

Slc19a3 Cas9-CKO Strategy

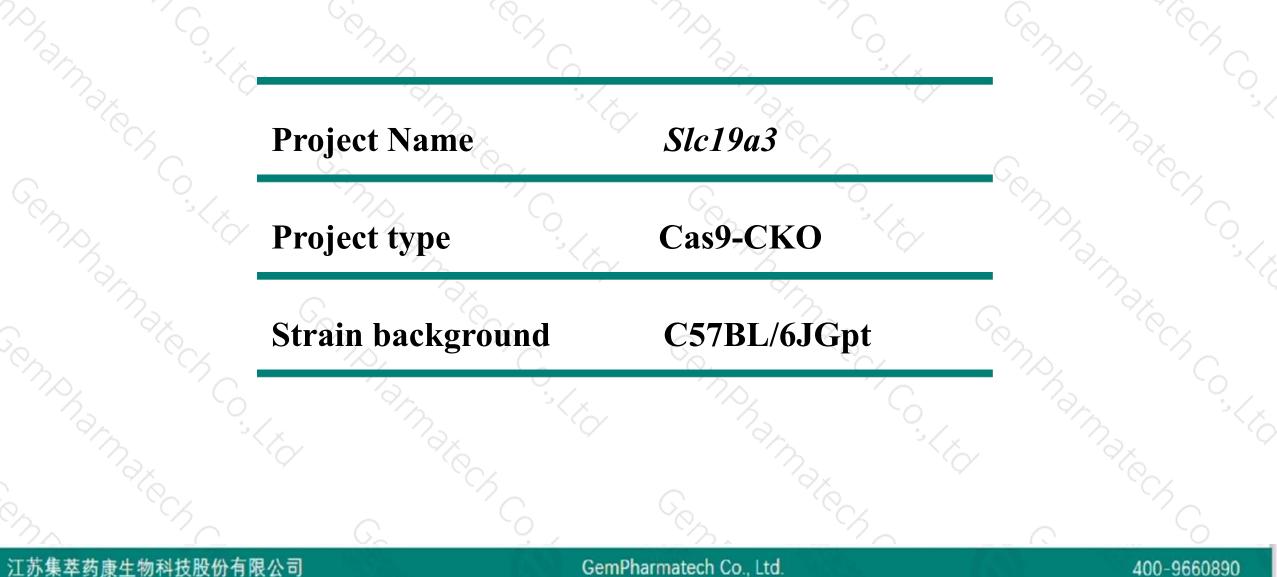
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Reviewer: Rui Xiong

Design Date: 2020-6-9

Project Overview





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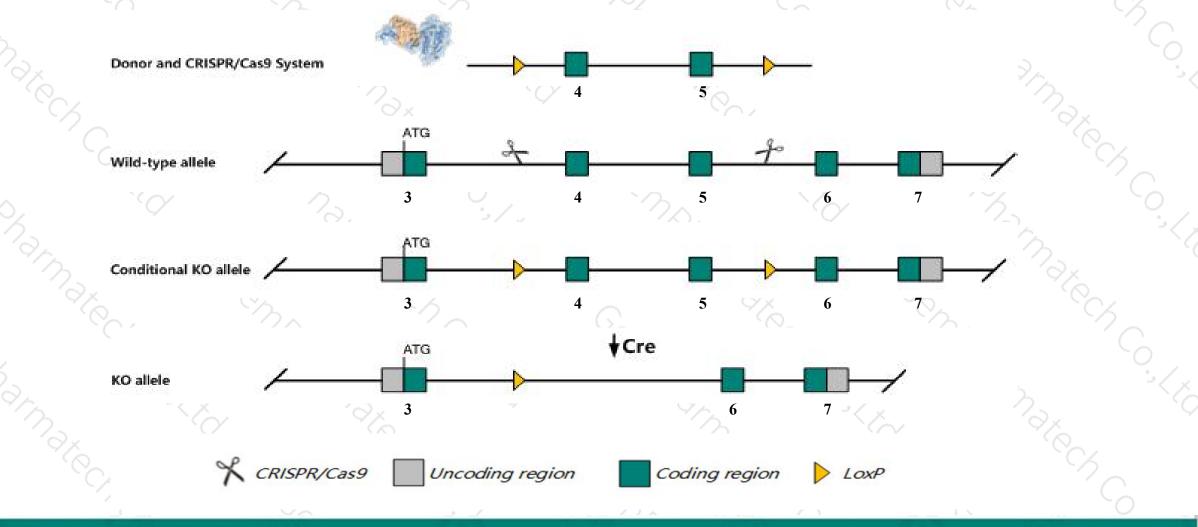
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Conditional Knockout strategy



400-9660890

This model will use CRISPR/Cas9 technology to edit the *Slc19a3* gene. The schematic diagram is as follows:



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The Slc19a3 gene has 3 transcripts. According to the structure of Slc19a3 gene, exon4-exon5 of Slc19a3-201 (ENSMUST00000045560.14) transcript is recommended as the knockout region. The region contains 1001bp coding sequence. Knock out the region will result in disruption of protein function.

In this project we use CRISPR/Cas9 technology to modify *Slc19a3* 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,mice homozygous for a knock-out allele exhibit premature death within a year of age, impaired thiamin uptake, lethargy, cachexia, injured liver parenchyma, hepatic necrosis, liver and kidney inflammmation, and nephrosclerosis.
- The Slc19a3 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 loxp insertion on gene transcription, RNA splicing and protein translation cannot be predicted at existing technological level.

Gene information (NCBI)



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SIc19a3 solute carrier family 19, member 3 [Mus musculus (house mouse)]

Gene ID: 80721, updated on 13-Mar-2020

Summary

Official Symbol	SIc19a3 provided by MGI
Official Full Name	solute carrier family 19, member 3 provided by MGI
Primary source	MGI:MGI:1931307
See related	Ensembl:ENSMUSG0000038496
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	A230084E24Rik, AI788884, ThTr2
Expression	Biased expression in kidney adult (RPKM 6.0), duodenum adult (RPKM 1.7) and 10 other tissuesSee more
Orthologs	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		
Slc19a3-201	ENSMUST0000045560.14	2974	<u>488aa</u>	Protein coding	CCDS15101	<u>Q99PL8</u>	TSL:1 GENCODE basic APPRIS P1		
Slc19a3-203	ENSMUST00000164473.1	1550	<u>488aa</u>	Protein coding	CCDS15101	<u>Q99PL8</u>	TSL:1 GENCODE basic APPRIS P1		
Slc19a3-202	ENSMUST00000142805.1	1892	No protein	Processed transcript	2	2	TSL:1		

The strategy is based on the design of *Slc19a3-201* transcript, the transcription is shown below:

< Slc19a3-201 protein coding Reverse strand

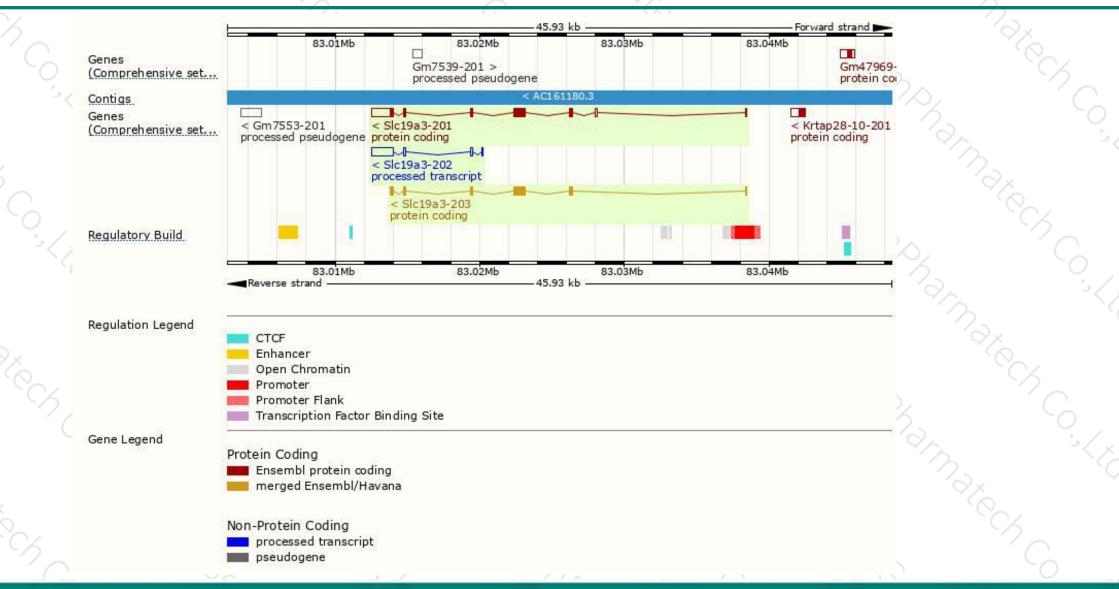
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Genomic location distribution





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Protein domain



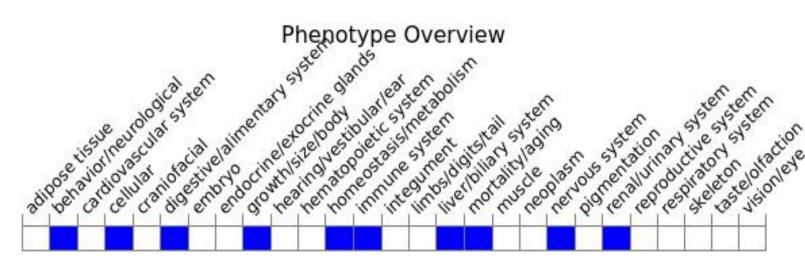
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PIRSF	Thiamine transporter 2						
PANTHER.	Reduced folate carrier PTHR10686:SF31 Reduced folate carrier						
CDD	cd17325	5					
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Mouse phenotype description(MGI)





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 homozygous for a knock-out allele exhibit premature death within a year of age, impaired thiamin uptake, lethargy, cachexia, injured liver parenchyma, hepatic necrosis, liver and kidney inflammation, and nephrosclerosis.

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



