

# Npr1 Cas9-KO Strategy

Designer: Huimin Su

## **Project Overview**



Project Name Npr1

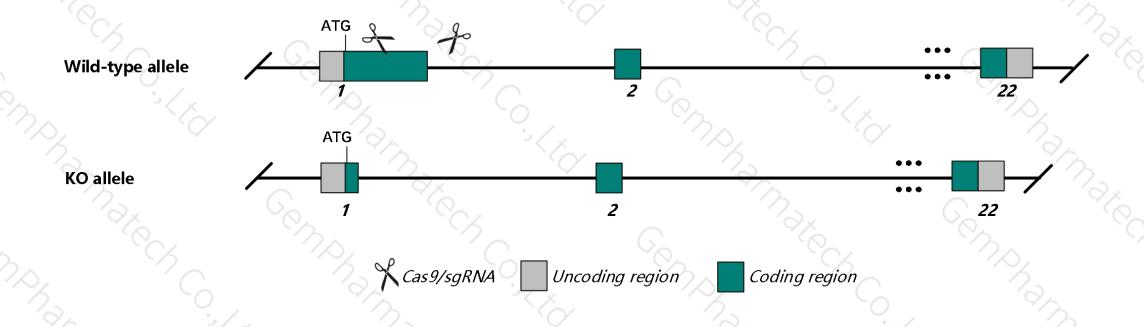
Project type Cas9-KO

Strain background C57BL/6J

## **Knockout strategy**



This model will use CRISPR/Cas9 technology to edit the *Npr1* gene. The schematic diagram is as follows:



### **Technical routes**



- ➤ The *Npr1* gene has 5 transcripts. According to the structure of *Npr1* gene, exon1 of *Npr1-201*(ENSMUST00000029540.12) transcript is recommended as the knockout region. Part of exon1 and intron1 was deleted, which will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Npr1* gene. The brief process is as follows: sgRNA was transcribed in vitro.Cas9 and sgRNA were microinjected into the fertilized eggs of C57BL/6J 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/6J mice.

### **Notice**



- ➤ According to the existing MGI data, Homozygous inactivation of this gene can lead to hypertension, cardiac hypertrophy, lethal vascular events, congestive heart failure in response to volume overload, reduced serum testosterone levels, altered steroidogenesis, and reduced myocardial PMN infiltration and infarct size after I/R injury.
- ➤ The *Npr1* gene is located on the Chr3. 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 the gene knockout on gene transcription, RNA splicing and protein translation cannot be predicted at the existing technology level.

## Gene information (NCBI)



#### Npr1 natriuretic peptide receptor 1 [Mus musculus (house mouse)]

Gene ID: 18160, updated on 3-Feb-2019

#### Summary

☆ ?

Official Symbol Npr1 provided by MGI

Official Full Name natriuretic peptide receptor 1 provided by MGI

Primary source MGI:MGI:97371

See related Ensembl:ENSMUSG00000027931

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 Al893888, GC-A, NPR-A, NPRA, Pndr

Expression Biased expression in adrenal adult (RPKM 198.6), ovary adult (RPKM 80.4) and 6 other tissuesSee more

Orthologs <u>human all</u>

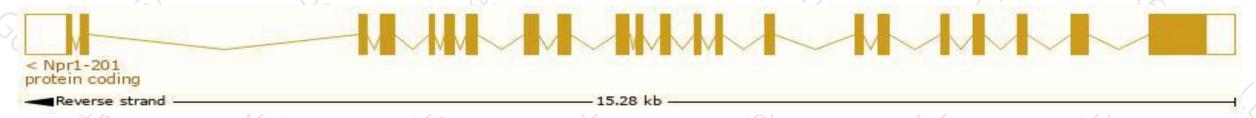
## Transcript information (Ensembl)



The gene has 5 transcripts, all transcripts are shown below:

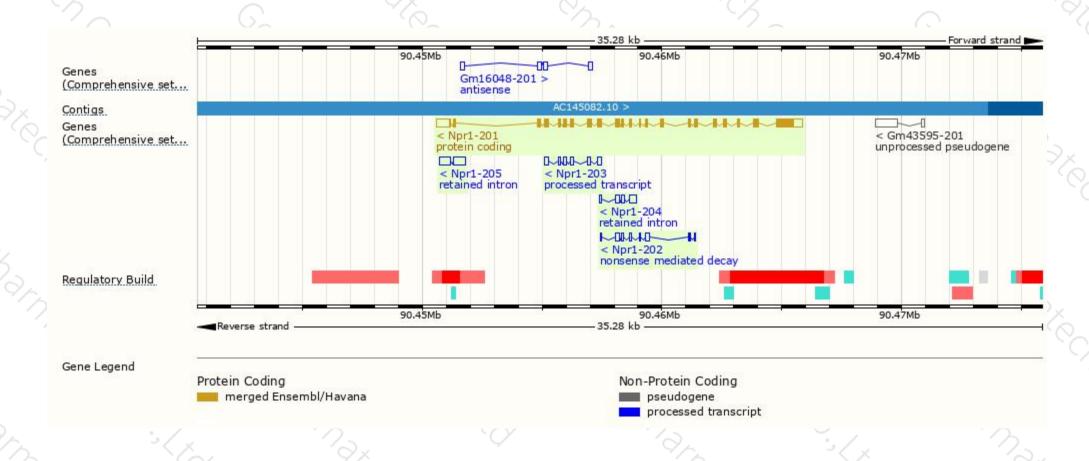
Name 🌲	Transcript ID 👙	bp 🍦	Protein 🍦	Biotype	CCDS 🍦	UniProt 🍦	Flags
Npr1-201	ENSMUST00000029540.12	4066	1057aa	Protein coding	CCDS17529₽	P18293 ₽ Q2TAY4 ₽	TSL:1 GENCODE basic APPRIS P1
Npr1-202	ENSMUST00000124760.1	790	<u>51aa</u>	Nonsense mediated decay	=	F6W125₽	CDS 5' incomplete TSL:5
Npr1-203	ENSMUST00000142243.1	767	No protein	Processed transcript	-	-	TSL:5
Npr1-205	ENSMUST00000152510.1	949	No protein	Retained intron	-	0-0	TSL:2
Npr1-204	ENSMUST00000146991.1	643	No protein	Retained intron	=	0.00	TSL:3

The strategy is based on the design of Npr1-201 transcript, The transcription is shown below



### Genomic location distribution





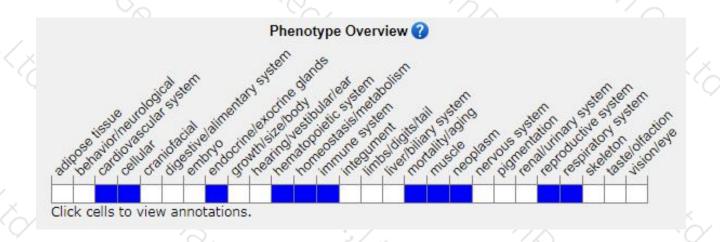
### Protein domain





## 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 can lead to hypertension, cardiac hypertrophy, lethal vascular events, congestive heart failure in response to volume overload, reduced serum testosterone level altered steroidogenesis, and reduced myocardial PMN infiltration and infarct size after I/R injury.



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

Tel: 025-5864 1534





