

Lta4h Cas9-KO Strategy

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Project Overview



Project Name Lta4h

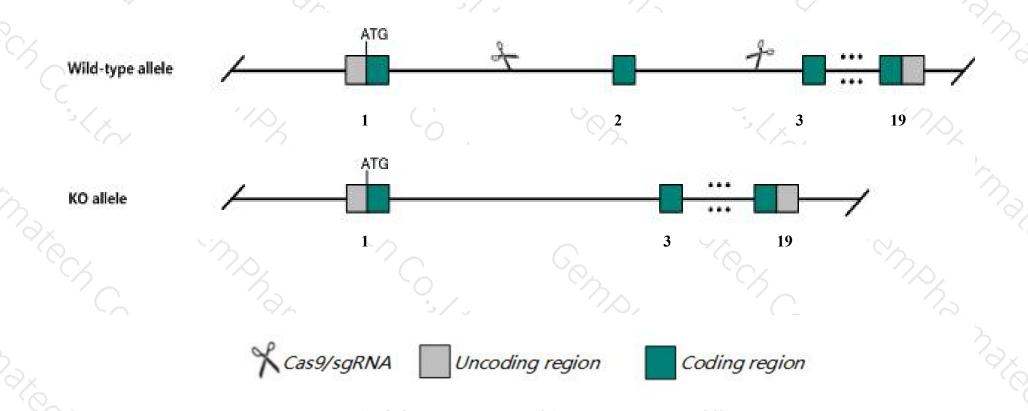
Project type Cas9-KO

Strain background C57BL/6J

Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Lta4h* gene. The schematic diagram is as follows:



Technical routes



- ➤ The *Lta4h* gene has 8 transcripts. According to the structure of *Lta4h* gene, exon2 of *Lta4h-201*(ENSMUST00000016033.8) transcript is recommended as the knockout region. The region contains 131bp coding sequence.

 Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Lta4h* gene. The brief process is as follows: sgRNA was transcribed in vitro. Cas9 and sgRNA were microinjected into the fertilized eggs of C57BL/6J 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/6J mice.

Notice



- ➤ According to the existing MGI data, Mice homozygous for disruptions in this gene have grossly normal phenotypes. Inflammatory reactions are reduced as are some other immunological responses.
- The *Lta4h* gene is located on the Chr10. 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)



Lta4h leukotriene A4 hydrolase [Mus musculus (house mouse)]

Gene ID: 16993, updated on 19-Mar-2019

Summary

↑ ?

Official Symbol Lta4h provided by MGI

Official Full Name leukotriene A4 hydrolase provided by MGI

Primary source MGI:MGI:96836

See related Ensembl:ENSMUSG00000015889

Gene type protein coding
RefSeq status REVIEWED
Organism Mus musculus

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

Muroidea; Muridae; Murinae; Mus; Mus

Summary The protein encoded by this gene is an enzyme that contains both hydrolase and aminopeptidase activities. The hydrolase activity is used

in the final step of the biosynthesis of leukotriene B4, a proinflammatory mediator. The aminopeptidase activity has been shown to degrade proline-glycine-proline (PGP), a neutrophil chemoattractant and biomarker for chronic obstructive pulmonary disease (COPD). Two transcript

variants encoding different isoforms have been found for this gene. [provided by RefSeq, Sep 2015]

Expression Ubiquitous expression in large intestine adult (RPKM 63.2), duodenum adult (RPKM 45.9) and 28 other tissuesSee more

Orthologs <u>human all</u>

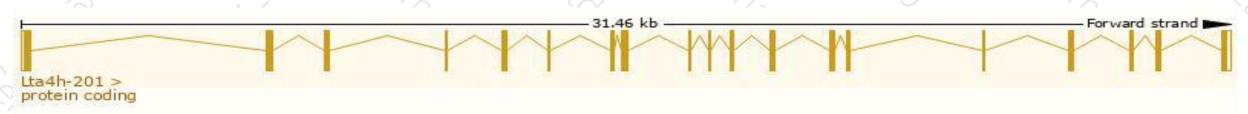
Transcript information (Ensembl)



The gene has 8 transcripts, all transcripts are shown below:

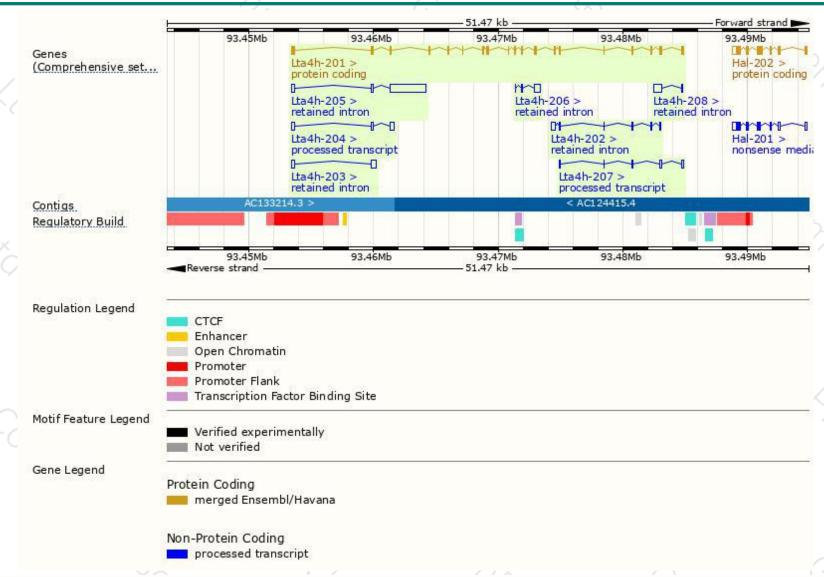
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Lta4h-201	ENSMUST00000016033.8	2029	<u>611aa</u>	Protein coding	CCDS24125	P24527	TSL:1 GENCODE basic APPRIS P1
Lta4h-204	ENSMUST00000215419.1	687	No protein	Processed transcript	.		TSL:1
Lta4h-207	ENSMUST00000216931.1	540	No protein	Processed transcript	48	-	TSL:2
Lta4h-205	ENSMUST00000216146.1	3225	No protein	Retained intron	20	-	TSL:1
Lta4h-208	ENSMUST00000217556.1	752	No protein	Retained intron	5 4	a l	TSL:2
Lta4h-202	ENSMUST00000214527.1	690	No protein	Retained intron	.		TSL:3
Lta4h-203	ENSMUST00000215224.1	629	No protein	Retained intron	49	2	TSL:2
Lta4h-206	ENSMUST00000216174.1	585	No protein	Retained intron	26	-	TSL:1

The strategy is based on the design of *Lta4h-201* transcript, The transcription is shown below



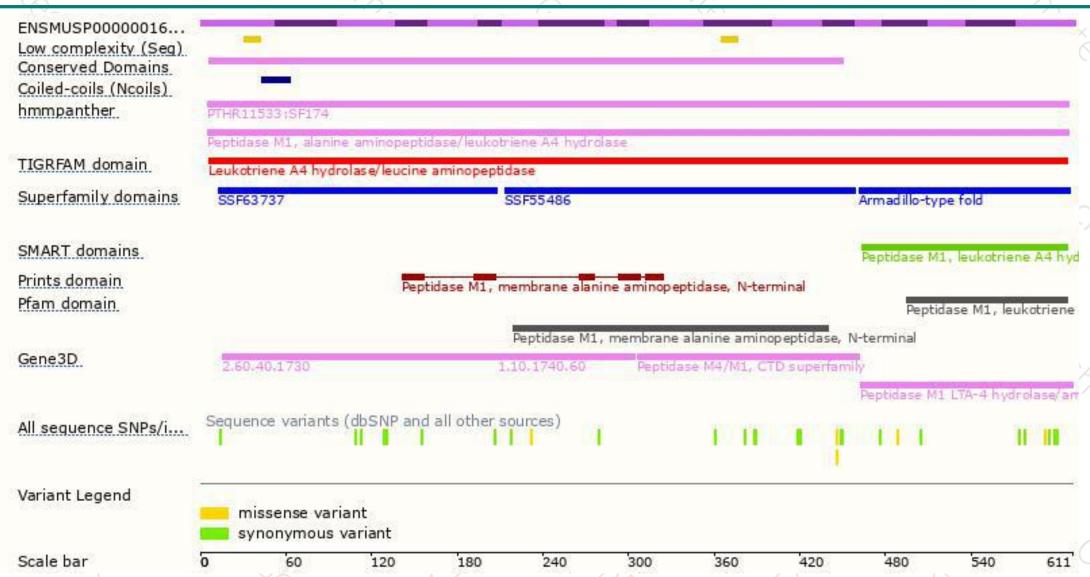
Genomic location distribution





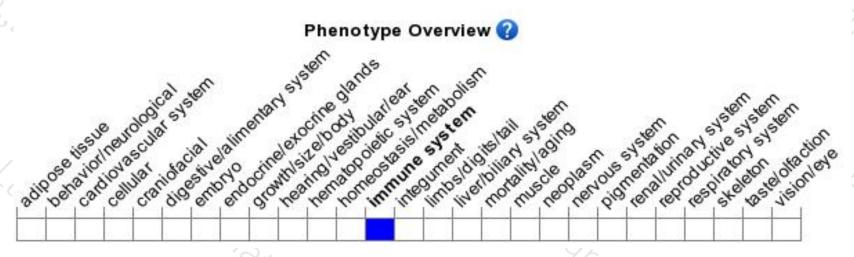
Protein domain





Mouse phenotype description(MGI)





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

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

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