

# Ttr Cas9-KO Strategy

Designer: Xiaojing Li

Design Date: 2019-9-16

Reviewer: JiaYu

## **Project Overview**



Project Name Ttr

**Project type** 

Cas9-KO

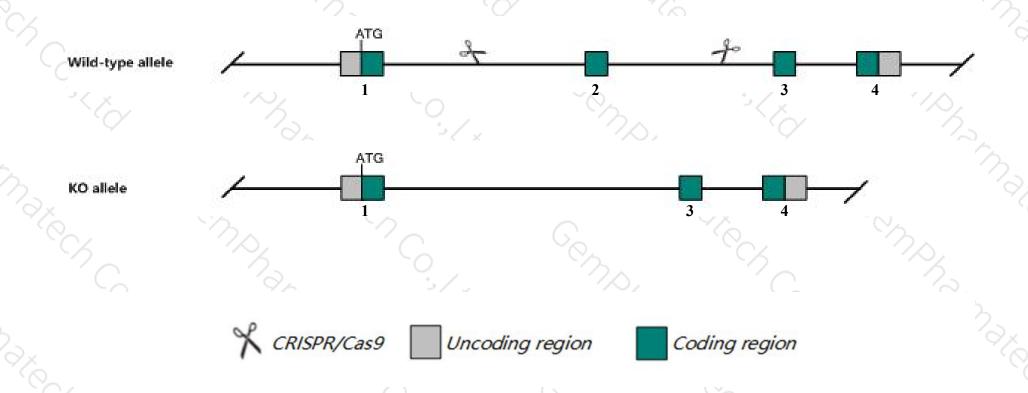
Strain background

C57BL/6JGpt

## **Knockout strategy**



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



### **Technical routes**



- ➤ The *Ttr* gene has 1 transcript. According to the structure of *Ttr* gene, exon2 of *Ttr-201*(ENSMUST00000075312.4) 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 *Ttr* gene. The brief process is as follows: CRISPR/Cas9 system we

### **Notice**



- > According to the existing MGI data, Homozygous mutation of this gene results in decreased circulating thyroxine, triiodothyronine, and retinol levels.
- The *Ttr* gene is located on the Chr18. 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)



#### Ttr transthyretin [Mus musculus (house mouse)]

Gene ID: 22139, updated on 2-Apr-2019

#### Summary

↑ ?

Official Symbol Ttr provided by MGI

Official Full Name transthyretin provided by MGI

Primary source MGI:MGI:98865

See related Ensembl:ENSMUSG00000061808

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 AA408768, AI787086, D17860, prealbumin

**Summary** This gene encodes a carrier protein responsible for the transport of thyroid hormones and retinol. The protein consists of a tetramer of

identical subunits. Due to increased stability of the tetramer form of this encoded protein in mouse, compared to the human protein, this gene product has a reduced tendency to form amyloid fibrils. In humans, this protein binds beta-amyloid preventing its aggregation and providing

a neuroprotective role in Alzheimer's disease. [provided by RefSeq, Mar 2010]

Expression Biased expression in liver adult (RPKM 3655.3), liver E18 (RPKM 2930.2) and 7 other tissues See more

Orthologs <u>human</u> all

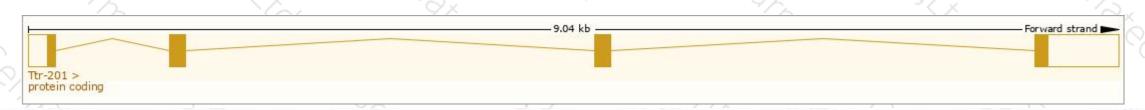
## Transcript information (Ensembl)



The gene has 1 transcript, and the transcript is shown below:

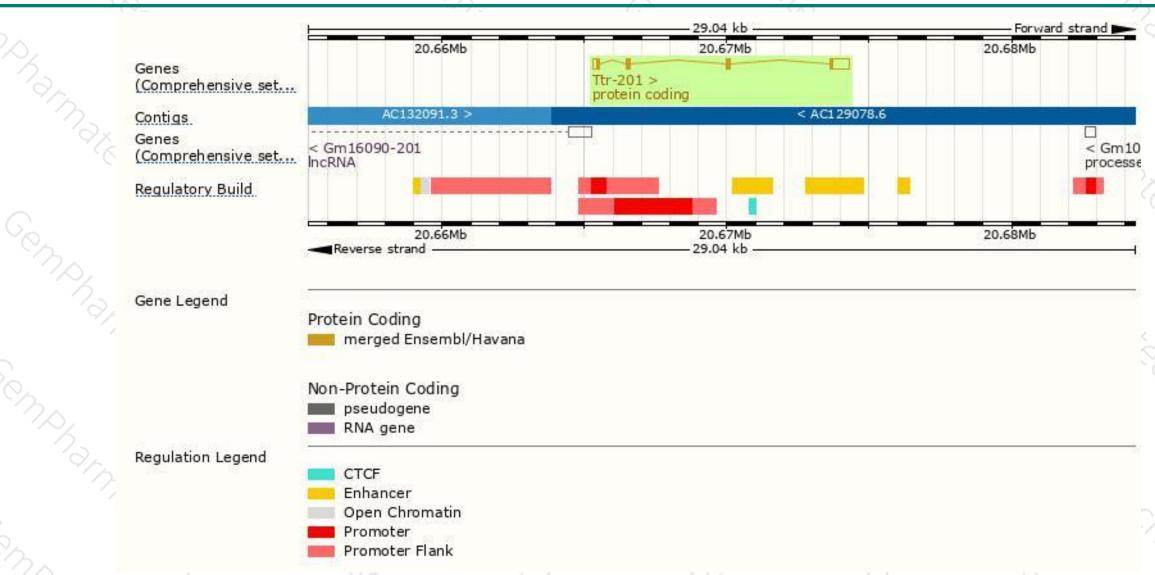
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags	
Ttr-201	ENSMUST00000075312.4	1186	<u>147aa</u>	Protein coding	CCDS29085	P07309 Q5M9K1	TSL:1 GENCODE basic APPRIS P1	L

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



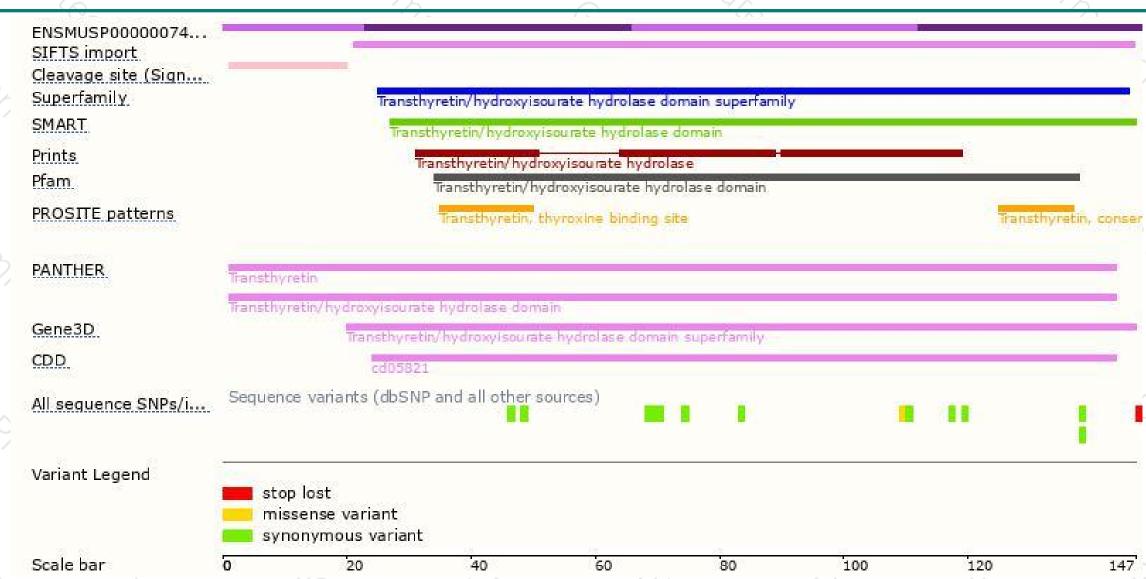
### Genomic location distribution





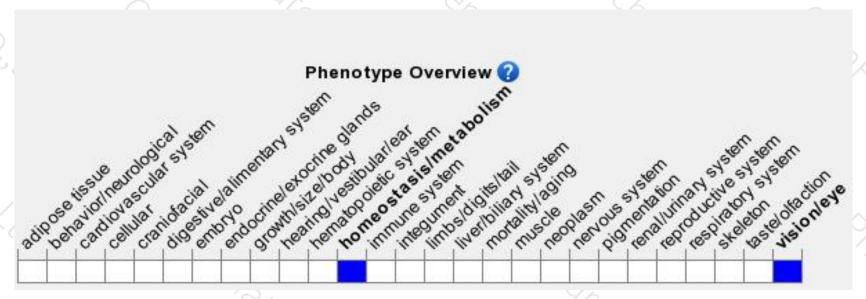
### Protein domain





## 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, Homozygous mutation of this gene results in decreased circulating thyroxine, triiodothyronine, and retinol levels.



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





