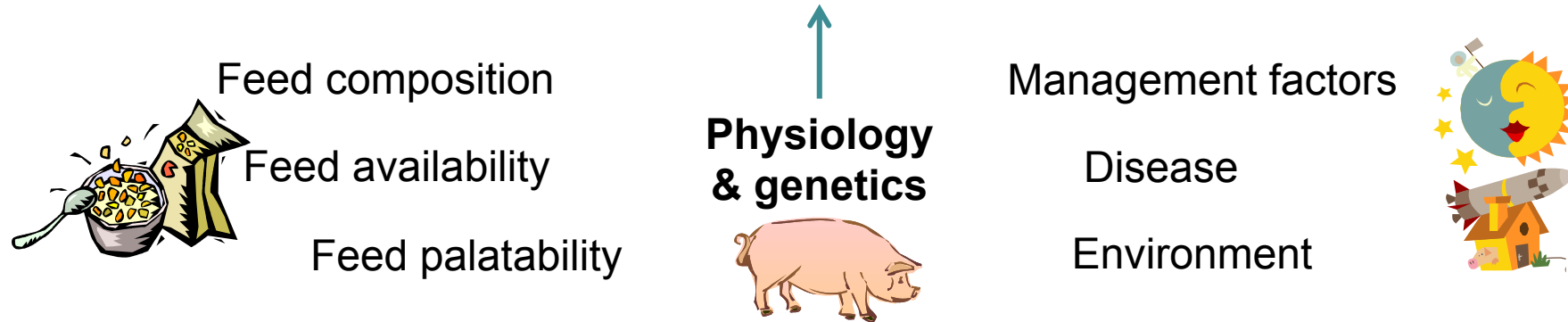


EAAP 2014: Appetite control –
mechanisms and comparative aspects



Animal feed efficiency (FE): a trait with a large economic impact for producers

Typical calculation of FE: **feed intake**/average daily gain



Residual feed intake (RFI) :

observed feed intake – expected feed intake

$BW^{0.60}$; ADG ; Backfat

maintenance *production*

RFI: ~ one third to half of the total variation in feed intake in growing pigs
(Knapp, 2009; Young & Dekkers, 2012)

- Context

Variations in residual feed intake (RFI):

Feed consumption & behavior (number of visits, duration of meal intake, time spent eating, etc...)

Young et al., 2011

Meunier-Salaün et al., 2012

Digestion and utilization of nutrients

Barea et al., 2010

Organ physiology : thermoregulation

basal metabolic rate

energy efficiency of mitochondria

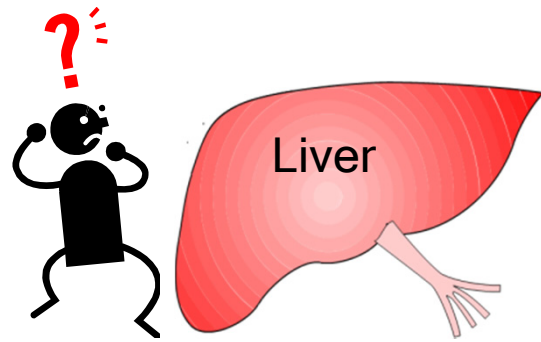
protein turnover rate

immunity, etc.

Skeletal
muscle



Le Naou et al., 2012; Gabler et al., 2013; Grubbs et al., 2013



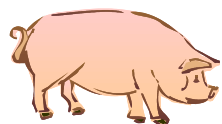
The liver is a complex and unique organ responsible for a breadth of functions that are crucial for sustaining life



We propose to use a proteomic differential approach to identify the modulation of **hepatic** metabolic processes participating **to feed efficiency**

- Objectives

6th generation of a **divergent selection for RFI** in pigs



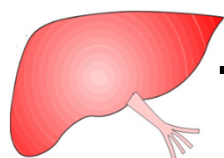
Large White pure genotype 10 wks of age
(30 kg BW)

RFI-
(low feed intake, high FE)

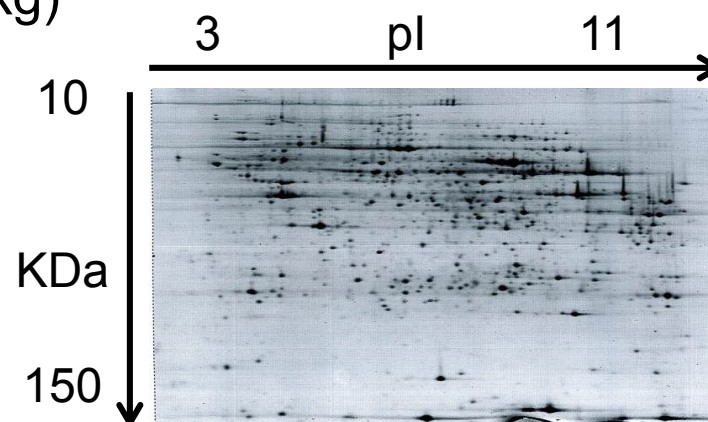
RFI⁺_R
(pair-fed to RFI- pigs)

RFI+
(high feed intake, low FE)

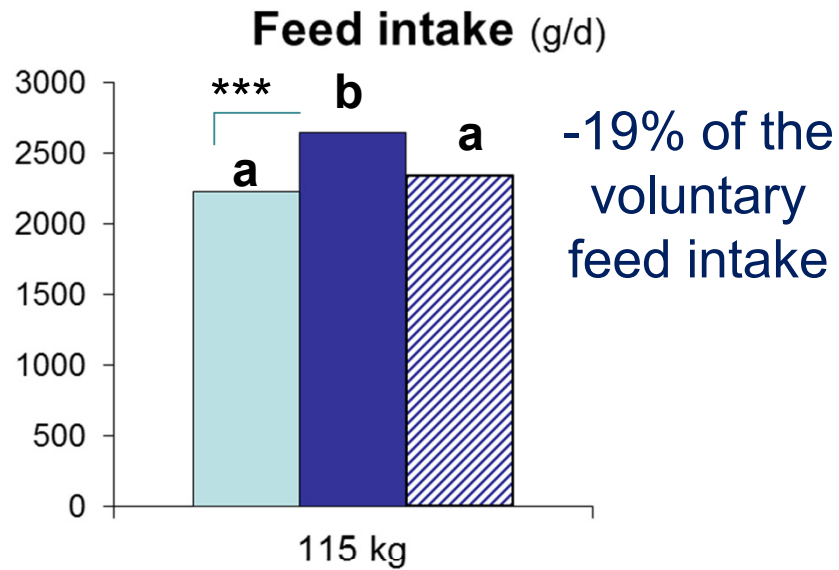
Slaughtered
at the same BW (115 kg)
(n = 8 / group)



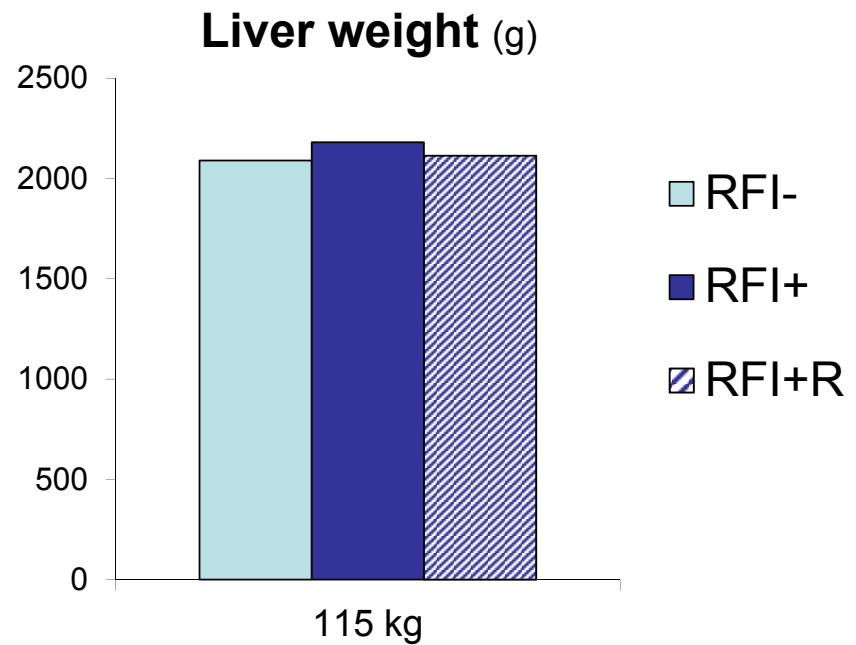
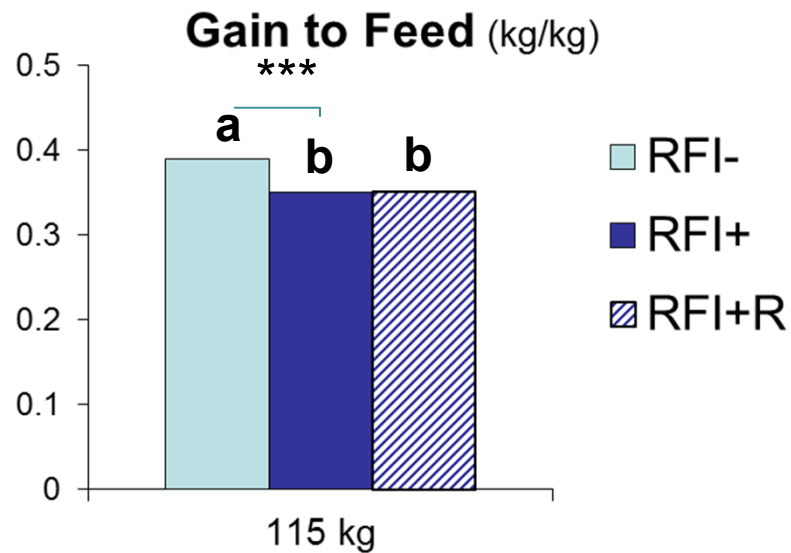
2D-electrophoresis of soluble
proteins
+ MS/MS identification of
differentially-abundant spots



- Experimental design



RFI corresponded to a biologically imposed feed restriction



- Results: Animal performance

A total of 50 protein spots in the liver had a differential abundance between pigs

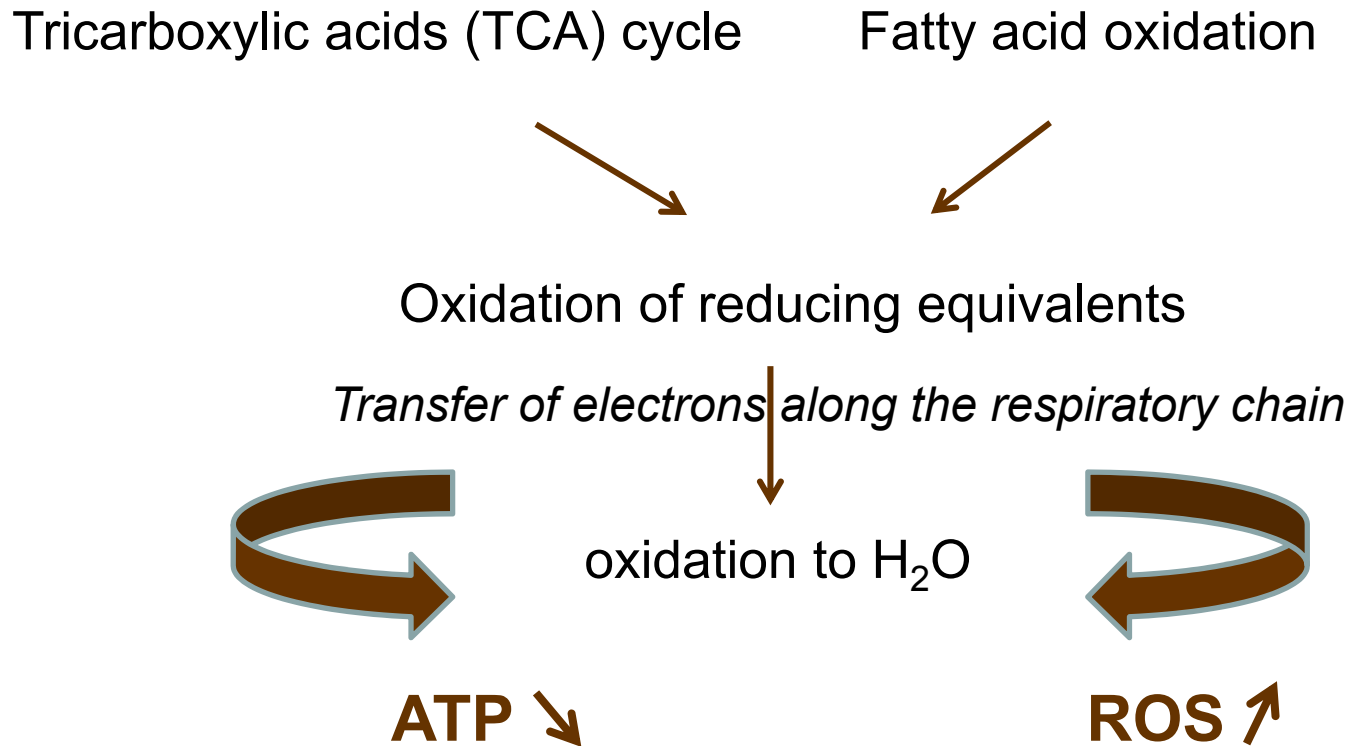
Proteins were included in pathways related to AA metabolism, glycolysis/gluconeogenesis, ion binding, purine binding, and **oxido-reduction**

11 unique proteins belong to a top cluster related to cell oxido-reduction processes
(enrichment score = 2.92)

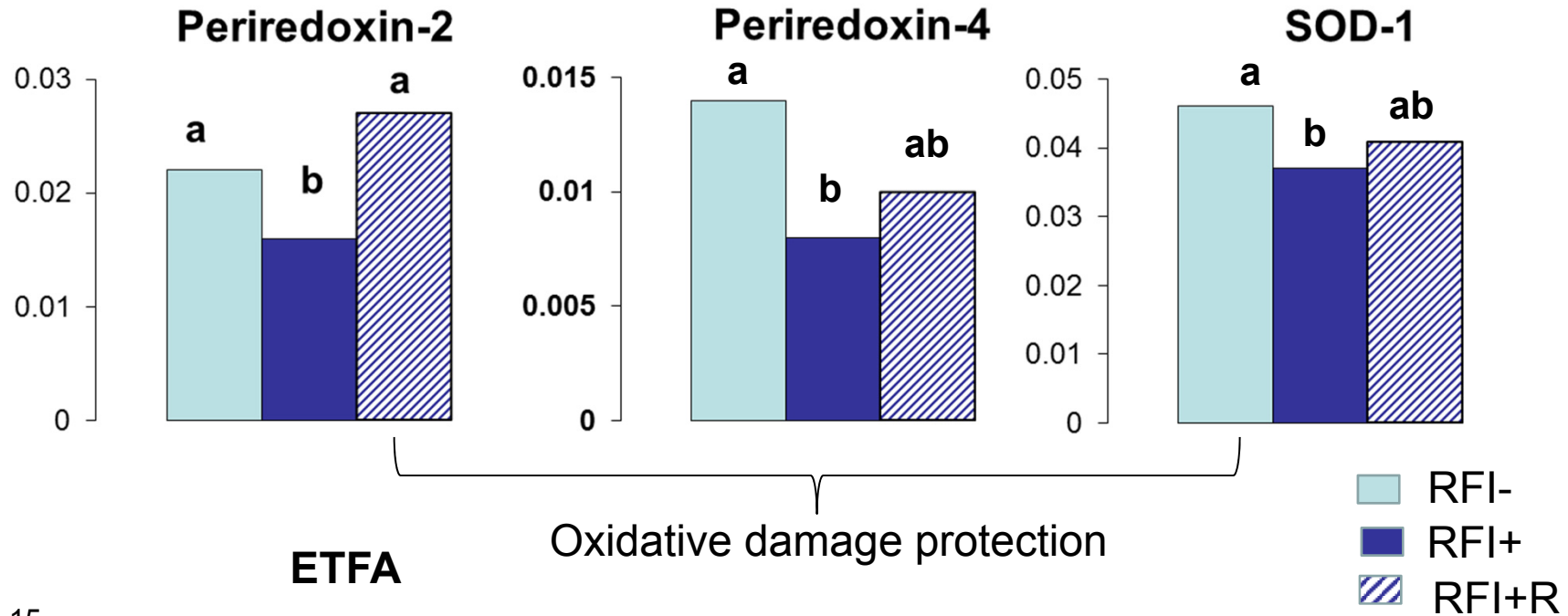
Response to oxidative stress	PRDX2, PRDX4, PRDX5, PRDX6, SOD1
Cell redox homeostasis	PRDX2, PRDX4, PRDX5, PRDX6, PDIA3, PDIA6, GSTO1
Oxidation reduction	PRDX2, PRDX4, PRDX5, PRDX6, SOD1, HAAO, DMGDH, ETFA

- Results: differential proteins in liver

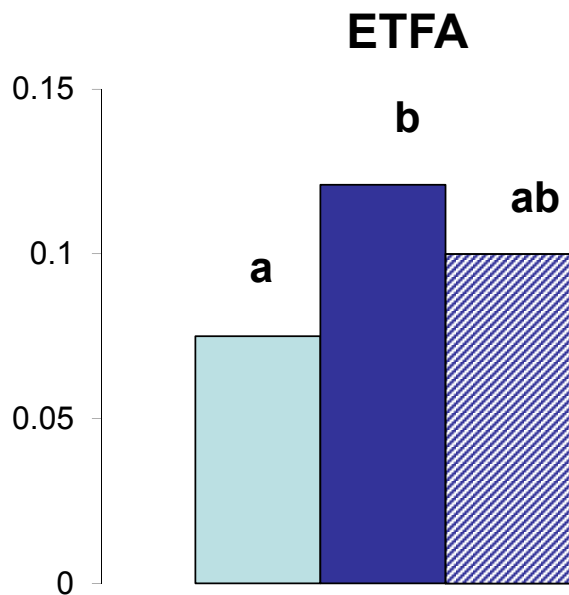
Liver mitochondria and Reactive Oxygen Species (ROS) production



- Liver mitochondria are a main site of ROS production



Oxidative damage protection



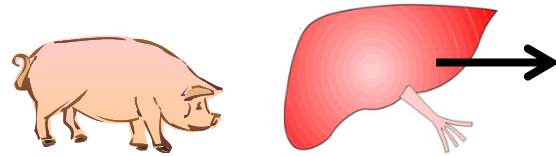
A lower abundance in the **electron transfer flavoprotein (ETF A)** is considered as leading to ROS elevation

Levels in feed-restricted high RFI pigs were similar or intermediary to levels observed in low RFI pigs

- Results: greater anti-oxidant enzymes in low RFI pigs

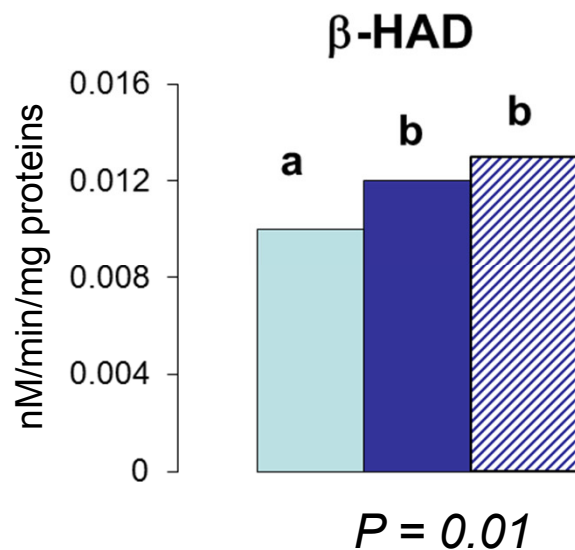
A low abundance of ETFA signs a low activity of the respiratory chain

A severe inhibition of the respiratory chain is considered as inhibiting lipid oxidation process in rats (*Pande, 1971*)

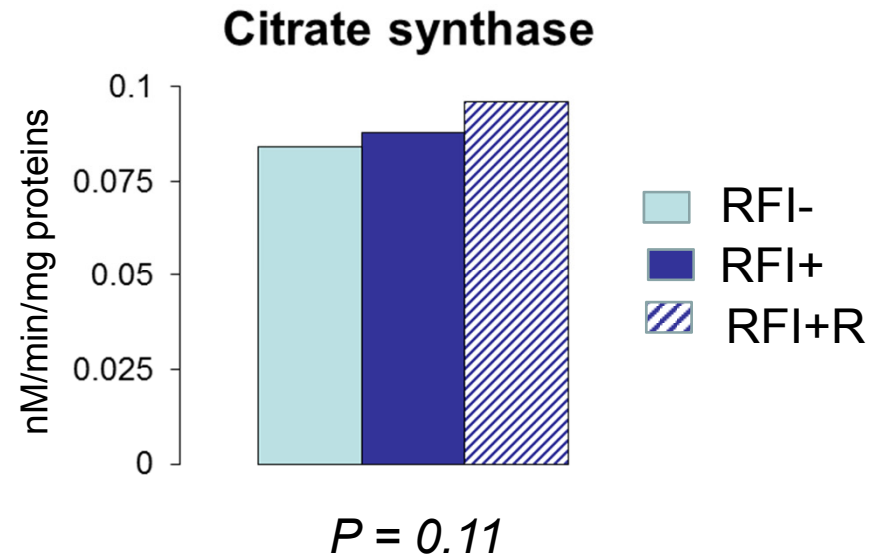


Monitoring oxidative enzyme activities in the 3 groups

Fatty acid oxidation



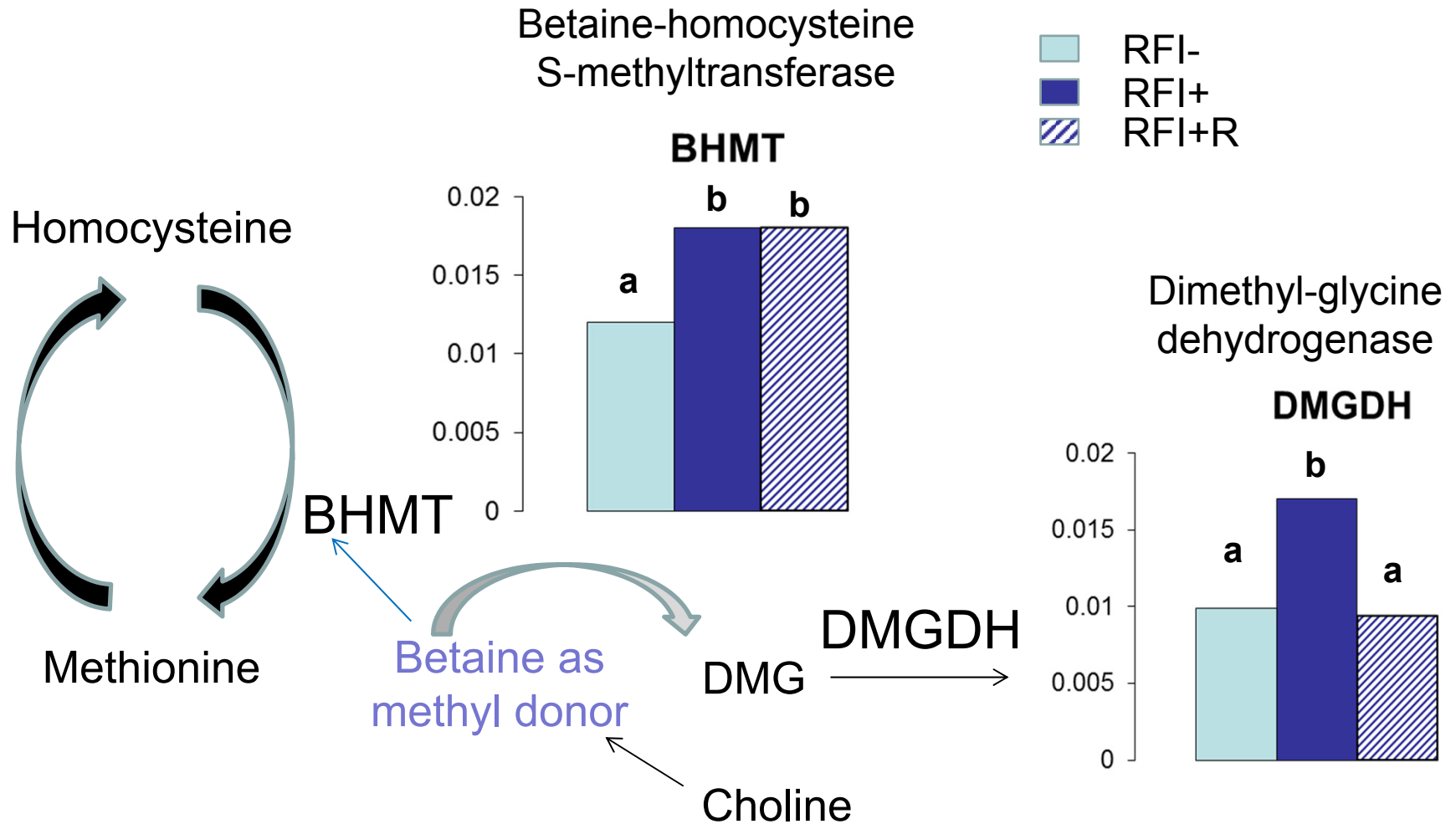
TCA cycle



Lower fatty acid oxidation in the low RFI pigs, independently of feed intake

- Results: lower lipid oxidation pathway in low RFI pigs

Balance in labile methyl groups (methionine, betaine, choline) is involved in maintaining normal lipid functions and health (*Mato et al., 2008*)



- Results: homocysteine/methionine/betaine equilibrium ?

❖ Pigs selected for a low residual feed intake had eaten less during the growing period

❖ Higher anti-oxidant capacities in the liver of the most efficient pigs, due to (?) greater ROS generation

Consequences on the redox state of the animals remains to be determined (other tissues)

❖ Levels of anti-oxidants were lower with feed restriction.

Relationships between feed intake and oxidative stress/redox metabolism in pigs deserve further studies

caloric restriction in aging mice decreased hydrogen peroxide production from the mitochondria.

Conclusions

Special thanks to:



The staffs of:

- UMR GenEsi (Poitou-Charente, France) for the divergent selection experiment
- UMR Pegase for the feeding trials, animal slaughtering and lab analyses



The financial support of:

- Program PigFeed (ANR-08-GANI-038)



High-proteomic core facilities (Rennes, France)

Acknowledgements