

# Angptl6 Cas9-KO Strategy

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Reviewer: Huimin Su

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## **Project Overview**



Project Name Angptl6

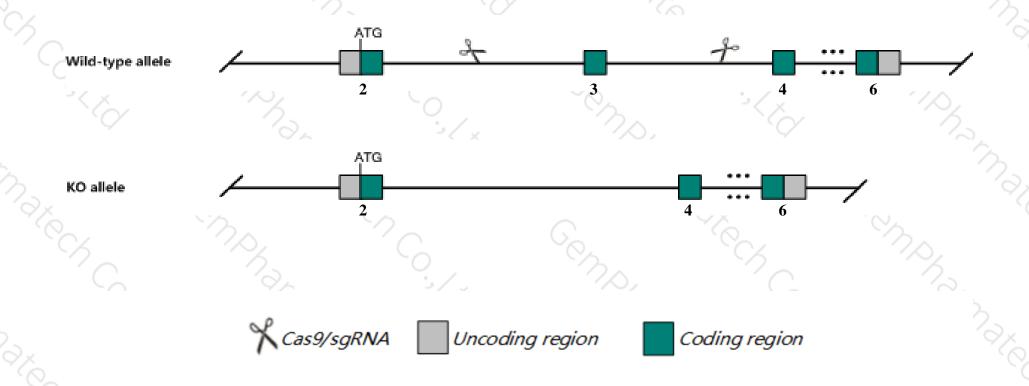
Project type Cas9-KO

Strain background C57BL/6JGpt

## **Knockout strategy**



This model will use CRISPR/Cas9 technology to edit the *Angptl6* gene. The schematic diagram is as follows:



### **Technical routes**



- ➤ The *Angptl6* gene has 1 transcript. According to the structure of *Angptl6* gene, exon3 of *Angptl6-201*(ENSMUST00000043726.7) transcript is recommended as the knockout region. The region contains 178bp coding sequence.

  Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Angptl6* gene. The brief process is as follows: sgRNA was transcribed in vitro.Cas9 and sgRNA 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, Most mice homozygous for a knock-out allele die around E13 with cardiovascular defects; survivors develop obesity, lipid accumulation in skeletal muscle and liver, hyperglycemia and hyperinsulinemia, and insulin resistance accompanied by reduced energy expenditure and whole-body oxygen consumption.
- ➤ The *Angptl6* gene is located on the Chr9. 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)



#### Angptl6 angiopoietin-like 6 [Mus musculus (house mouse)]

Gene ID: 70726, updated on 31-Jan-2019

#### Summary

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Official Symbol Angptl6 provided by MGI

Official Full Name angiopoietin-like 6 provided by MGI

Primary source MGI:MGI:1917976

See related Ensembl:ENSMUSG00000038742

Gene type protein coding
RefSeq status PROVISIONAL
Organism Mus musculus

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

Muroidea; Muridae; Murinae; Mus; Mus

Also known as 6330404E11Rik, AGF, ARP3, Arp5

Expression Ubiquitous expression in liver adult (RPKM 15.9), placenta adult (RPKM 7.6) and 27 other tissuesSee more

Orthologs <u>human</u> all

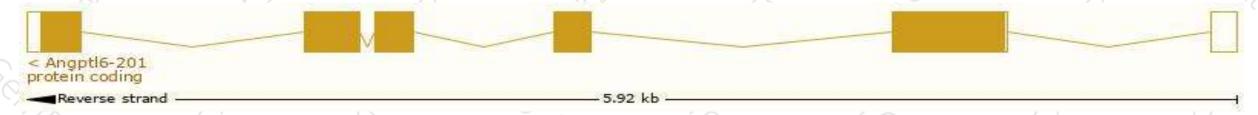
## Transcript information (Ensembl)



The gene has 1 transcript, and the transcript is shown below:

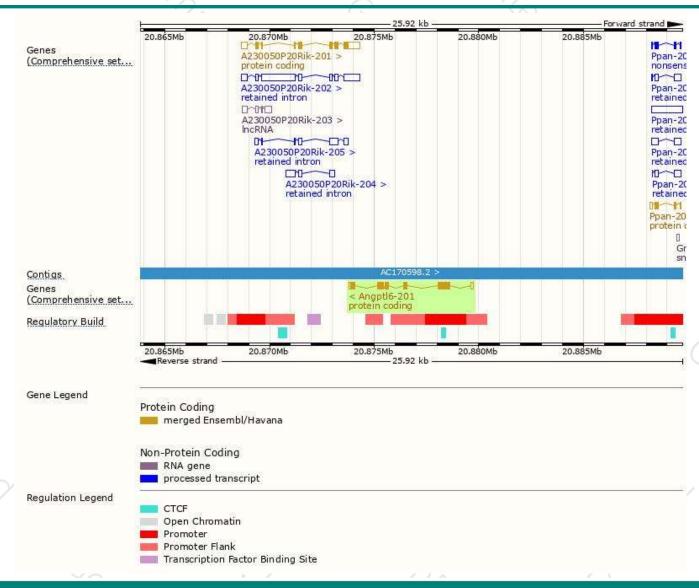
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Angptl6-201	ENSMUST00000043726.7	1582	<u>457aa</u>	Protein coding	CCDS22886	Q8R0Z6	TSL:1 GENCODE basic APPRIS P1

The strategy is based on the design of Angptl6-201 transcript, The transcription is shown below



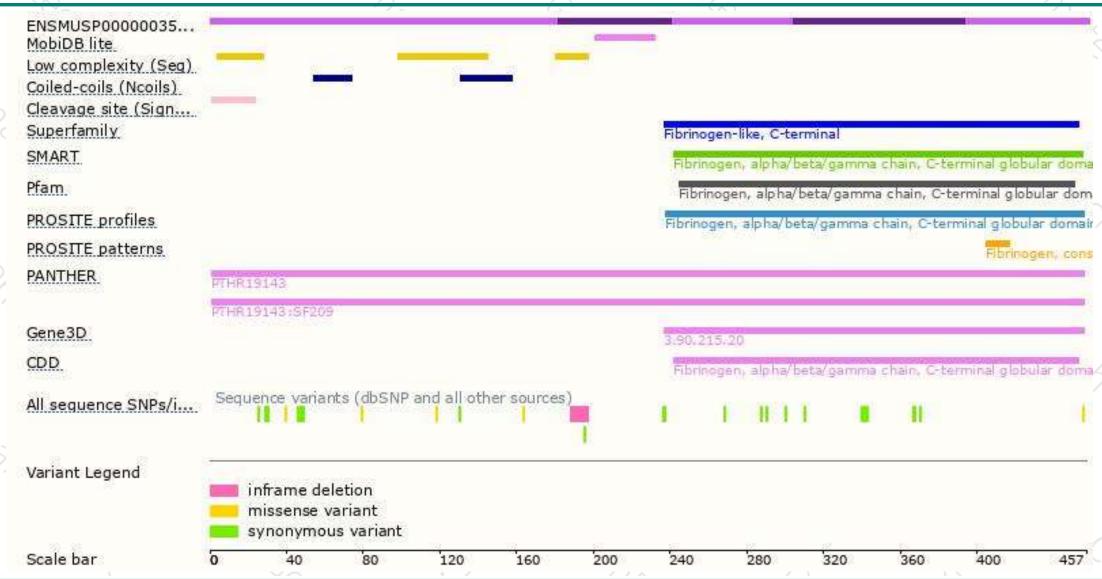
### Genomic location distribution





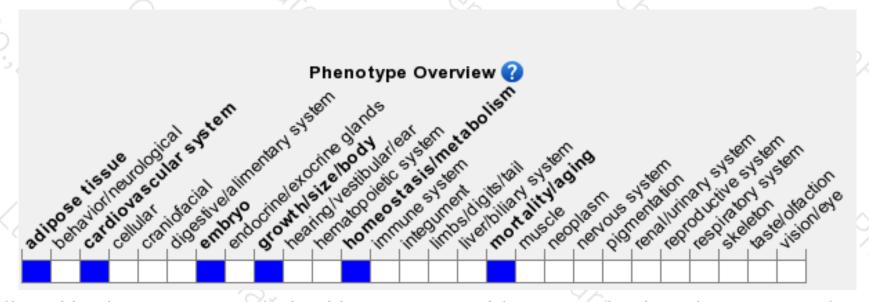
### **Protein domain**





## 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, Most mice homozygous for a knock-out allele die around E13 with cardiovascular defects; survivors develop obesity, lipid accumulation in skeletal muscle and liver, hyperglycemia and hyperinsulinemia, and insulin resistance accompanied by reduced energy expenditure and whole-body oxygen consumption.



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

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