

Lrp6 Cas9-KO Strategy

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Project Overview

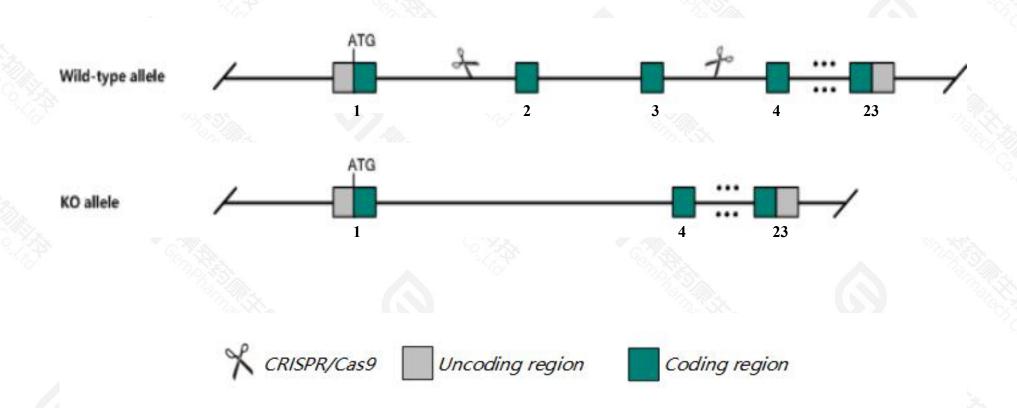


Project Name	Lrp6
Project type	Cas9-KO
Strain background	C57BL/6JGpt

Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Lrp6* gene. The schematic diagram is as follows:



Technical routes



- ➤ The *Lrp6* gene has 6 transcripts. According to the structure of *Lrp6* gene, exon2-exon3 of *Lrp6*201(ENSMUST00000032322.15) transcript is recommended as the knockout region. The region contains 592bp coding sequence. Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Lrp6* gene. The brief process is as follows: CRISPR/Cas9 system 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.

Notice



- > According to the existing MGI data, animals homozygous for this mutation exhibit partial embryonic lethality, growth retardation, crooked tail, abnormal vertebrae, small skull with occasional bent nose, absence of the third molars and small and/or unerupted lower incisors. Heterozygotes exhibit a crooked tail and abnormal vertebrae.
- > The *Lrp6* gene is located on the Chr6. 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)



Lrp6 low density lipoprotein receptor-related protein 6 [Mus musculus (house mouse)]

Gene ID: 16974, updated on 22-Mar-2020

Summary



Official Symbol Lrp6 provided by MGI

Official Full Name low density lipoprotein receptor-related protein 6 provided by MGI

Primary source MGI:MGI:1298218

See related Ensembl: ENSMUSG00000030201

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 C030016K15Rik, Cd, Gw, ska26, ska, skax26

Expression Ubiquitous expression in bladder adult (RPKM 13.4), kidney adult (RPKM 13.4) and 28 other tissuesSee more

Orthologs human all

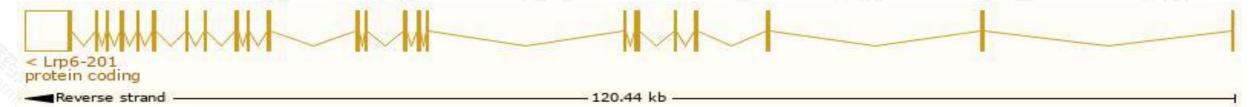
Transcript information (Ensembl)



The gene has 6 transcripts, all transcripts are shown below:

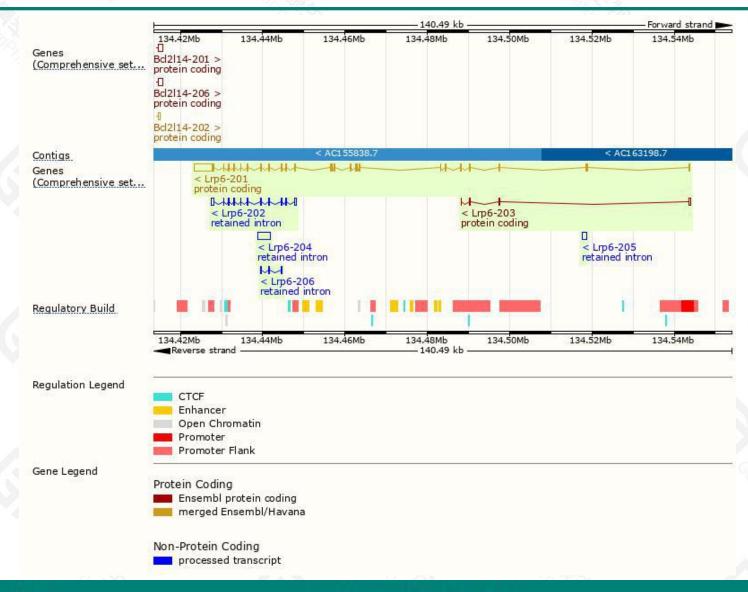
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Lrp6-201	ENSMUST00000032322.14	9368	<u>1613aa</u>	Protein coding	CCDS39678	A0A0R4J0A9	TSL:1 GENCODE basic APPRIS is a system to annotate alternatively spliced transcripts based on a range of computational methods to identify the most functionally important transcript(s) of a gene. APPRIS P1
Lrp6-203	ENSMUST00000126836.3	722	<u>131aa</u>	Protein coding	-	A0A0N4SUL9	CDS 3' incomplete TSL:5
Lrp6-204	ENSMUST00000203016.1	3246	No protein	Retained intron	-	-	TSL:NA
Lrp6-202	ENSMUST00000111950.3	2756	No protein	Retained intron	2	24	TSL:1
Lrp6-205	ENSMUST00000203235.1	941	No protein	Retained intron	-	-	TSL:NA
Lrp6-206	ENSMUST00000203773.1	727	No protein	Retained intron	-	-	TSL:3

The strategy is based on the design of Lrp6-201 transcript, the transcription is shown below:



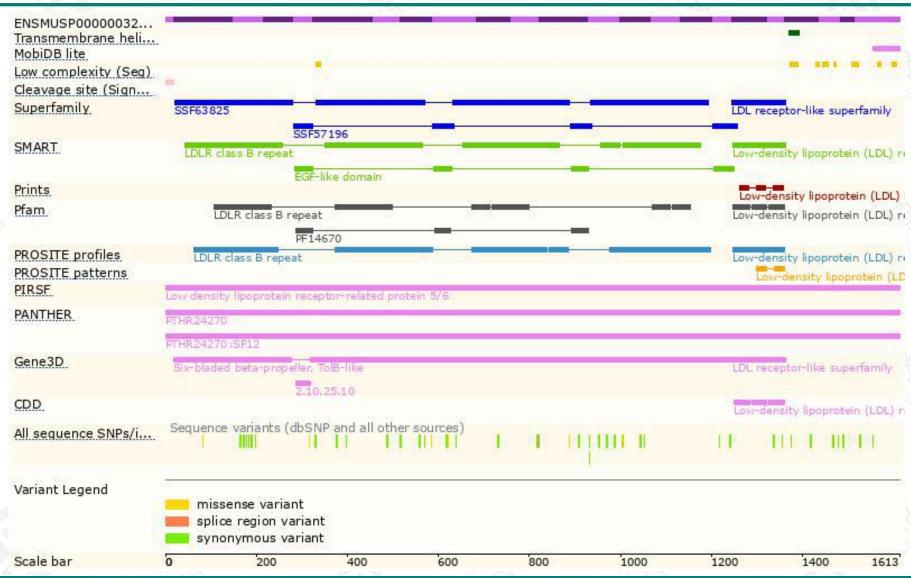
Genomic location distribution





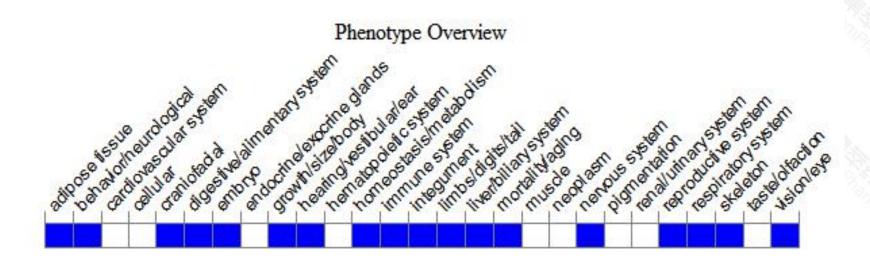
Protein domain





Mouse phenotype description(MGI)





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

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

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