

Erbb4 Cas9-CKO Strategy

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Project Overview

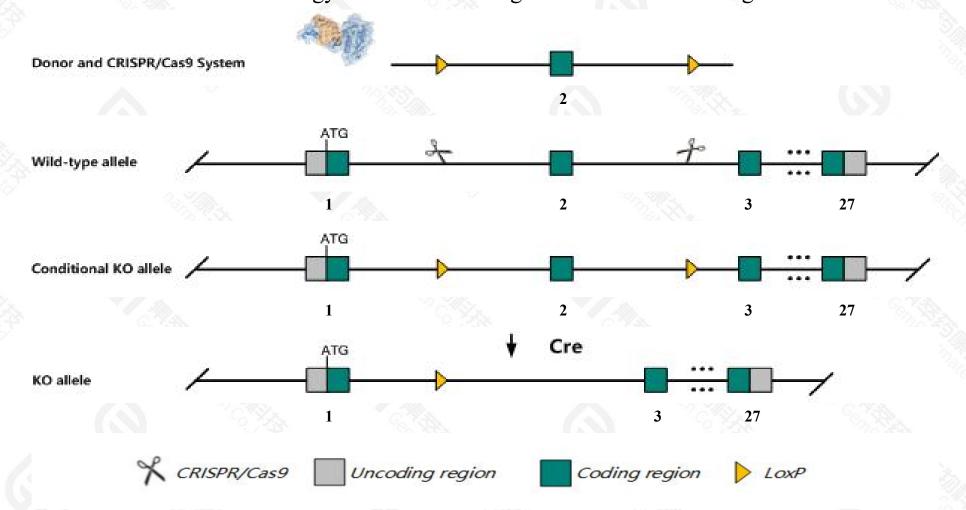


Project Name	Erbb4		
Project type	Cas9-CKO		
Strain background	C57BL/6JGpt		

Conditional Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Erbb4* gene. The schematic diagram is as follows:



Technical routes



- ➤ The *Erbb4* gene has 5 transcripts. According to the structure of *Erbb4* gene, exon2 of *Erbb4*202(ENSMUST00000121473.8) transcript is recommended as the knockout region. The region contains 152bp coding sequence.

 Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Erbb4* 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.

Notice



- > According to the existing MGI data, homozygotes for a targeted null mutation exhibit cardiac defects, alterations in hindbrain development, and midgestational lethality. Heterozygotes show schizophrenia-like behavior. Genetically rescued females show mammary defects.
- > The *Erbb4* gene is located on the Chr1. 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)



Erbb4 erb-b2 receptor tyrosine kinase 4 [Mus musculus (house mouse)]

Gene ID: 13869, updated on 22-Mar-2020

Summary

☆ ?

Official Symbol Erbb4 provided by MGI

Official Full Name erb-b2 receptor tyrosine kinase 4 provided by MGI

Primary source MGI:MGI:104771

See related Ensembl: ENSMUSG00000062209

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 Her4, c-erbB-4

Expression Biased expression in CNS E18 (RPKM 2.3), frontal lobe adult (RPKM 2.1) and 8 other tissuesSee more

Orthologs <u>human all</u>

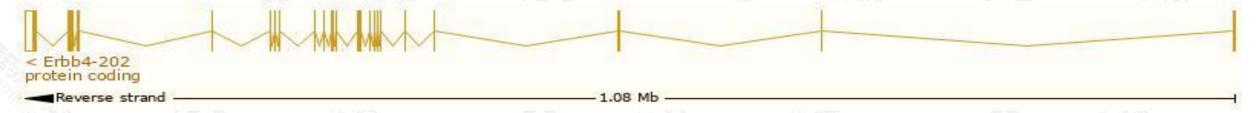
Transcript information (Ensembl)



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

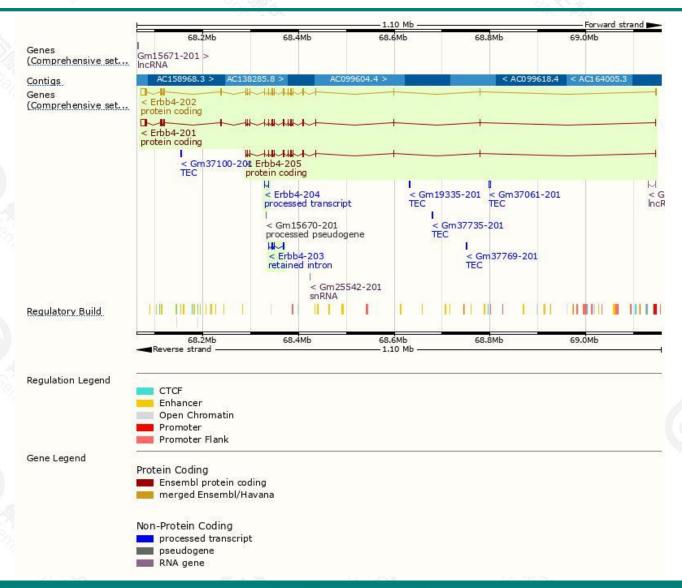
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Erbb4-202	ENSMUST00000121473.7	12037	<u>1292aa</u>	Protein coding	CCDS48285	Q61527	TSL:1 GENCODE basic
Erbb4-201	ENSMUST00000119142.7	11782	1308aa	Protein coding	-	Q61527	TSL:5 GENCODE basic APPRIS is a system to annotate alternatively spliced transcripts based on a range of computational methods to identify the most functionally important transcript(s) of a gene. APPRIS P
Erbb4-205	ENSMUST00000153432.1	2196	732aa	Protein coding	12	B2KGF7	CDS 3' incomplete TSL:5
Erbb4-204	ENSMUST00000131148.1	240	No protein	Processed transcript	2	2	TSL:5
Erbb4-203	ENSMUST00000126682.1	744	No protein	Retained intron	-		TSL:5

The strategy is based on the design of *Erbb4-202* transcript, the transcription is shown below:



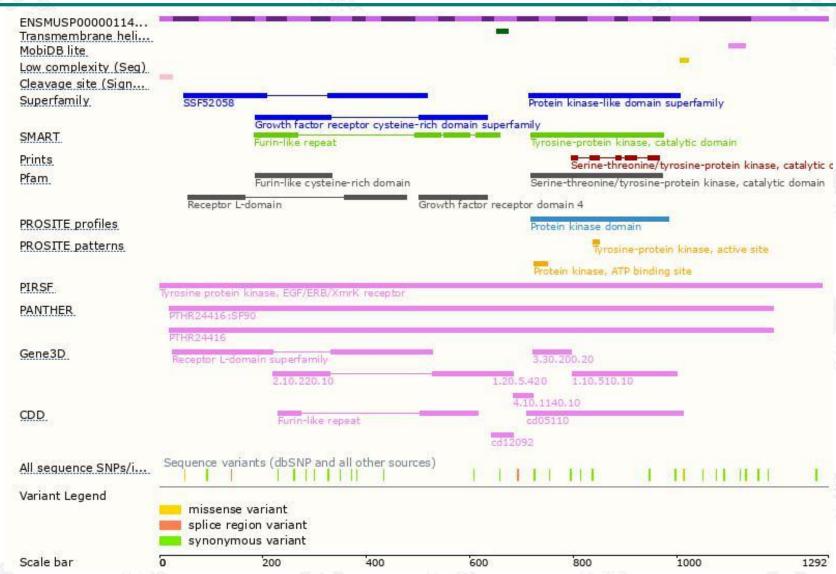
Genomic location distribution





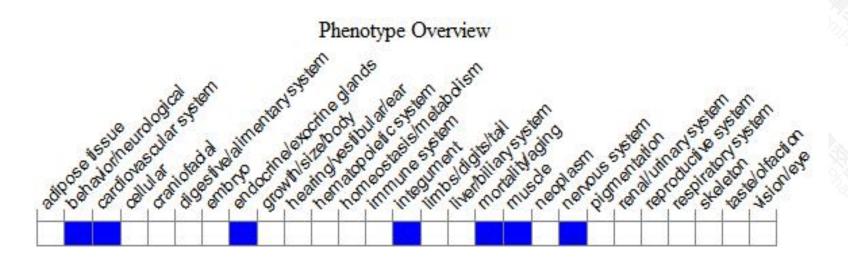
Protein domain





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 a targeted null mutation exhibit cardiac defects, alterations in hindbrain development, and midgestational lethality. Heterozygotes show schizophrenia-like behavior. Genetically rescued females show mammary defects.



If you have any questions, you are welcome to inquire.

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