

Irs1 Cas9-CKO Strategy

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Reviewer:

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Design Date:

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Project Overview

Project Name

Irs1

Project type

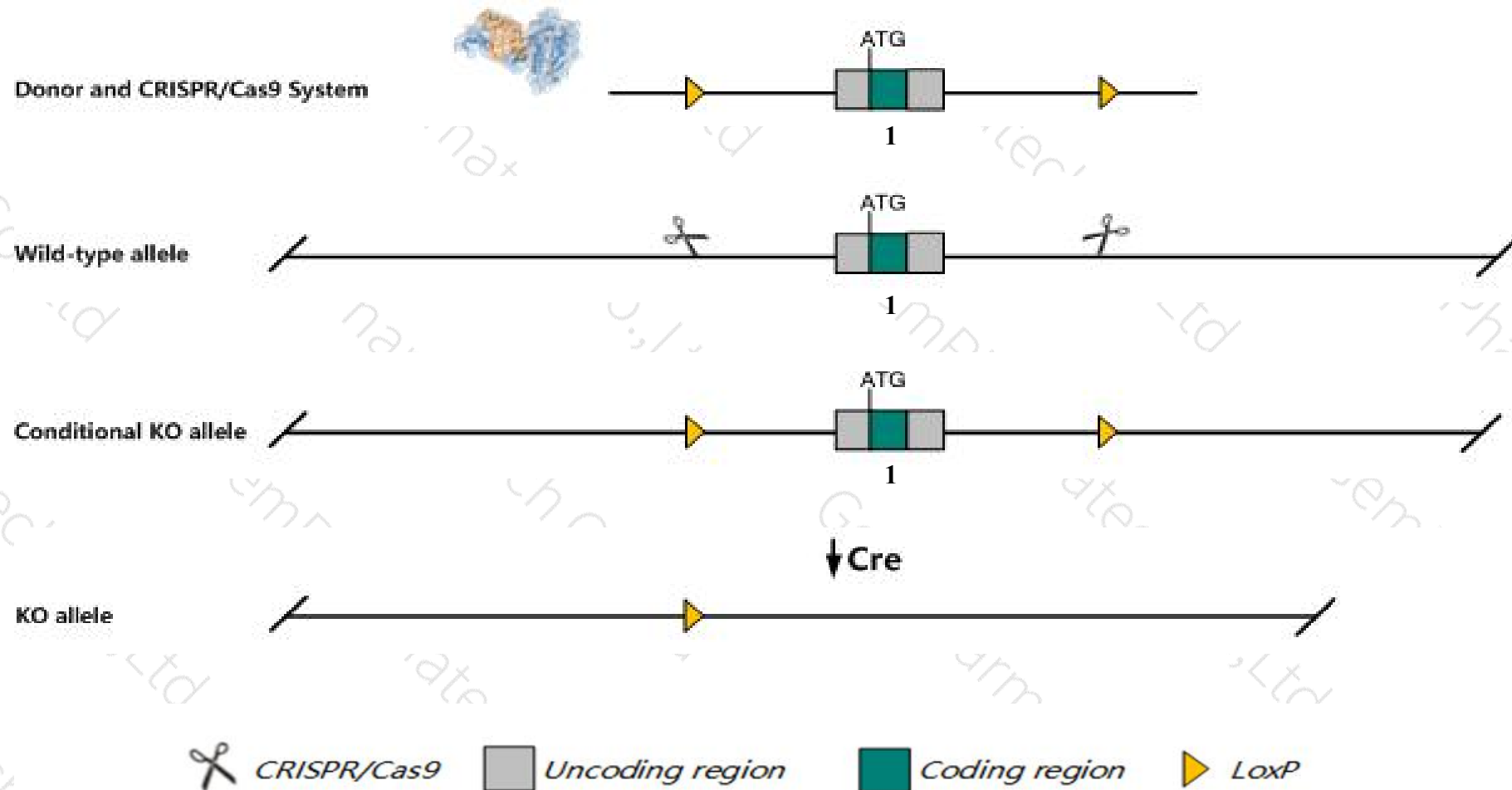
Cas9-CKO

Strain background

C57BL/6JGpt

Conditional Knockout strategy

This model will use CRISPR/Cas9 technology to edit the *Irs1* gene. The schematic diagram is as follows:



Technical routes

- The *Irs1* gene has 1 transcript. According to the structure of *Irs1* gene, exon1 of *Irs1-201* (ENSMUST00000069799.2) transcript is recommended as the knockout region. The region contains all 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 *Irs1* 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.

- According to the existing MGI data, Homozygotes for targeted null mutations exhibit 50 percent reductions in body weights at birth and at 4 months of age, impaired glucose tolerance, and mild insulin and IGF-1 resistance.
- The *Irs1* 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 loxp insertion on gene transcription, RNA splicing and protein translation cannot be predicted at existing technological level.

Gene information (NCBI)

Irs1 insulin receptor substrate 1 [Mus musculus (house mouse)]

Gene ID: 16367, updated on 2-Apr-2019

Summary



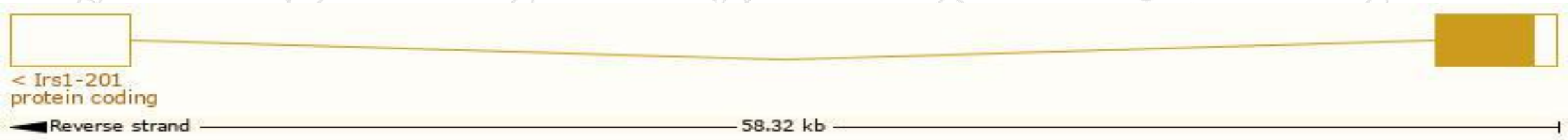
Official Symbol	Irs1 provided by MGI
Official Full Name	insulin receptor substrate 1 provided by MGI
Primary source	MGI:MGI:99454
See related	Ensembl:ENSMUSG00000055980
Gene type	protein coding
RefSeq status	VALIDATED
Organism	Mus musculus
Lineage	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha; Muroidea; Muridae; Murinae; Mus; Mus
Also known as	G972R, IRS-1
Expression	Broad expression in ovary adult (RPKM 15.6), subcutaneous fat pad adult (RPKM 14.2) and 20 other tissues See more
Orthologs	human all

Transcript information (Ensembl)

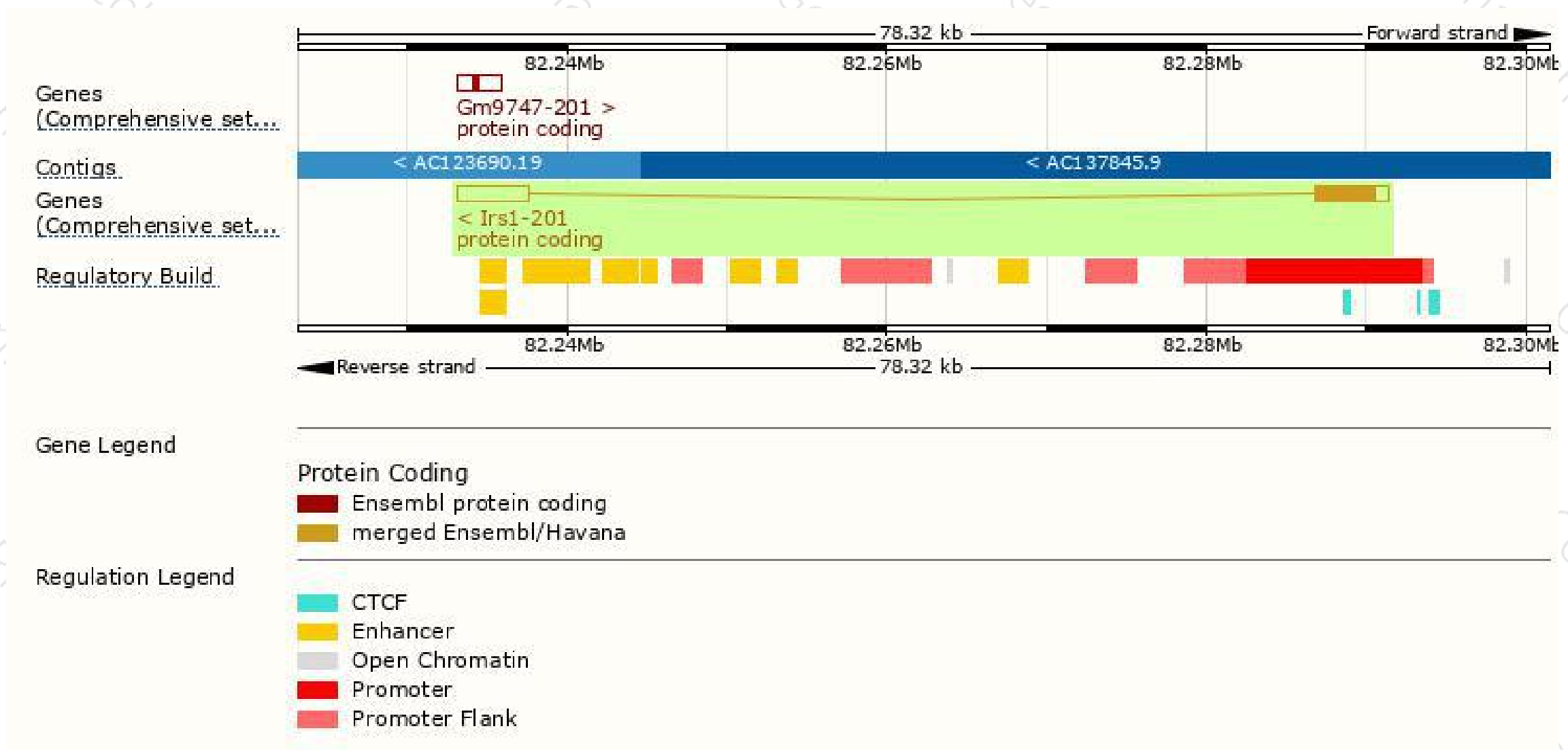
The gene has 1 transcript, and the transcript is shown below:

Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Irs1-201	ENSMUST00000069799.2	9144	1231aa	Protein coding	CCDS15096	Q543V3	TSL:1 GENCODE basic APPRIS P1

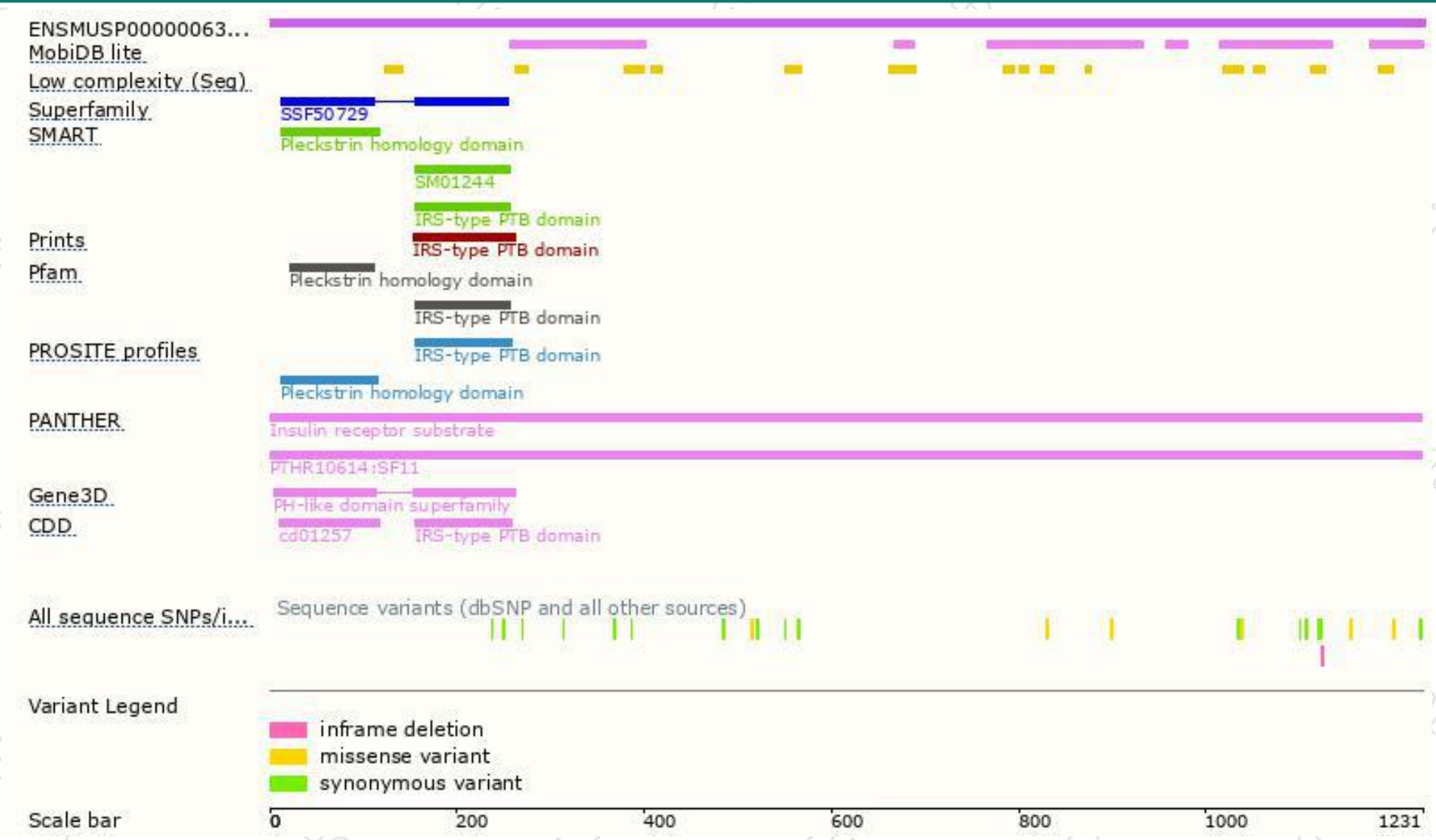
The strategy is based on the design of *Irs1-201* transcript, The transcription is shown below



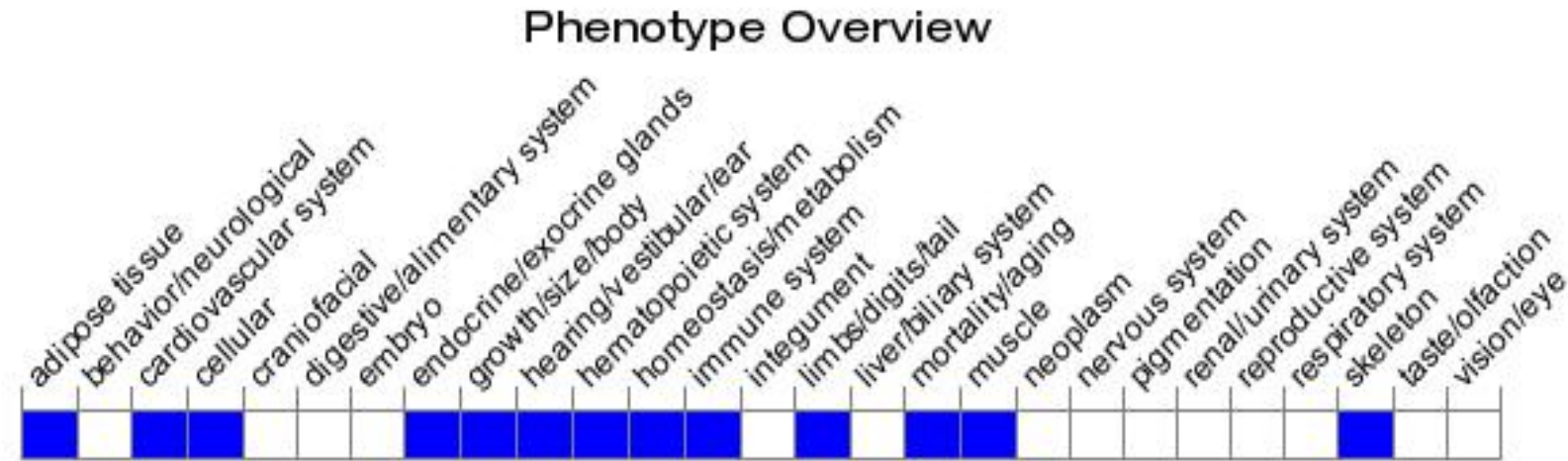
Genomic location distribution



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 targeted null mutations exhibit 50 percent reductions in body weights at birth and at 4 months of age, impaired glucose tolerance, and mild insulin and IGF-1 resistance.

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

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