



# Unraveling the genetic background of $\alpha_{s1}$ - and $\alpha_{s2}$ -casein phosphorylation in Dutch Holstein Friesian

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nederlandse zuivel organisatie



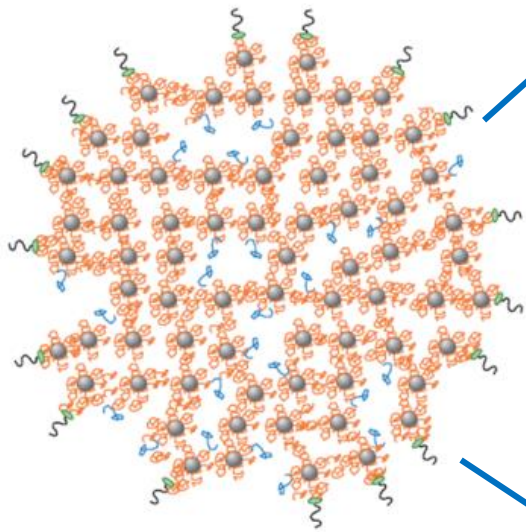
# Acknowledgements

This study was a part of Dutch Milk Genomic Initiative

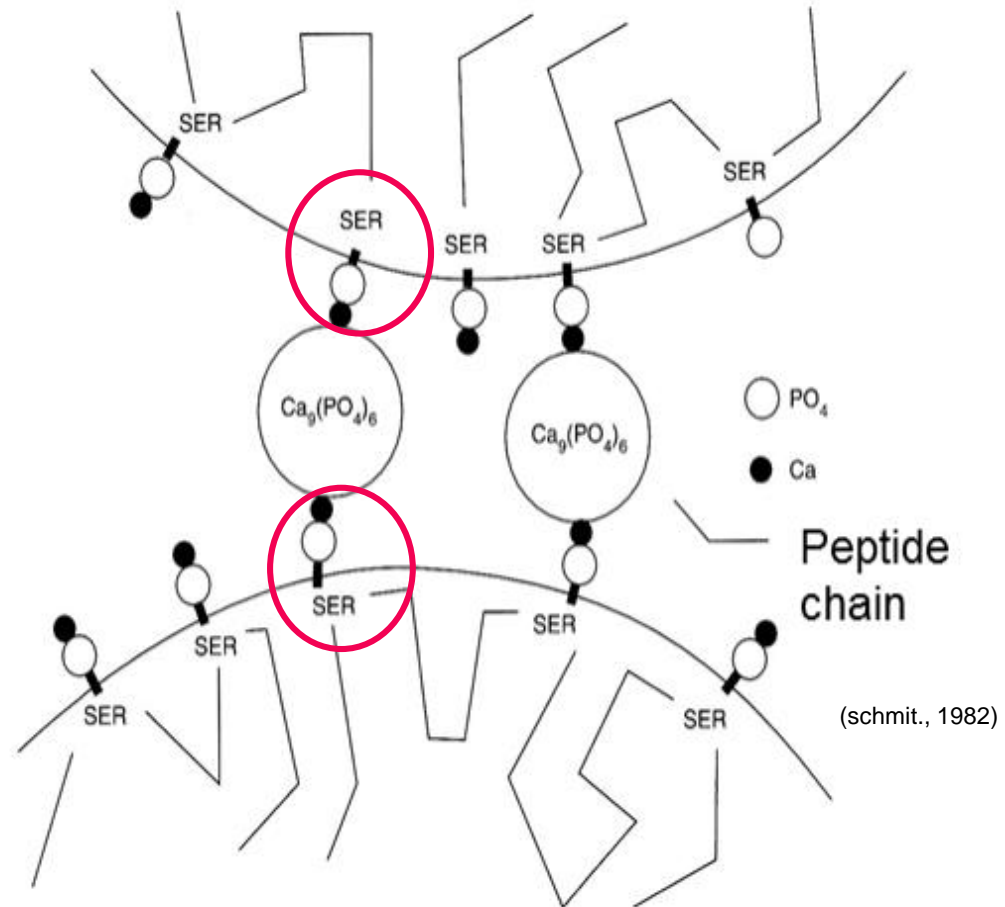
# $\alpha_{s1}$ - and $\alpha_{s2}$ -casein

- Proportion in cow's milk protein (%wt/wt):  
 $\alpha_{s1}$ -CN: ~40%,  $\alpha_{s2}$ -CN: ~10%
- Variation in phosphorylation levels (P)
  - $\alpha_{s1}$ -CN: 8P and 9P
  - $\alpha_{s2}$ -CN: from 10P to 15P

# Casein micelle structure



(Dalglish et al., 2012)





# $\alpha_{S1}$ - and $\alpha_{S2}$ -casein

Trait	Mean	$h^2$
$\alpha_{S1}$ -CN	total	33.64
	8P	21.26
	9P	7.42
$\alpha_{S2}$ -CN	total	6.67
	10P	0.99
	11P	3.44
	12P	2.24
$\alpha_{S1}$ -CN PD	25.79	
$\alpha_{S2}$ -CN PD	34.01	



# $\alpha_{S1}$ - and $\alpha_{S2}$ -casein

Trait	Mean	$h^2$
$\alpha_{S1}$ -CN	total 33.64	0.52 (0.11)
	8P 21.26	0.48 (0.10)
	9P 7.42	0.76 (0.12)
$\alpha_{S2}$ -CN	total 6.67	0.94 (0.12)
	10P 0.99	0.54 (0.11)
	11P 3.44	0.89 (0.12)
	12P 2.24	0.71 (0.12)
$\alpha_{S1}$ -CN PD	25.79	0.78 (0.12)
$\alpha_{S2}$ -CN PD	34.01	0.64 (0.11)

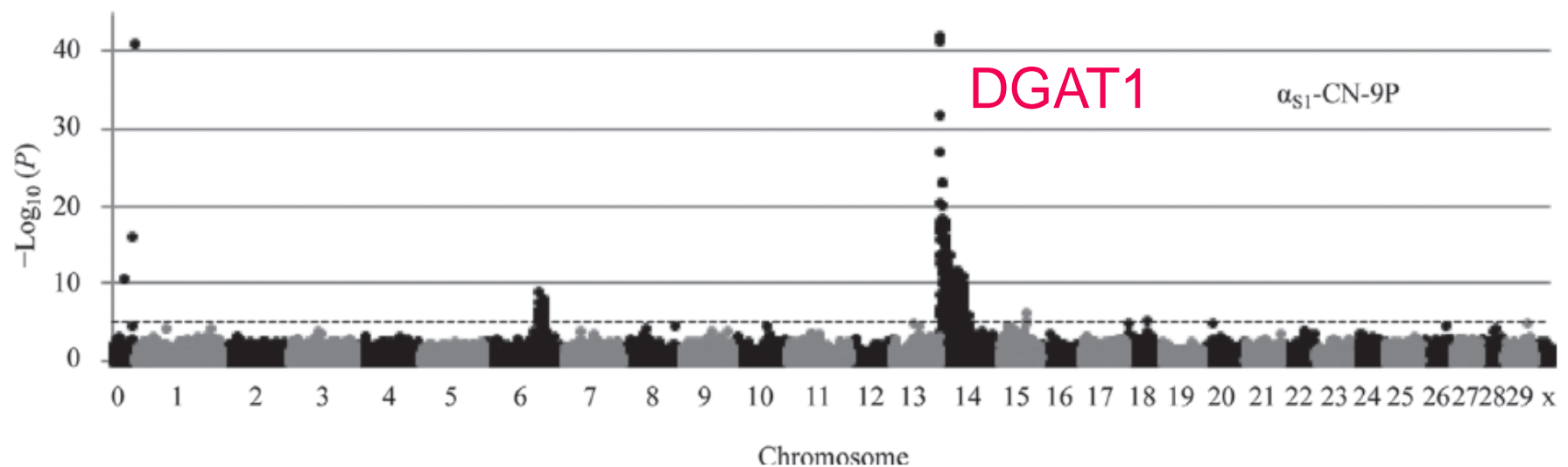
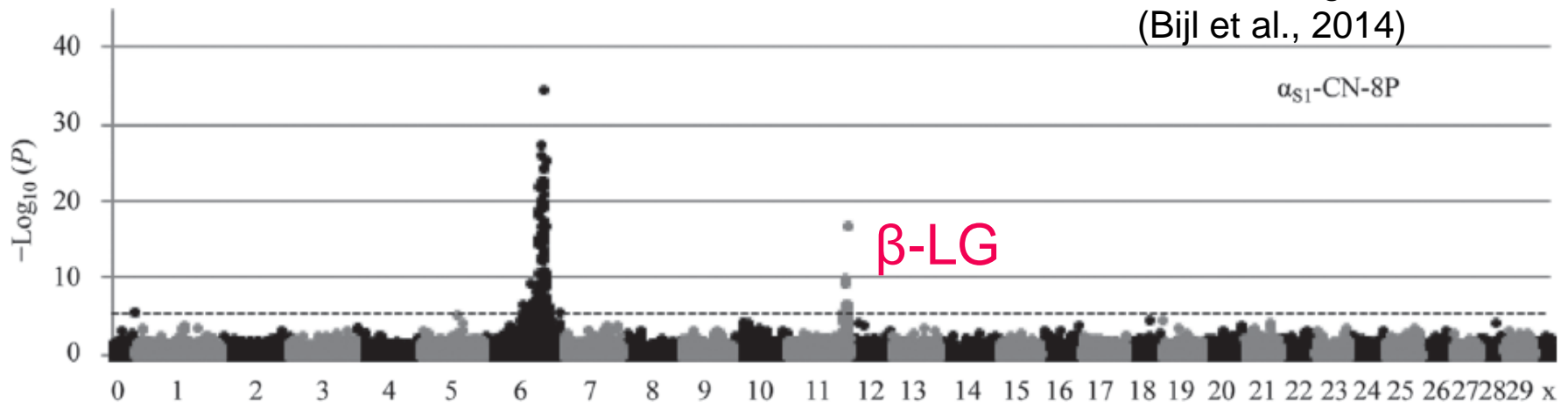
**Phosphorylation degree:**

$\alpha_{S1}$ -CN PD =  $\alpha_{S1}$ -CN-9P/ total  $\alpha_{S1}$ -CN X 100%

$\alpha_{S2}$ -CN PD =  $\alpha_{S2}$ -CN-12P/ total  $\alpha_{S2}$ -CN X 100%

# GWAS for $\alpha_{s1}$ -casein phosphoforms

Dutch milk genomics initiative  
(Bijl et al., 2014)

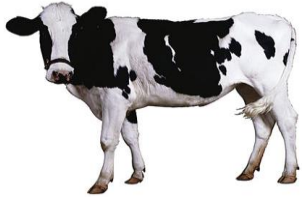


# Objectives

- Investigate the genetic background of  $\alpha_{s2}$ -CN phosphoforms
- Shared genetic control between  $\alpha_{s1}$ -CN PD and  $\alpha_{s2}$ -CN PD?



# Phenotypes



~2000 primiparous, mid-late lactation  
from ~400 herds

- Relative concentrations of  $\alpha_{s2}$ -CN-10P,  $\alpha_{s2}$ -CN-11P and  $\alpha_{s2}$ -CN-12P (determined by capillary zone electrophoresis)
- Phosphorylation degree of  $\alpha_{s1}$ -CN and  $\alpha_{s2}$ -CN ( $\alpha_{s1}$ -CN PD and  $\alpha_{s2}$ -CN PD)

# Genotypes

- 50K SNP chip

# Single-SNP associations

$$y = \beta_1 \text{dim} + \beta_2 e^{-0.05 \cdot \text{dim}} + \beta_3 \text{ca} + \beta_4 \text{ca}^2 + \text{season} + \text{scode} + \text{SNP} + \text{herd} + a + e$$

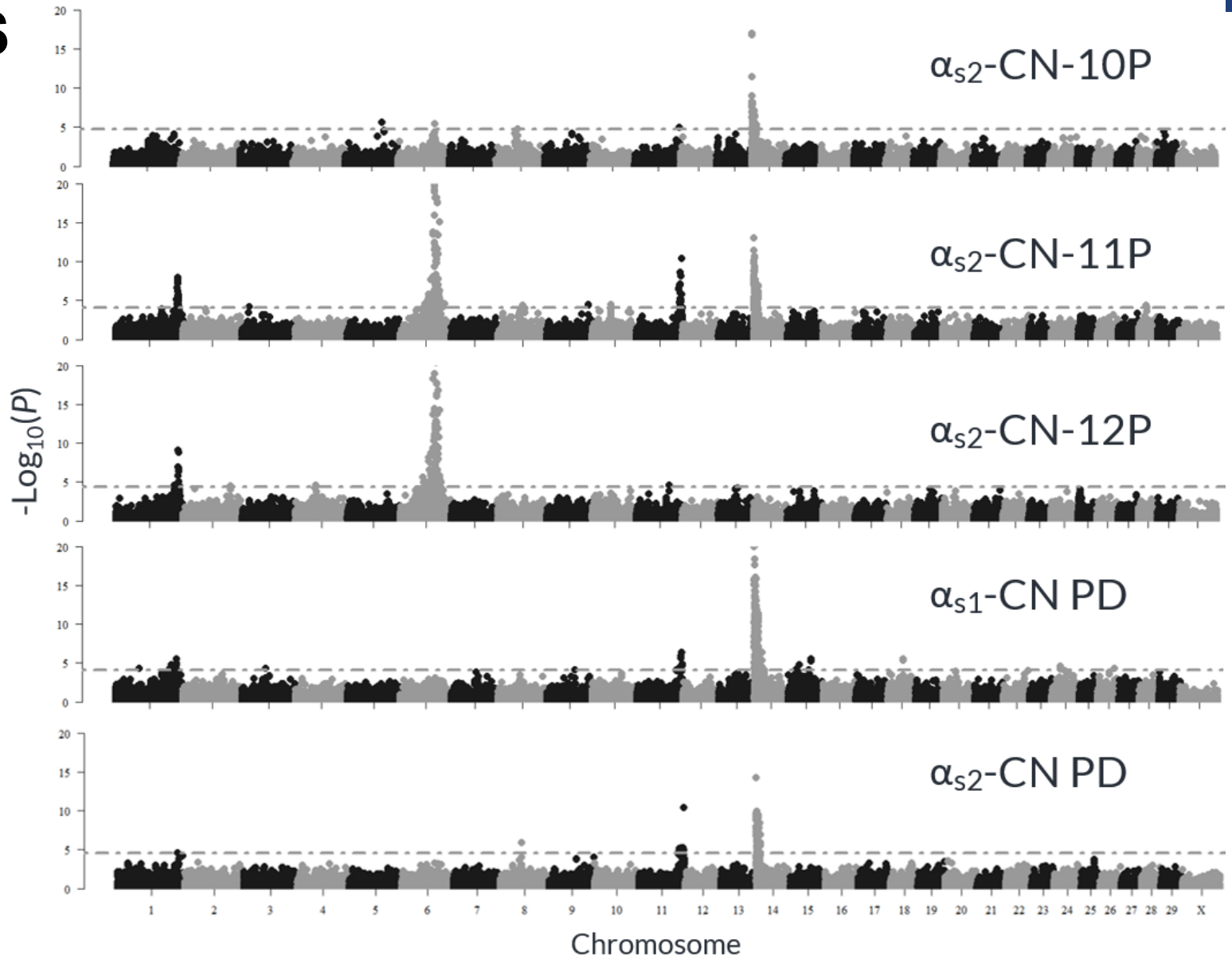
**Fixed effects**

- dim**: days in milk
- ca**: age at the first calving
- season**: calving season
- scode**: proven vs young bulls
- SNP**: effect of SNP genotypes

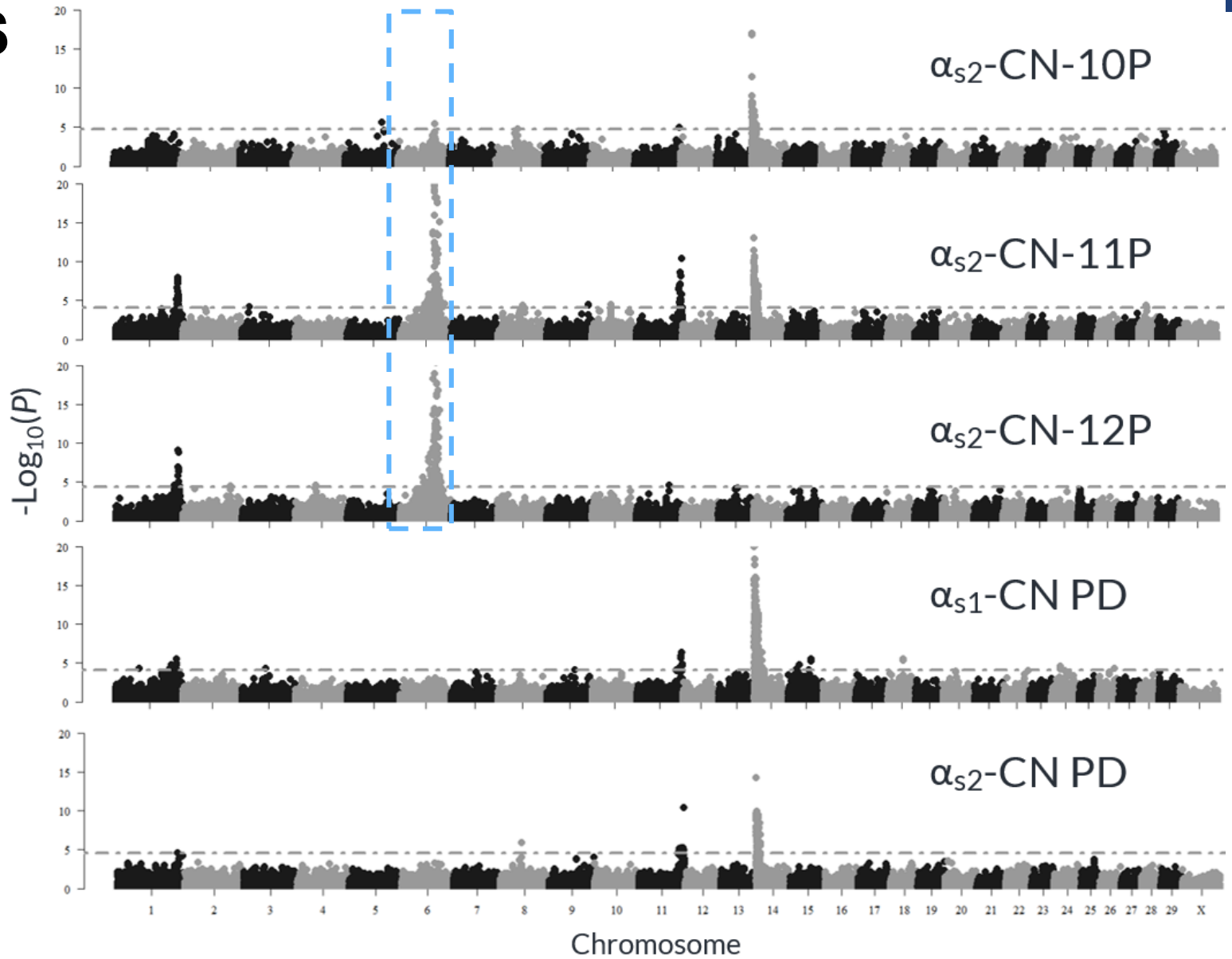
**Random effects**

- herd**: effect of herd
- a**: additive genetic effect
- e**: residual

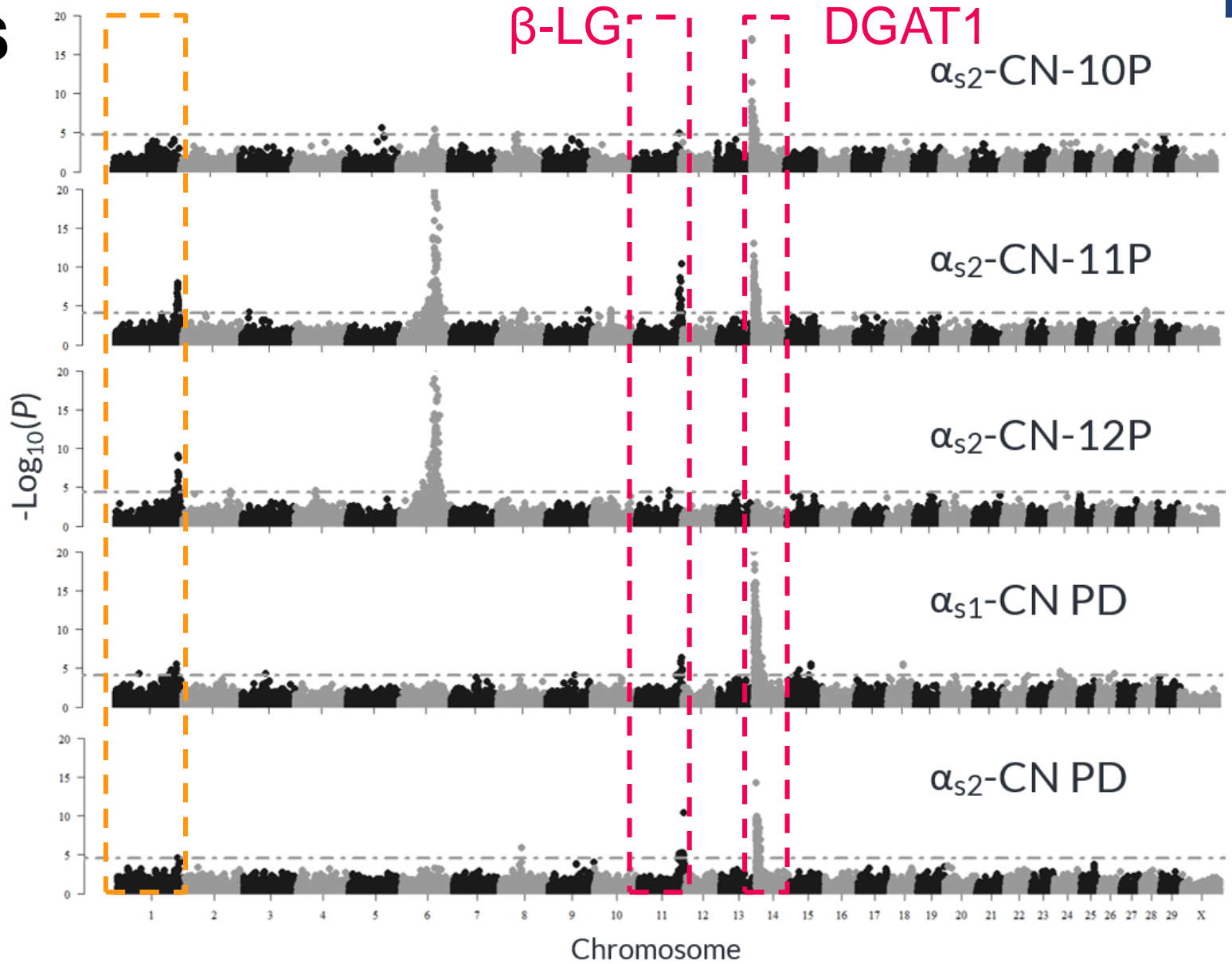
## GWAS



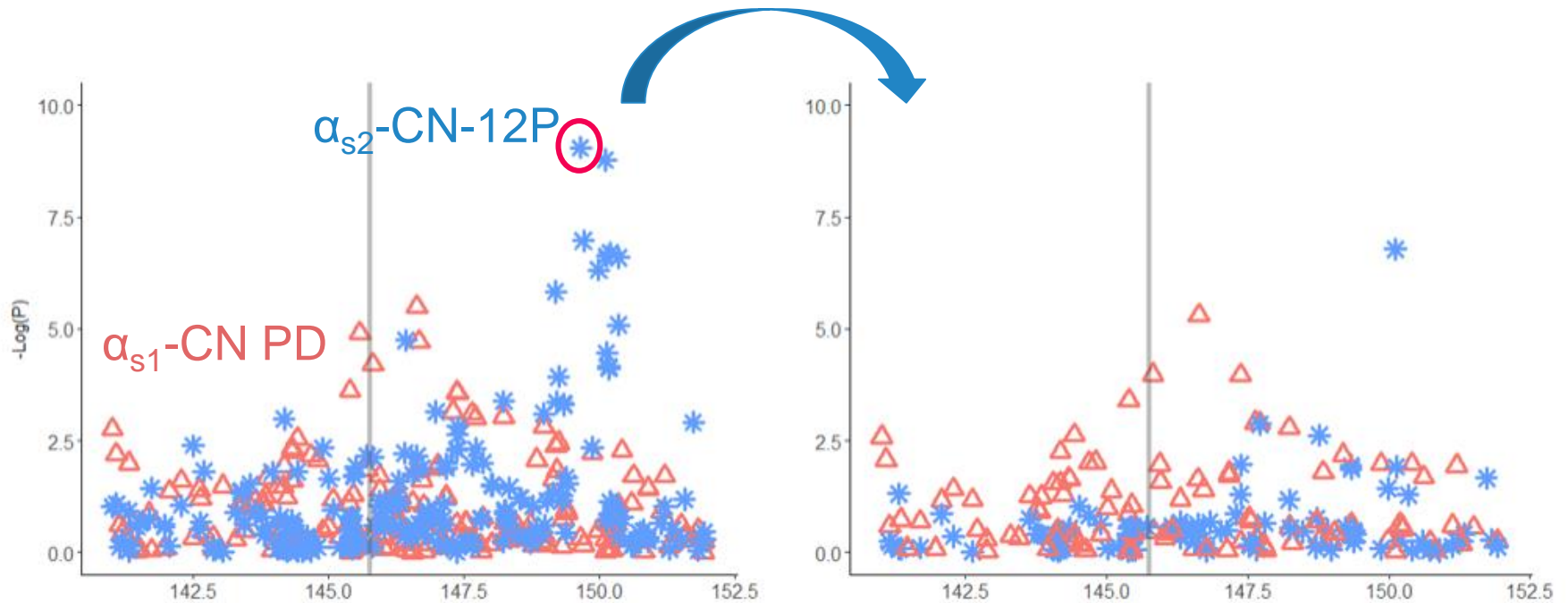
## GWAS



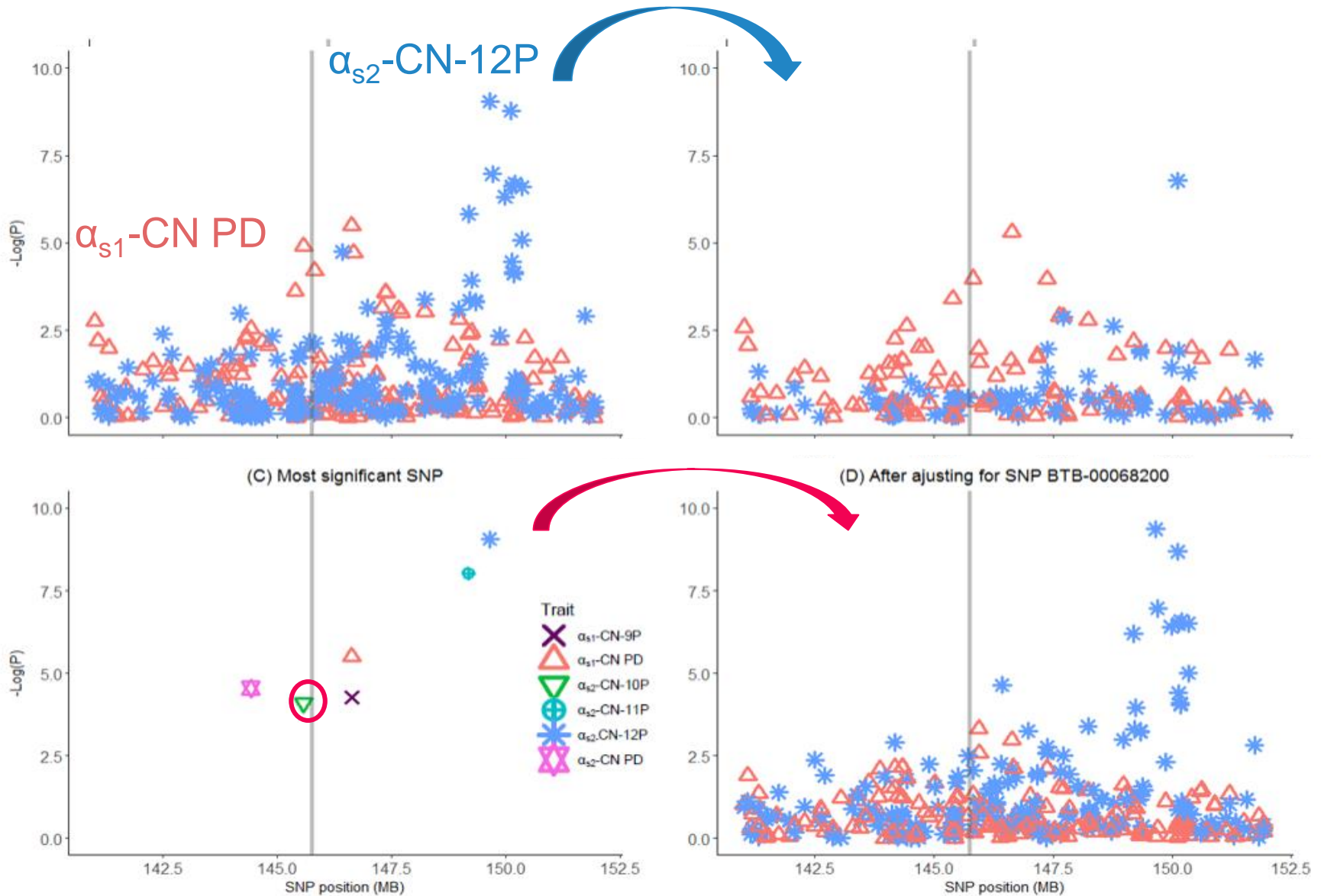
## GWAS



# QTL on chromosome 1



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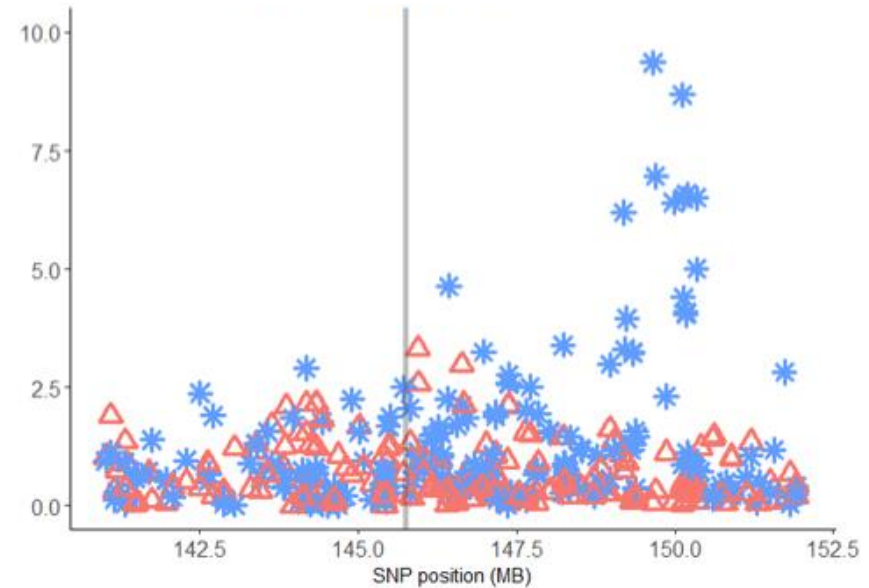
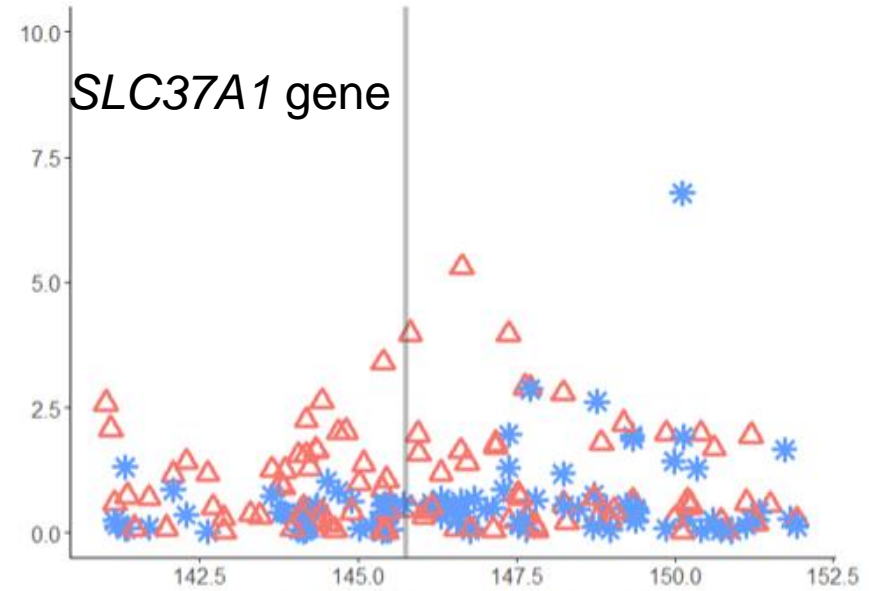
$\alpha_{s1}$ -CN PD  
&  
 $\alpha_{s2}$ -CN PD



2 QTL



$\alpha_{s2}$ -CN production





# Conclusions

- Three genomic regions affect phosphorylation degree of  $\alpha_{s1}$ - and  $\alpha_{s2}$ -CN:

chromosome1 : SLC37A1 (translocate phosphorus)

chromosome11 :  $\beta$ -LG

chromosome14 : DGAT1

- Actual roles of these proteins need further investigation