

# *Celsr1* Cas9-CKO Strategy

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**Reviewer:**

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**Design Date:**

**2018-6-28**

# Project Overview

**Project Name**

*Celsr1*

**Project type**

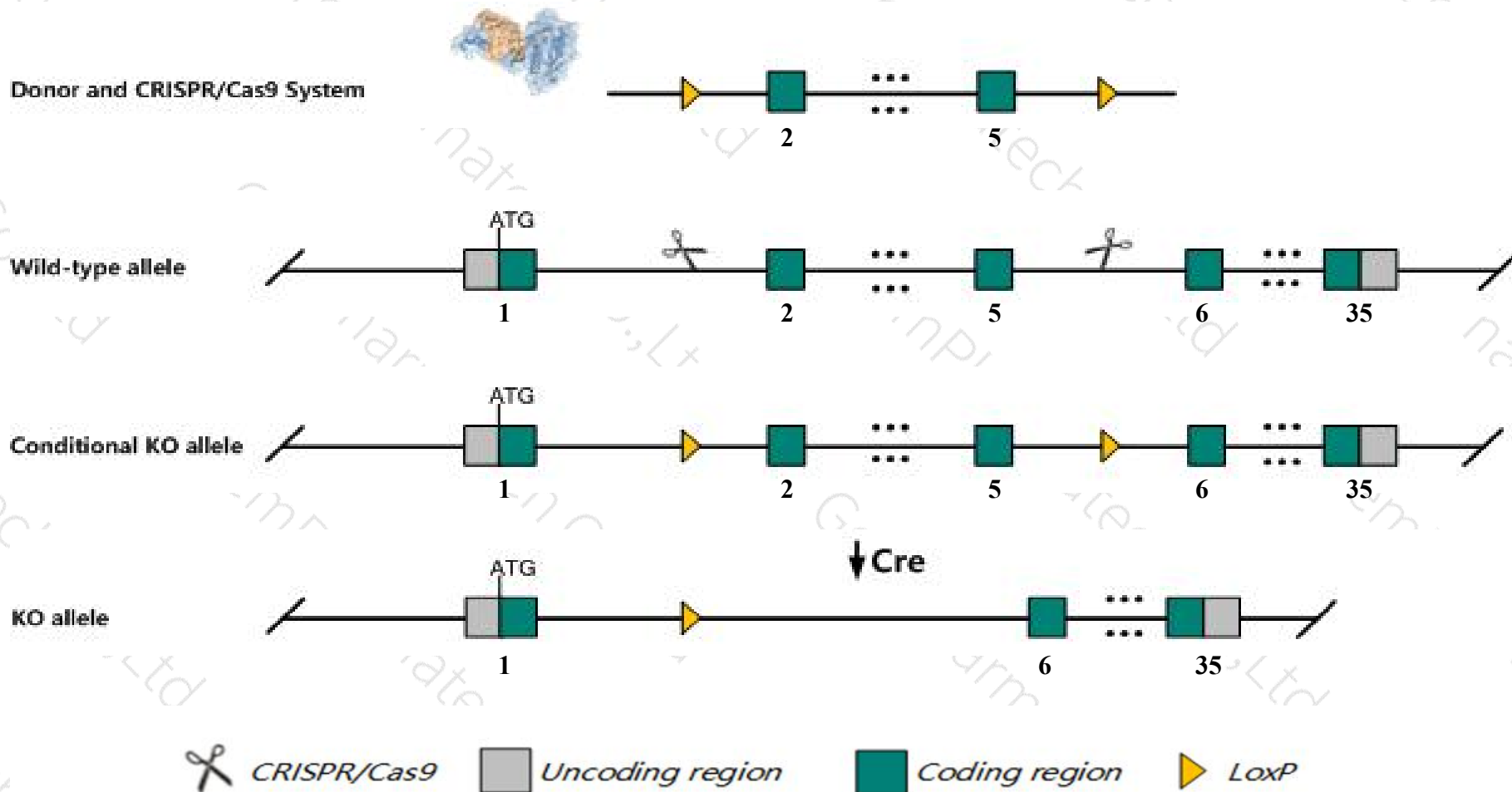
**Cas9-CKO**

**Strain background**

**C57BL/6JGpt**

# Conditional Knockout strategy

This model will use CRISPR/Cas9 technology to edit the *Celsr1* gene. The schematic diagram is as follows:



# Technical routes

- The *Celsr1* gene has 4 transcripts. According to the structure of *Celsr1* gene, exon2-exon5 of *Celsr1-201* (ENSMUST00000016172.8) transcript is recommended as the knockout region. The region contains 1067bp coding sequence. Knock out the region will result in disruption of protein function.
- In this project we use CRISPR/Cas9 technology to modify *Celsr1* gene. The brief process is as follows: CRISPR/Cas9 system and Donor were microinjected into the fertilized eggs of C57BL/6JGpt mice. Fertilized eggs were transplanted to obtain positive F0 mice which were confirmed by PCR and sequencing. A stable F1 generation mouse model was obtained by mating positive F0 generation mice with C57BL/6JGpt mice.
- The flox mice will be knocked out after mating with mice expressing Cre recombinase, resulting in the loss of function of the target gene in specific tissues and cell types.

- According to the existing MGI data, Nullizygous mice exhibit kinky tails, variable neural tube defects, abnormal hair follicle orientation, whorl-like hair patterns, and partial prenatal lethality. ENU-induced mutants show defects in planar polarity of inner ear hair cells and complete perinatal lethality due to craniorachischisis.
- The *Celsr1* gene is located on the Chr15. If the knockout mice are crossed with other mice strains to obtain double gene positive homozygous mouse offspring, please avoid the two genes on the same chromosome.
- This Strategy is designed based on genetic information in existing databases. Due to the complexity of biological processes, all risk of loxp insertion on gene transcription, RNA splicing and protein translation cannot be predicted at existing technological level.



# Gene information (NCBI)

## Celsr1 cadherin, EGF LAG seven-pass G-type receptor 1 [ *Mus musculus* (house mouse) ]

Gene ID: 12614, updated on 12-Aug-2019

### Summary



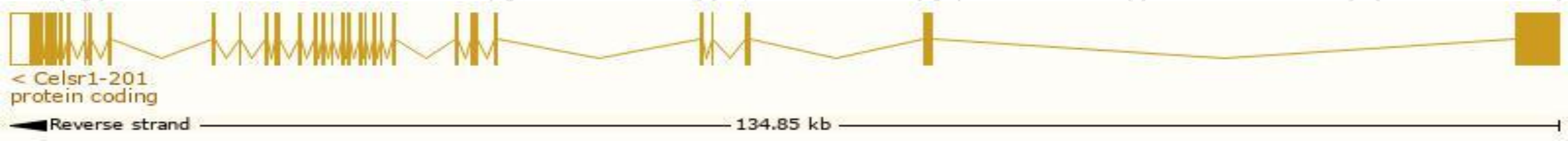
|                           |                                                                                                                                                                           |
|---------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <b>Official Symbol</b>    | Celsr1 provided by <a href="#">MGI</a>                                                                                                                                    |
| <b>Official Full Name</b> | cadherin, EGF LAG seven-pass G-type receptor 1 provided by <a href="#">MGI</a>                                                                                            |
| <b>Primary source</b>     | <a href="#">MGI:MGI:1100883</a>                                                                                                                                           |
| <b>See related</b>        | <a href="#">Ensembl:ENSMUSG00000016028</a>                                                                                                                                |
| <b>Gene type</b>          | protein coding                                                                                                                                                            |
| <b>RefSeq status</b>      | VALIDATED                                                                                                                                                                 |
| <b>Organism</b>           | <a href="#">Mus musculus</a>                                                                                                                                              |
| <b>Lineage</b>            | Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha; Muroidea; Muridae; Murinae; Mus; Mus |
| <b>Also known as</b>      | Scy; Crsh; crash; Adgrc1                                                                                                                                                  |
| <b>Expression</b>         | Broad expression in colon adult (RPKM 9.9), ovary adult (RPKM 9.8) and 18 other tissues <a href="#">See more</a>                                                          |
| <b>Orthologs</b>          | <a href="#">human</a> <a href="#">all</a>                                                                                                                                 |

# Transcript information (Ensembl)

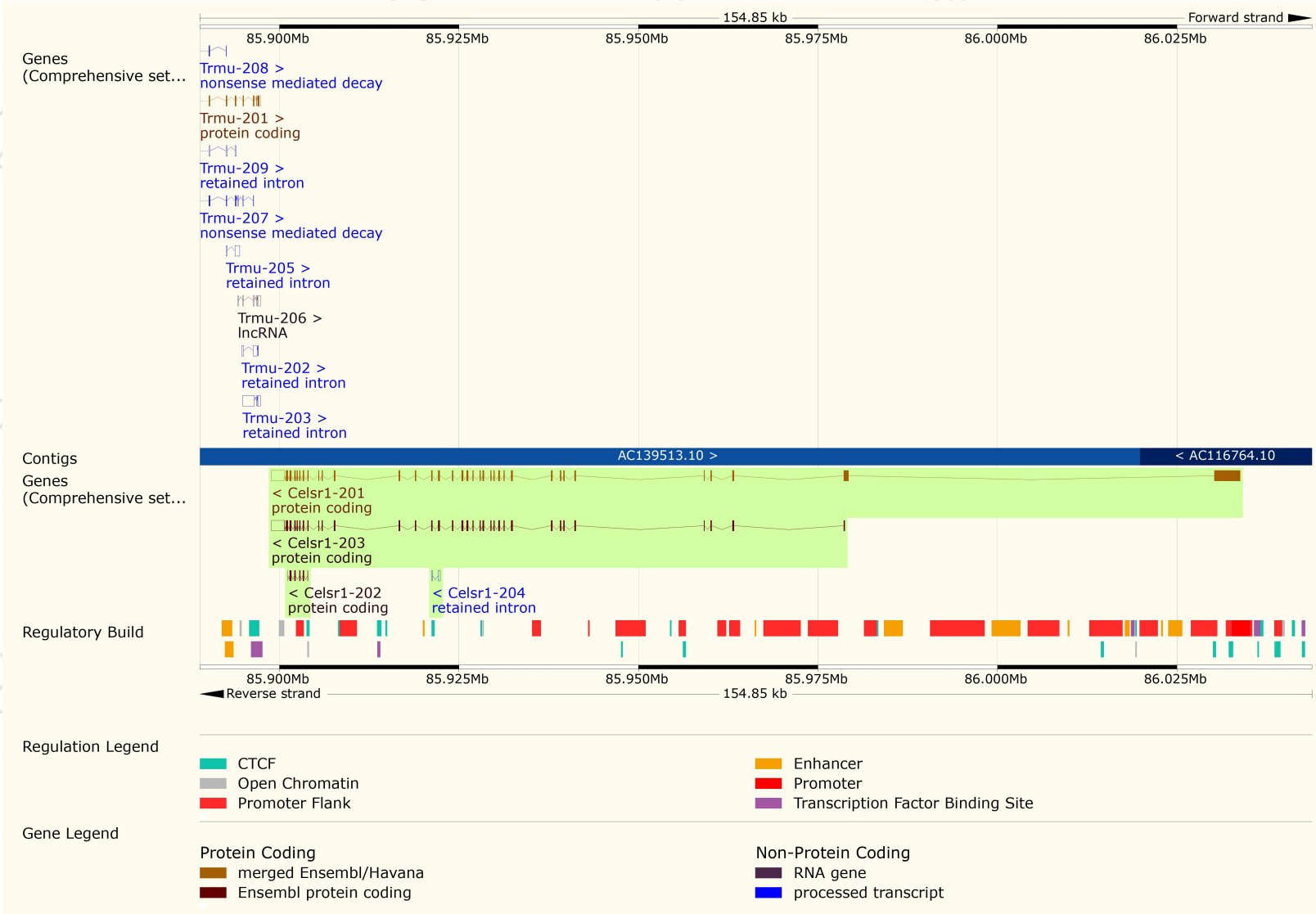
The gene has 4 transcripts,all transcripts are shown below:

| Name       | Transcript ID                        | bp    | Protein                | Biotype         | CCDS                      | UniProt                    | Flags                                                                                                                         |
|------------|--------------------------------------|-------|------------------------|-----------------|---------------------------|----------------------------|-------------------------------------------------------------------------------------------------------------------------------|
| Celsr1-201 | <a href="#">ENSMUST00000016172.8</a> | 10879 | <a href="#">3034aa</a> | Protein coding  | <a href="#">CCDS37172</a> | <a href="#">Q35161</a>     | TSL:1 GENCODE basic APPRIS P1                                                                                                 |
| Celsr1-203 | <a href="#">ENSMUST00000226840.1</a> | 6802  | <a href="#">1672aa</a> | Protein coding  | -                         | <a href="#">A0A2I3BPU9</a> | CDS 5' incomplete                                                                                                             |
| Celsr1-202 | <a href="#">ENSMUST00000226204.1</a> | 646   | <a href="#">216aa</a>  | Protein coding  | -                         | <a href="#">A0A2I3BQH0</a> | 5' and 3' truncations in transcript evidence prevent annotation of the start and the end of the CDS. CDS 5' and 3' incomplete |
| Celsr1-204 | <a href="#">ENSMUST00000227450.1</a> | 377   | No protein             | Retained intron | -                         | -                          |                                                                                                                               |

The strategy is based on the design of *Celsr1-201* transcript,The transcription is shown below

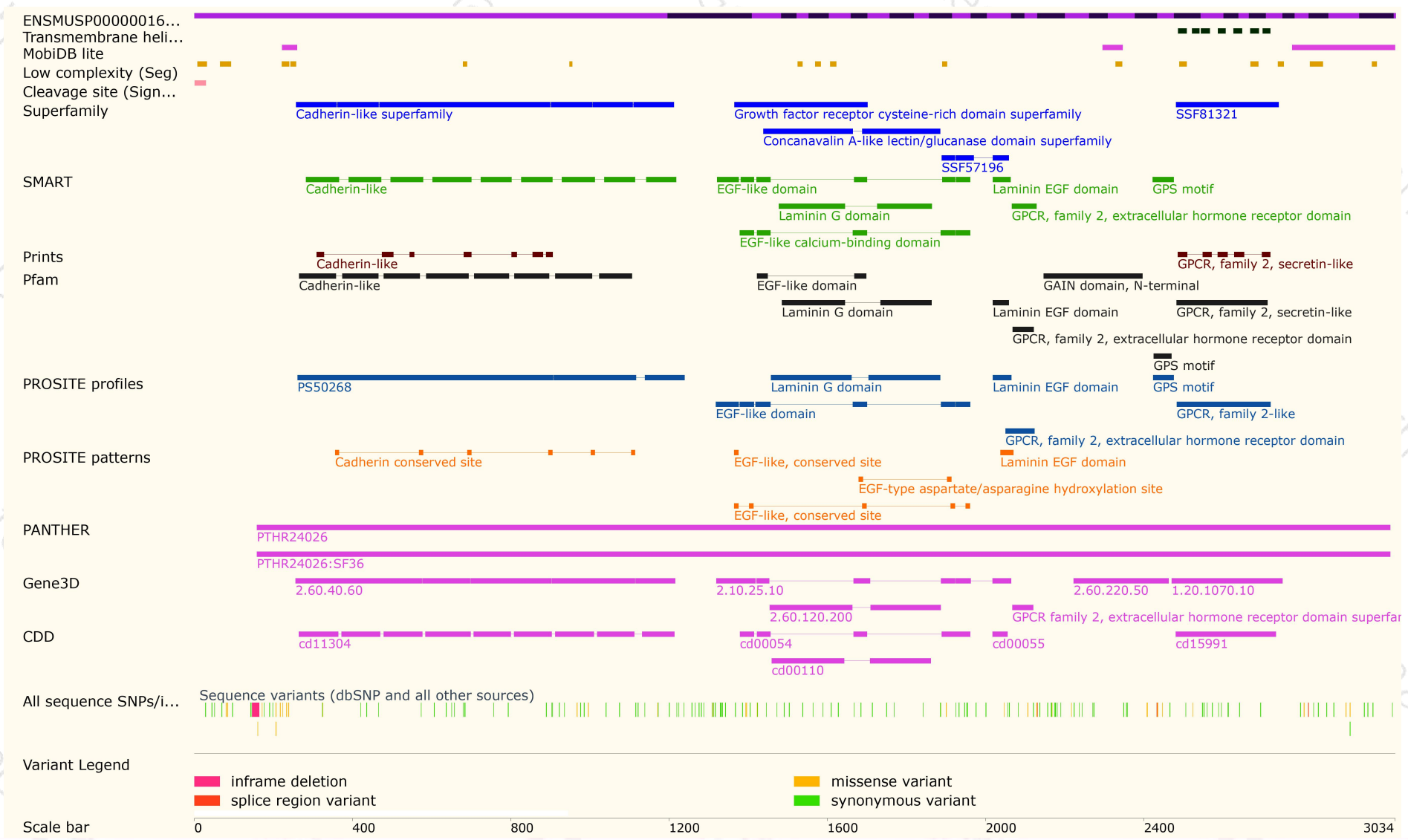


# Genomic location distribution

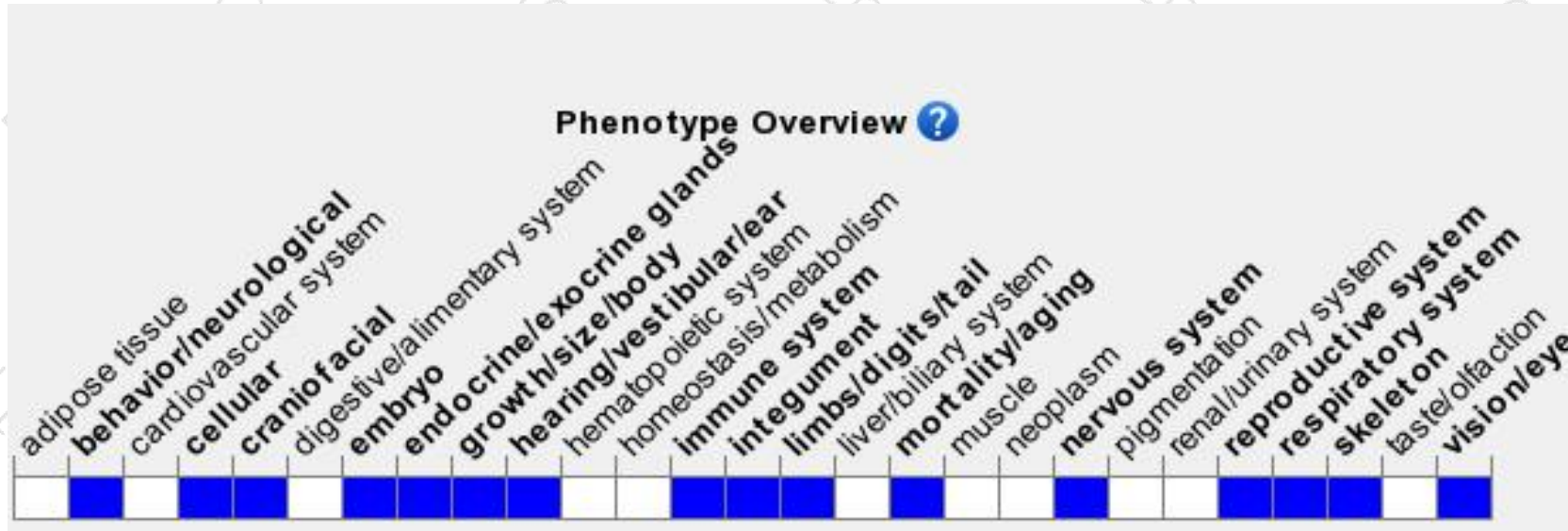




# Protein domain



# Mouse phenotype description(MGI )



*Phenotypes affected by the gene are marked in blue. Data quoted from MGI database(<http://www.informatics.jax.org/>).*

According to the existing MGI data, Nullizygous mice exhibit kinky tails, variable neural tube defects, abnormal hair follicle orientation, whorl-like hair patterns, and partial prenatal lethality. ENU-induced mutants show defects in planar polarity of inner ear hair cells and complete perinatal lethality due to craniorachischisis.

If you have any questions, you are welcome to inquire.

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