

Mecom Cas9-KO Strategy

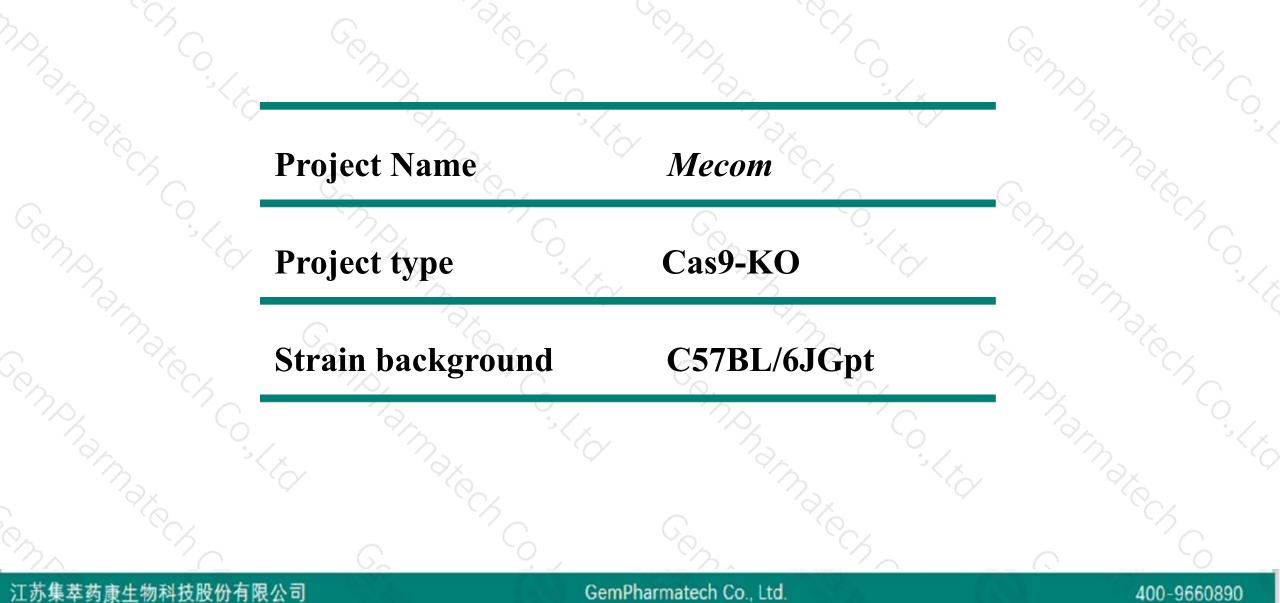
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Reviewer: Richeng Wang

Design Date: 2019-4-15

Project Overview

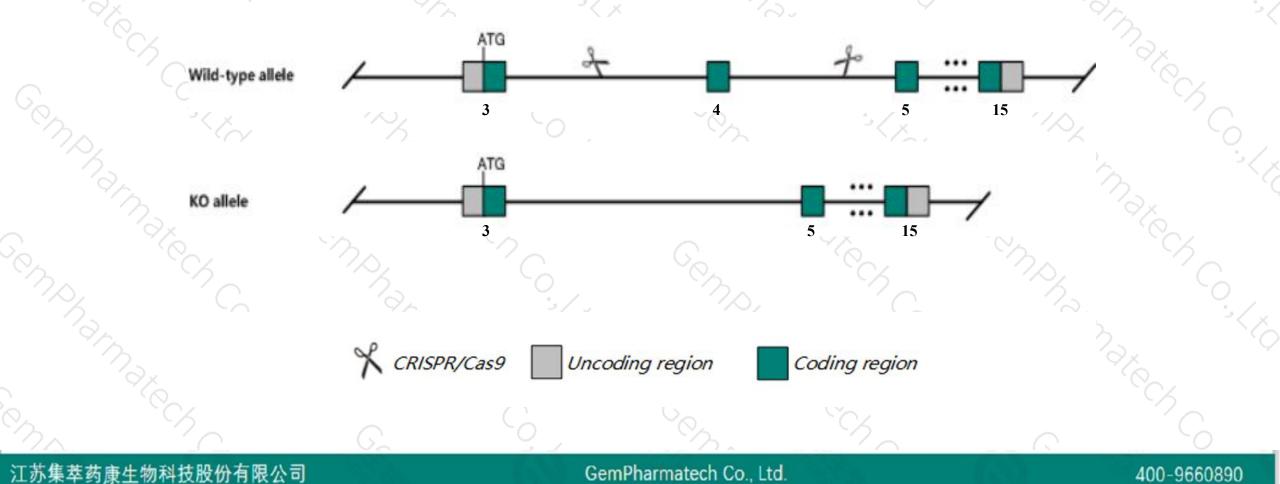




Knockout strategy



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





- The Mecom gene has 17 transcripts. According to the structure of Mecom gene, exon4 of Mecom-206 (ENSMUST00000172694.7) transcript is recommended as the knockout region. The region contains 217bp coding sequence. Knock out the region will result in disruption of protein function.
- > In this project we use CRISPR/Cas9 technology to modify Mecom gene. The brief process is as follows: CRISPR/Cas9 system

- According to the existing MGI data, embryos homozygous for a targeted null mutation die at 10.5 dpc displaying widespread hypocellularity, hemorrhage, and disruption in the development of the heart, somites, and neural crest-derived cells.
- The *Mecom* 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 the gene knockout on gene transcription, RNA splicing and protein translation cannot be predicted at the existing technology level.

Notice

Gene information (NCBI)



☆ ?

Mecom MDS1 and EVI1 complex locus [Mus musculus (house mouse)]

Gene ID: 14013, updated on 20-Mar-2020

- Summary

Official SymbolMecom provided by MGIOfficial Full NameMDS1 and EVI1 complex locus provided by MGIPrimary sourceMGI:MGI:95457See relatedEnsembl:ENSMUSG0000027684Gene typeprotein codingRefSeq statusVALIDATEDOrganismMus musculusLineageEukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha;
Muroidea; Murinae; Mus; MusAlso known asD630039M04Rik, Evi-1, Evi1, Jbo, Mds, Mds1, Mds1-Evi1, Prdm3, Znfpr1b1ExpressionBroad expression in bladder adult (RPKM 7.5), limb E14.5 (RPKM 5.2) and 15 other tissues
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Transcript information (Ensembl)



The gene has 17 transcripts, all transcripts are shown below:

| Name | Transcript ID | bp | Protein | Biotype | CCDS | UniProt | Flags |
|-----------|----------------------|------|--------------|-------------------------|-----------|---------------|---|
| Aecom-206 | • | 4381 | 718aa | Protein coding | CCDS17281 | <u>G3UZ13</u> | TSL:1 GENCODE basic |
| | ENSMUST00000108270.9 | | | Protein coding | - | H9T841 P14404 | TSL:5 GENCODE basic APPRIS is a system to annotate alternatively spliced transcripts based on a range of computational methods to identify the most functionally important transcript(s) of a gene. APPRIS AL |
| | ENSMUST00000173495.7 | | | Protein coding | - | G3UZT5 | TSL:5 GENCODE basic APPRIS is a system to annotate alternatively spliced transcripts based on a range of computational methods to identify the most functionally important transcript(s) of a gene. APPRIS F |
| ecom-203 | ENSMUST00000108271.9 | 3851 | <u>795aa</u> | Protein coding | - | Q8CCA6 | TSL:1 GENCODE basic APPRIS is a system to annotate alternatively spliced transcripts based on a range of computational methods to identify the most functionally important transcript(s) of a gene. APPRIS AL |
| ecom-207 | ENSMUST00000172697.7 | 3204 | <u>949aa</u> | Protein coding | - | G3UYK2 | CDS 3' incomplete TSL:5 |
| ecom-213 | ENSMUST00000173899.7 | 3031 | <u>994aa</u> | Protein coding | | - | CDS 3' incomplete TSL:5 |
| ecom-204 | ENSMUST00000166001.7 | 2914 | <u>949aa</u> | Protein coding | 12 | <u>O35700</u> | TSL:1 GENCODE basic APPRIS is a system to annotate alternatively spliced transcripts based on a range of computational methods to identify the most functionally important transcript(s) of a gene. APPRIS A |
| ecom-210 | ENSMUST00000173059.7 | 2680 | <u>843aa</u> | Protein coding | <u></u> | <u>G3UWJ0</u> | CDS 5' incomplete TSL:5 |
| ecom-215 | ENSMUST00000174413.7 | 1942 | <u>606aa</u> | Protein coding | | G3UYY9 | CDS 5' incomplete TSL:1 |
| ecom-214 | ENSMUST00000174406.1 | 524 | <u>52aa</u> | Protein coding | - | <u>G3UY46</u> | CDS 3' incomplete TSL:5 |
| ecom-201 | ENSMUST0000061088.6 | 348 | <u>116aa</u> | Protein coding | 12 | F6SZX1 | TSL:NA GENCODE basic |
| ecom-208 | ENSMUST00000172754.7 | 3182 | <u>45aa</u> | Nonsense mediated decay | <u></u> | G3UYN3 | TSL:5 |
| ecom-205 | ENSMUST00000170212.7 | 1586 | No protein | Processed transcript | a | 5 | TSL:1 |
| ecom-211 | ENSMUST00000173411.7 | 1302 | No protein | Processed transcript | - | - | TSL:5 |
| ecom-209 | ENSMUST00000173022.1 | 1075 | No protein | Processed transcript | 12 | - | TSL:1 |
| lecom-216 | ENSMUST00000174428.1 | 2803 | No protein | Retained intron | <u></u> | 2 | TSL:1 |
| lecom-217 | ENSMUST00000195690.1 | 2518 | No protein | Retained intron | | - | TSL:NA |

The strategy is based on the design of *Mecom-206* transcript, the transcription is shown below:

< Mecom-206 protein coding

Reverse strand -

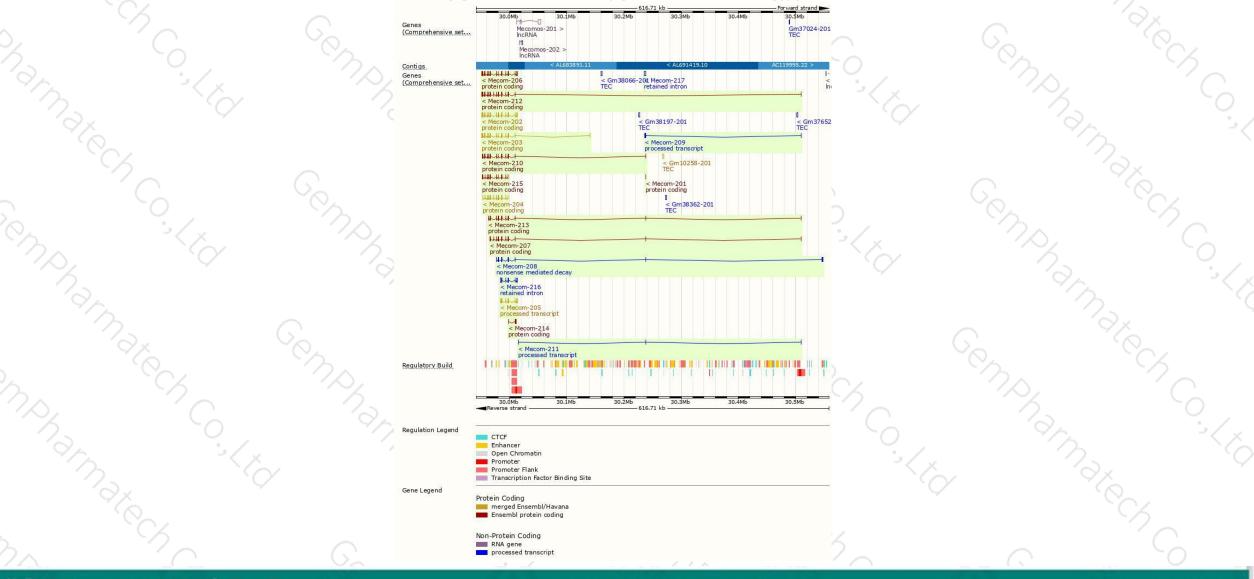
- 61.86 kb -

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Genomic location distribution





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Protein domain



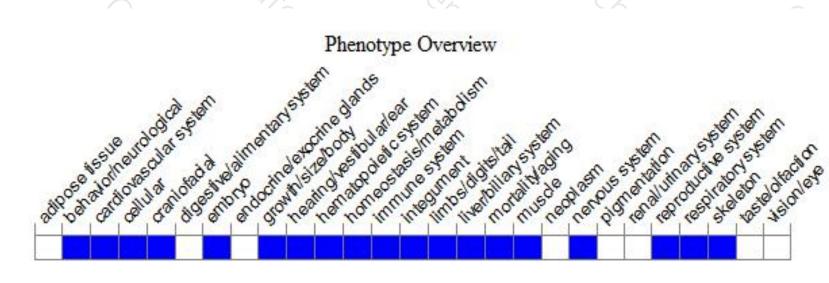


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Mouse phenotype description(MGI)





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, embryos homozygous for a targeted null mutation die at 10.5 dpc displaying widespread hypocellularity, hemorrhage, and disruption in the development of the heart, somites, and neural crest-derived cell

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



