

# Cdk12 Cas9-CKO Strategy

**Designer:** 

**Huan Wang** 

**Reviewer:** 

**Huan Fan** 

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## **Project Overview**



**Project Name** 

*Cdk12* 

**Project type** 

Cas9-CKO

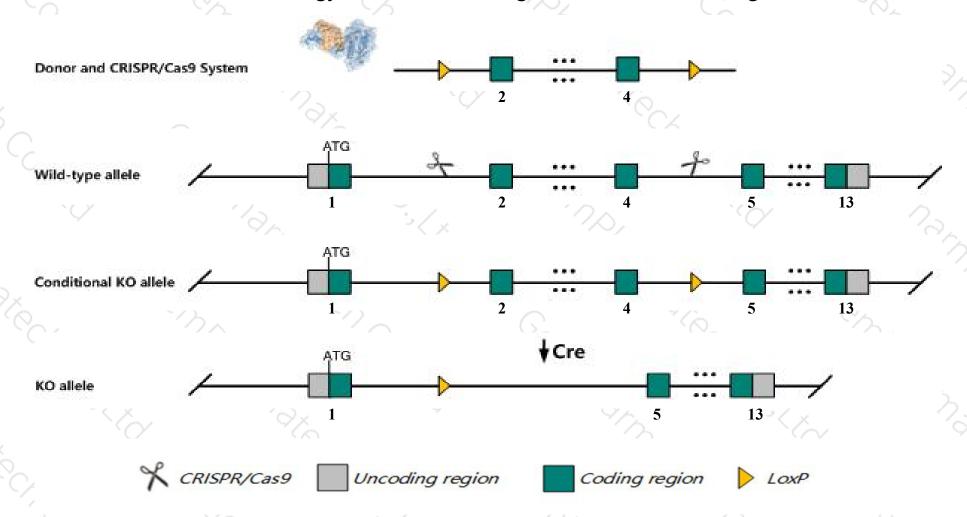
Strain background

C57BL/6JGpt

## Conditional Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Cdk12* gene. The schematic diagram is as follows:



### Technical routes



- ➤ The *Cdk12* gene has 6 transcripts. According to the structure of *Cdk12* gene, exon2-exon4 of *Cdk12-201* (ENSMUST0000003203.13) transcript is recommended as the knockout region. The region contains 1193bp coding sequence. Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Cdk12* gene. The brief process is as follows:gRNA was transcribed in vitro, donor was constructed.Cas9, gRNA 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, Mice homozygous for a knock-out allele exhibit lethality between implantation and E6.5 with decreased proliferation and increased apoptosis in the inner cell mass.
- The *Cdk12* gene is located on the Chr11. 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)



#### Cdk12 cyclin-dependent kinase 12 [Mus musculus (house mouse)]

Gene ID: 69131, updated on 31-Jan-2019

#### Summary

☆ ?

Official Symbol Cdk12 provided by MGI

Official Full Name cyclin-dependent kinase 12 provided by MGI

Primary source MGI:MGI:1098802

See related Ensembl: ENSMUSG00000003119

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 1810022J16Rik, Al646528, Crk7, Crkrs, D11Ertd752e, Pksc

Expression Ubiquitous expression in CNS E11.5 (RPKM 3.5), limb E14.5 (RPKM 3.3) and 28 other tissuesSee more

Orthologs <u>human</u> all

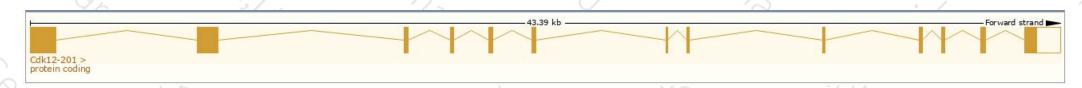
## Transcript information (Ensembl)



The gene has 6 transcripts, all transcripts are shown below:

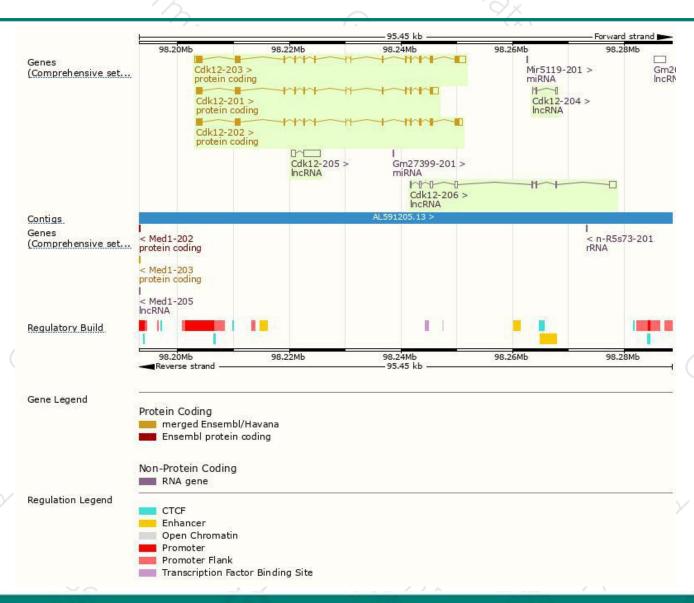
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Cdk12-203	ENSMUST00000107539.7	5912	<u>1475aa</u>	Protein coding	CCDS48902	Q14AX6	TSL:5 GENCODE basic APPRIS ALT2
Cdk12-202	ENSMUST00000107538.1	5143	<u>1484aa</u>	Protein coding	CCDS48901	Q14AX6	TSL:1 GENCODE basic APPRIS ALT2
Cdk12-201	ENSMUST00000003203.13	4837	<u>1258aa</u>	Protein coding	CCDS25342	Q14AX6	TSL:1 GENCODE basic APPRIS P3
Cdk12-205	ENSMUST00000145113.1	3648	No protein	IncRNA	-	62	TSL:1
Cdk12-206	ENSMUST00000147441.7	2605	No protein	IncRNA	a	85	TSL:2
Cdk12-204	ENSMUST00000125152.1	443	No protein	IncRNA	-		TSL:2

The strategy is based on the design of *Cdk12-201* transcript, The transcription is shown below



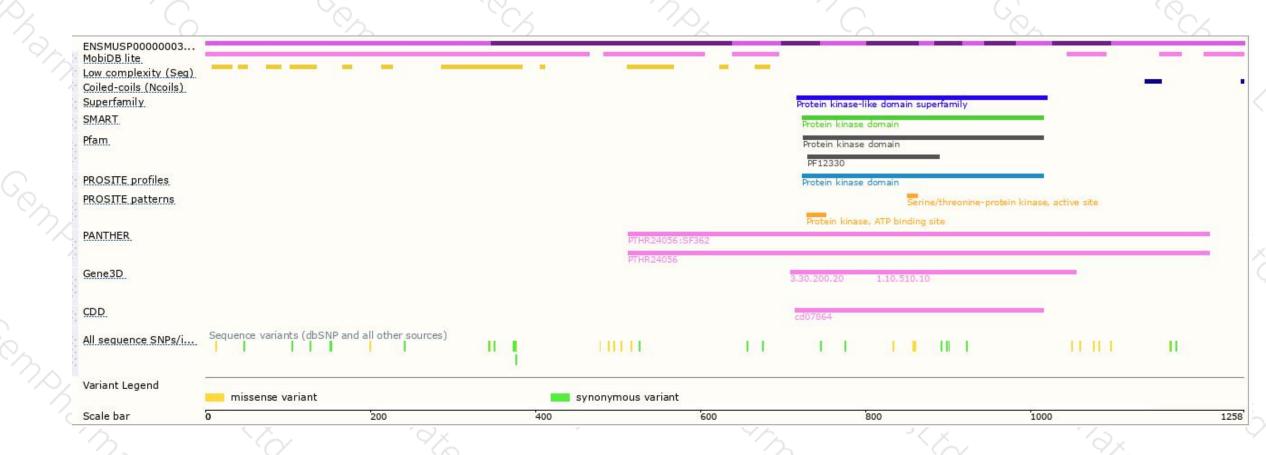
### Genomic location distribution





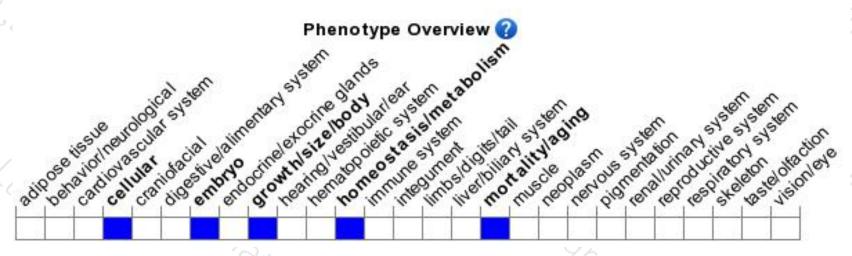
## Protein domain





## 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 knock-out allele exhibit lethality between implantation and E6.5 with decreased proliferation and increased apoptosis in the inner cell mass.



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





