

Bmper Cas9-CKO Strategy

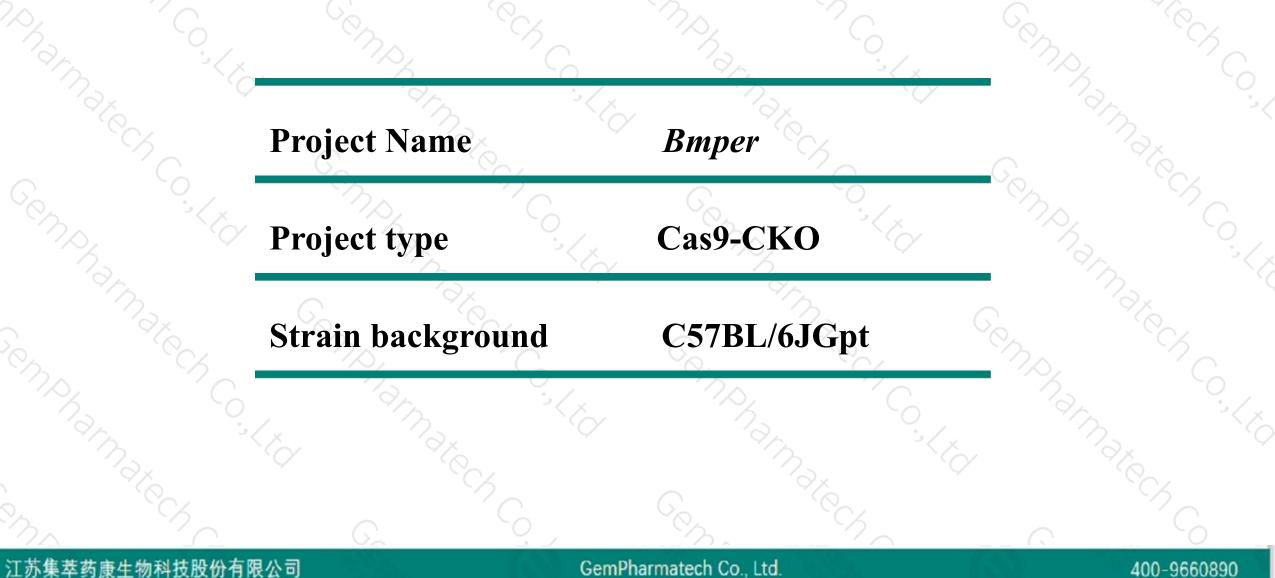
Designer: Huimin Su

Reviewer: Ruiuri Zhang

Design Date: 2020-6-28

Project Overview





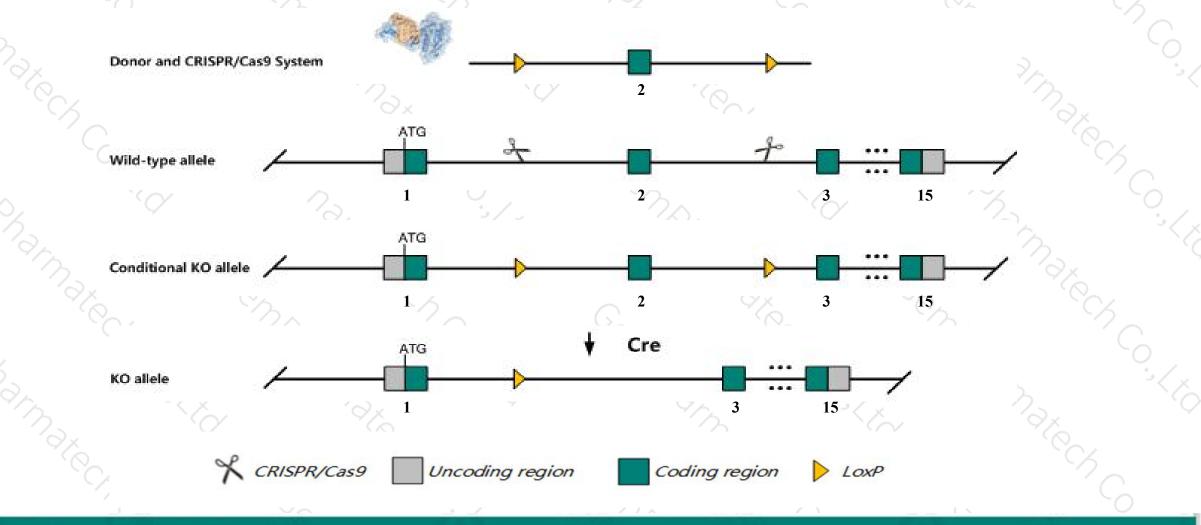
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Conditional Knockout strategy



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



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The *Bmper* gene has 3 transcripts. According to the structure of *Bmper* gene, exon2 of *Bmper* 201(ENSMUST00000071982.6) transcript is recommended as the knockout region. The region contains 86bp coding sequence.
Knock out the region will result in disruption of protein function.

➤ In this project we use CRISPR/Cas9 technology to modify *Bmper* 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,mice homozygous for a knock-out mutation exhibit neonatal lethality associated with abnormal lung and skeleton development. Mice heterozygous for a null allele exhibit abnromal lung development.
Transcript *Bmper-202* is incomplete, so the effect on it is unknown.
The *Bmper* gene is located on the Chr9. 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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Bmper BMP-binding endothelial regulator [Mus musculus (house mouse)]

Gene ID: 73230, updated on 26-Jun-2020

Summary

Official Symbol	Bmper provided by MGI	
Official Full Name	BMP-binding endothelial regulator provided by MGI	
Primary source	MGI:MGI:1920480	
See related	Ensembl:ENSMUSG0000031963	
Gene type	protein coding	
RefSeq status	REVIEWED	
Organism	Mus musculus	
Lineage	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha; Muroidea; Muridae; Murinae;	
	Mus; Mus	
Also known as	Cv2; CV-2; Crim3; 3110056H04Rik	
Summary	This gene encodes a secreted protein that contains five Von Willebrand factor type C domains and a Von Willebrand factor type D domain and a trypsin inhibitory-like	
	domain. The encoded protein binds to bone morphogenetic proteins (BMP) and regulates their activity. Mutation of the related gene in humans causes	
	diaphanospondylodysostosis. [provided by RefSeq, Mar 2013]	P
Expression	Broad expression in lung adult (RPKM 4.5), limb E14.5 (RPKM 4.1) and 20 other tissues See more	
Orthologs	human all	
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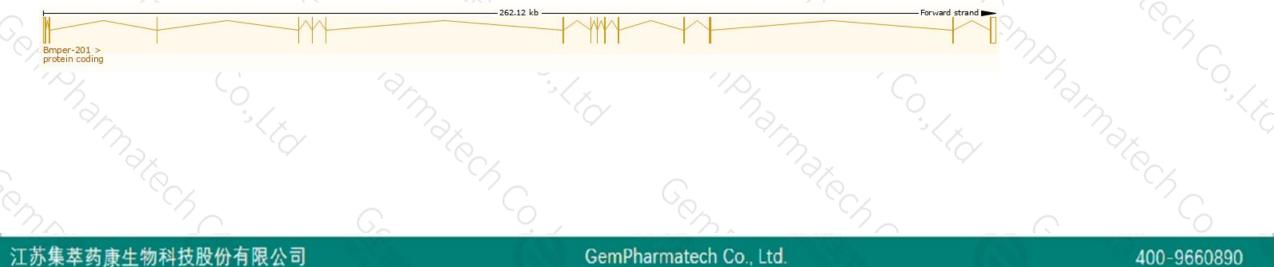
Transcript information (Ensembl)



The gene has 3 transcripts, all transcripts are shown below:

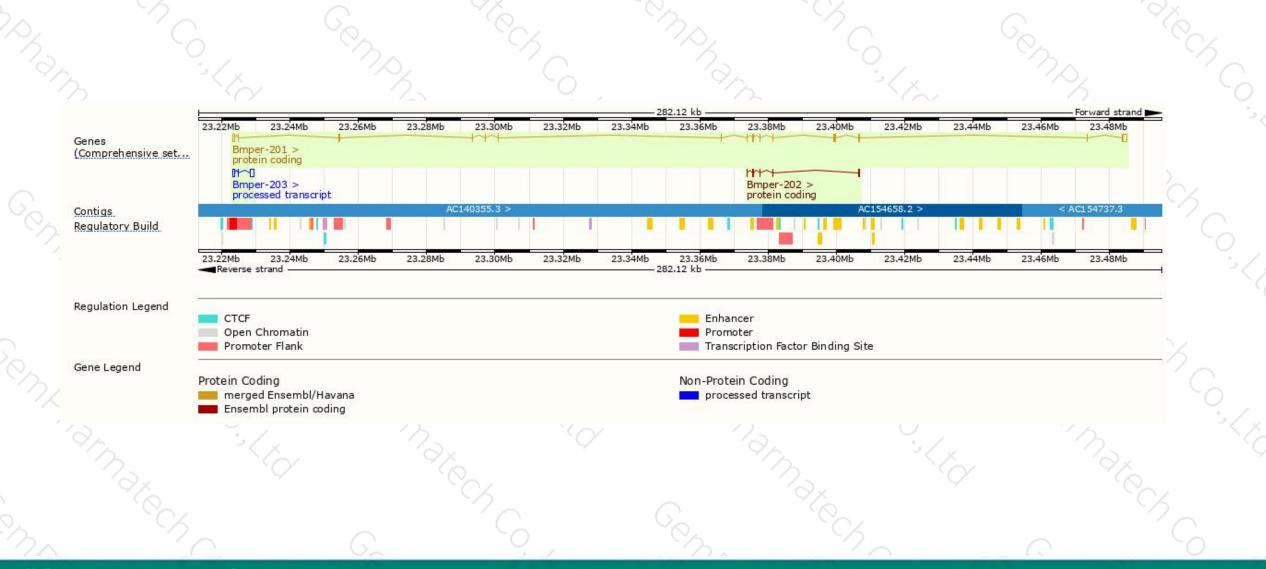
Name 🍦	Transcript ID	bp 🖕	Protein A	Biotype	CCDS	UniProt 🖕	Flags 🍦
Bmper-202	ENSMUST00000214050.1	640	<u>213aa</u>	Protein coding	-	A0A1L1SS01@	CDS 5' and 3' incomplete TSL:5
Bmper-201	ENSMUST0000071982.6	3791	<u>685aa</u>	Protein coding	<u>CCDS22929</u> &	Q8CJ69@	TSL:1 GENCODE basic APPRIS P1
Bmper-203	ENSMUST00000217648.1	1616	No protein	Processed transcript	-	(14)	TSL:1

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



Genomic location distribution





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Protein domain



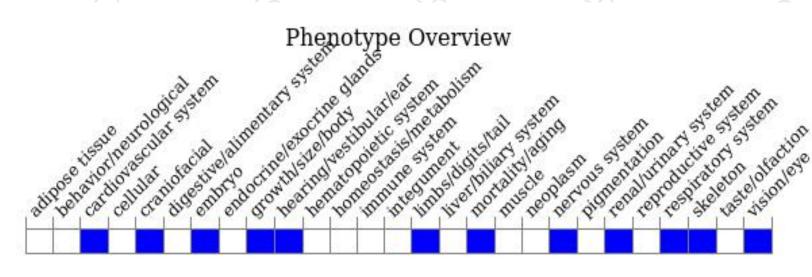


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Mouse phenotype description(MGI)





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,mice homozygous for a knock-out mutation exhibit neonatal lethality associated with abnormal lung and skeleton development. Mice heterozygous for a null allele exhibit abnromal lung development.

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



