

Zdhhc8 Cas9-CKO Strategy

Designer: Xueting Zhang

Reviewer: Yanhua Shen

Design Date: 2020-2-6

Project Overview

Project Name

Zdhhc8

Project type

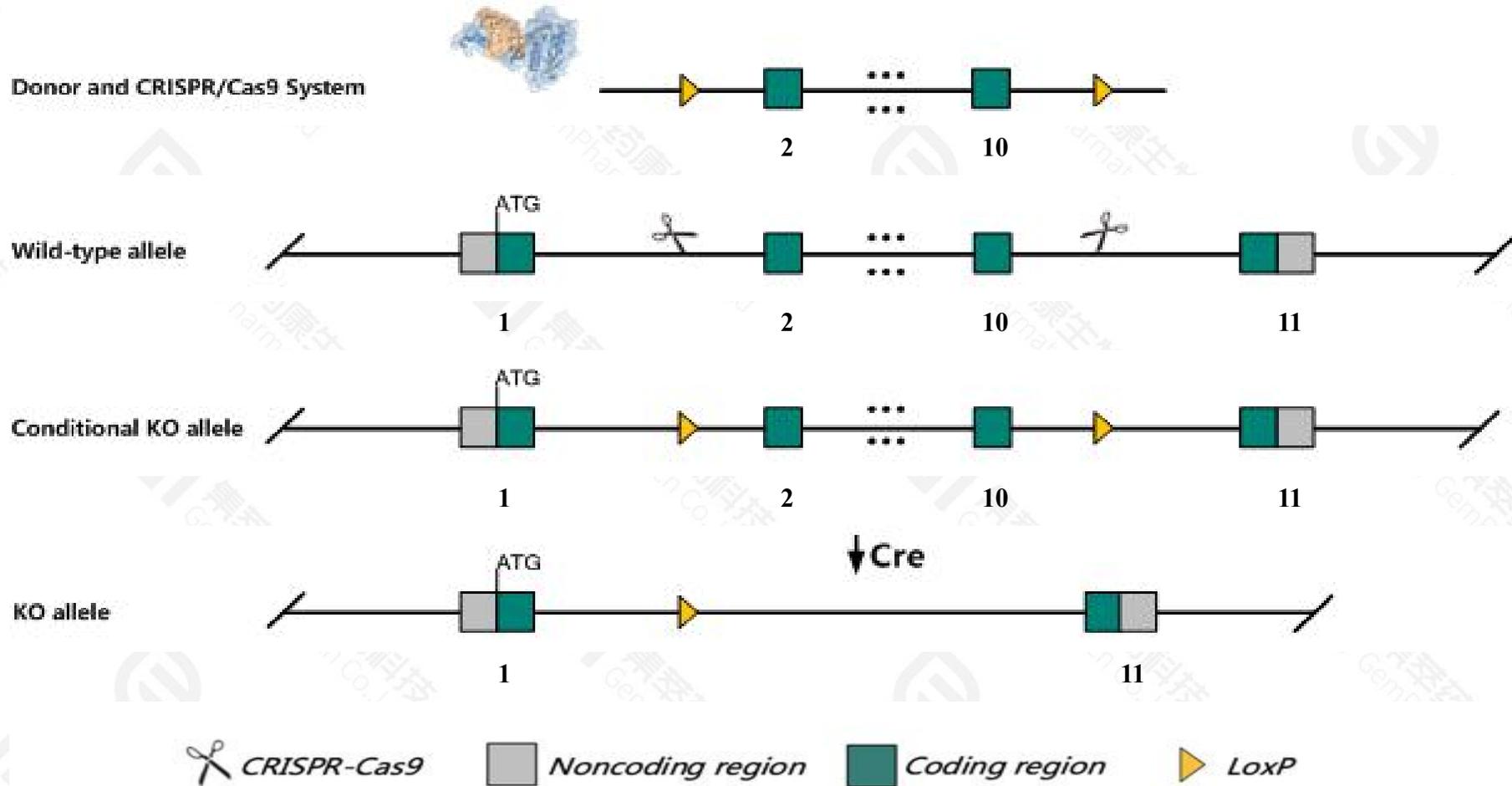
Cas9-CKO

Strain background

C57BL/6JGpt

Conditional Knockout strategy

This model will use CRISPR-Cas9 technology to edit the *Zdhhc8* gene. The schematic diagram is as follows:



- The *Zdhhc8* gene has 2 transcripts. According to the structure of *Zdhhc8* gene, exon2-exon10 of *Zdhhc8-201*(ENSMUST00000076957.7) transcript is recommended as the knockout region. The region contains most of the coding sequence. Knock out the region will result in disruption of protein function.
- In this project we use CRISPR-Cas9 technology to modify *Zdhhc8* 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, homozygous null females display impaired prepulse inhibition and reduced exploration in new environments. Homozygous null males display normal prepulse inhibition and only a slight decrease in exploration.
- The effect on transcript *Zdhhc8-202* is unknown.
- The *Zdhhc8* gene is located on the Chr16. 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)

Zdhhc8 zinc finger, DHHC domain containing 8 [Mus musculus (house mouse)]

Gene ID: 27801, updated on 31-Jan-2019

Summary



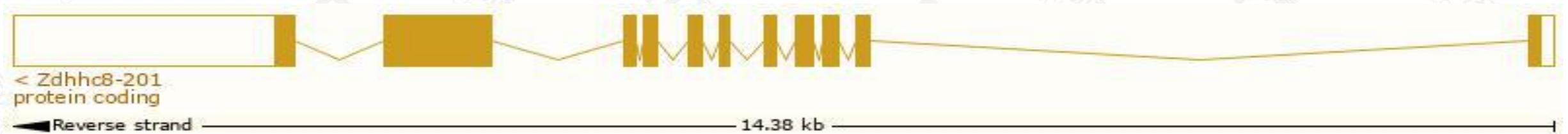
| | |
|---------------------------|---|
| Official Symbol | Zdhhc8 provided by MGI |
| Official Full Name | zinc finger, DHHC domain containing 8 provided by MGI |
| Primary source | MGI:MGI:1338012 |
| See related | Ensembl:ENSMUSG00000060166 |
| Gene type | protein coding |
| RefSeq status | VALIDATED |
| Organism | Mus musculus |
| Lineage | Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha; Muroidea; Muridae; Murinae; Mus; Mus |
| Also known as | D16H22S1738E, DHHC-8, E330009O14Rik, Op53c05, ZDHHCL1 |
| Expression | Ubiquitous expression in thymus adult (RPKM 67.5), ovary adult (RPKM 44.5) and 24 other tissues See more |
| Orthologs | human all |

Transcript information (Ensembl)

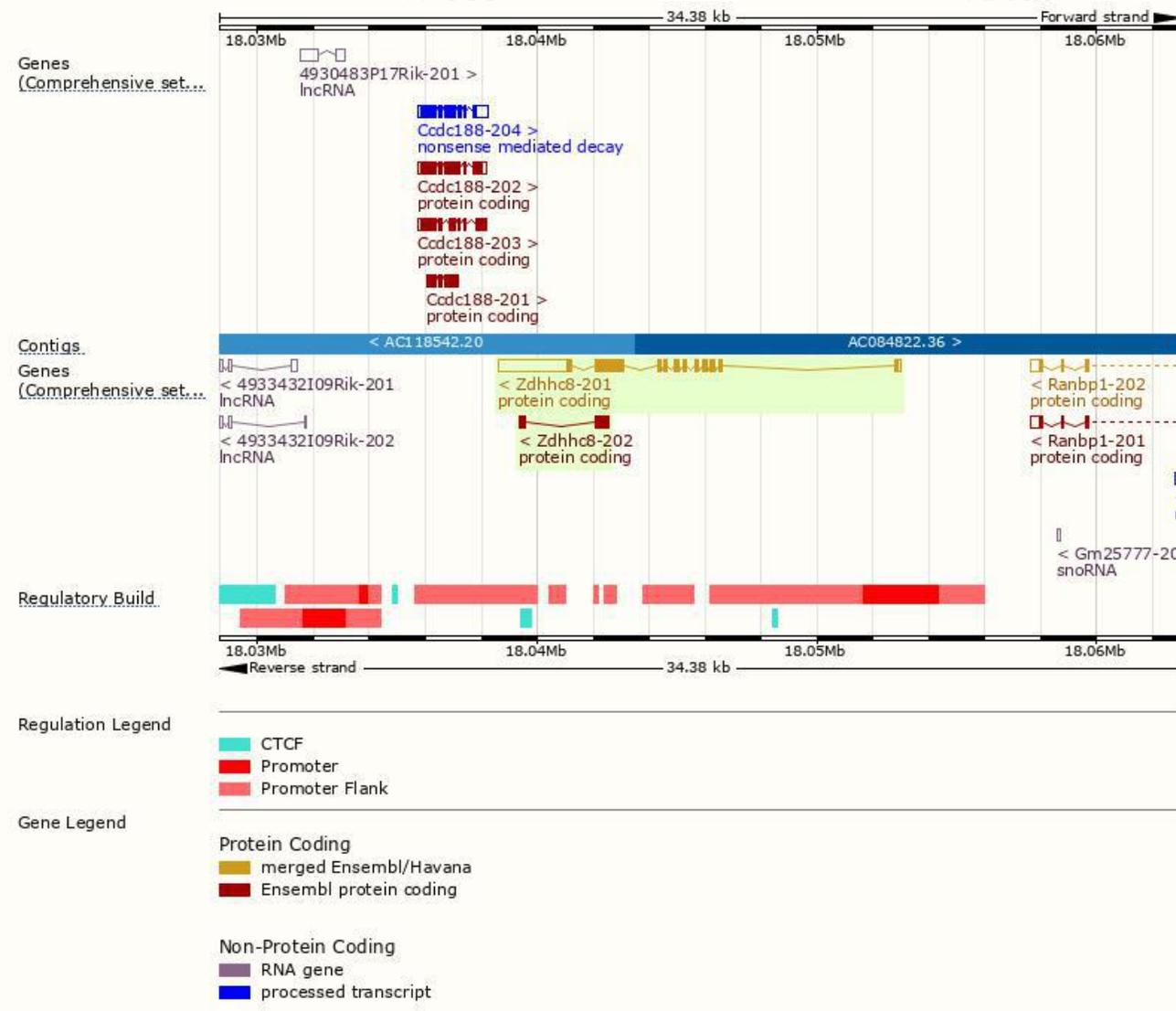
The gene has 2 transcripts, all transcripts are shown below:

| Name | Transcript ID | bp | Protein | Biotype | CCDS | UniProt | Flags |
|------------|--------------------------------------|------|-----------------------|----------------|---------------------------|----------------------------|---|
| Zdhhc8-201 | ENSMUST00000076957.6 | 4868 | 762aa | Protein coding | CCDS28017 | Q5Y5T5 | TSL:1 GENCODE basic APPRIS P1 |
| Zdhhc8-202 | ENSMUST00000231412.1 | 652 | 218aa | Protein coding | - | A0A338P749 | 5' and 3' truncations in transcript evidence prevent annotation of the start and the end of the CDS. CDS 5' and 3' incomplete |

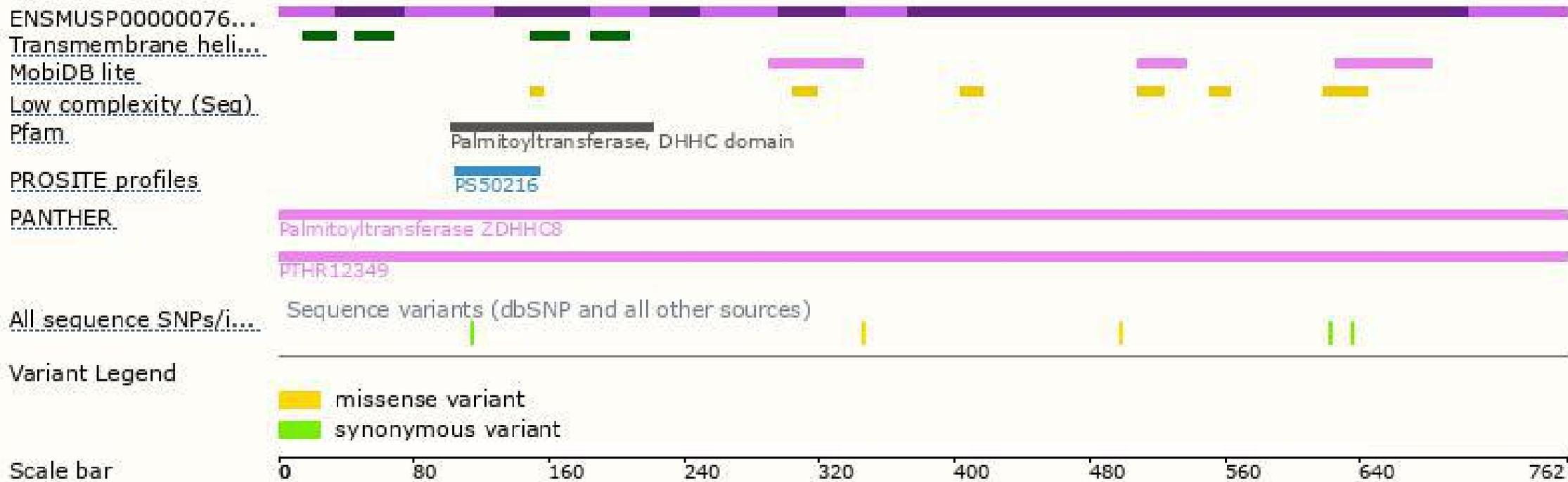
The strategy is based on the design of *Zdhhc8-201* transcript, the transcription is shown below:



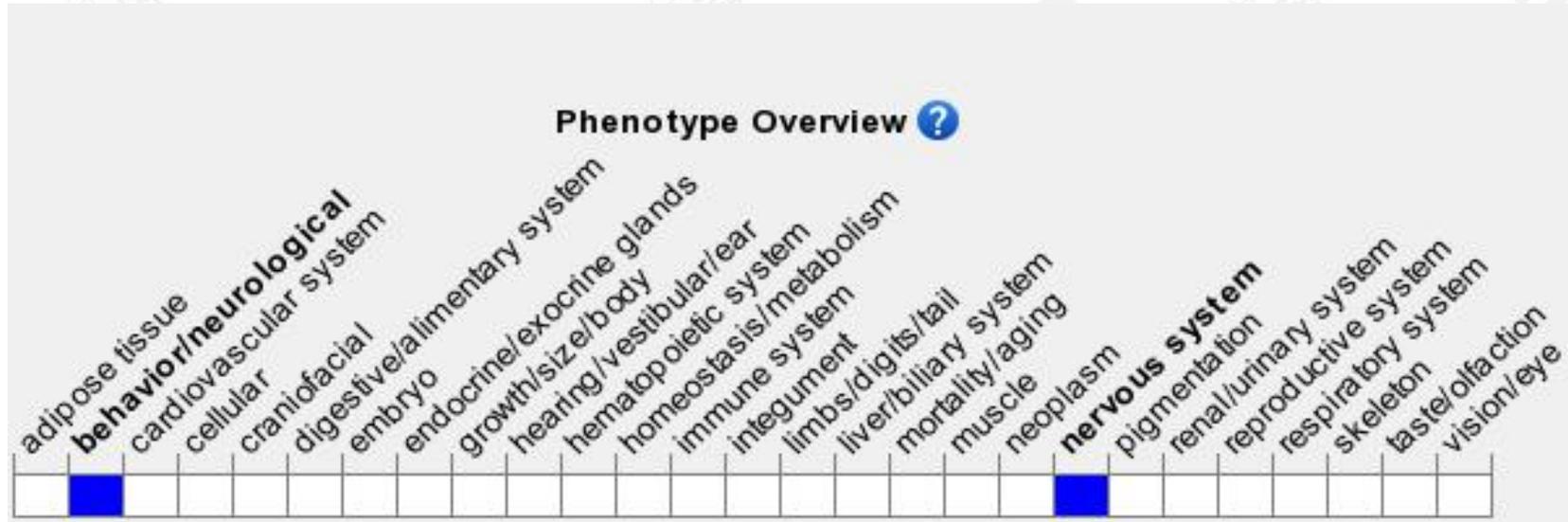
Genomic location distribution



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, homozygous null females display impaired prepulse inhibition and reduced exploration in new environments. Homozygous null males display normal prepulse inhibition and only a slight decrease in exploration.

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

Tel: 400-9660890

