

Hic2 Cas9-CKO Strategy

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Reviewer: Yumeng Wang

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Project Overview



Project Name Hic2

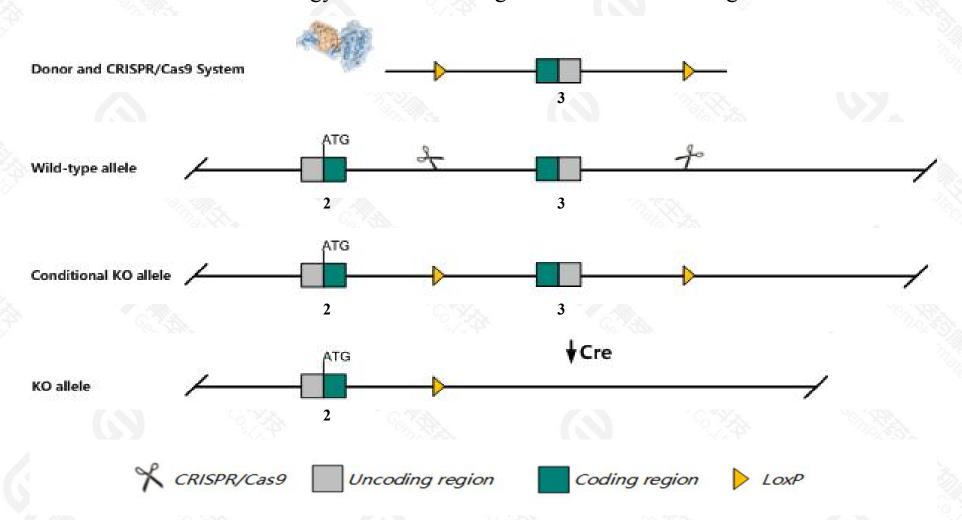
Project type Cas9-CKO

Strain background C57BL/6JGpt

Conditional Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Hic2* gene. The schematic diagram is as follows:



Technical routes



- The *Hic2* gene has 2 transcripts. According to the structure of *Hic2* gene, exon3 of *Hic2-201*(ENSMUST00000090190.13) transcript is recommended as the knockout region. The region contains 1834bp coding sequence. Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Hic2* 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 inactivation of this gene results in embryonic lethality. A subset of mice heterozygous for a knock-out allele exhibit prenatal lethality and cardiac defects including a ventricular septal defect with overriding aortic valve, and thin myocardial and trabecular layers.
- > The KO region contains intron of the Gm49573 gene. Knockout the region may affect the function of Gm49573 gene.
- > The *Hic2* 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)



Hic2 hypermethylated in cancer 2 [Mus musculus (house mouse)]

Gene ID: 58180, updated on 13-Mar-2020

Summary

☆ ?

Official Symbol Hic2 provided by MGI

Official Full Name hypermethylated in cancer 2 provided by MGI

Primary source MGI:MGI:1929869

See related Ensembl:ENSMUSG00000050240

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 AA409108, HRG22, mKIAA1020

Expression Broad expression in placenta adult (RPKM 11.4), CNS E11.5 (RPKM 5.1) and 26 other tissuesSee more

Orthologs <u>human all</u>

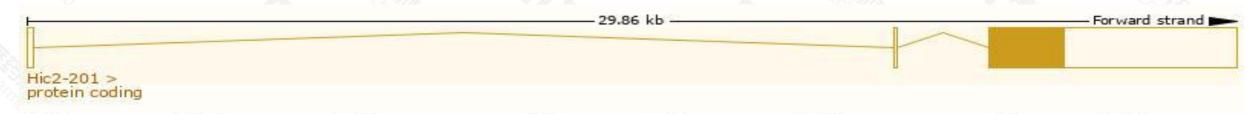
Transcript information (Ensembl)



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

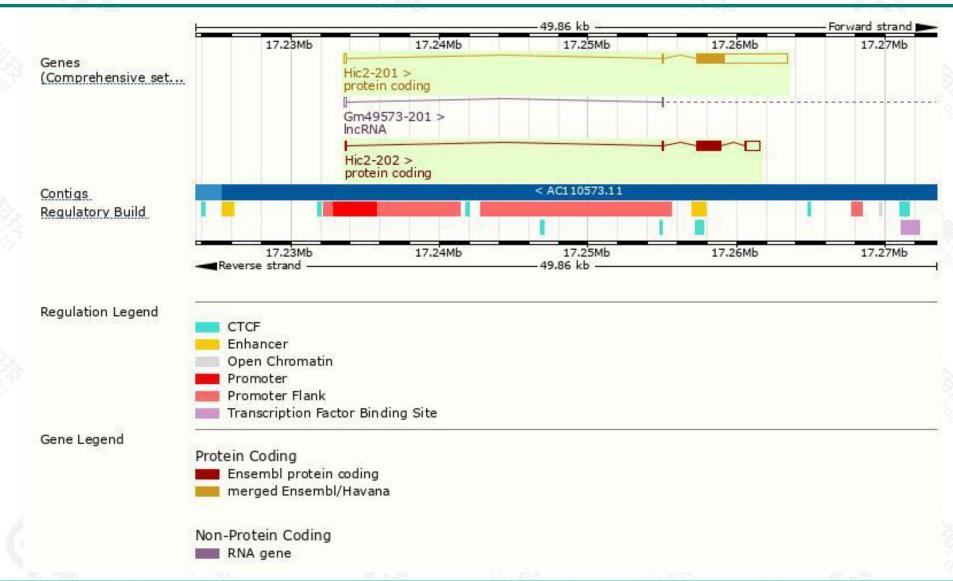
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Hic2-201	ENSMUST00000090190.13	6354	619aa	Protein coding	CCDS27999	Q9JLZ6	TSL:1 GENCODE basic APPRIS P1
Hic2-202	ENSMUST00000232082.1	2765	<u>566aa</u>	Protein coding	-	Q9JLZ6	GENCODE basic

The strategy is based on the design of *Hic2-201* transcript, the transcription is shown below:



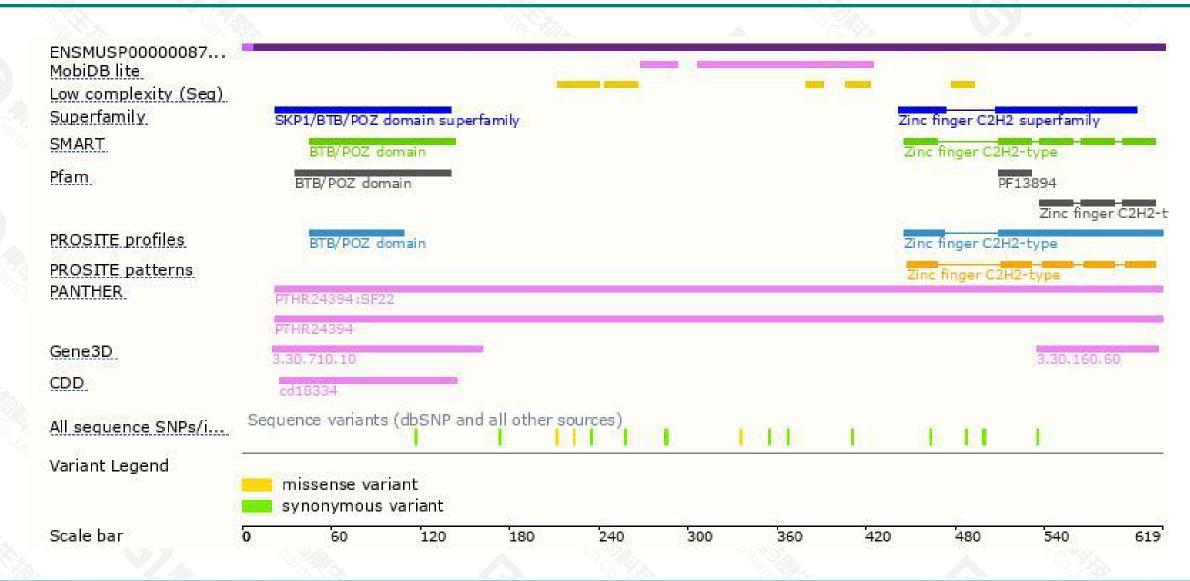
Genomic location distribution





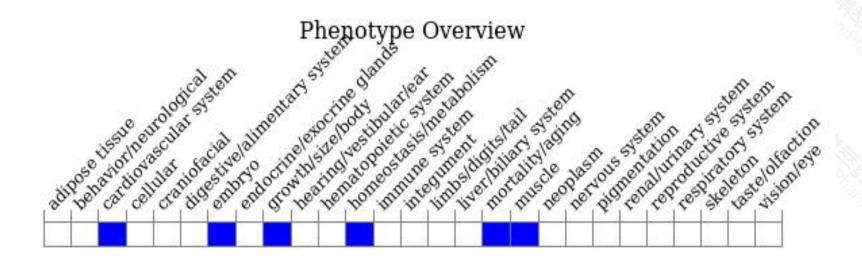
Protein domain





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 inactivation of this gene results in embryonic lethality. A subset of mice heterozygous for a knock-out allele exhibit prenatal lethality and cardiac defects including a ventricular septal defect with overriding aortic valve, and thin myocardial and trabecular layers.



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

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