

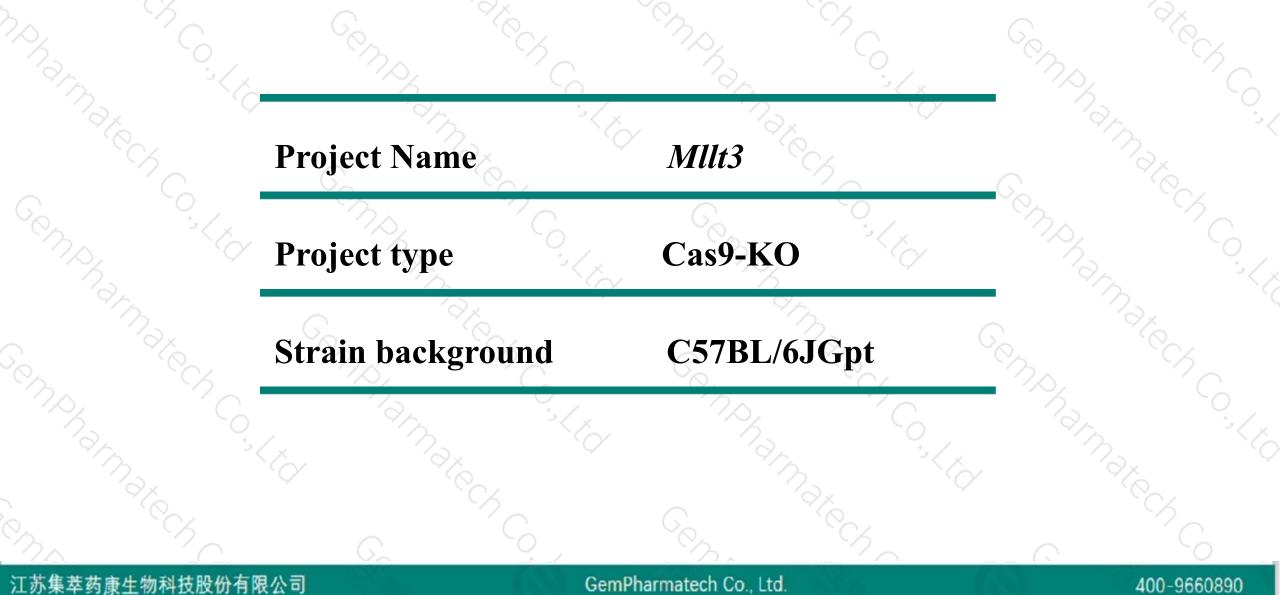
Mllt3 Cas9-KO Strategy

empharmatect

Designer: Yanhua Shen Design Date: 2019-7-24

Project Overview

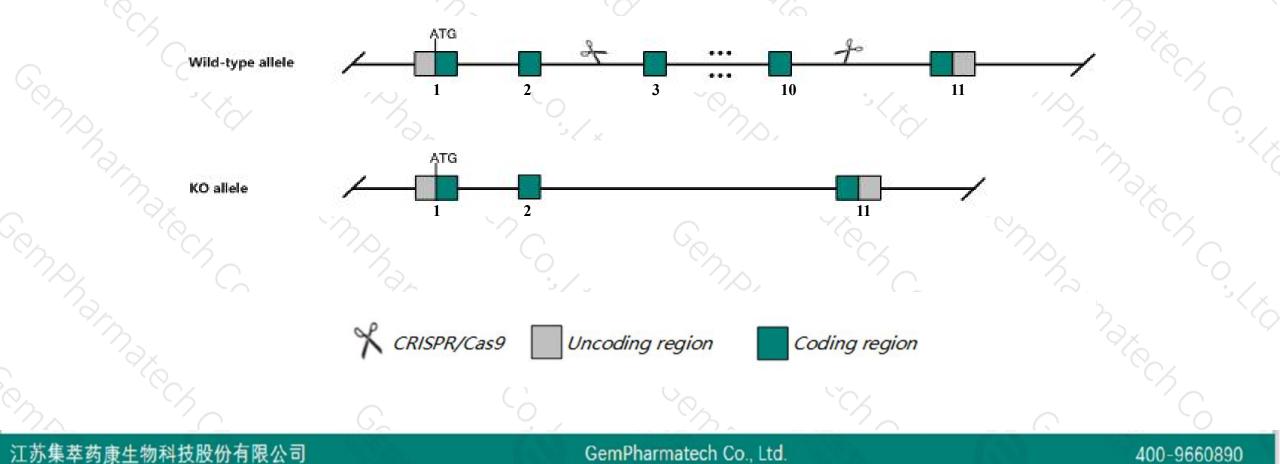




Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Mllt3* gene. The schematic diagram is as follows:





- The *Mllt3* gene has 8 transcripts. According to the structure of *Mllt3* gene, exon3-exon10 of *Mllt3-201* (ENSMUST00000078090.11) transcript is recommended as the knockout region. The region contains 1385bp coding sequence Knock out the region will result in disruption of protein function.
- > In this project we use CRISPR/Cas9 technology to modify *Mllt3* gene. The brief process is as follows: CRISPR/Cas9 system



- According to the existing MGI data, About 50% of homozygous null mice die perinatally while the remaining 50% become runted and die within two weeks of birth with severe defects of the axial skeleton, including anterior homeotic transformation of the cervical and thoracic regions, a deformed atlas and an extra cervical vertebra.
- The *Mllt3* gene is located on the Chr4. 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.

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Gene information (NCBI)



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MIIt3 myeloid/lymphoid or mixed-lineage leukemia; translocated to, 3 [Mus musculus (house mouse)]

Gene ID: 70122, updated on 31-Jan-2019

Summary

Official Symbol	MIIt3 provided by MGI
Official Full Name	myeloid/lymphoid or mixed-lineage leukemia; translocated to, 3 provided byMGI
Primary source	MGI:MGI:1917372
See related	Ensembl:ENSMUSG0000028496
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	2210011H10Rik, 2610012I03Rik, 3830408D16Rik, Af9, D4Ertd321e
Expression	Broad expression in whole brain E14.5 (RPKM 9.1), CNS E14 (RPKM 9.1) and 26 other tissues See more
Orthologs	human all

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Transcript information (Ensembl)



The gene has 8 transcripts, all transcripts are shown below:

Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
ENSMUST00000078090.11	6069	<u>569aa</u>	Protein coding	CCDS38796	A2AM29	TSL:1 GENCODE basic APPRIS P1
ENSMUST00000126353.7	1956	<u>138aa</u>	Protein coding	CCDS71418	Q9D2P1	TSL:1 GENCODE basic
ENSMUST00000149357.7	794	<u>162aa</u>	Protein coding	CCDS38797		TSL:1 GENCODE basic
ENSMUST00000148059.1	422	<u>54aa</u>	Protein coding	2	E9PXP8	TSL:5 GENCODE basic
ENSMUST00000142454.1	966	<u>73aa</u>	Nonsense mediated decay	51	<u>E9Q3B8</u>	TSL:1
ENSMUST00000134555.8	3159	No protein	Processed transcript	-		TSL:1
ENSMUST00000128930.7	1293	No protein	Processed transcript	2	40	TSL:5
ENSMUST00000141526.7	4556	No protein	Retained intron	2	20 20	TSL:1
	ENSMUST0000078090.11 ENSMUST00000126353.7 ENSMUST00000149357.7 ENSMUST00000148059.1 ENSMUST00000142454.1 ENSMUST00000134555.8 ENSMUST00000128930.7	ENSMUST0000078090.11 6069 ENSMUST00000126353.7 1956 ENSMUST0000149357.7 794 ENSMUST0000148059.1 422 ENSMUST0000142454.1 966 ENSMUST0000134555.8 3159 ENSMUST0000128930.7 1293	ENSMUST0000078090.11 6069 569aa ENSMUST0000126353.7 1956 138aa ENSMUST0000149357.7 794 162aa ENSMUST0000148059.1 422 54aa ENSMUST0000134555.8 966 73aa ENSMUST0000134555.8 3159 No protein	ENSMUST0000078090.116069569aaProtein codingENSMUST00000126353.71956138aaProtein codingENSMUST00000149357.7794162aaProtein codingENSMUST00000148059.142254aaProtein codingENSMUST00000142454.196673aaNonsense mediated decayENSMUST00000134555.83159No proteinProcessed transcriptENSMUST00000128930.71293No proteinProcessed transcript	ENSMUST0000078090.116069569aaProtein codingCCDS38796ENSMUST00000126353.71956138aaProtein codingCCDS71418ENSMUST00000149357.7794162aaProtein codingCCDS38797ENSMUST00000148059.142254aaProtein coding-ENSMUST00000142454.196673aaNonsense mediated decay-ENSMUST00000134555.83159No proteinProcessed transcript-ENSMUST00000128930.71293No proteinProcessed transcript-	ENSMUST0000078090.116069569aaProtein codingCCDS38796A2AM29ENSMUST00000126353.71956138aaProtein codingCCDS71418Q9D2P1ENSMUST00000149357.7794162aaProtein codingCCDS38797Q3UIA3ENSMUST00000148059.142254aaProtein coding-E9PXP8ENSMUST00000142454.196673aaNonsense mediated decayENSMUST00000134555.83159No proteinProcessed transcriptENSMUST00000128930.71293No proteinProcessed transcript

The strategy is based on the design of *Mllt3-201* transcript, The transcription is shown below

< Mllt3-201 protein coding

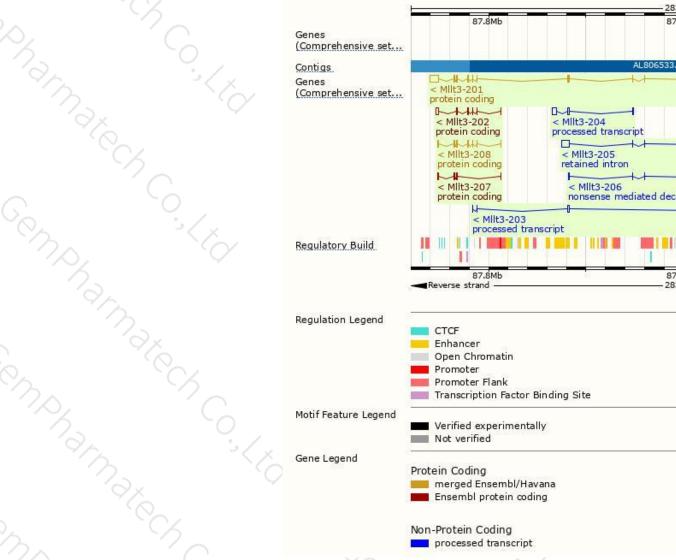
Reverse strand

_____´__ ^

263.44 kb

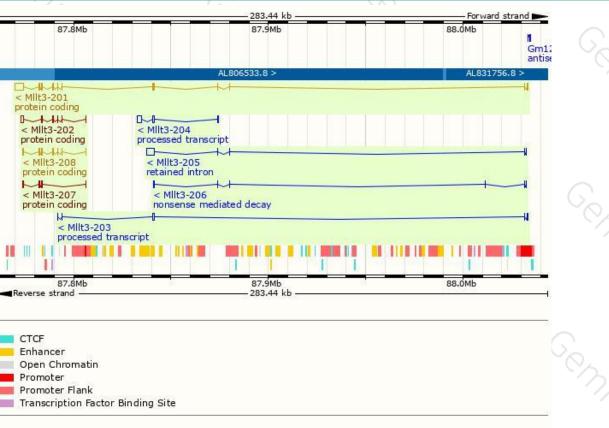
Genomic location distribution





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Protein domain



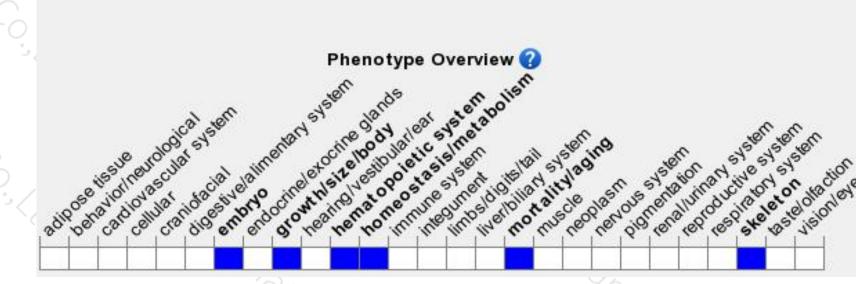


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Mouse phenotype description(MGI)





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, About 50% of homozygous null mice die perinatally while the remaining 50% become runted and die within two weeks of birth with severe defects of the axial skeleton, including anterior homeotic transformation of the cervical and thoracic regions, a deformed atlas and an extra cervical vertebra.

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



