

GM2.7

New breeding tools for
improving mastitis resistance
in European dairy cattle



johanna.vilkki@mtt.fi

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MASTITIS RESISTANCE

- MTT (Finland)/ Johanna Vilkki, Sirja Viitala, Nina Schulman
- SLU (Sweden)/ Lena Andersson-Eklund, Leif Andersson, Ana Fernandez
- EAU (Estonia)/ Haldja Viinalass, Sirje Värvi
- DIAS (Denmark)/ Mogens Sando Lund, Bo Thomsen, Ayman Sabry
- ROSLIN (UK)/ John Williams, John Woolliams, Nicola Hastings
- ID-Lelystad (NL)/ Lucia Kaal, Luc Janss, Herwin Eding

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• Mastitis resistance - aiming at MAS!



- the most common disease in dairy cattle (up to 40%)
- low heritability
- difficult to improve by selection
- difficult to measure
- veterinary records available only in Nordic countries

Objectives

- Combine different genome scans into a multi-population, multiple trait analysis
- Improve statistical methodology for fine-mapping
- Identify marker haplotypes associated with QTL alleles (-> MAS)
- Dissect the genetic basis of mastitis by :
 - (i) disease etiology correlated to specific QTL alleles and pathogens,
 - (ii) positional candidate gene sequence variation,
 - (iii) pleiotropic effects of each QTL
 - (iv) gene expression

Material

- Nordic QTL scans (old & new families)
- DYDs for clinical mastitis and SCC
- pathogen data of milk samples
- bovine RH-panel and BAC libraries (Roslin, DIAS)

<i>BREED</i>	<i>N sires</i>	<i>N bulls</i>
<i>"Red" group:</i>		
<i>Ayrshire</i>	<i>24</i>	<i>900</i>
<i>Swedish R&W</i>	<i>20</i>	<i>750</i>
<i>Red Danish</i>	<i>9</i>	<i>300</i>
<i>Holstein-Friesian</i>	<i>25</i>	<i>2100</i>
<i>Total</i>	<i>78</i>	<i>4050</i>

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Statistical developments

- Multiple trait analysis (CM and SCC)
 - to increase power and accuracy
 - distinction between pleiotropy and linkage
- Longitudinal analysis (SCC)
 - non-constant QTL effects
 - information from fitting time dependent QTL (Lund et al. 2002)

QTL regions for fine-mapping

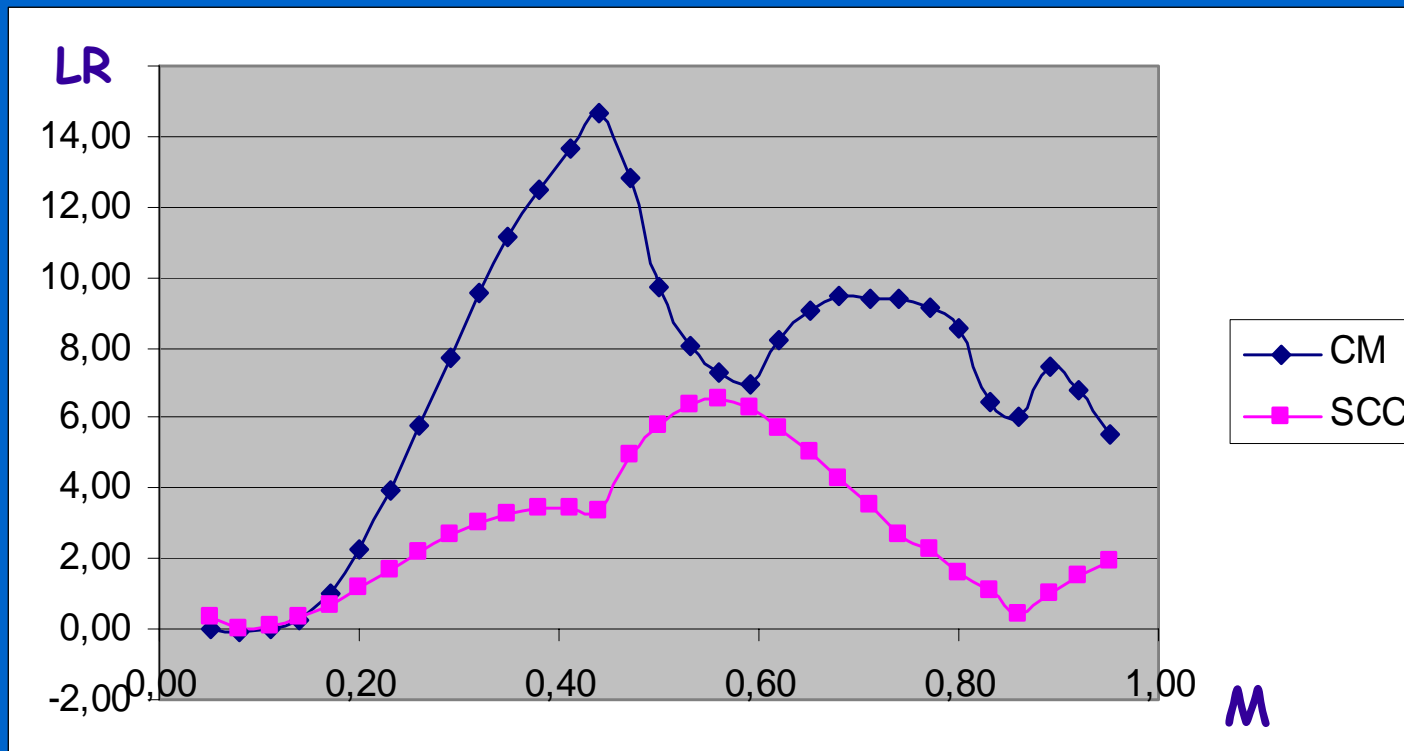
- 6 chromosome regions with best support from previous QTL scans (40 families, 3 countries) were chosen
- 78 families genotyped with common marker set at 20 cM intervals
- segregating families ($p < 0.1$, single trait least squares analysis) genotyped at 5 cM marker intervals for best regions - 15 to 25 families/ region

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Statistical analyses

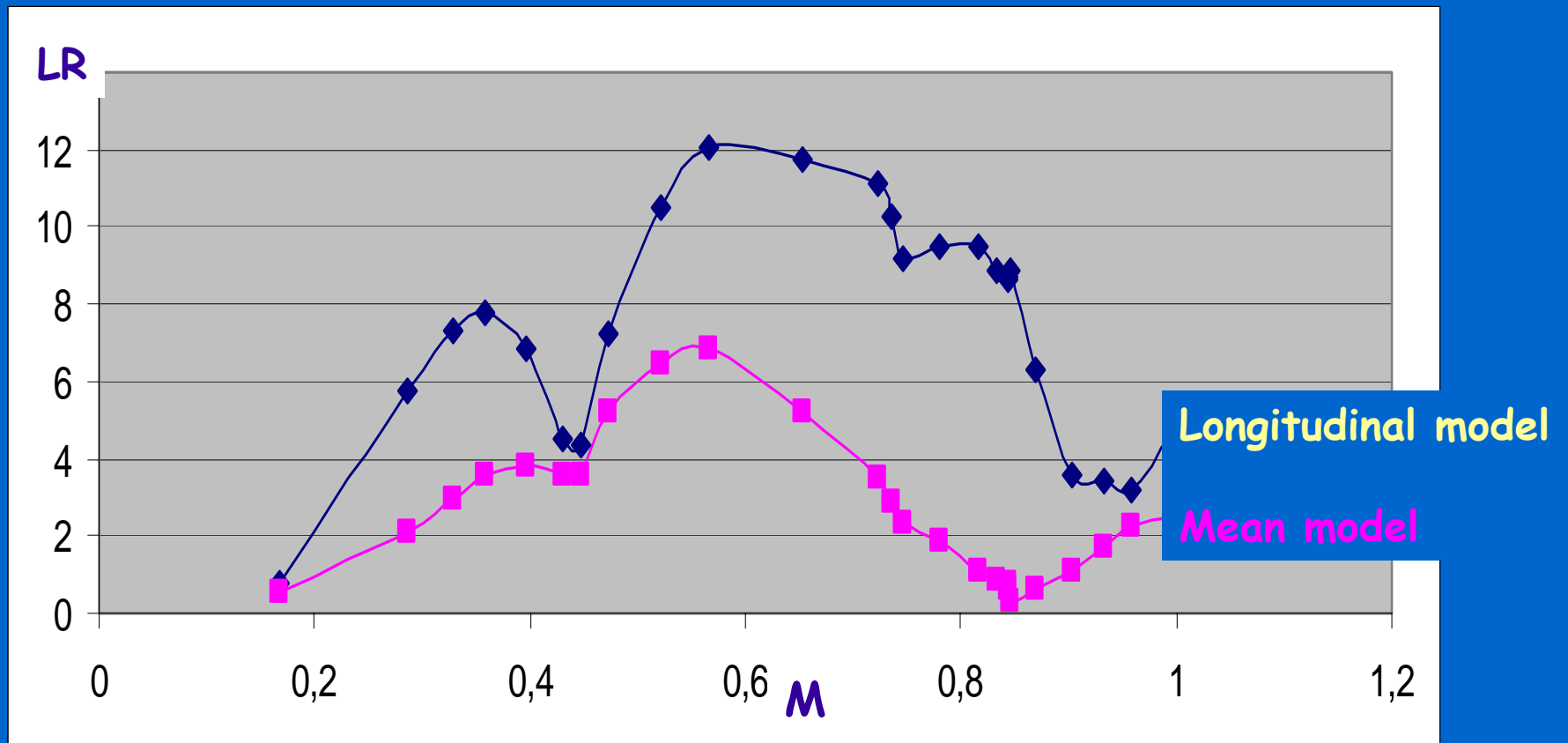
- Least squares analyses within and across families
- Variance components analyses:
 - linkage analysis (LA)
 - LA/LD
 - Multitrait: CM + SCC
 - Longitudinal data: SCC

"Multitrait" analysis of CM and SCC



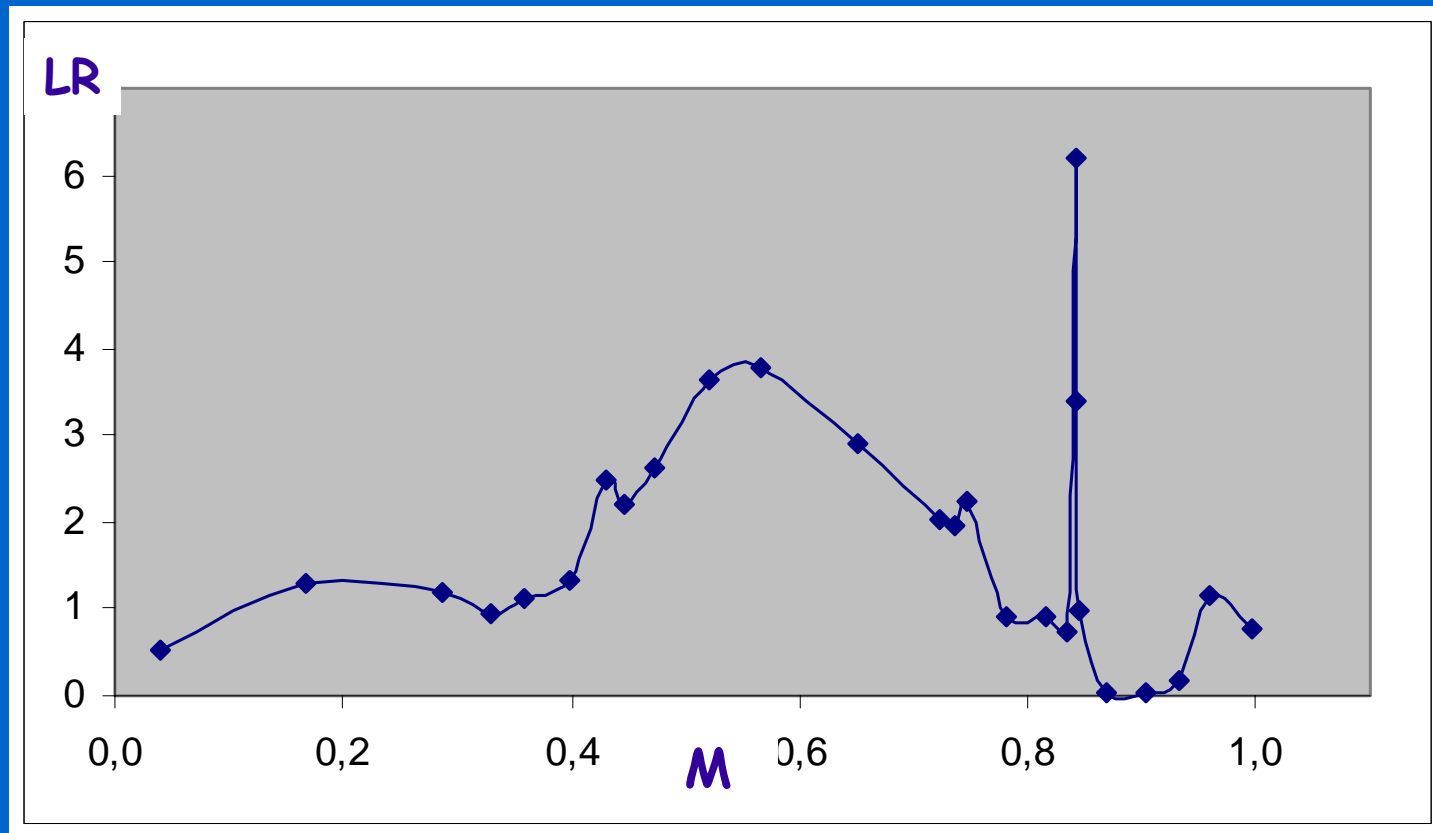
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SCC field data - preliminary results



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Linkage/Linkage disequilibrium



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Fine-mapping

Development of a dense marker map for the defined short regions:

- new microsatellites from bovine sequence trace archives
- comparative mapping of BACs in QTL regions to human map
- sequencing of genes (BACs & human orthologues in bovine WGS) -> SNPs

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Basis for further studies

- A list of candidate genes, based on their position, including knowledge of sequence variation (SNPs)
- Animals with known QTL genotypes
- Knowledge of differences in pathogen specificity of identified QTL alleles

→ Material for functional analysis

Strategy for use



- pleiotropic effects of each QTL on other important traits known
-> a strategy for the use of these QTL in breeding programs.
- developing a set of guidelines to aid introgression programmes