Npr1 Cas9-CKO Strategy

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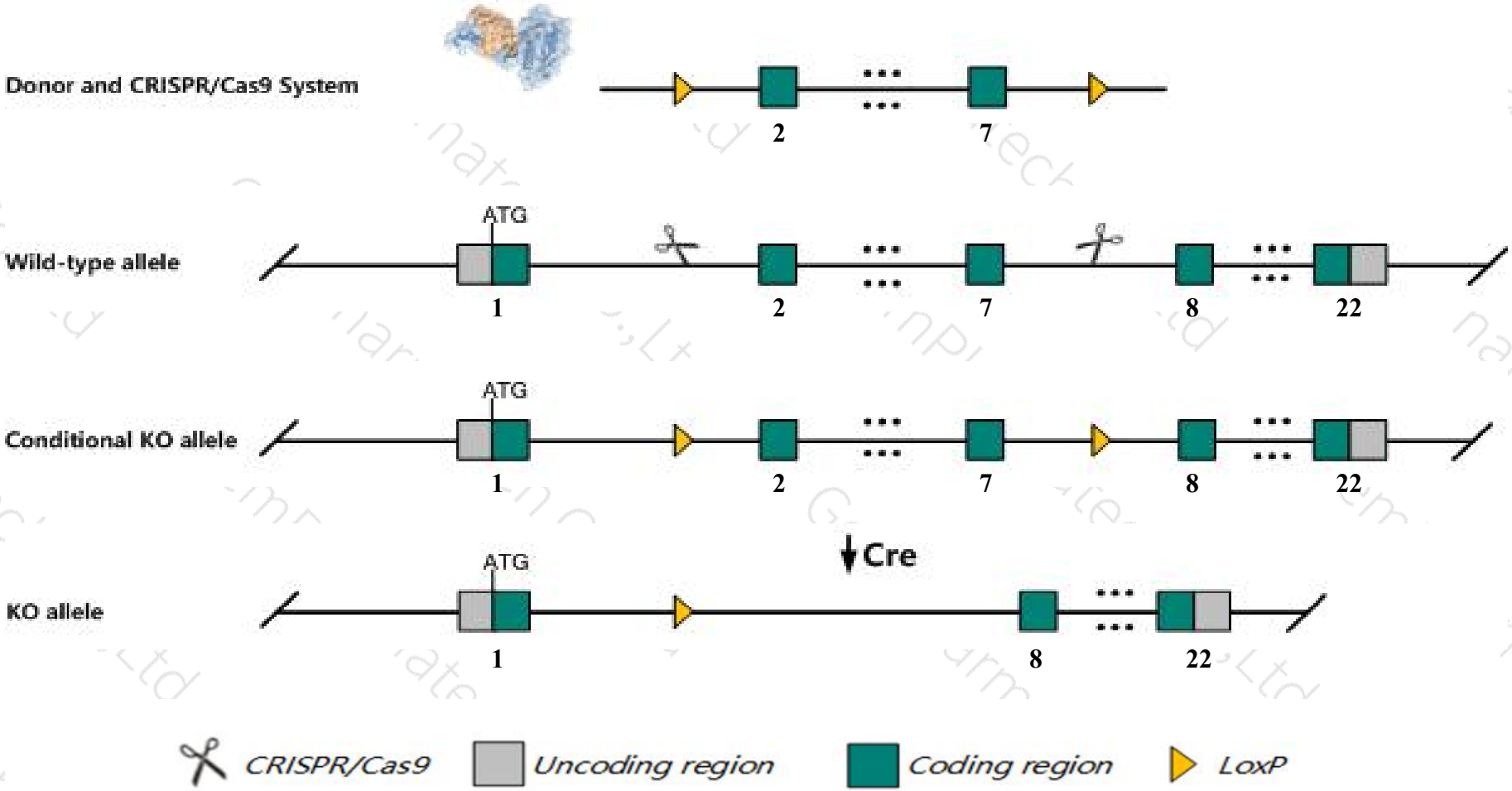
Project Overview

Project Name	***Npr1***
Project type	**Cas9-CKO**
Strain background	**C57BL/6JGpt**

Conditional Knockout strategy

This model will use CRISPR/Cas9 technology to edit the *Npr1* gene. The schematic diagram is as follows:



Technical routes

- The *Npr1* gene has 5 transcripts. According to the structure of *Npr1* gene, exon2-exon7 of *Npr1-201* (ENSMUST00000029540.12) transcript is recommended as the knockout region.The region contains 763bp coding sequence. Knock out the region will result in disruption of protein function.

- In this project we use CRISPR/Cas9 technology to modify *Npr1* 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,Homozygous inactivation of this gene can lead to hypertension, cardiac hypertrophy, lethal vascular events, congestive heart failure in response to volume overload, reduced serum testosterone levels, altered steroidogenesis, and reduced myocardial PMN infiltration and infarct size after I/R injury.

- The *Npr1* gene is located on the Chr3. 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）

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Npr1 natriuretic peptide receptor 1 [Mus musculus (house mouse)]

Gene ID: 18160, updated on 3-Feb-2019

Summary

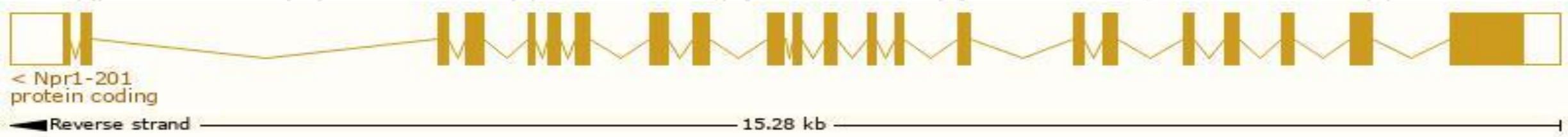
Official Symbol	Npr1 provided by MGI
Official Full Name	natriuretic peptide receptor 1 provided by MGI
Primary source	MGI:MGI:97371
See related	Ensembl:ENSMUSG00000027931
Gene type	protein coding
RefSeq status	VALIDATED
Organism	Mus musculus
Lineage	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha; Muroidea; Muridae; Murinae; Mus; Mus
Also known as	AI893888, GC-A, NPR-A, NPRA, Pndr
Expression	Biased expression in adrenal adult (RPKM 198.6), ovary adult (RPKM 80.4) and 6 other tissues See more
Orthologs	human all

Transcript information（Ensembl）

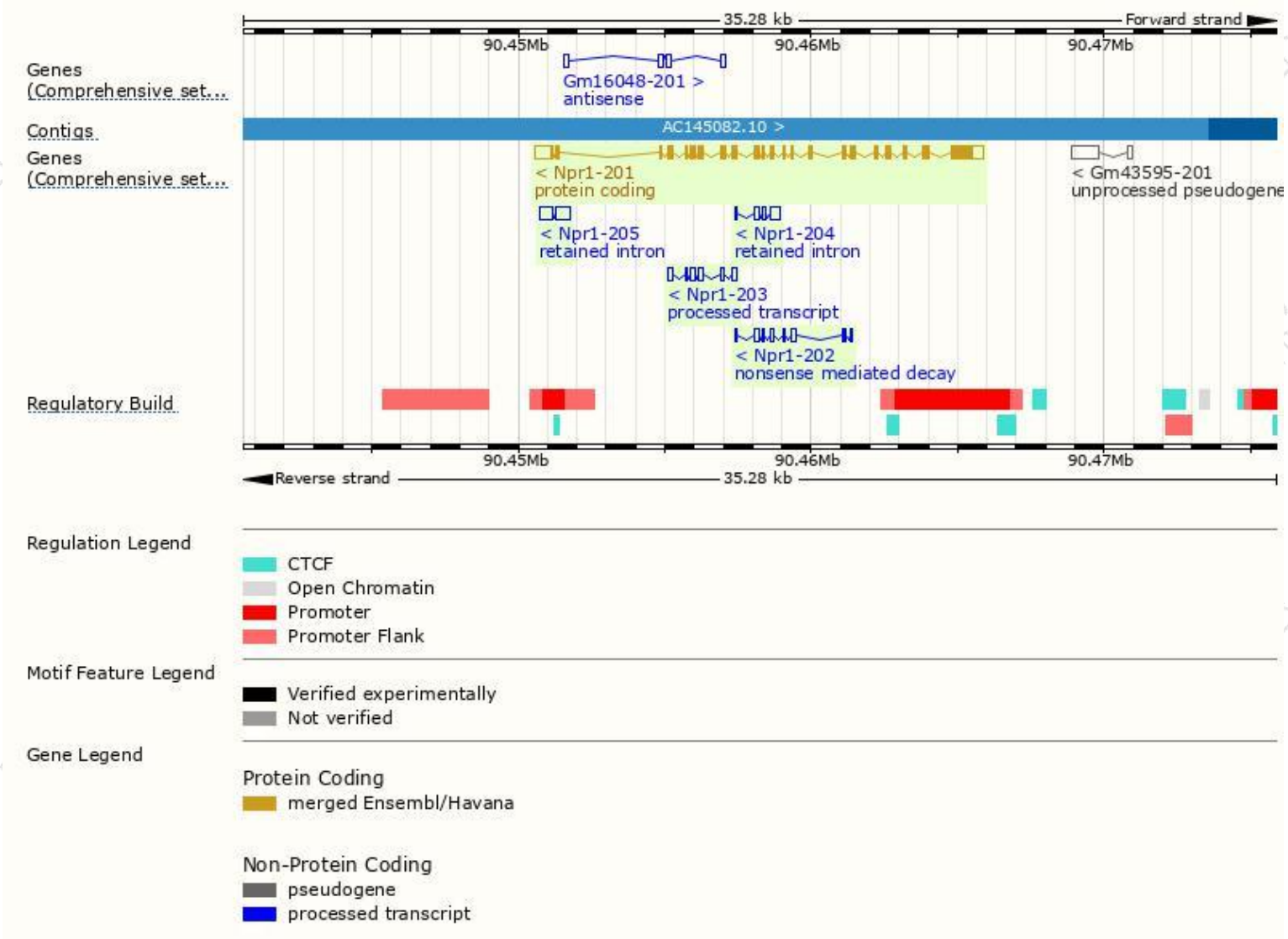
The gene has 5 transcripts,all transcripts are shown below:

Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Npr1-201	ENSMUST00000029540.12	4066	1057aa	Protein coding	CCDS17529	P18293 Q2TAY4	TSL:1 GENCODE basic APPRIS P1
Npr1-202	ENSMUST00000124760.1	790	51aa	Nonsense mediated decay	-	F6W125	CDS 5' incomplete TSL:5
Npr1-203	ENSMUST00000142243.1	767	No protein	Processed transcript	-	-	TSL:5
Npr1-205	ENSMUST00000152510.1	949	No protein	Retained intron	-	-	TSL:2
Npr1-204	ENSMUST00000146991.1	643	No protein	Retained intron	-	-	TSL:3

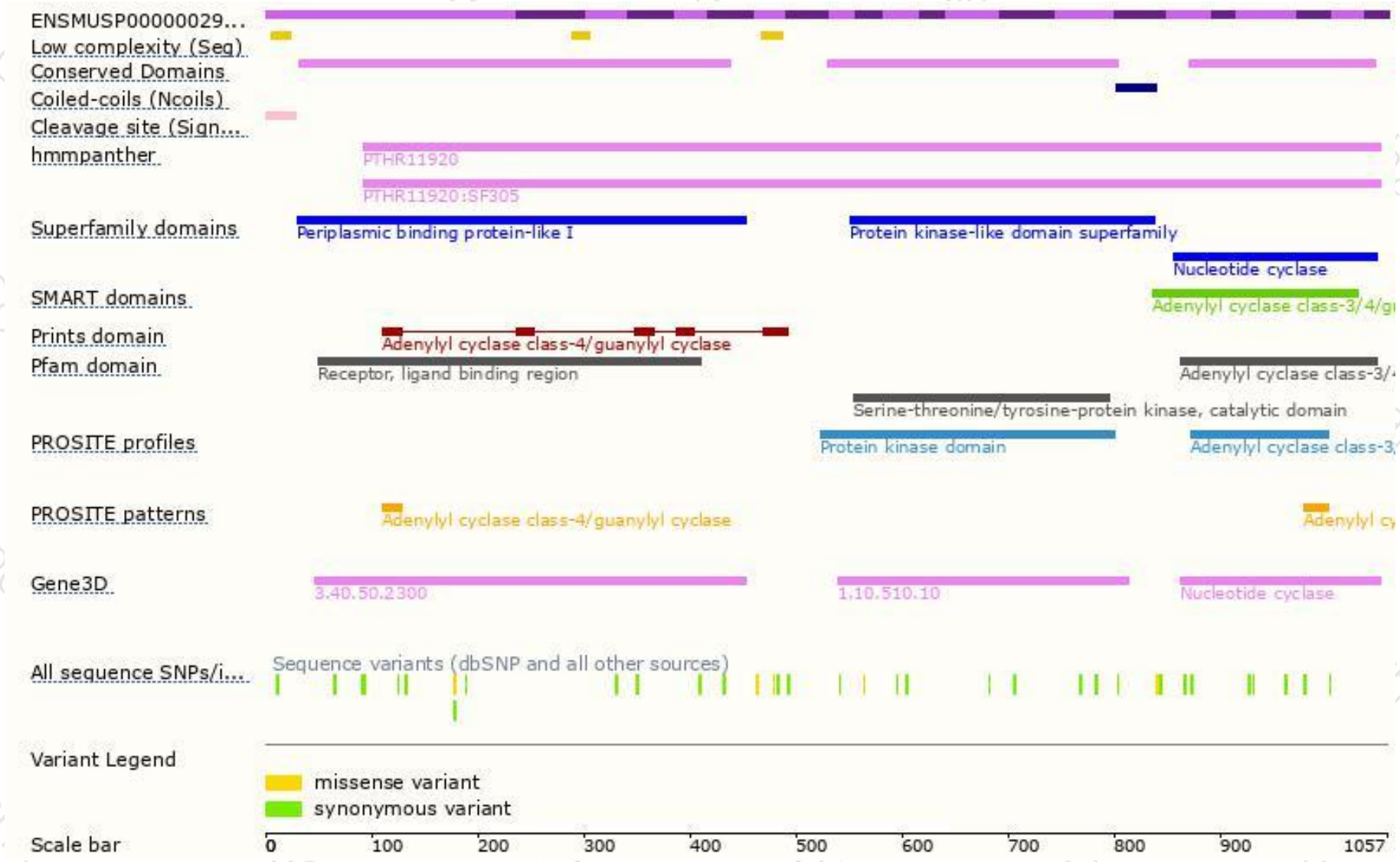
The strategy is based on the design of *Npr1-201* transcript,The transcription is shown below



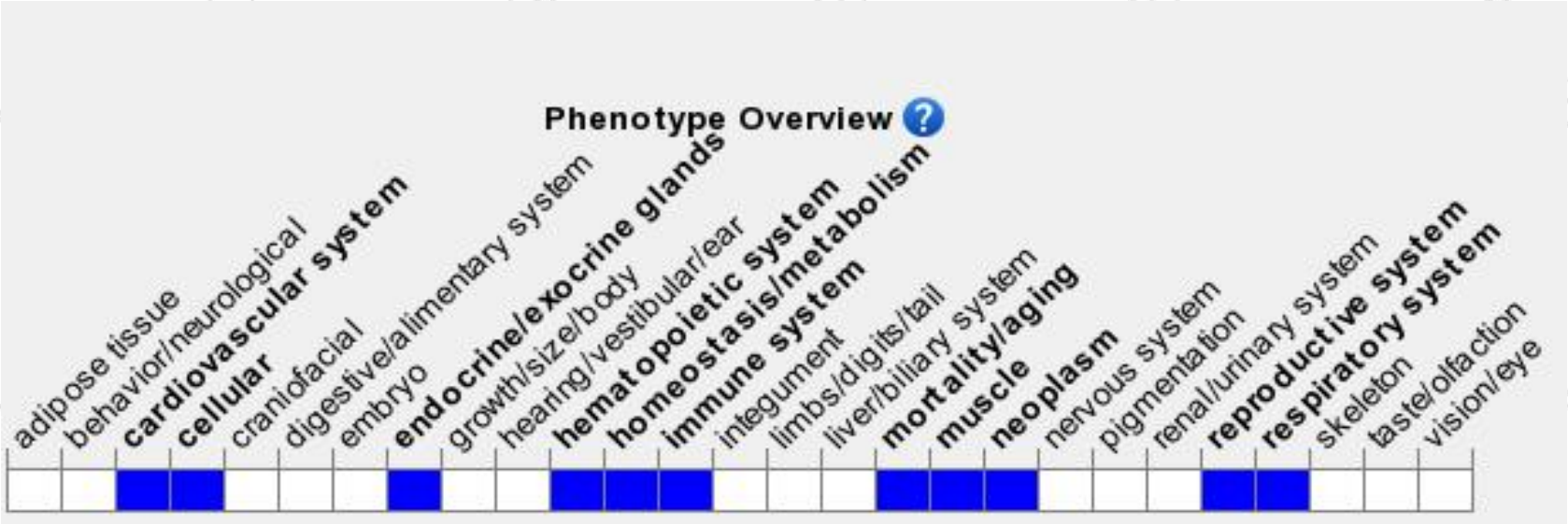
Genomic location distribution



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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