

# ***Fkbp1b*** Cas9-CKO Strategy

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# Project Overview

**Project Name**

***Fkbp1b***

**Project type**

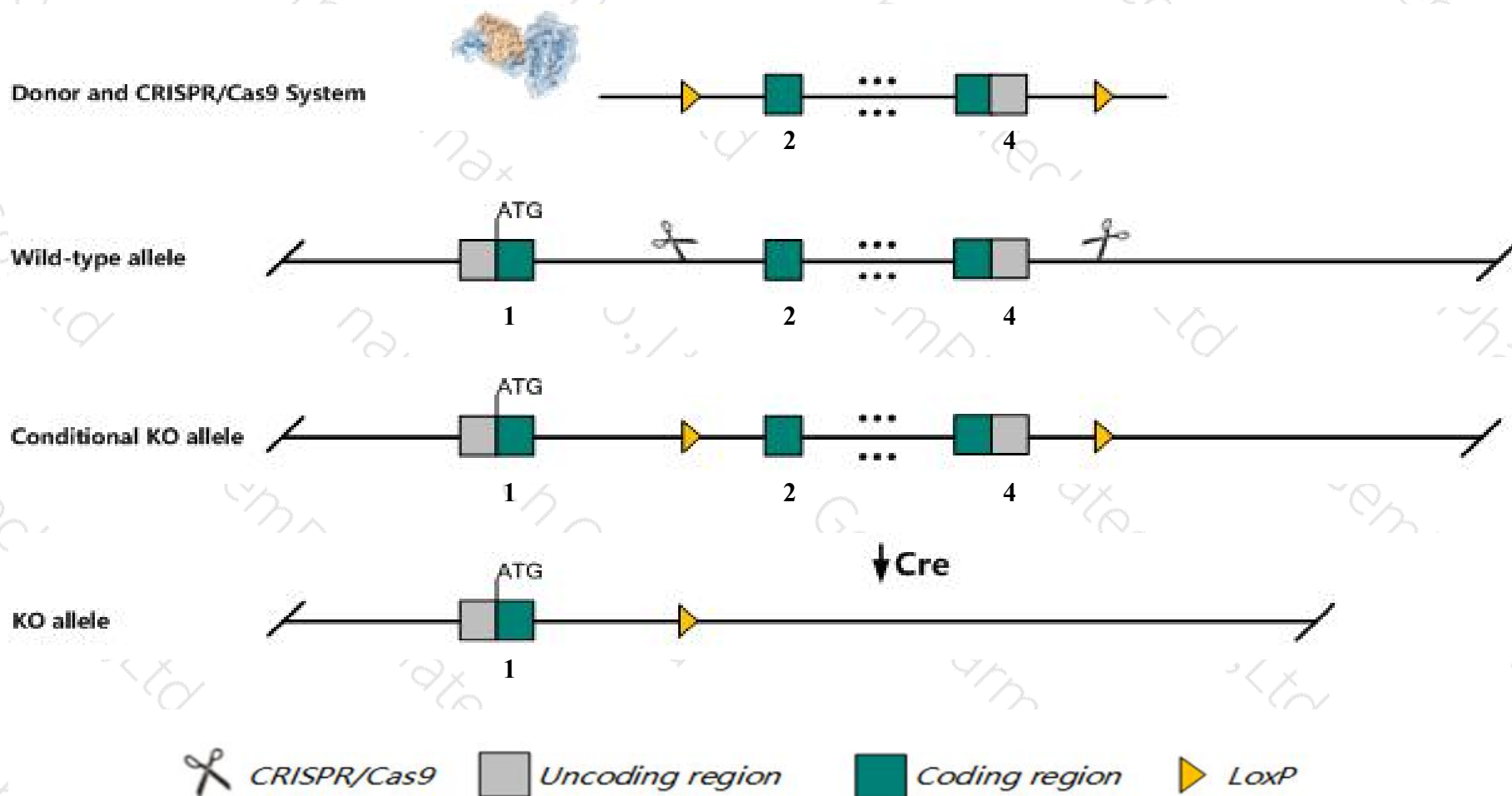
**Cas9-CKO**

**Strain background**

**C57BL/6JGpt**

# Conditional Knockout strategy

This model will use CRISPR/Cas9 technology to edit the *Fkbp1b* gene. The schematic diagram is as follows:



# Technical routes

- The *Fkbp1b* gene has 2 transcripts. According to the structure of *Fkbp1b* gene, exon2-exon4 of *Fkbp1b*-201 (ENSMUST00000020964.6) transcript is recommended as the knockout region. The region contains 290bp coding sequence. Knock out the region will result in disruption of protein function.
- In this project we use CRISPR/Cas9 technology to modify *Fkbp1b* 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.

- According to the existing MGI data, Homozygotes for a null allele display exercise-induced sudden cardiac death. Homozygotes for a second null allele show impaired glucose tolerance and glucose-driven insulin secretion. Homozygotes for a third null allele show  $\text{Ca}^{2+}$  dysregulation and male-specific cardiac hypertrophy and hypertension.
- The *Fkbp1b* gene is located on the Chr12. 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)

## Fkbp1b FK506 binding protein 1b [ *Mus musculus* (house mouse) ]

Gene ID: 14226, updated on 16-Dec-2019

### Summary

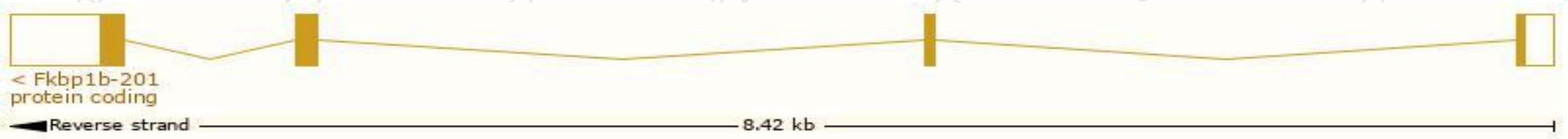
Official Symbol	Fkbp1b provided by <a href="#">MGI</a>
Official Full Name	FK506 binding protein 1b provided by <a href="#">MGI</a>
Primary source	<a href="#">MGI:MGI:1336205</a>
See related	<a href="#">Ensembl:ENSMUSG00000020635</a>
Gene type	protein coding
RefSeq status	VALIDATED
Organism	<a href="#">Mus musculus</a>
Lineage	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha; Muroidea; Muridae; Murinae; Mus; Mus
Also known as	12.6kDa; AW494148; FKBP12.6; calstabin2
Expression	Broad expression in frontal lobe adult (RPKM 19.1), CNS E18 (RPKM 18.2) and 21 other tissues <a href="#">See more</a>
Orthologs	<a href="#">human</a> <a href="#">all</a>

# Transcript information (Ensembl)

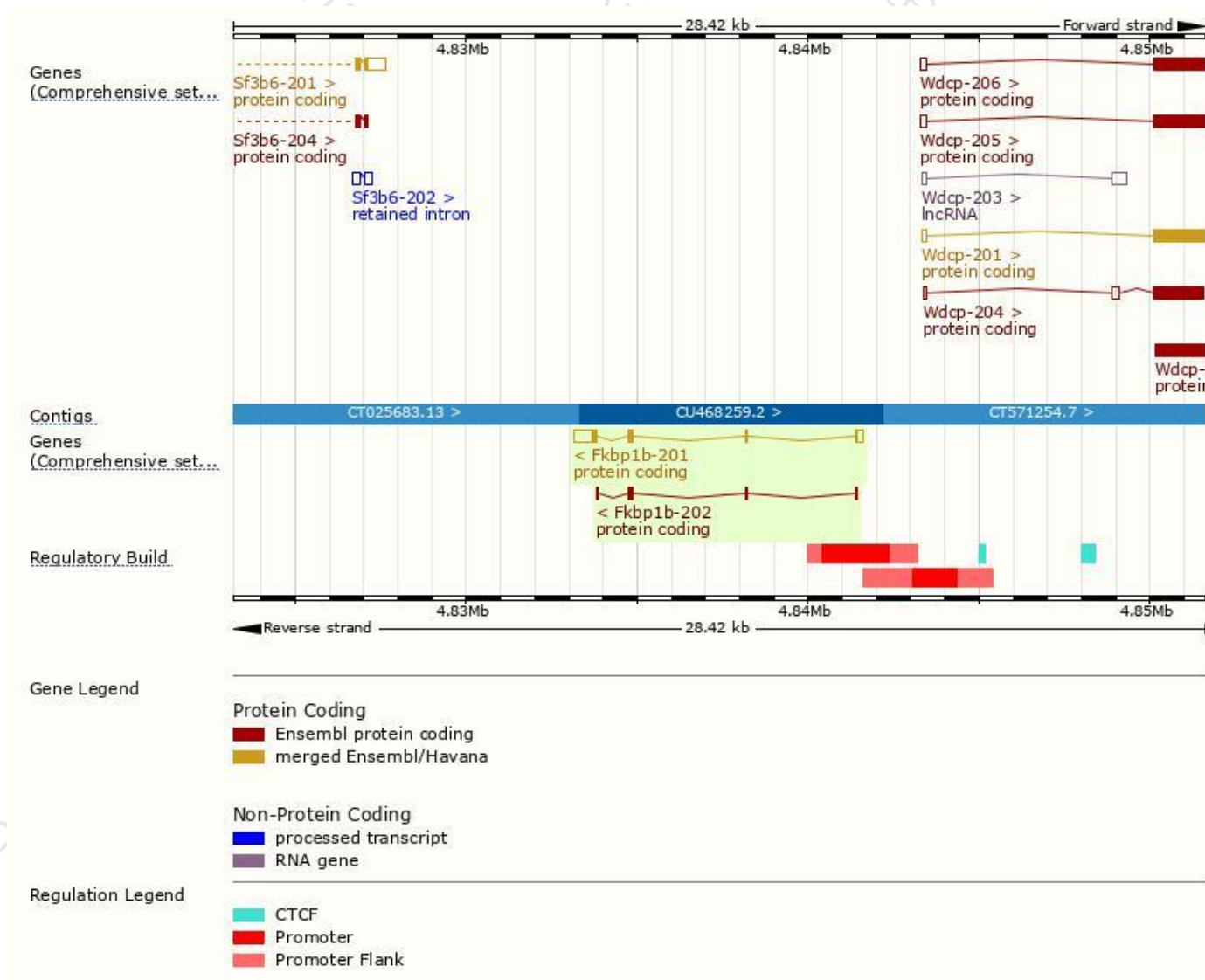
The gene has 2 transcripts,all transcripts are shown below:

Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Fkbp1b-201	<a href="#">ENSMUST00000020964.6</a>	976	<a href="#">108aa</a>	Protein coding	<a href="#">CCDS25792</a>	<a href="#">A9E3L2 Q9Z2I2</a>	TSL:1 GENCODE basic APPRIS P1
Fkbp1b-202	<a href="#">ENSMUST00000219880.1</a>	246	<a href="#">80aa</a>	Protein coding	-	<a href="#">Q80YE1</a>	TSL:1 GENCODE basic

The strategy is based on the design of *Fkbp1b-201* transcript,The transcription is shown below

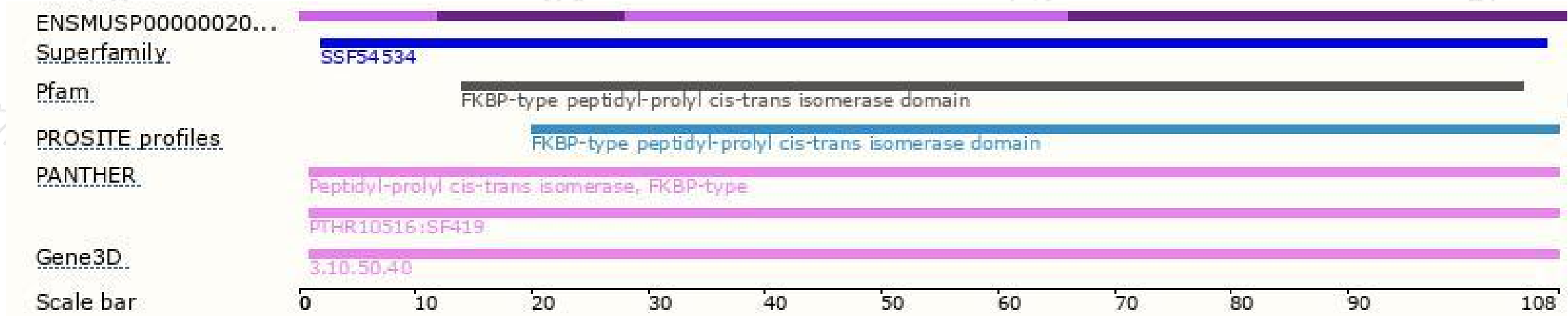


# Genomic location distribution

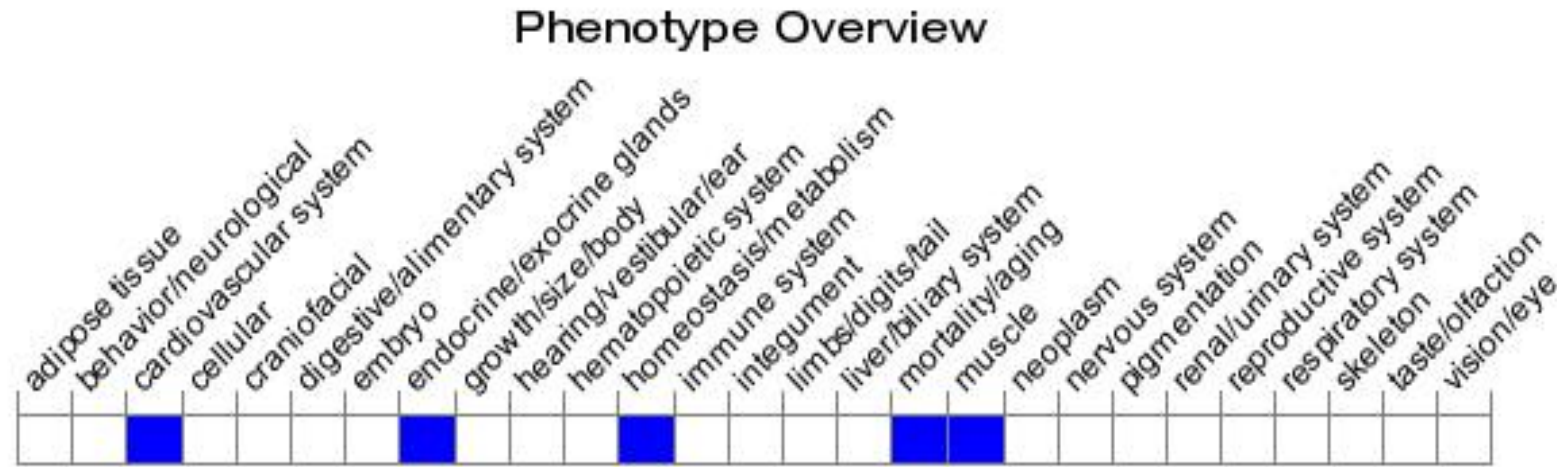




# Protein domain



# 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, Homozygotes for a null allele display exercise-induced sudden cardiac death.

Homozygotes for a second null allele show impaired glucose tolerance and glucose-driven insulin secretion. Homozygotes for third null allele show  $\text{Ca}^{2+}$  dysregulation and male-specific cardiac hypertrophy and hypertension.

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

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