

# Foxa3 Cas9-CKO Strategy

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Reviewer: Huan Wang

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## **Project Overview**



Project Name Foxa3

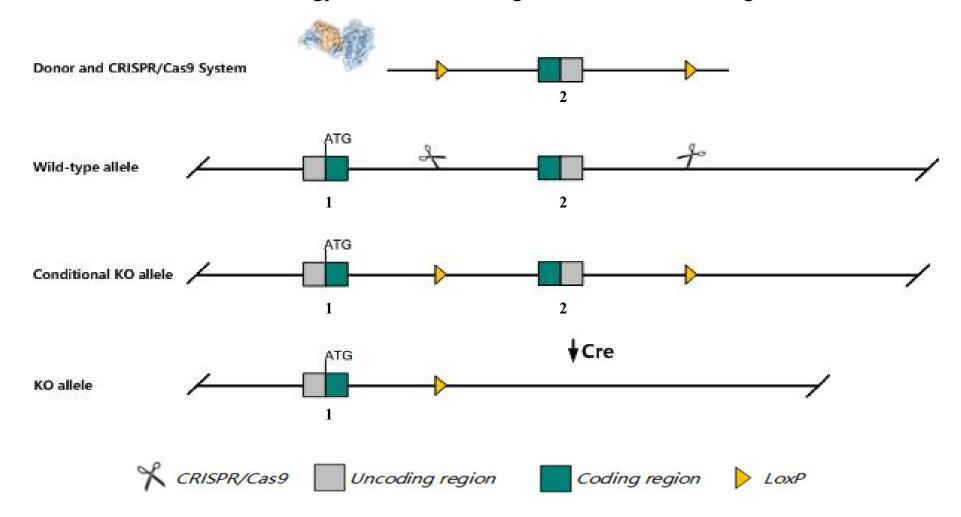
Project type Cas9-CKO

Strain background C57BL/6JGpt

## Conditional Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Foxa3* gene. The schematic diagram is as follows:



### **Technical routes**



The *Foxa3* gene has 1 transcript. According to the structure of *Foxa3* gene, exon2 of *Foxa3-201* (ENSMUST00000036018.5) transcript is recommended as the knockout region. The region contains most of the coding sequence. Knock out the region will result in disruption of protein function.

In this project we use CRISPR/Cas9 technology to modify *Foxa3* 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.

## **Notice**



According to the existing MGI data,homozygotes for a targeted null mutation exhibit reduced expression of several liver-specific and liver-enriched genes. homozygous mice exhibit abnormal glucose homeostasis and reduced male fertility with abnormal spermatogeneis. mice remodel their fat tissues on high fat diets, store less fat, and burn more energy as they age. mutant mice live significantly longer than controls.

The *Foxa3* gene is located on the Chr7. 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



#### Foxa3 forkhead box A3 [Mus musculus (house mouse)]

Gene ID: 15377, updated on 13-Mar-2020

#### Summary

☆ ?

Official Symbol Foxa3 provided by MGI

Official Full Name forkhead box A3 provided by MGI

Primary source MGI:MGI:1347477

See related Ensembl: ENSMUSG00000040891

Gene type protein coding
RefSeq status PROVISIONAL
Organism Mus musculus

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

Muroidea; Muridae; Murinae; Mus; Mus

Also known as Hnf-3g, Hnf3g, Tcf-3g, Tcf3g

Expression Biased expression in stomach adult (RPKM 32.1), colon adult (RPKM 31.9) and 10 other tissuesSee more

Orthologs <u>human all</u>

## Transcript information Ensembl

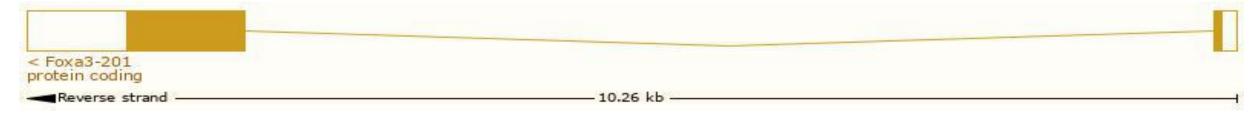




The gene has 1 transcript, and the transcript is shown below:

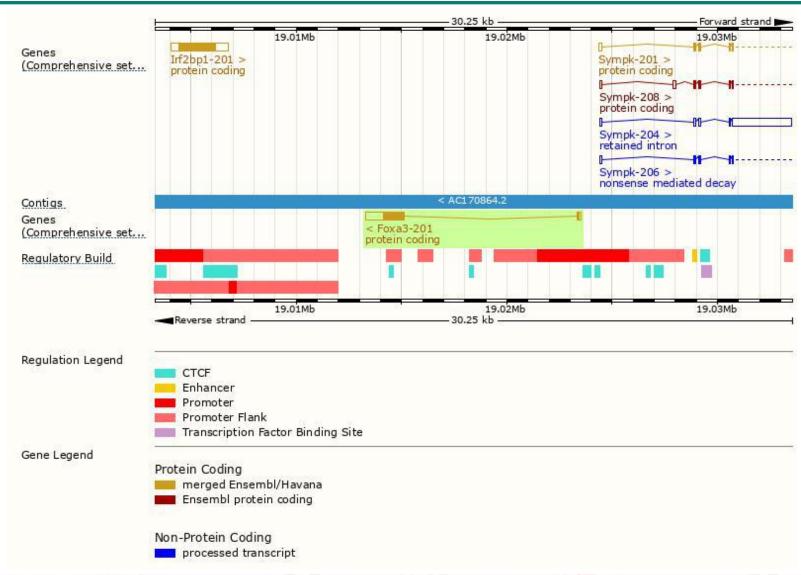
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Foxa3-201	ENSMUST00000036018.5	2037	353aa	Protein coding	CCDS20886	P35584	TSL:1 GENCODE basic APPRIS is a system to annotate alternatively spliced transcripts based on a range of computational methods to identify the most functionally important transcript(s) of a gene. APPRIS P

The strategy is based on the design of *Foxa3-201* transcript, the transcription is shown below:



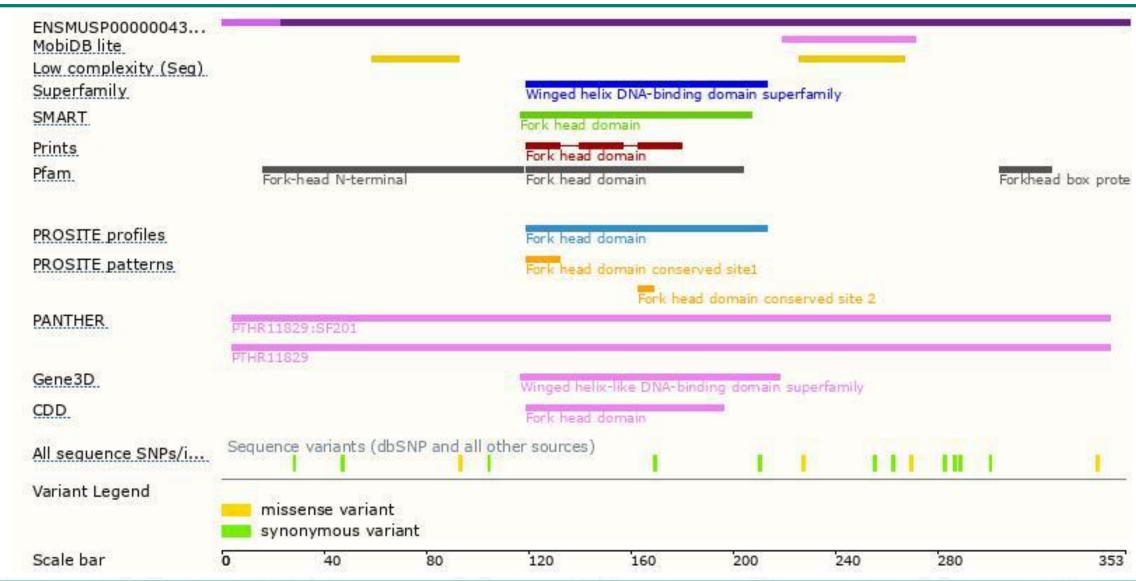
### Genomic location distribution





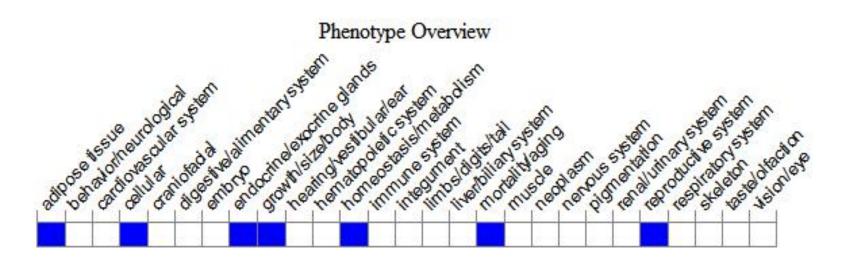
### Protein domain





# 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,homozygotes for a targeted null mutation exhibit reduced expression of several liver-specific and liver-enriched genes. Homozygous mice exhibit abnormal glucose homeostasis and reduced male fertility with abnormal spermatogeneis. Mice remodel their fat tissues on high fat diets, store less fat, and burn more energy as they age. Mutant mice live significantly longer than controls.



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





