

Cnotl Cas9-KO Strategy

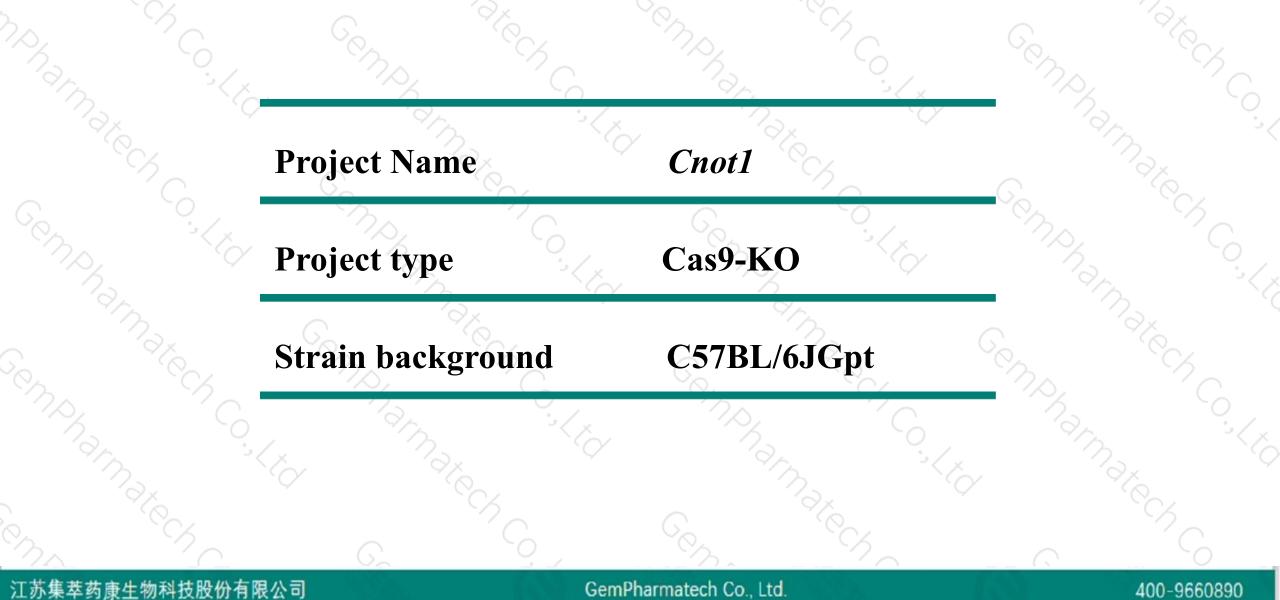
Designer: Huimin Su

Reviewer: Ruiuri Zhang

Design Date: 2020-6-16

Project Overview

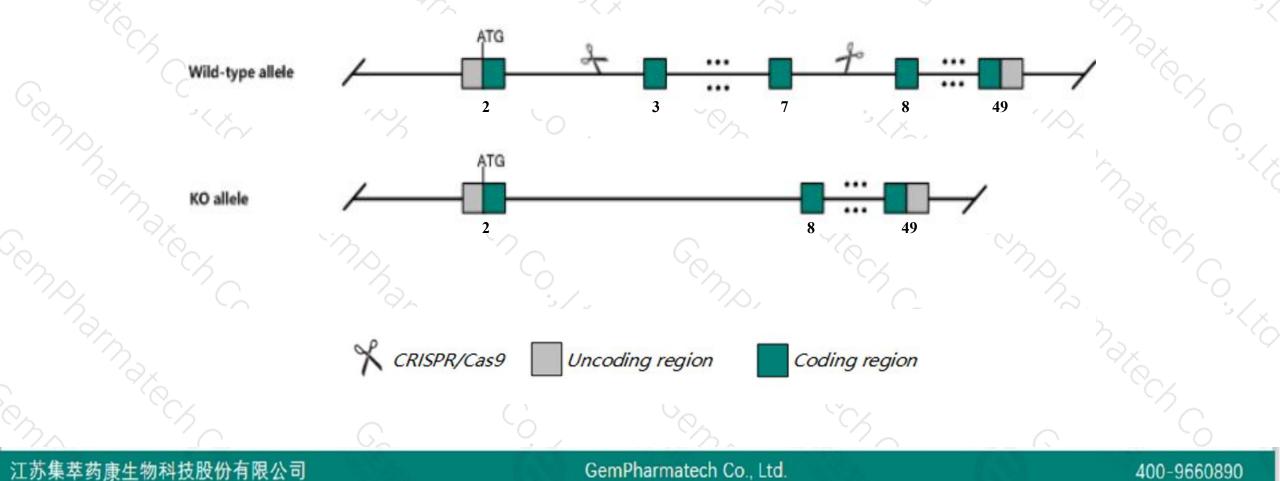




Knockout strategy



This model will use CRISPR/Cas9 technology to edit the Cnot1 gene. The schematic diagram is as follows:





- The Cnot1 gene has 17 transcripts. According to the structure of Cnot1 gene, exon3-exon7 of Cnot1-203 (ENSMUST00000211887.1) transcript is recommended as the knockout region. The region contains 529bp coding sequence. Knock out the region will result in disruption of protein function.
- > In this project we use CRISPR/Cas9 technology to modify Cnot1 gene. The brief process is as follows: CRISPR/Cas9 system

- According to the existing MGI data, mice hmozygous for a conditional allele activated in cardiomyocytes exhibit postnatal lethality, decreased cardiac muscle contractility, prolonged qt interval and cardiac muscle cell death.
- > The 3-regulation of Gm45762-201 may be affect.
- > Cnot1-217 and Cnot1-205 transcripts are unknown, and the effect on these two transcripts are unknown.
- The Cnot1 gene is located on the Chr8. 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.

Notice

Gene information (NCBI)



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Cnot1 CCR4-NOT transcription complex, subunit 1 [Mus musculus (house mouse)]

Gene ID: 234594, updated on 8-Jun-2020

Summary

 Official Symbol
 Cnot1 provided by MGI

 Official Full Name
 CCR4-NOT transcription complex, subunit 1 provided by MGI

 Primary source
 MGI:MGI:2442402

 See related
 Ensembl:ENSMUSG00000036550

 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
 AA815922; D830048B13; 6030411K04Rik

 Expression
 Ubiquitous expression in testis adult (RPKM 29.9), placenta adult (RPKM 21.5) and 28 other tissues <u>See more</u>

 Orthologs
 human all

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Transcript information (Ensembl)



The gene has 17 transcripts, all transcripts are shown below:

	Name 🖕	Transcript ID 💧	bp 👙	Protein 🖕	Biotype 🍦	CCDS 🍦	UniProt 🖕	Flags 🖕
20 N	Cnot1-203	ENSMUST00000211887.1	8364	<u>2369aa</u>	Protein coding	<u>CCDS85589</u> &	A0A1D5RMJ8	TSL:1 GENCODE basic APPRIS P2
	Cnot1-201	ENSMUST0000068452.9	8152	<u>2326aa</u>	Protein coding	CCDS57635@	<u>B7ZWL1</u> 교	TSL:1 GENCODE basic
	Cnot1-202	ENSMUST0000098473.10	8390	<u>2376aa</u>	Protein coding	870	<u>Q6ZQ08</u> &	TSL:5 GENCODE basic APPRIS ALT1
	Cnot1-205	ENSMUST00000211973.1	2469	<u>823aa</u>	Protein coding	373	A0A1D5RML9	CDS 5' and 3' incomplete TSL:5
	Cnot1-217	ENSMUST00000213046.1	550	<u>92aa</u>	Protein coding	870	<u>A0A1D5RMB6</u> ഗ്ര	CDS 3' incomplete TSL:3
	Cnot1-216	ENSMUST00000213006.1	7479	<u>1614aa</u>	Nonsense mediated decay	879	A0A1D5RMD8	TSL:1
	Cnot1-209	ENSMUST00000212323.1	3277	<u>213aa</u>	Nonsense mediated decay	(A0A1D5RM03 @	TSL:2
	Cnot1-212	ENSMUST00000212415.1	956	<u>122aa</u>	Nonsense mediated decay	870	A0A1D5RM04 @	CDS 5' incomplete TSL:5
4	Cnot1-213	ENSMUST00000212535.1	1546	No protein	Processed transcript	870		TSL:NA
	Cnot1-214	ENSMUST00000212556.1	521	No protein	Processed transcript	870	5	TSL:5
	Cnot1-208	ENSMUST00000212302.1	3809	No protein	Retained intron	870		TSL:1
	Cnot1-207	ENSMUST00000212228.1	2925	No protein	Retained intron	870		TSL:1
	Cnot1-210	ENSMUST00000212340.1	<mark>23</mark> 81	No protein	Retained intron	870		TSL:NA
	Cnot1-206	ENSMUST00000212195.1	696	No protein	Retained intron	870		TSL:3
	Cnot1-215	ENSMUST00000212712.1	656	No protein	Retained intron	870		TSL:3
	Cnot1-204	ENSMUST00000211937.1	500	No protein	Retained intron	8.00	5	TSL:3
	Cnot1-211	ENSMUST00000212369.1	479	No protein	Retained intron			TSL:2
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The strategy is based on the design of *Cnot1-203* transcript, the transcription is shown below:

protein coding

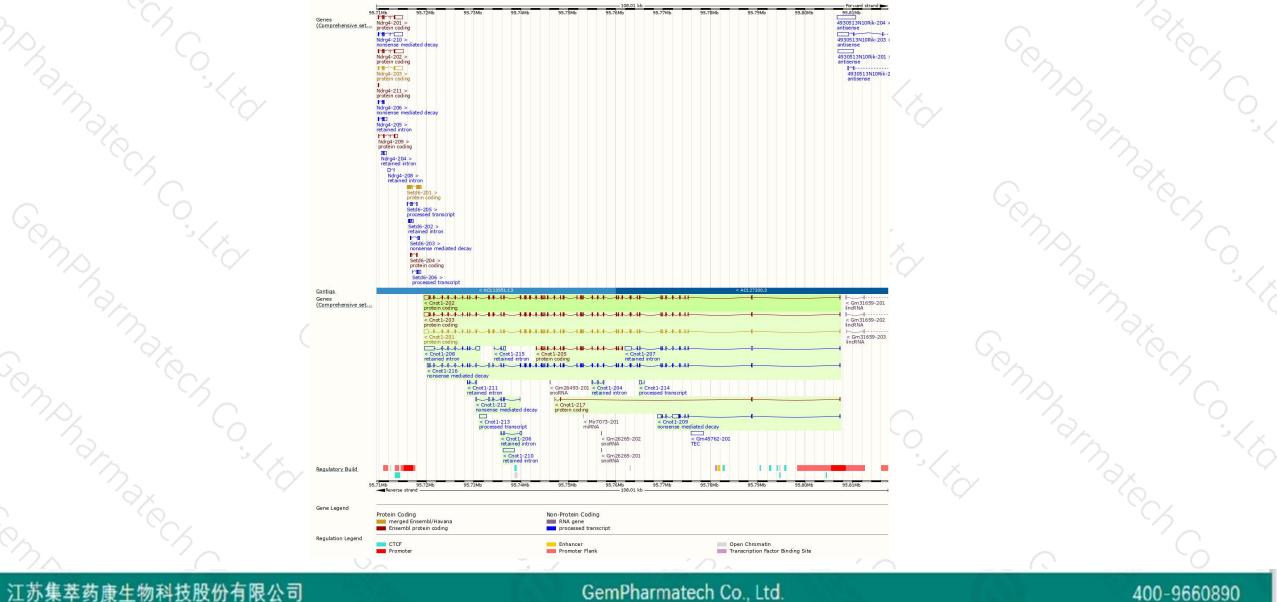
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Genomic location distribution





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Protein domain

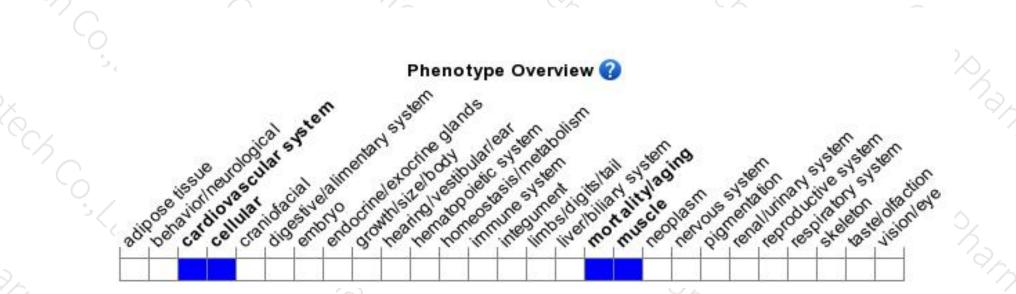




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Mouse phenotype description(MGI)





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 hmozygous for a conditional allele activated in cardiomyocytes exhibit postnatal lethality, decreased cardiac muscle contractility, prolonged QT interval and cardiac muscle cell death.



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



