

# Donal Day Co. Fus Cas9-KO Strategy

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# **Project Overview**



Project Name Fus

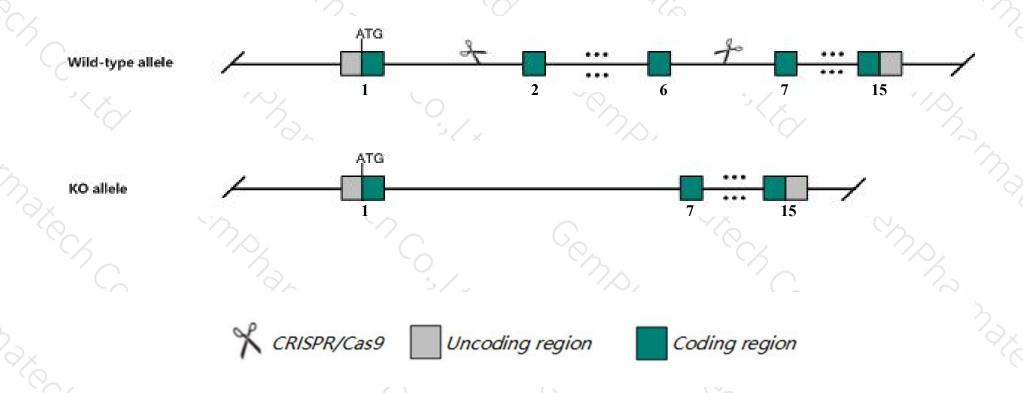
Project type Cas9-KO

Strain background C57BL/6JGpt

# **Knockout strategy**



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



## **Technical routes**



- ➤ The Fus gene has 16 transcripts. According to the structure of Fus gene, exon2-exon6 of Fus-203

  (ENSMUST00000106251.9) transcript is recommended as the knockout region. The region contains 730bp coding sequence.

  Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Fus* gene. The brief process is as follows: CRISPR/Cas9 system w

## **Notice**



- ➤ According to the existing MGI data, Homozygotes for targeted null mutations exhibit impaired lymphocyte development, chromosomal instability, increased cellular radiation sensitivity, high neonatal mortality, and male sterility associated with lack of chromosomal pairing.
- > The *Fus* 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.

## Gene information (NCBI)



#### Fus fused in sarcoma [Mus musculus (house mouse)]

Gene ID: 233908, updated on 7-Apr-2019

#### Summary

↑ ?

Official Symbol Fus provided by MGI

Official Full Name fused in sarcoma provided by MGI

Primary source MGI:MGI:1353633

See related Ensembl:ENSMUSG00000030795

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 D430004D17Rik, D930039C12Rik, Fus1, Tls

Expression Ubiquitous expression in limb E14.5 (RPKM 160.0), CNS E11.5 (RPKM 159.5) and 28 other tissuesSee more

Orthologs <u>human</u> all

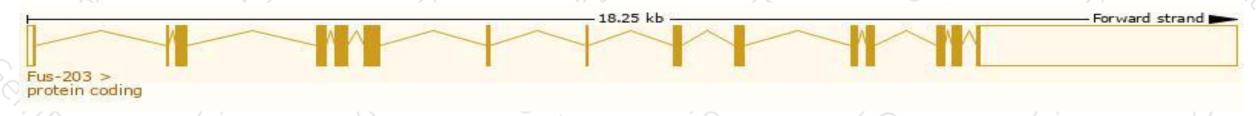
# Transcript information (Ensembl)



#### The gene has 16 transcripts, all transcripts are shown below:

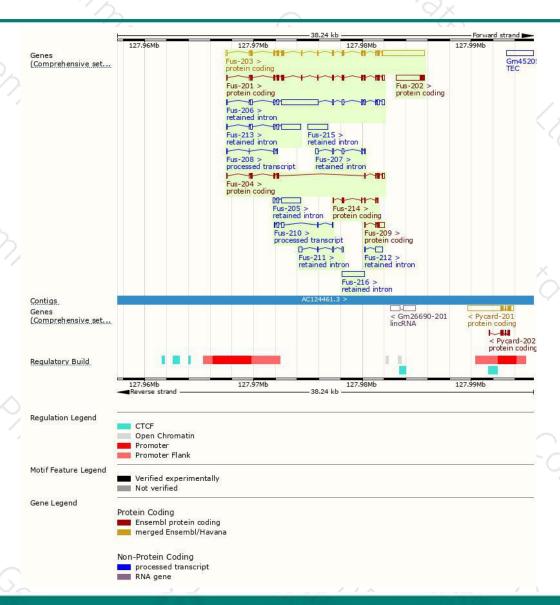
Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
ENSMUST00000106251.9	5536	518aa	Protein coding	CCDS21886	P56959 Q564D0	TSL:1 GENCODE basic APPRIS P3
ENSMUST00000077609.11	1831	<u>517aa</u>	Protein coding	CCDS85425	Q8CFQ9	TSL:1 GENCODE basic APPRIS ALT2
ENSMUST00000079045.2	2623	<u>122aa</u>	Protein coding	2	Q8BNR3	TSL:NA GENCODE basic
ENSMUST00000121616.8	1102	280aa	Protein coding		Q91VQ2	TSL:1 GENCODE basic
ENSMUST00000141997.1	761	<u>104aa</u>	Protein coding		G3UZD2	CDS 5' incomplete TSL:2
ENSMUST00000174632.7	390	130aa	Protein coding		G3UXT7	5' and 3' truncations in transcript evidence prevent annotation of the start and the end of the CDS. CDS 5' and 3' incomplete TSL:5
ENSMUST00000154843.7	600	No protein	Processed transcript	-		TSL:3
ENSMUST00000137464.6	354	No protein	Processed transcript	-	29	TSL:5
ENSMUST00000128851.7	4909	No protein	Retained intron			TSL:5
ENSMUST00000174196.7	2374	No protein	Retained intron			TSL:1
ENSMUST00000123151.7	2158	No protein	Retained intron	-	20	TSL:1
ENSMUST00000205351.1	2055	No protein	Retained intron	-	29	TSL:NA
ENSMUST00000205261.1	1859	No protein	Retained intron			TSL:NA
ENSMUST00000172755.1	670	No protein	Retained intron		-8	TSL:5
ENSMUST00000136289.1	600	No protein	Retained intron	-	20	TSL:3
ENSMUST00000155941.7	516	No protein	Retained intron	2	29	TSL:3
	ENSMUST00000106251.9 ENSMUST00000077609.11 ENSMUST00000079045.2 ENSMUST00000121816.8 ENSMUST00000141997.1 ENSMUST00000174632.7 ENSMUST00000154843.7 ENSMUST00000137464.6 ENSMUST00000137464.6 ENSMUST00000174196.7 ENSMUST00000174196.7 ENSMUST00000123151.7 ENSMUST00000123151.7 ENSMUST00000123151.7 ENSMUST0000012351.1 ENSMUST00000172755.1 ENSMUST00000136289.1	ENSMUSTO0000106251.9 5536 ENSMUSTO0000077609.11 1831 ENSMUSTO0000077609.11 18031 ENSMUSTO00000121616.8 1102 ENSMUSTO0000121616.8 1102 ENSMUSTO0000141997.1 761 ENSMUSTO0000174632.7 390 ENSMUSTO0000154843.7 600 ENSMUSTO0000137464.6 354 ENSMUSTO0000137464.6 354 ENSMUSTO0000174196.7 2374 ENSMUSTO0000174196.7 2374 ENSMUSTO0000123151.7 2158 ENSMUSTO0000205351.1 2055 ENSMUSTO0000205261.1 1859 ENSMUSTO0000172755.1 670 ENSMUSTO0000136289.1 600	ENSMUST00000106251.9         5536         518aa           ENSMUST00000077609.11         1831         517aa           ENSMUST00000079045.2         2623         122aa           ENSMUST00000121616.8         1102         280aa           ENSMUST00000141997.1         761         104aa           ENSMUST00000174632.7         390         130aa           ENSMUST00000154843.7         600         No protein           ENSMUST00000137464.6         354         No protein           ENSMUST00000128851.7         4909         No protein           ENSMUST00000174196.7         2374         No protein           ENSMUST00000123151.7         2158         No protein           ENSMUST00000205261.1         1859         No protein           ENSMUST00000172755.1         670         No protein           ENSMUST00000136289.1         600         No protein	ENSMUST00000106251.9         5536         518aa         Protein coding           ENSMUST00000077609.11         1831         517aa         Protein coding           ENSMUST00000079045.2         2623         122aa         Protein coding           ENSMUST00000121616.8         1102         280aa         Protein coding           ENSMUST00000141997.1         761         104aa         Protein coding           ENSMUST00000174632.7         390         130aa         Protein coding           ENSMUST00000154843.7         600         No protein         Processed transcript           ENSMUST00000137464.6         354         No protein         Retained intron           ENSMUST00000128851.7         4909         No protein         Retained intron           ENSMUST00000123151.7         2158         No protein         Retained intron           ENSMUST00000205351.1         2055         No protein         Retained intron           ENSMUST00000172755.1         670         No protein         Retained intron           ENSMUST00000136289.1         600         No protein         Retained intron	ENSMUST00000106251.9         5536         518aa         Protein coding         CCDS21886           ENSMUST00000077609.11         1831         517aa         Protein coding         CCDS85425           ENSMUST00000079045.2         2623         122aa         Protein coding         -           ENSMUST00000121616.8         1102         280aa         Protein coding         -           ENSMUST000001441997.1         761         104aa         Protein coding         -           ENSMUST00000174632.7         390         130aa         Protein coding         -           ENSMUST00000154843.7         600         No protein         Processed transcript         -           ENSMUST00000137464.6         354         No protein         Retained intron         -           ENSMUST00000128851.7         4909         No protein         Retained intron         -           ENSMUST00000174196.7         2374         No protein         Retained intron         -           ENSMUST00000205351.1         2055         No protein         Retained intron         -           ENSMUST00000172755.1         670         No protein         Retained intron         -           ENSMUST00000136289.1         600         No protein         Retained intron         - </td <td>ENSMUST00000106251.9         5536         518aa         Protein coding         CCDS21886         P56959 Q564D0           ENSMUST00000077609.11         1831         517aa         Protein coding         CCDS85425         Q8CFQ9           ENSMUST00000079045.2         2623         122aa         Protein coding         -         Q8BNR3           ENSMUST00000121616.8         1102         280aa         Protein coding         -         Q91VQ2           ENSMUST00000174632.7         390         130aa         Protein coding         -         G3UXT7           ENSMUST00000154843.7         600         No protein         Processed transcript         -         -           ENSMUST00000137464.6         354         No protein         Retained intron         -         -           ENSMUST00000174196.7         2374         No protein         Retained intron         -         -           ENSMUST00000123151.7         2158         No protein         Retained intron         -         -           ENSMUST00000205351.1         2055         No protein         Retained intron         -         -           ENSMUST00000172755.1         670         No protein         Retained intron         -         -           ENSMUST00000136289.1         600</td>	ENSMUST00000106251.9         5536         518aa         Protein coding         CCDS21886         P56959 Q564D0           ENSMUST00000077609.11         1831         517aa         Protein coding         CCDS85425         Q8CFQ9           ENSMUST00000079045.2         2623         122aa         Protein coding         -         Q8BNR3           ENSMUST00000121616.8         1102         280aa         Protein coding         -         Q91VQ2           ENSMUST00000174632.7         390         130aa         Protein coding         -         G3UXT7           ENSMUST00000154843.7         600         No protein         Processed transcript         -         -           ENSMUST00000137464.6         354         No protein         Retained intron         -         -           ENSMUST00000174196.7         2374         No protein         Retained intron         -         -           ENSMUST00000123151.7         2158         No protein         Retained intron         -         -           ENSMUST00000205351.1         2055         No protein         Retained intron         -         -           ENSMUST00000172755.1         670         No protein         Retained intron         -         -           ENSMUST00000136289.1         600

The strategy is based on the design of Fus-203 transcript, The transcription is shown below



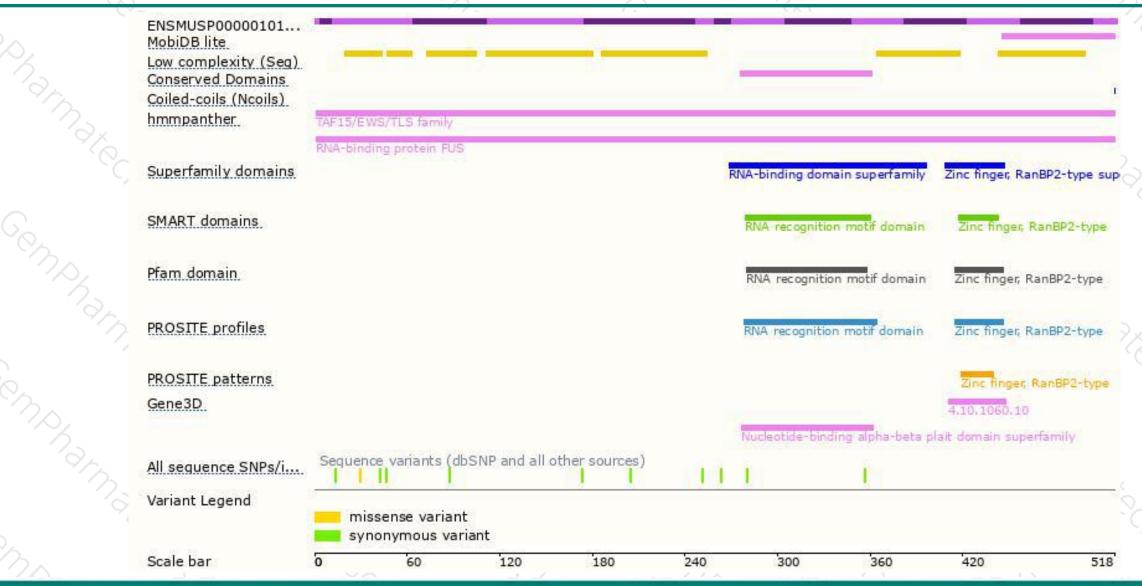
## Genomic location distribution





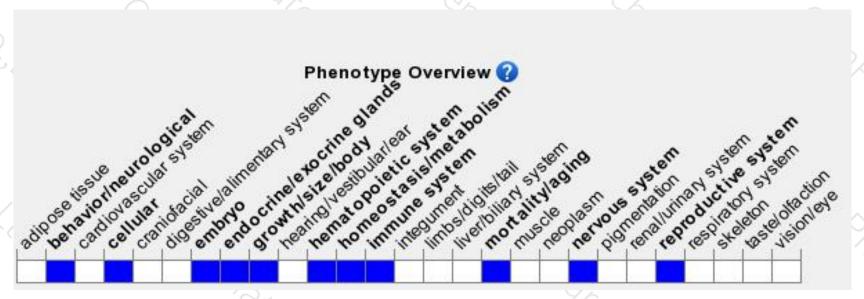
## Protein domain





## 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, Homozygotes for targeted null mutations exhibit impaired lymphocyte development, chromosomal instability, increased cellular radiation sensitivity, high neonatal mortality, and male sterility associated with lack of chromosomal pairing.



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





