

# Dsg2 Cas9-CKO Strategy

Designer: Reviewer:

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**Design Date:** 

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## **Project Overview**



**Project Name** 

Dsg2

**Project type** 

Cas9-CKO

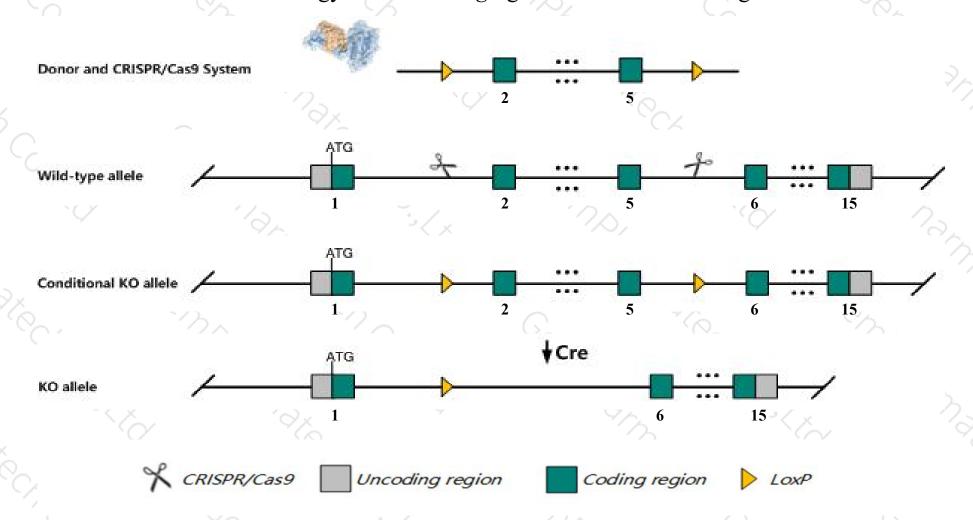
Strain background

C57BL/6JGpt

## Conditional Knockout strategy



This model will use CRISPR/Cas9 technology to edit the Dsg2 gene. The schematic diagram is as follows:



## Technical routes



- ➤ The *Dsg2* gene has 3 transcripts. According to the structure of *Dsg2* gene, exon2-exon5 of *Dsg2-201*(ENSMUST00000059787.14) transcript is recommended as the knockout region. The region contains 478bp coding sequence. Knock out the region will result in disruption of protein function.
- In this project we use CRISPR/Cas9 technology to modify *Dsg2* 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.

### **Notice**



- ➤ According to the existing MGI data, Homozygous mutation of this gene results in embryonic lethality before somite formation, impaired cell proliferation, and increased apoptosis. Heterozygous mutation of this gene also results in embryonic lethality before somite formation with partial penetrance.
- > The *Dsg2* gene is located on the Chr18. 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)



#### Dsg2 desmoglein 2 [Mus musculus (house mouse)]

Gene ID: 13511, updated on 2-Apr-2019

#### Summary

☆ ?

Official Symbol Dsg2 provided by MGI

Official Full Name desmoglein 2 provided by MGI

Primary source MGI:MGI:1196466

See related Ensembl:ENSMUSG00000044393

Gene type protein coding
RefSeq status REVIEWED

Organism Mus musculus

Lineage Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha;

Muroidea; Muridae; Murinae; Mus; Mus

Also known as AA408168, D18Ertd293e

Summary This gene encodes a member of the cadherin family of proteins that forms an integral transmembrane component of desmosomes, the

multiprotein complexes involved in cell adhesion, organization of the cytoskeleton, cell sorting and cell signaling. The encoded preproprotein undergoes proteolytic processing to generate a mature, functional protein. Mice lacking the encoded protein die in utero. Mutant mice lacking a part of the extracellular adhesive domain of the encoded protein develop cardiac fibrosis and dilation. This gene is located in a cluster of

desmosomal cadherin genes on chromosome 18. [provided by RefSeq, Jan 2016]

Expression Broad expression in large intestine adult (RPKM 25.2), colon adult (RPKM 24.6) and 16 other tissuesSee more

Orthologs <u>human all</u>

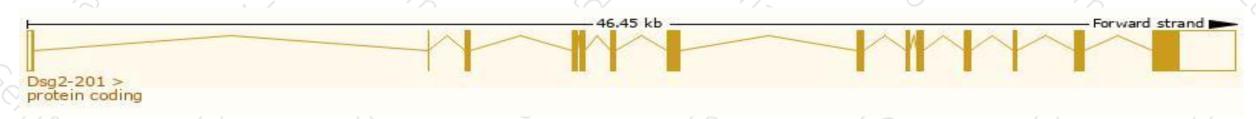
## Transcript information (Ensembl)



The gene has 3 transcripts, all transcripts are shown below:

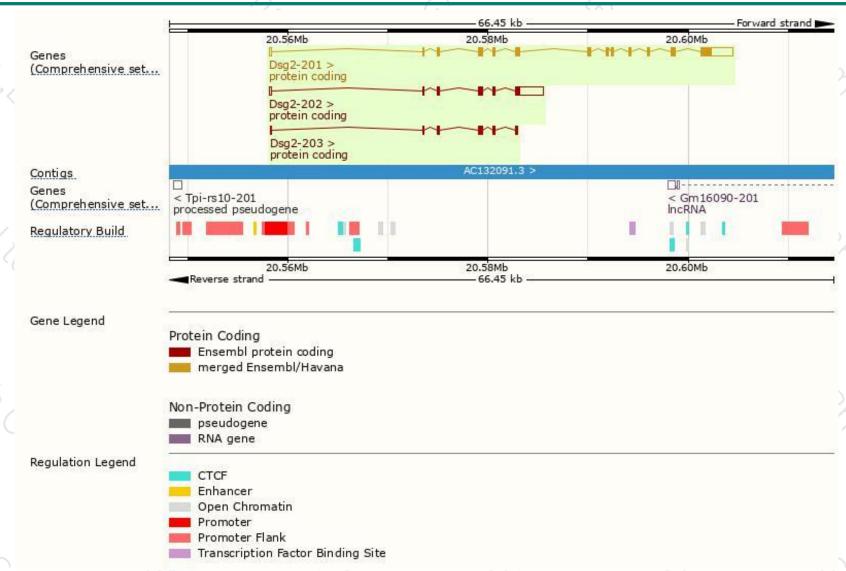
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Dsg2-201	ENSMUST00000059787.14	5766	<u>1122aa</u>	Protein coding	CCDS29084	055111	TSL:1 GENCODE basic APPRIS P1
Dsg2-202	ENSMUST00000120102.7	3590	360aa	Protein coding	19 <del>1</del>	Q811I1	TSL:1 GENCODE basic
Dsg2-203	ENSMUST00000121837.1	953	292aa	Protein coding	84	Q8VCE3	TSL:1 GENCODE basic

The strategy is based on the design of *Dsg2-201* transcript, The transcription is shown below



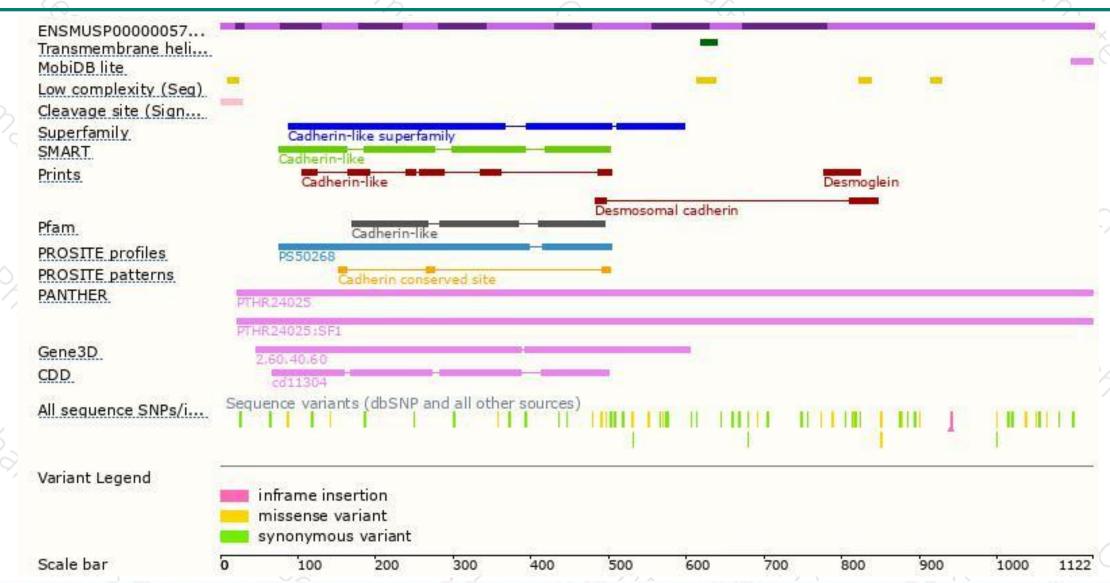
## Genomic location distribution





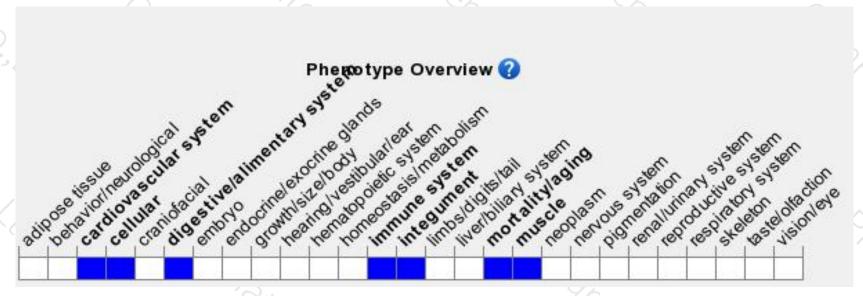
### Protein domain





## 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 mutation of this gene results in embryonic lethality before somite formation, impaired cell proliferation, and increased apoptosis. Heterozygous mutation of this gene also results in embryonic lethality before somite formation with partial penetrance.



If you have any questions, you are welcome to inquire. Tel: 400-9660890





