

# Casq1 Cas9-KO Strategy

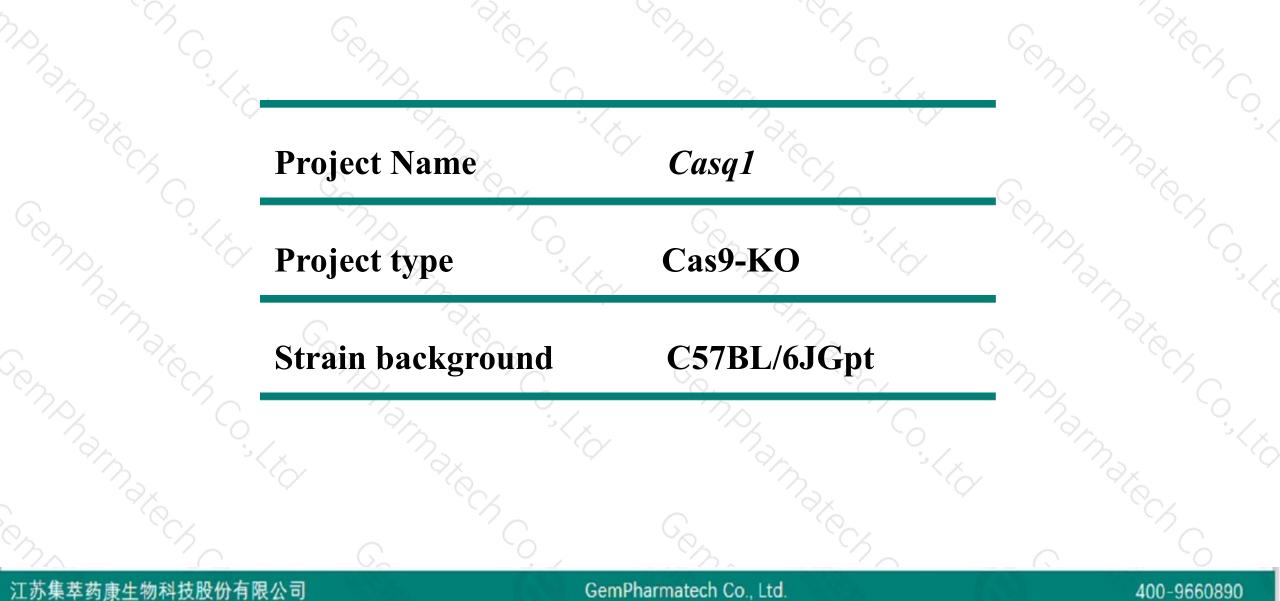
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**Reviewer: Lingyan Wu** 

**Design Date: 2020-7-6** 

## **Project Overview**

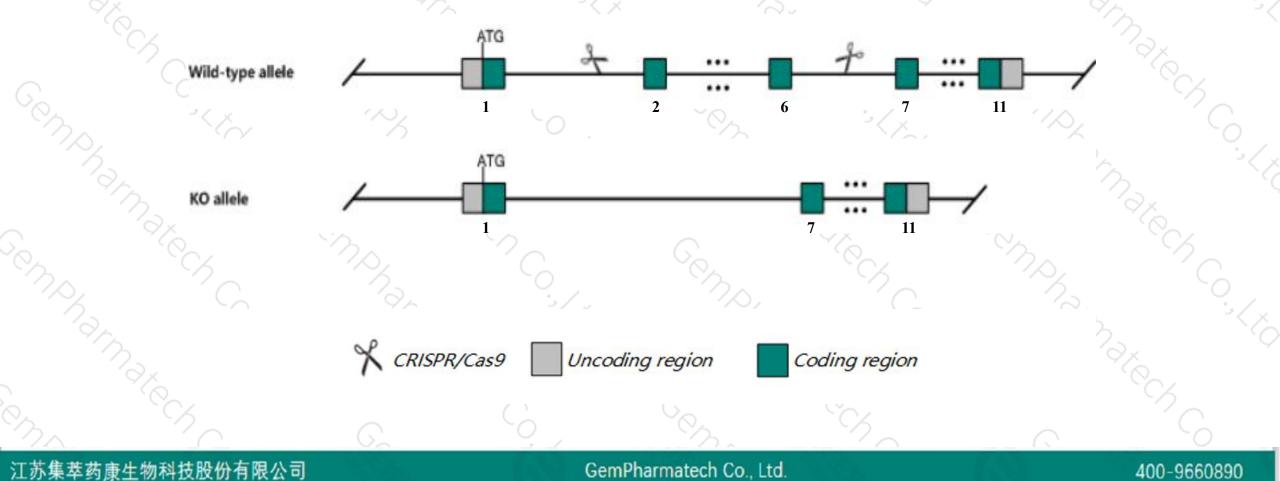




## **Knockout strategy**



This model will use CRISPR/Cas9 technology to edit the Casq1 gene. The schematic diagram is as follows:





> The *Casq1* gene has 3 transcripts. According to the structure of *Casq1* gene, exon2-exon6 of *Casq1*-201(ENSMUST00000003554.10) transcript is recommended as the knockout region. The region contains 503bp coding sequence. Knock out the region will result in disruption of protein function.

➤ In this project we use CRISPR/Cas9 technology to modify *Casq1* 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.



- > According to the existing MGI data, mice homozygous for an insertional mutation that inactivates the gene exhibit structural alterations of the Ca2+ release units, an increased frequency of mitochondria, and significantly impaired calcium handling in skeletal muscle.
- > The *Casq1* gene is located on the Chr1. 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)**



☆ ?

## Casq1 calsequestrin 1 [Mus musculus (house mouse)]

Gene ID: 12372, updated on 13-Mar-2020

### - Summary

Official SymbolCasq1 provided by MGIOfficial Full Namecalsequestrin 1 provided byMGIPrimary sourceMGI:MGI:1309468See relatedEnsembl:ENSMUSG0000007122Gene typeprotein codingRefSeq statusVALIDATEDOrganismMus musculusLineageEukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha;<br/>Muroidea; Murinae; Mus; MusAlso known asCSQ, CSQ-1, CSQ1, sCSQExpressionBiased expression in mammary gland adult (RPKM 118.1), heart adult (RPKM 12.4) and 1 other tissue<u>See more</u><br/>human all

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## **Transcript information (Ensembl)**



## The gene has 3 transcripts, all transcripts are shown below:

Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Casq1-201	ENSMUST0000003554.10	1847	<u>405aa</u>	Protein coding	CCDS35781	<u>009165</u>	TSL:1 GENCODE basic APPRIS P1
Casq1-203	ENSMUST00000170700.1	<mark>4</mark> 93	<u>156aa</u>	Protein coding	-	E9Q489	CDS 3' incomplete TSL:5
Casq1-202	ENSMUST00000170638.1	990	No protein	Retained intron	2	2	TSL:1

The strategy is based on the design of Casq1-201 transcript, the transcription is shown below:

#### < Casq1-201 protein coding

Reverse strand

- 9.97 kb -

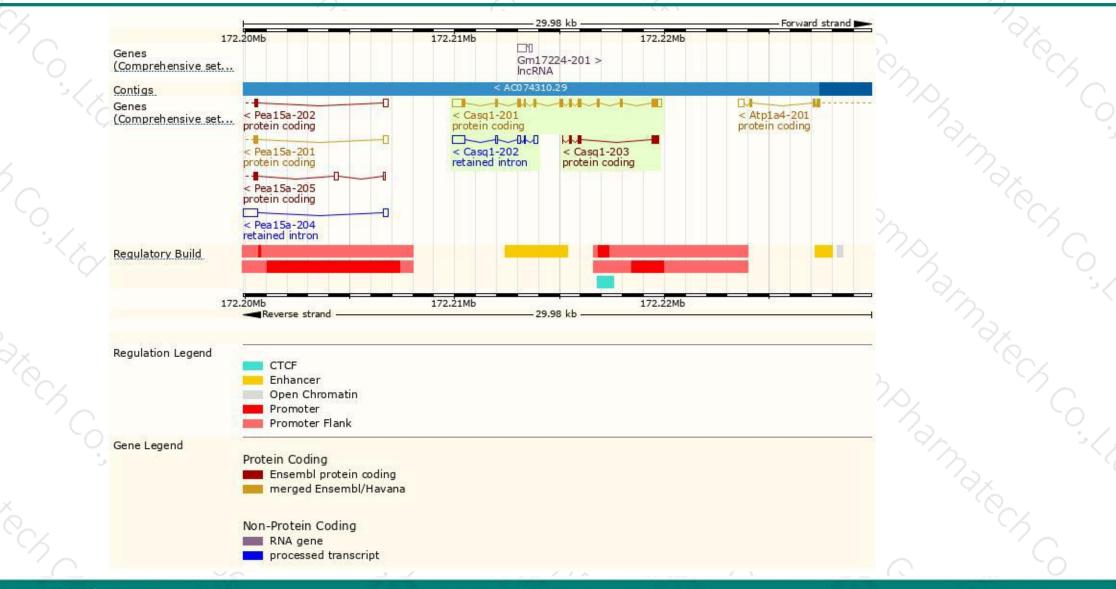
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## **Genomic location distribution**



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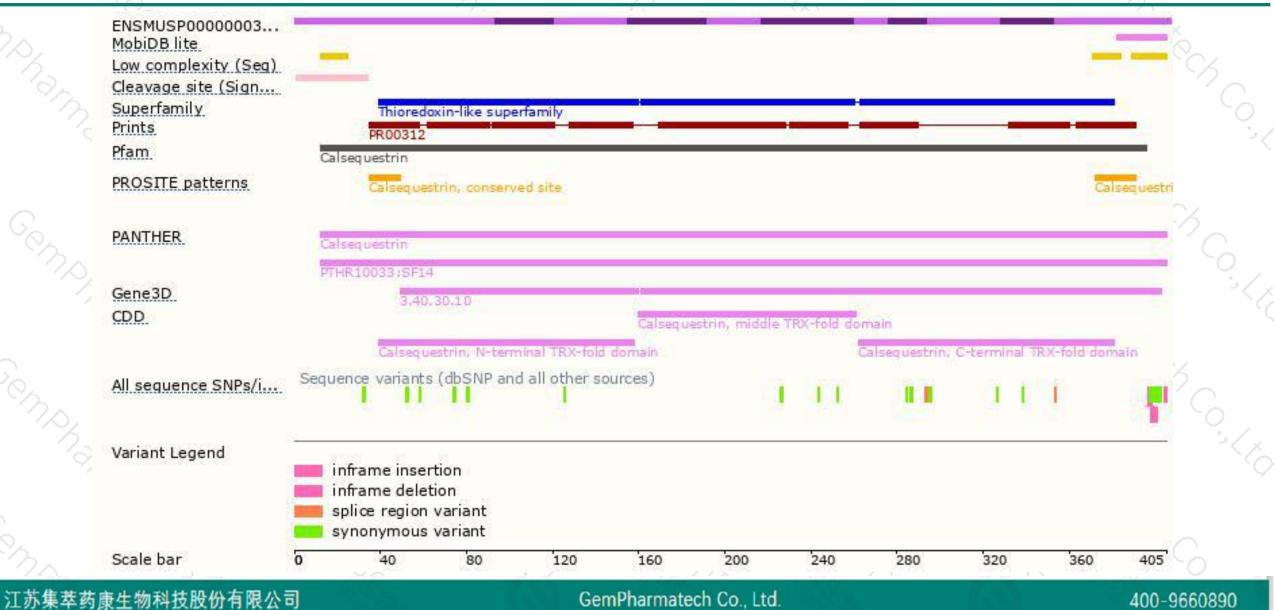


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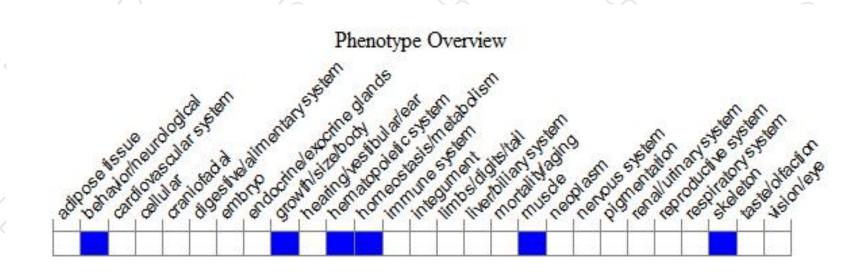
## **Protein domain**





## 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,mice homozygous for an insertional mutation that inactivates the gene exhibit structural alterations of the Ca2+ release units, an increased frequency of mitochondria, and significantly impaired calcium handling in skeletal muscle

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If you have any questions, you are welcome to inquire. Tel: 400-9660890



