

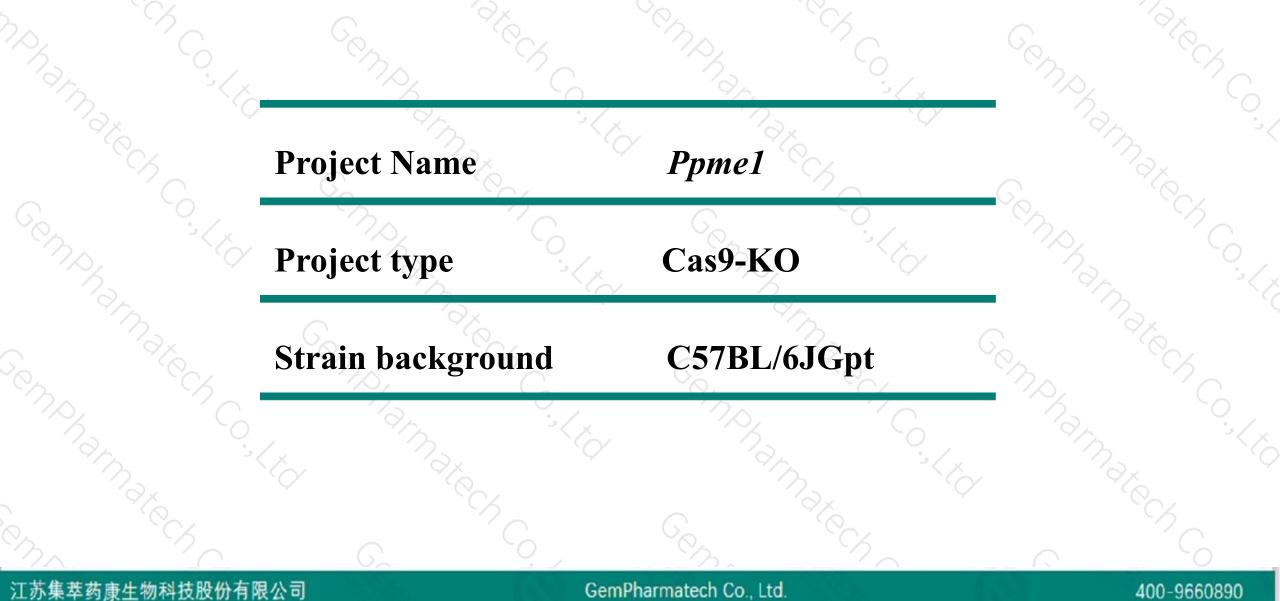
# **Ppmel Cas9-KO Strategy**

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Designer: Xueting Zhang Reviewer:Yanhua Shen Date:2020-03-11

## **Project Overview**

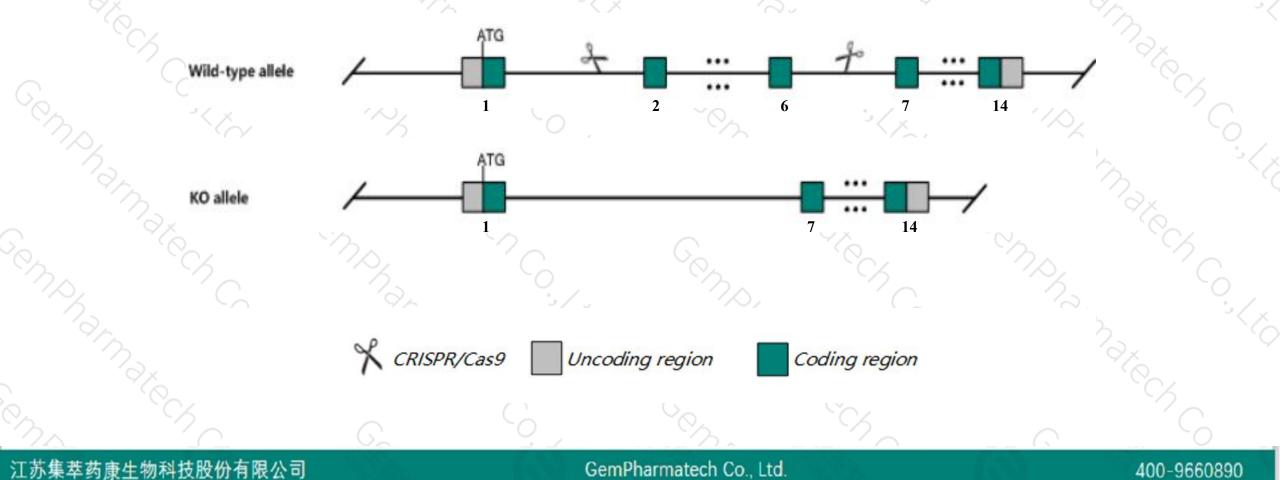




# **Knockout strategy**



This model will use CRISPR/Cas9 technology to edit the *Ppme1* gene. The schematic diagram is as follows:





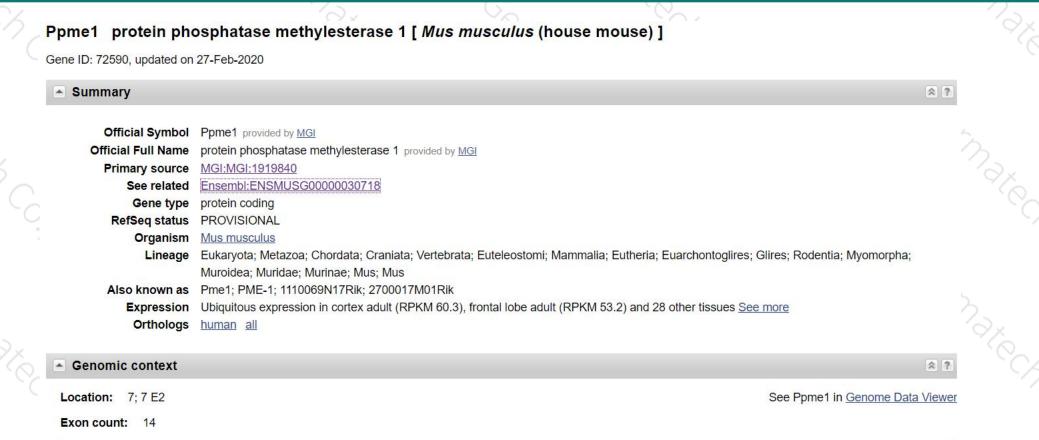
- The Ppme1 gene has 5 transcripts. According to the structure of Ppme1 gene, exon2-exon6 of Ppme1-201 (ENSMUST00000032963.9) transcript is recommended as the knockout region. The region contains 452bp coding sequence. Knock out the region will result in disruption of protein function.
- > In this project we use CRISPR/Cas9 technology to modify *Ppme1* gene. The brief process is as follows: CRISPR/Cas9 system

- According to the existing MGI data, Targeted disruption of this gene causes virtual loss of the demethylated form of phosphoprotein phosphatase 2A in the nervous system and peripheral tissues. Homozygous null mice fail to initiate normal breathing or suckling behavior and die within the first day of life.
- > The effect on transcript Ppme1-204 is unknown.
- ➤ Transcript *Ppme1*-203 may not be affected.
- The *Ppme1* 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 the gene knockout on gene transcription, RNA splicing and protein translation cannot be predicted at the existing technology level.

Notice

# Gene information (NCBI)





Annotation release	Status	Assembly	Chr	Location
<u>108</u>	current	GRCm38.p6 (GCF_000001635.26)	7	NC_000073.6 (100326737100371896, complement)
Build 37.2	previous assembly	MGSCv37 (GCF_000001635.18)	7	NC_000073.5 (107475247107520406, complement)

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# **Transcript information (Ensembl)**



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

Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags	
Ppme1-201	ENSMUST0000032963.9	2772	<u>386aa</u>	Protein coding	CCDS40036	<u>Q8BVQ5</u>	TSL:1 GENCODE basic APPRIS P1	
Ppme1-204	ENSMUST00000207634.1	673	<u>182aa</u>	Protein coding	-	A0A140LI84	CDS 5' incomplete TSL:5	
Ppme1-203	ENSMUST00000207622.1	942	No protein	Retained intron	84	1	TSL:2	
Ppme1-205	ENSMUST00000208168.1	474	No protein	Retained intron	62	2	TSL:2	
Ppme1-202	ENSMUST00000207092.1	754	No protein	IncRNA	15	-	TSL:3	

The strategy is based on the design of *Ppme1-201* transcript, The transcription is shown below

#### < Ppme1-201 protein coding

Reverse strand -

- 45.57 kb -

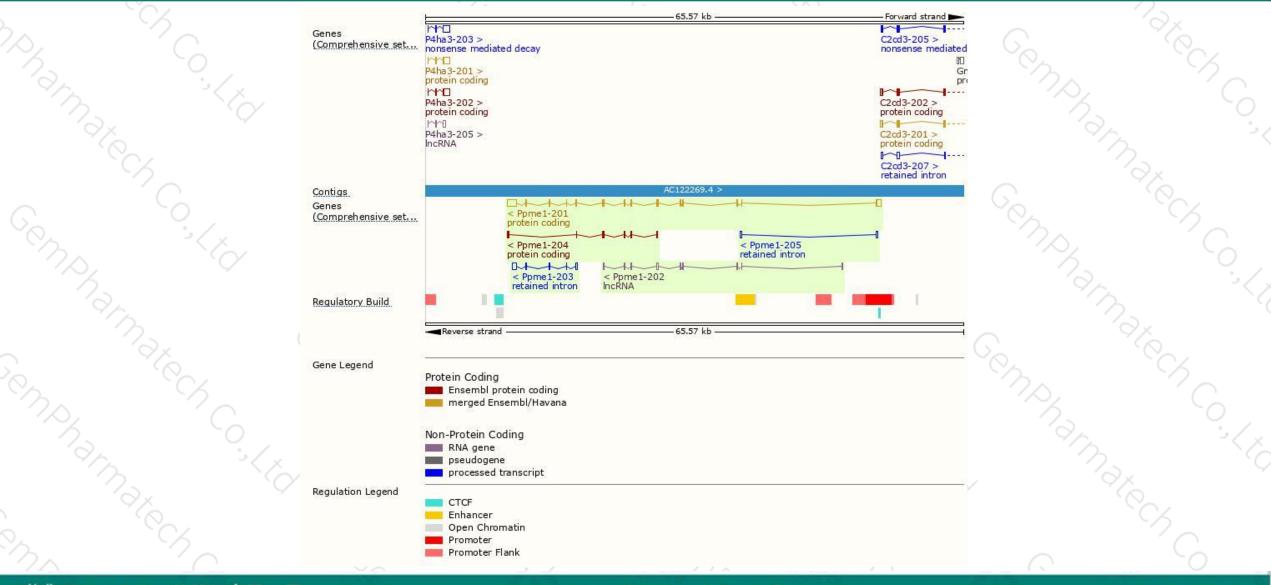
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### **Genomic location distribution**



400-9660890



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### **Protein domain**



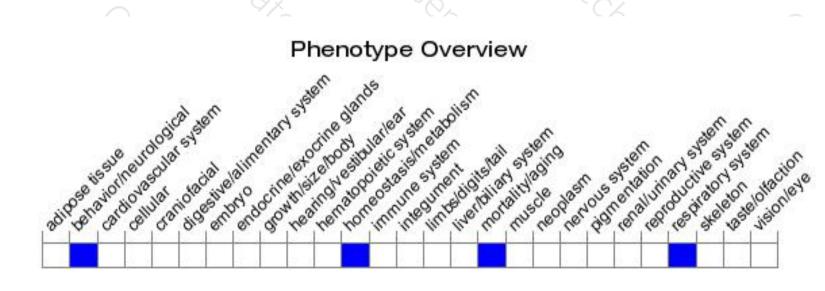
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ENSMUSP00000032 MobiDB lite							
Low complexity (Seg)							
Superfamily		Alpha/Beta hydrolase fold	and the state				
Prints			ydrolase fold-1				
Pfam.		Epoxide hydro Alpha/beta hydrolase	and the second				- C
PIRSF	Protein phosphatase n	nethylesterase, eukaryotic	lense levelse i				
PANTHER	Protein phosphata	ise methylesterase, eukaryo	otic				
All sequence SNPs/i	Sequence variants (	(dbSNP and all other sou	irces)	í -		1	(II)
Variant Legend	missense varia splice region v synonymous vi	/ariant					
Scale bar	<b>0</b> 40	80 120	160	200 240	280	320	386
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### 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, Targeted disruption of this gene causes virtual loss of the demethylated form of phosphoprotein phosphatase 2A in the nervous system and peripheral tissues. Homozygous null mice fail to initiate normal breathing or suckling behavior and die within the first day of life.

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



