

Dio1 Cas9-KO Strategy

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Design Date: 2019-7-26

Project Overview



Project Name

Project type

Cas9-KO

Dio1

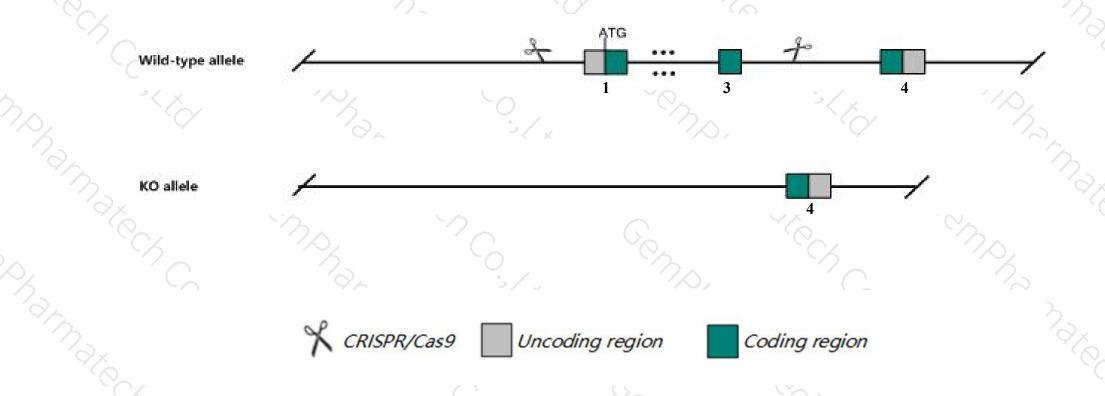
Strain background

C57BL/6JGpt

Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Dio1* gene. The schematic diagram is as follows:



Technical routes



- ➤ The *Dio1* gene has 8 transcripts. According to the structure of *Dio1* gene, exon1-exon3 of *Dio1-201* (ENSMUST00000082426.9) transcript is recommended as the knockout region. The region contains start codon ATG. Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Dio1* gene. The brief process is as follows: CRISPR/Cas9 system v

Notice



- ➤ According to the existing MGI data, Mice homozygous for a disruption in this gene display elevated thyroxine (T4) and reverse triiodothyronine (rT3) levels and changes in the metabolism and excretion of iodothyronines.
- The *Dio1* gene is located on the Chr4. 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)



Dio1 deiodinase, iodothyronine, type I [Mus musculus (house mouse)]

Gene ID: 13370, updated on 19-Feb-2019

Summary

△ ?

Official Symbol Dio1 provided by MGI

Official Full Name deiodinase, iodothyronine, type I provided by MGI

Primary source MGI:MGI:94896

See related Ensembl:ENSMUSG00000034785

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

Also known as 5DI, D1, ITDI1, TXDI1

Summary The protein encoded by this gene belongs to the iodothyronine deiodinase family. It catalyzes the activation, as well as the inactivation of

thyroid hormone by outer and inner ring deiodination, respectively. The activation reaction involves the conversion of the prohormone thyroxine

(3,5,3',5'-tetraiodothyronine, T4), secreted by the thyroid gland, to the bioactive thyroid hormone (3,5,3'-triiodothyronine, T3) by 5'-

deiodination. This protein is expressed predominantly in the liver and kidney and provides most of the circulating T3, which is essential for growth, differentiation and basal metabolism in vertebrates. This protein is a selenoprotein, containing the rare amino acid selenocysteine (Sec) at its active site. Sec is encoded by the UGA codon, which normally signals translation termination. The 3' UTRs of selenoprotein mRNAs contain a conserved stem-loop structure, designated the Sec insertion sequence (SECIS) element, that is necessary for the

recognition of UGA as a Sec codon, rather than as a stop signal. [provided by RefSeq, Apr 2016]

Expression Biased expression in liver adult (RPKM 54.1), kidney adult (RPKM 25.9) and 5 other tissuesSee more

Orthologs human all

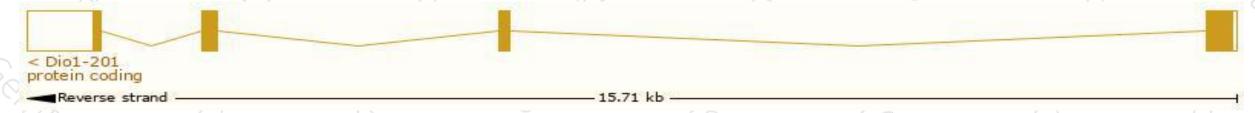
Transcript information (Ensembl)



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

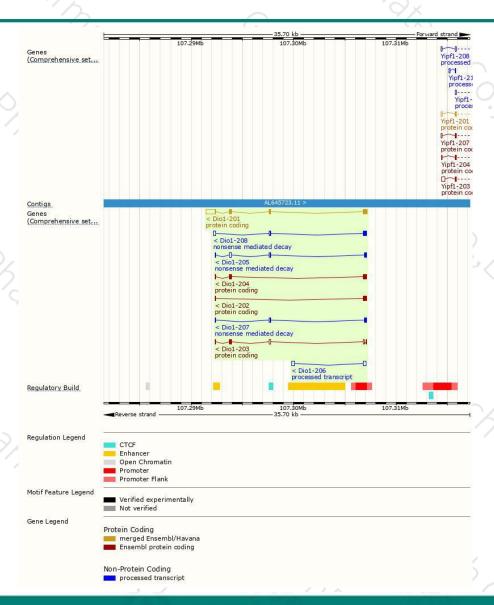
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Dio1-201	ENSMUST00000082426.9	1690	257aa	Protein coding	CCDS18436	A0A0R4J110	TSL:1 GENCODE basic APPRIS P1
Dio1-204	ENSMUST00000129138.1	632	208aa	Protein coding		D3YX15	CDS 3' incomplete TSL:5
Dio1-203	ENSMUST00000126291.1	616	<u>61aa</u>	Protein coding	-	D6RDQ6	CDS 3' incomplete TSL:5
Dio1-202	ENSMUST00000106748.1	396	<u>113aa</u>	Protein coding	-	D3YVM7	TSL:5 GENCODE basic
Dio1-208	ENSMUST00000150974.1	761	<u>125aa</u>	Nonsense mediated decay	5	A0A0R4J1U2	TSL:5
Dio1-205	ENSMUST00000134366.7	671	82aa	Nonsense mediated decay	-	D6RCH6	TSL:5
Dio1-207	ENSMUST00000147709.1	431	<u>82aa</u>	Nonsense mediated decay	÷	D6RCH6	TSL:5
Dio1-206	ENSMUST00000145332.1	565	No protein	Processed transcript	-	727	TSL:5
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The strategy is based on the design of *Dio1-201* transcript, The transcription is shown below



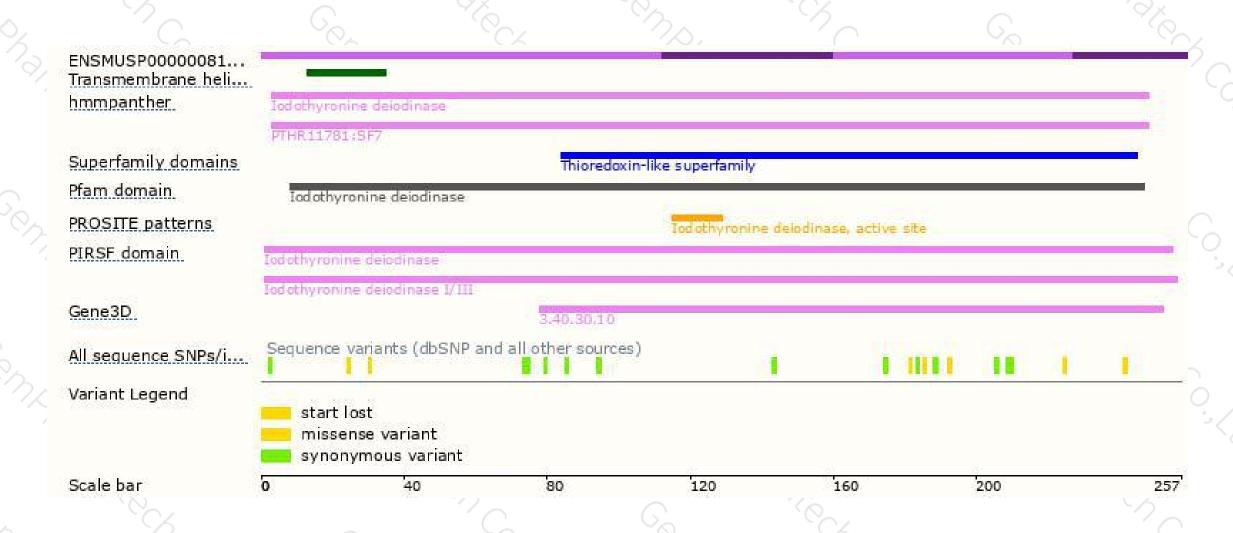
Genomic location distribution





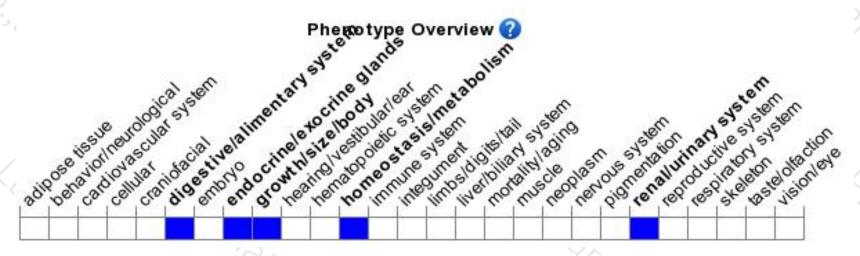
Protein domain





Mouse phenotype description(MGI)





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

According to the existing MGI data, Mice homozygous for a disruption in this gene display elevated thyroxine (T4) and reverse triiodothyronine (rT3) levels and changes in the metabolism and excretion of iodothyronines.



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





