

Acadl Cas9-KO Strategy

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Project Overview



Project Name Acadl

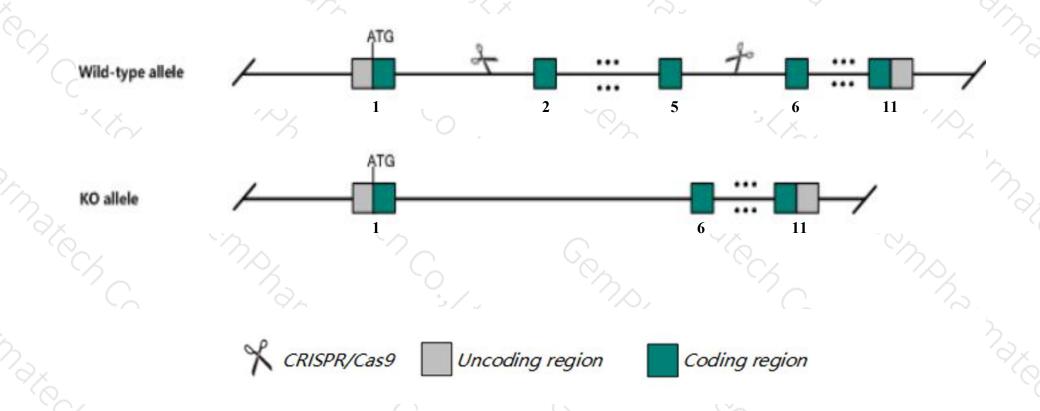
Project type Cas9-KO

Strain background C57BL/6JGpt

Knockout strategy



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



Technical routes



- ➤ The *Acadl* gene has 2 transcripts. According to the structure of *Acadl* gene, exon2-exon5 of *Acadl-201*(ENSMUST00000027153.5) transcript is recommended as the knockout region. The region contains 526bp coding sequence.

 Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Acadl* gene. The brief process is as follows: CRISPR/Cas9 system

Notice



- According to the existing MGI data, Homozygous mutation of this gene results in reduced litter size, sudden death between 2-14 weeks of age, reduced serum glucose levels, lipid accumulation in the liver and heart, and cardiomyopathy. Heterozygous mutant animals exhibit reduced litter size.
- > Gm27934 and Gm15793 gene will be deleted.
- The *Acadl* 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)



Acadl acyl-Coenzyme A dehydrogenase, long-chain [Mus musculus (house mouse)]

Gene ID: 11363, updated on 13-Mar-2020

Summary

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Official Full Name acyl-Coenzyme A dehydrogenase, long-chain provided by MGI

Primary source MGI:MGI:87866

Official Symbol Acadl provided by MGI

See related Ensembl:ENSMUSG00000026003

Gene type protein coding
RefSeq status REVIEWED
Organism <u>Mus musculus</u>

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

Murinae; Mus; Mus

Also known as LCAD; C79855; AA960361; AU018452

Summary This gene encodes a homotetrameric mitochondrial flavoprotein and is a member of the acyl-CoA dehydrogenase family. Members of this family catalyze the first step

of fatty acid beta-oxidation, forming a C2-C3 trans-double bond in a FAD-dependent reaction. As beta-oxidation cycles through its four steps, each member of the acyl-CoA dehydrogenase family works at an optimum fatty acid chain-length. This enzyme has its optimum length between C12- and C16-acylCoA. In mice, deficiency

of this gene can cause sudden death, cardiomyopathy as well as fasting and cold intolerance. [provided by RefSeq, Nov 2012]

Expression Broad expression in heart adult (RPKM 89.2), liver E18 (RPKM 50.6) and 22 other tissues See more

Orthologs human all

Genomic context

2 7

Location: 1 C3; 1 33.64 cM

See Acadl in Genome Data Viewer

Exon count: 11

Annotation release	Status	Assembly	Chr	Location	
108	current	GRCm38.p6 (GCF_000001635.26)	1	NC_000067.6 (6683083966863309, complement)	
Build 37.2	previous assembly	MGSCv37 (GCF_000001635.18)	1	NC_000067.5 (6687742766909841, complement)	

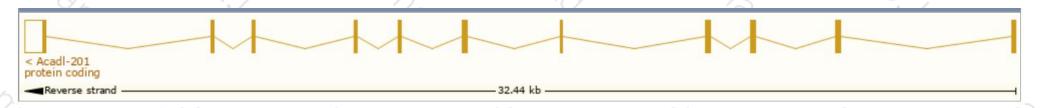
Transcript information (Ensembl)



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

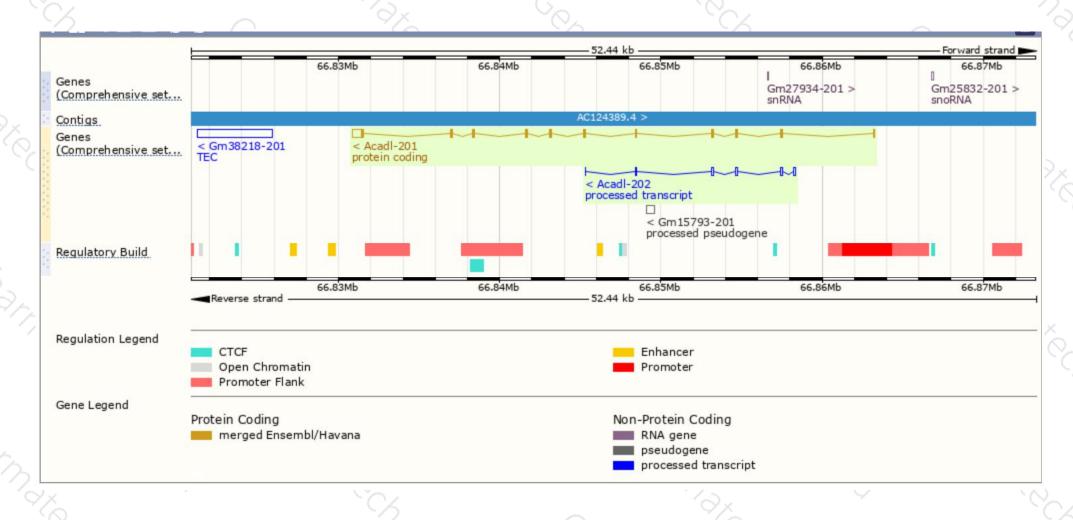
Name	Transcript ID 🗼	bp 🍦	Protein	Biotype	CCDS	UniProt 🍦		Flags	\$
AcadI-201	ENSMUST00000027153.5	1916	<u>430aa</u>	Protein coding	CCDS15023&	A0A0R4J083 怪	TSL:1	GENCODE basic	APPRIS P1
AcadI-202	ENSMUST00000139208.1	649	No protein	Processed transcript	-	÷	TSL:3		

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



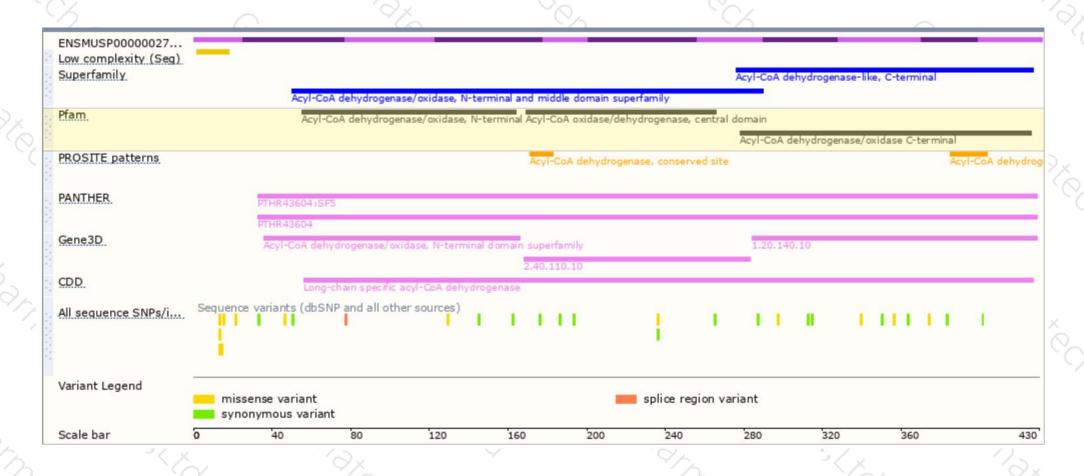
Genomic location distribution





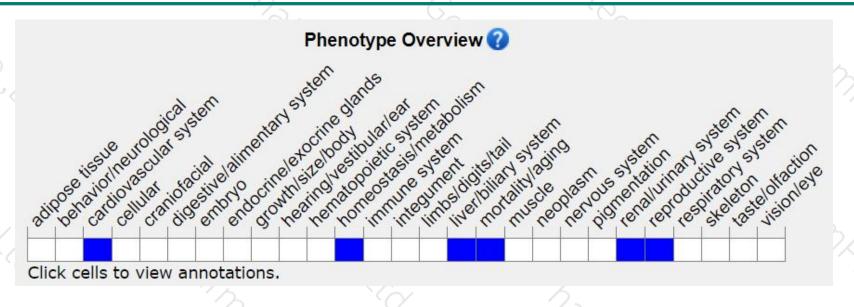
Protein domain





Mouse phenotype description(MGI)





Phenotypes affected by the gene are marked in blue.Data quoted from MGI database(http://www.informatics.jax.org/).

Homozygous mutation of this gene results in reduced litter size, sudden death between 2-14 weeks of age, reduced serum glucose levels, lipid accumulation in the liver and heart, and cardiomyopathy. Heterozygous mutant animals exhibit reduced litter size.



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





