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Fine Mapping a QTL for Somatic Cell Score in the German Holstein

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Introduction

Somatic Cell Score (SCS)

- indicator trait for mastitis
- $h^2 = 16 - 17\%$
- marker assisted selection

😊 Over 50 QTL for SCS reported

☹ Large confidence intervals



Objectives

- Investigate chromosomal region on BTA27 affecting SCS:
 - ↑ marker density
 - ↑ number of families
 - linkage disequilibrium
- Identify haplotypes associated with increases or decreases in SCS



Experimental Design

Pedigree

- GDD (6 families)
- 492 sires → **492 DYD for SCS**
- 4,622 animals

Linkage Map

- 19 Microsatellites (~95 SNP) spanning 33.8 cM
- 6.84 alleles / marker
- Marker interval: 1.78 cM
- Putative QTL at midpoint of every marker interval

$\mathbf{y}_{i(m \times 1)}$

Observations for sire i
(DYD for SCS)

$\mathbf{X}_{(m \times k)} \mathbf{Z}_{(m \times n)} \mathbf{W}_{(m \times \text{gam})}$

Design matrices


$$\mathbf{y}_i = \mathbf{X}\boldsymbol{\beta} + \mathbf{Z}\mathbf{u} + \mathbf{W}\mathbf{v} + \mathbf{e}$$

$\boldsymbol{\beta}_{(k \times 1)}$

Fixed
Average

$\mathbf{u}_{(n \times 1)}$

Random polygenic effects
which have nothing
to do with the QTL

$\mathbf{v}_{(\text{gam} \times 1)}$

Random gametic
(QTL) effects

$\mathbf{e}$

Random
residual

Analysis

Restricted log likelihood ratio test statistic
for all QTL positions ($pos=1-18$):

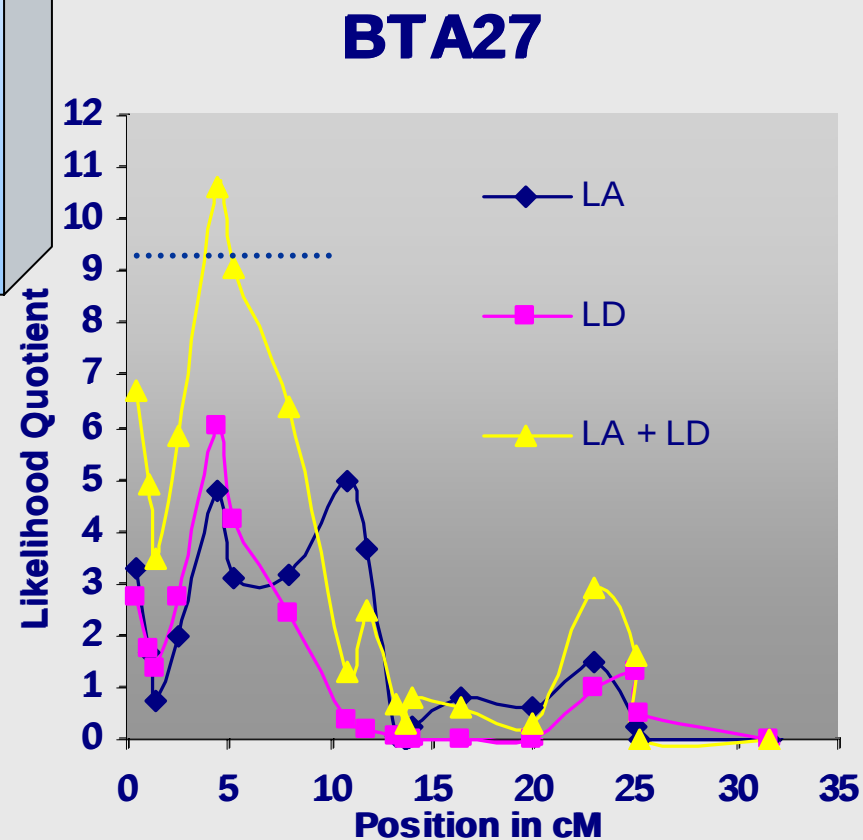
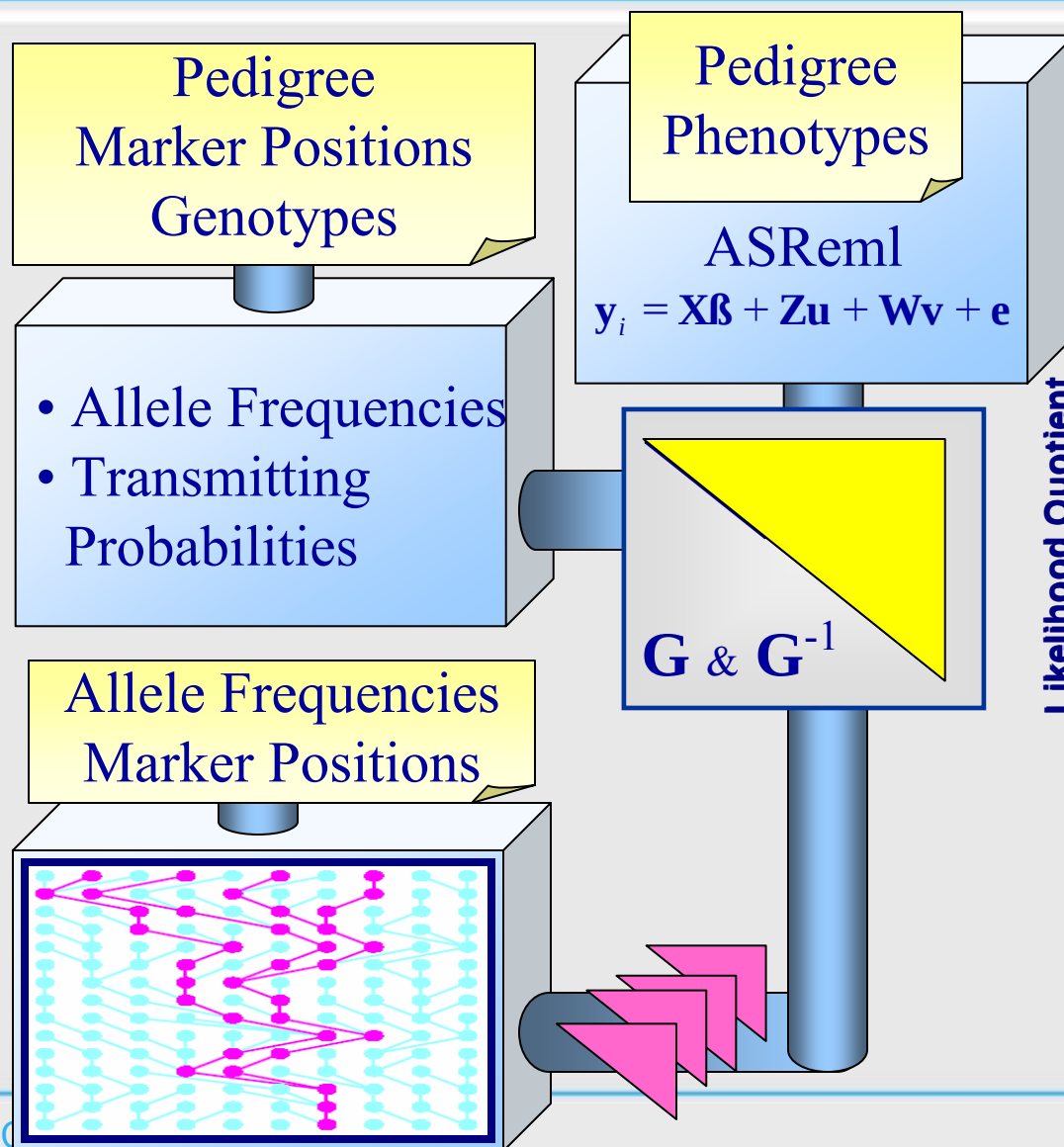
$$RLRT^{Pos} = -2 \left[\ln(L_0) - \ln(L_1^{Pos}) \right]$$



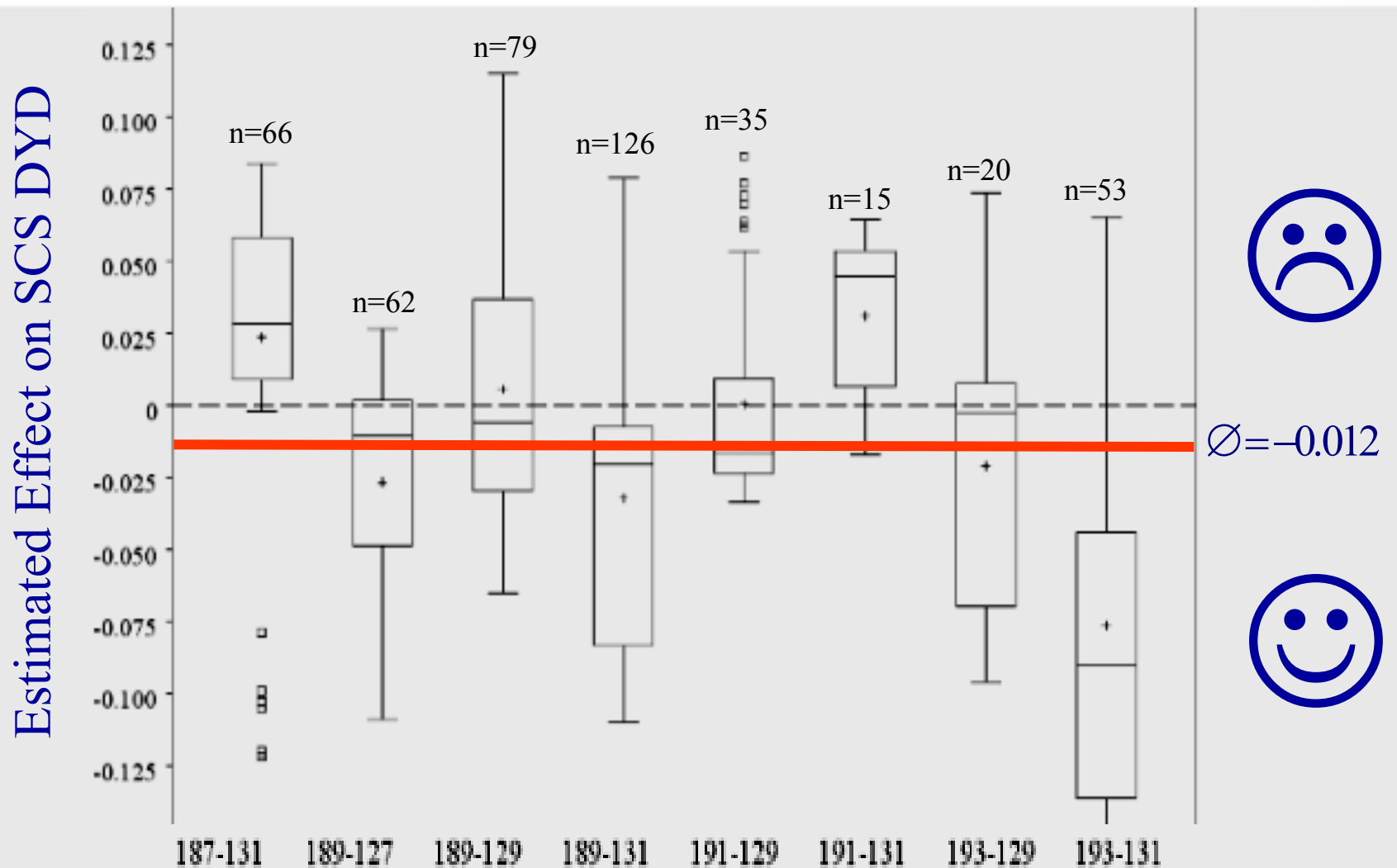
$$L_0 \longrightarrow y_i = X\beta + Zu + e$$

$$L_1^{Pos} \longrightarrow y_i = X\beta + Zu + Wv + e$$

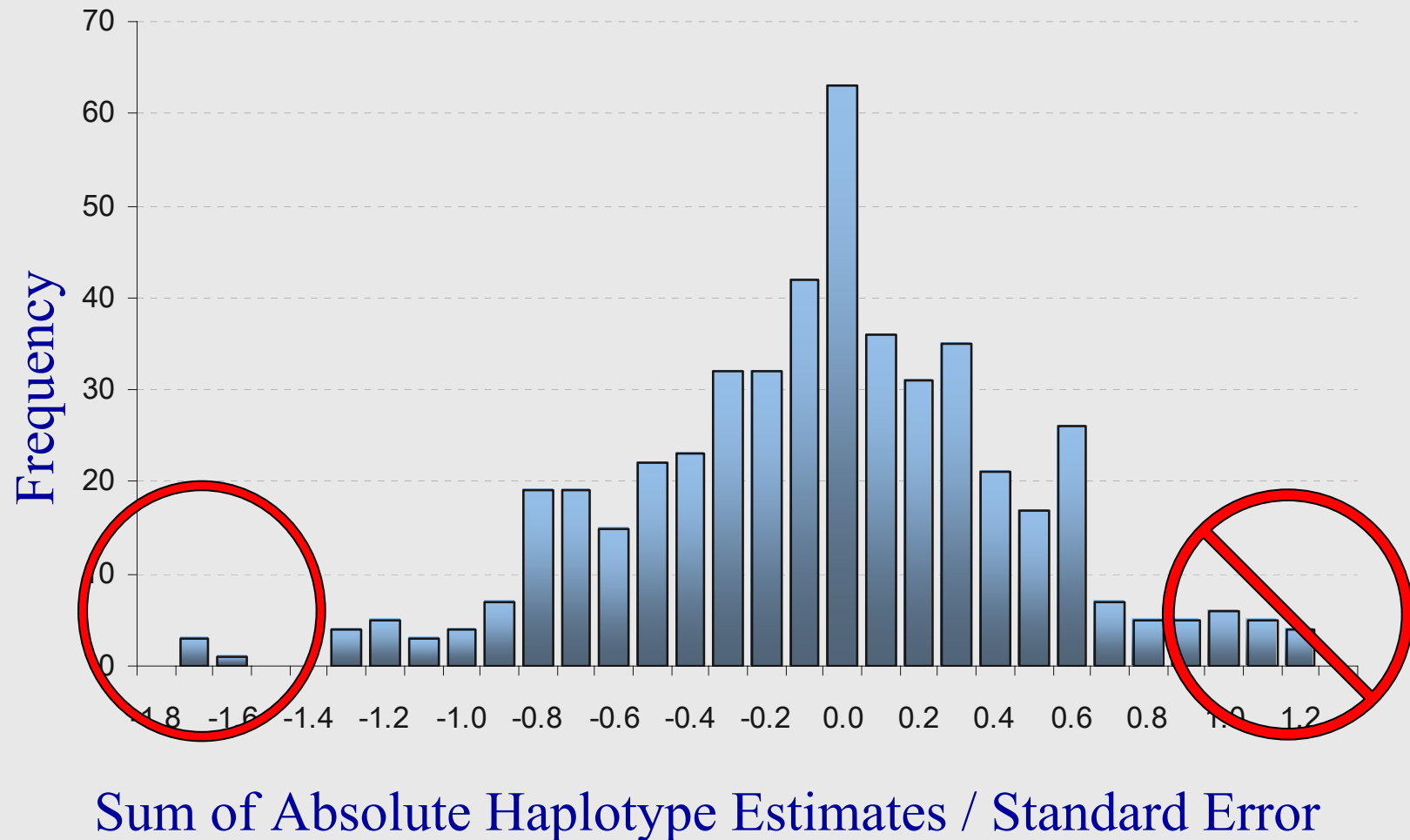
Analysis



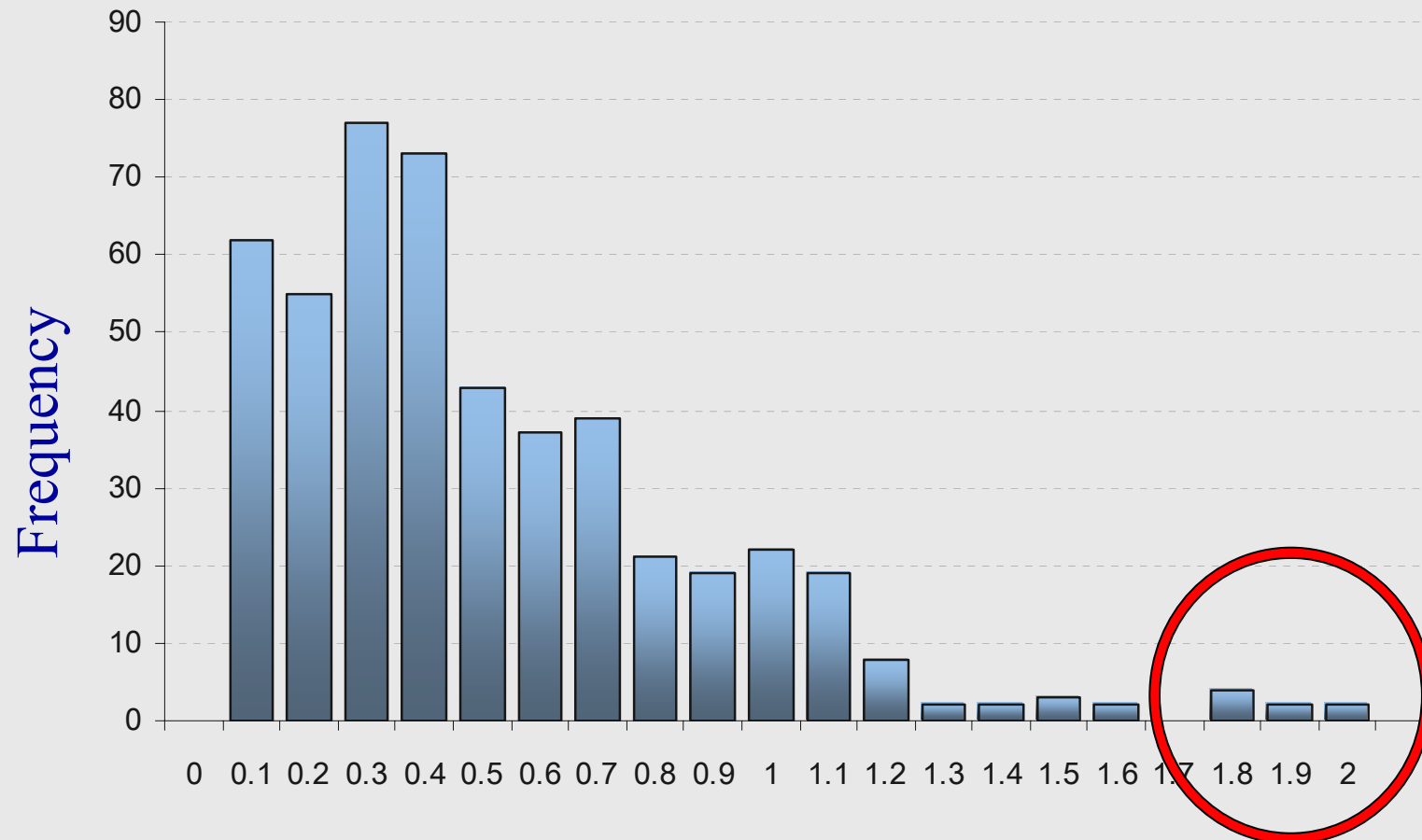
Analysis of Haplotype Effects



Analysis of Haplotype Effects



Analysis of Haplotype Effects



Difference in Absolute Haplotype Estimates / Standard Error

Summary

- Chromosomal region on BTA27 affecting udder health further investigated
- Sharper curves and higher peaks
- Specific haplotypes identified
- Basis for comparative sequencing

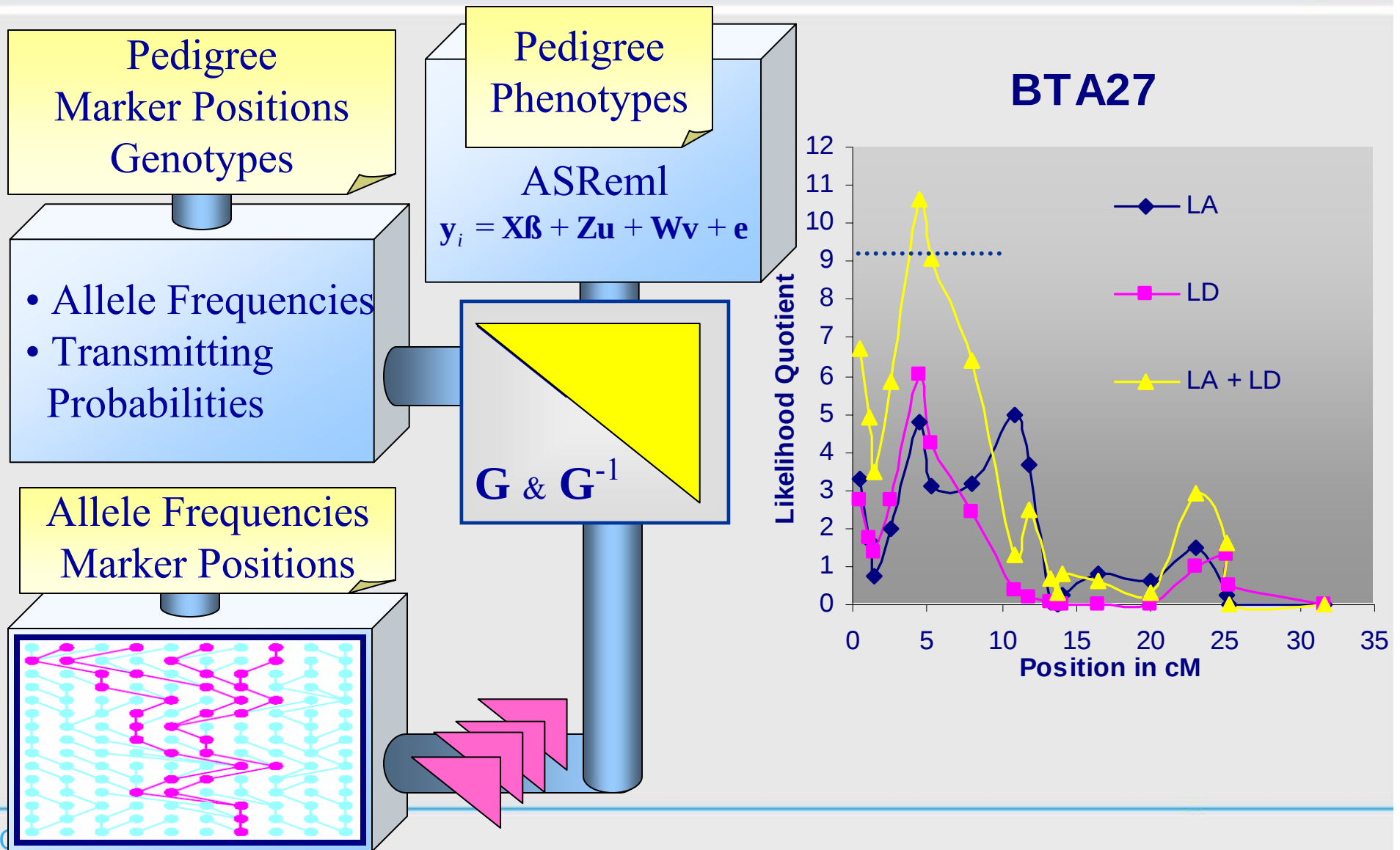


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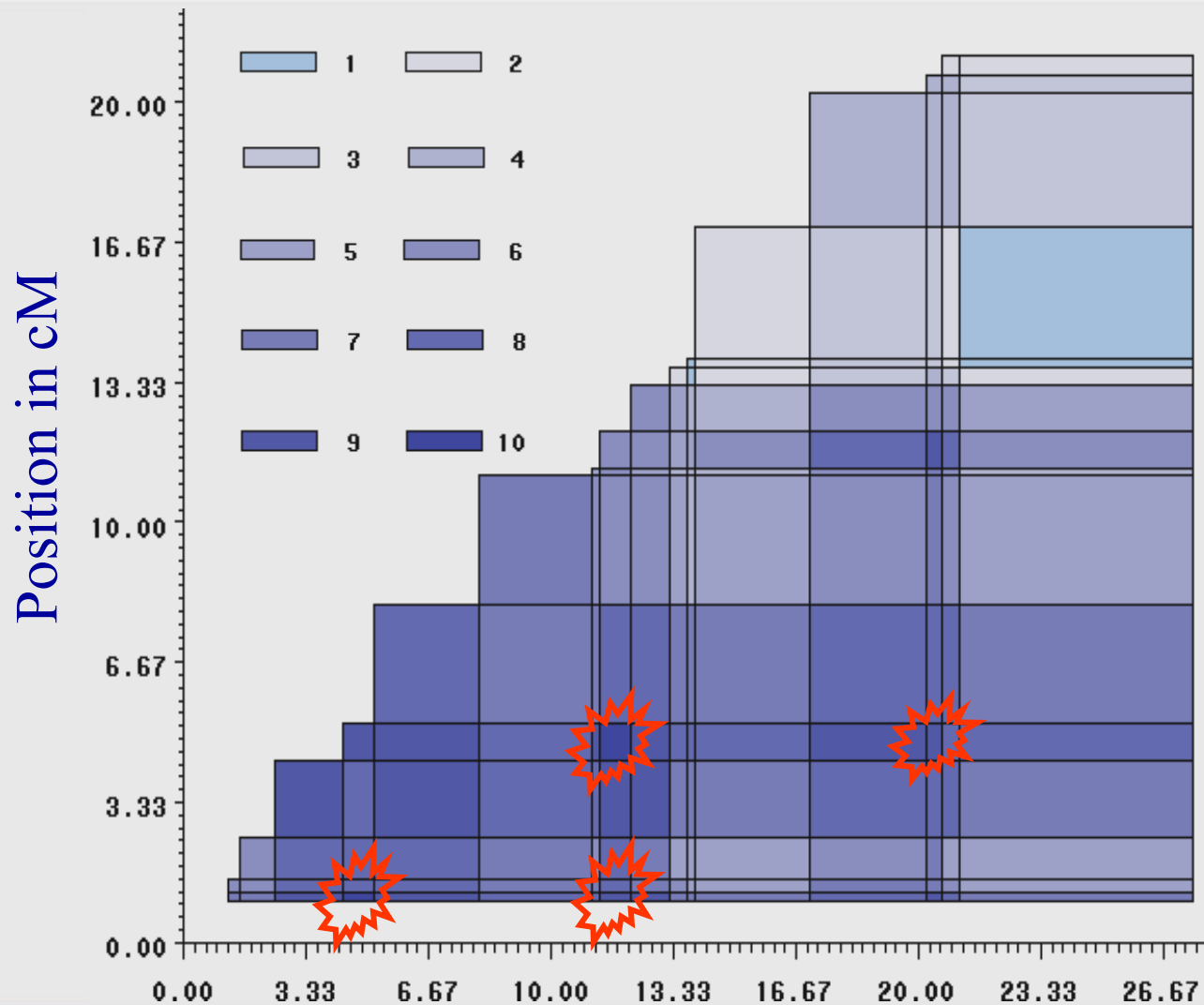


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Analysis



Analysis: 2-QTL Model



Pos1	Pos2	LogL
1.00	4.34	10.78
1.00	12.16	10.62
4.34	12.16	10.70
4.34	20.20	10.08