

**SABRE**  
 CUTTING EDGE GENOMICS FOR SUSTAINABLE ANIMAL BREEDING

*WP5 Mammary Function*

**Understanding the Genetics of Mastitis**

Partners:  
 MTT (Finland)  
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**Genetics of Mastitis**

- **Population level**
  - low heritable trait ~ 5-15%
  - clinical mastitis and somatic cell score correlated ~ 0.6-0.7
  - correlated to milk yield ~ 0.4-0.6
- **QTL studies**
  - Several QTL identified
  - BUT no causative genes
- **Variation on different levels**
  - Genome
  - Transcriptome
  - Proteome
  - Phenotype

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**Objectives WP5**

- **QTL studies**
  - Finemapping of QTLs for mastitis/SCS
  - Assess pathogen specificity of mastitis QTL
- **Expression studies**
  - Identify candidate genes/pathways associated with infection
  - Identify candidate genes/pathways associated with mastitis QTL
- **Association study**
  - Identify SNPs in candidate genes associated with mastitis in population

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**Combine QTL and Expression studies**

**QTL study** identifying a chromosomal region associated with mastitis

**Expression study** comparing expression profiles of animals carrying high (H) or low (L) resistant haplotypes

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**Combine QTL and Expression studies**

DNA: L1, L2, L3, L4

RNA: R1, R2, R3, R4

Environment: E1, E2

Complex trait: C1, C2

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**QTL study**

Mastitis QTL finemapped in Danish Holstein population

Pathogen specificity of haplotypes

H | L

High resistant | Low resistant

E-coli

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### Combine QTL and Expression studies

Mastitis QTL finemapped in Danish Holstein population

Expression profiling of H/L haplotypes

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### Expression studies

Experiment 1: Cows challenged with LPS (from E-coli)

Experiment 2: High/Low cows challenged with E-coli

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### Cows challenged with LPS

- Mastitis QTL associated E-coli
- Lipopolysaccharide (LPS) - from E.coli
- Initiating factor of acute phase response:
  - A systemic response with physiological phenomena like fever, leukocytosis and altered serum concentrations of acute phase proteins (APP)
- Altered gene expression in liver is thought to be responsible for altered serum concentrations of APP

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### Cows challenged with LPS

→
→

8

Challenge with LPS
Liver
Gene expression

(-22, 3, 6, 9, 12, 48)

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### Cows challenged with LPS

RNA

Clinical traits

before/after LPS

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### Cows challenged with LPS

- immune response
- metabolism

Time


LPS

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## Expression studies

Experiment 1: Cows challenged with LPS


Experiment 2: High/Low cows challenged with E-coli




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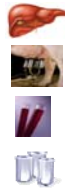
## High/Low cows challenged with E-coli


18 H




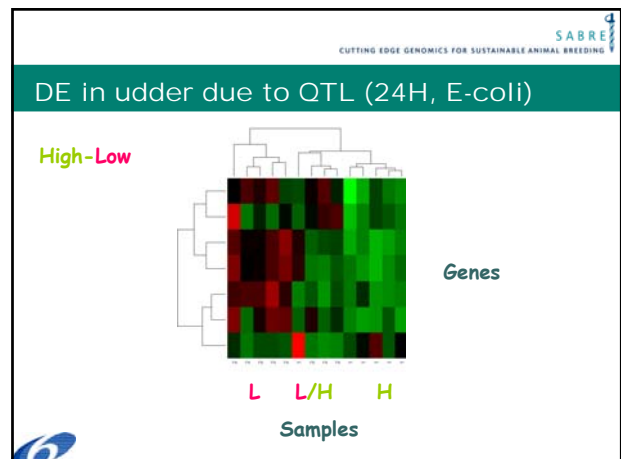
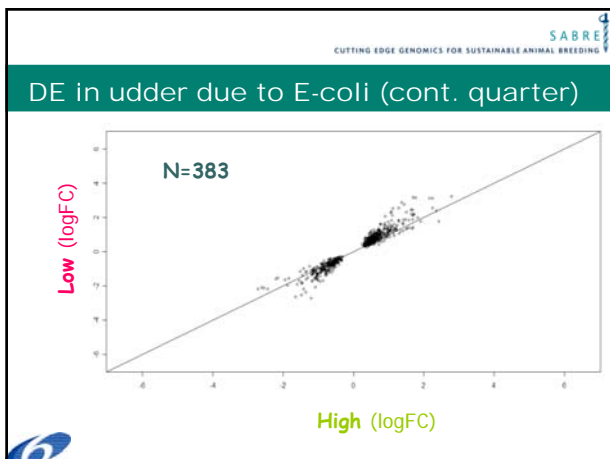
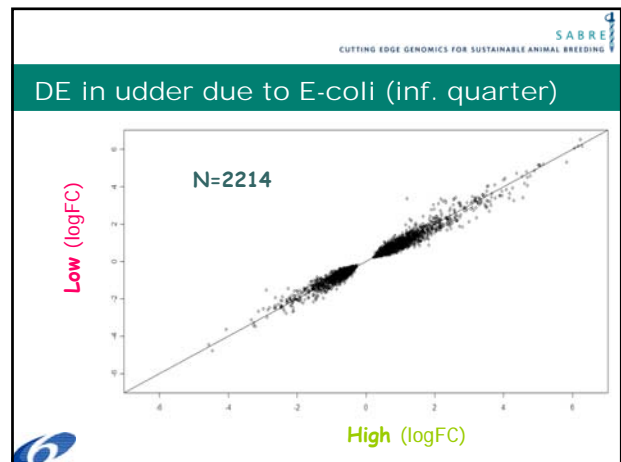
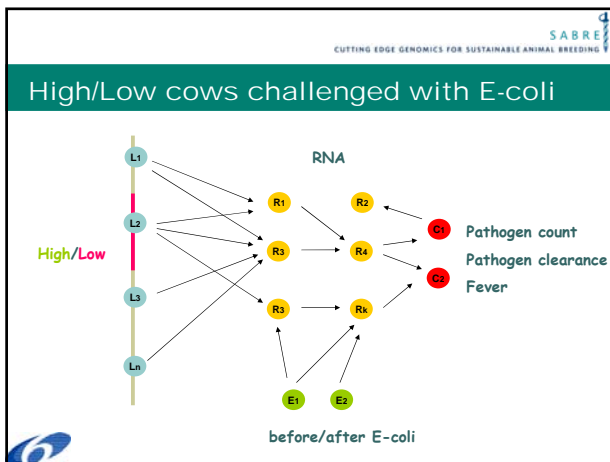
14 L








Challenge with E-coli      Liver/Udder/Blood/Milk      Gene expression  
 (-144, 12, 24, 192)

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## Summary

- **QTL studies:**
  - Mastitis QTL finemapped in Holstein population
  - Mastitis QTL associated with E-coli
- **Expression studies:**
  - LPS induced mastitis coordinately altered the expression of immune response and metabolism genes in the bovine liver
  - E-coli induced mastitis indicate a different expression profile in the udder of high/low resistant cows
- **Next steps**
  - Bioinformatic approaches for integrated analyses of linkage and expression data for ranking candidate genes within linkage region
  - Identify SNPs in candidate genes and associate with mastitis phenotypes in population



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## Acknowledgements

Mogens S. Lund  
Christian Bendixen  
Goutam Sahana  
Lars P. Sørensen  
Jørn R. Thomassen  
Johanna Vilkki  
Li Jiang  
Axel Skarman  
Christine Marie Røntved  
Klaus L. Ingvarstsen

