

Edn1 Cas9-CKO Strategy

Designer:

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Design Date:

2019-7-17

Project Overview



Project Name

Edn1

Project type

Cas9-CKO

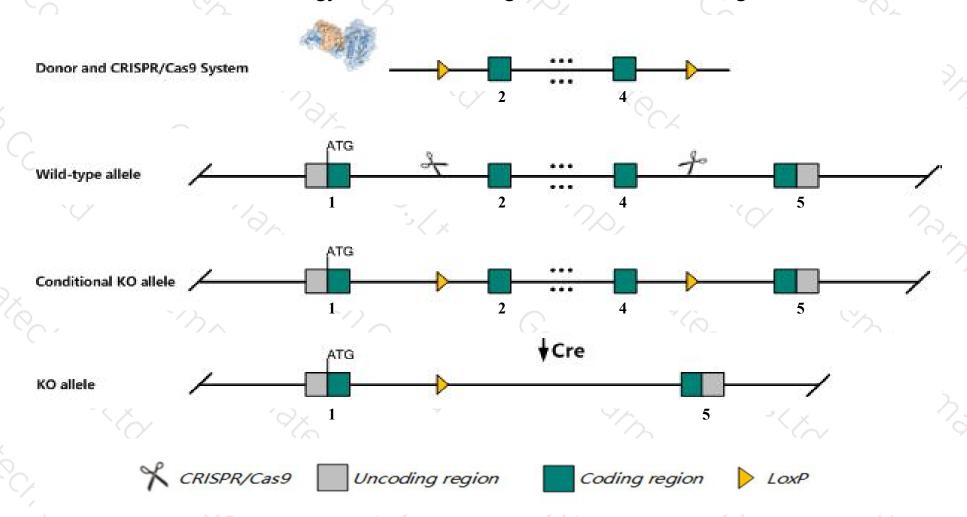
Strain background

C57BL/6JGpt

Conditional Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Edn1* gene. The schematic diagram is as follows:



Technical routes



- The *Edn1* gene has 1 transcript. According to the structure of *Edn1* gene, exon2-exon4 of *Edn1-201* (ENSMUST00000021796.8) transcript is recommended as the knockout region. The region contains 442bp coding sequence. Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Edn1* 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 cardiovascular malformations, craniofacial abnormalities, and lethality due to respiratory failure at birth. Heterozygotes develop elevated arterial blood pressure.
- > The *Edn1* gene is located on the Chr13. 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)



Edn1 endothelin 1 [Mus musculus (house mouse)]

Gene ID: 13614, updated on 25-Mar-2019

Summary

☆ [?]

Official Symbol Edn1 provided by MGI

Official Full Name endothelin 1 provided by MGI

Primary source MGI:MGI:95283

See related Ensembl:ENSMUSG00000021367

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 ET-1, PPET1, preproET

Summary This gene encodes a member of the endothelin family of peptides. The encoded preproprotein undergoes proteolytic processing to generate

a peptide before secretion by the vascular endothelial cells. The mature peptide has various biological activities such as vasoconstriction, cell proliferation, stimulation of hormone release and modulation of central nervous activity. Mice lacking the encoded protein exhibit neonatal lethality accompanied with numerous craniofacial and cardiovascular defects due to disruption in cranial and cardiac neural crest cell

patterning during early embryogenesis. [provided by RefSeq, Feb 2016]

Expression Broad expression in lung adult (RPKM 5.9), colon adult (RPKM 3.5) and 20 other tissuesSee more

Orthologs <u>human</u> all

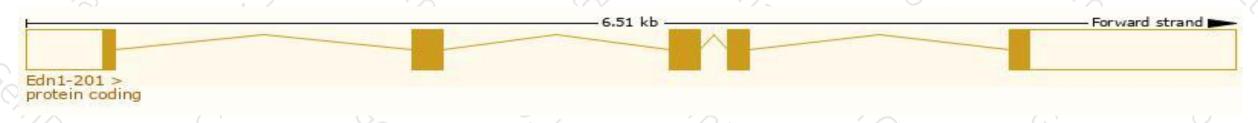
Transcript information (Ensembl)



The gene has 1 transcript, and the transcript is shown below:

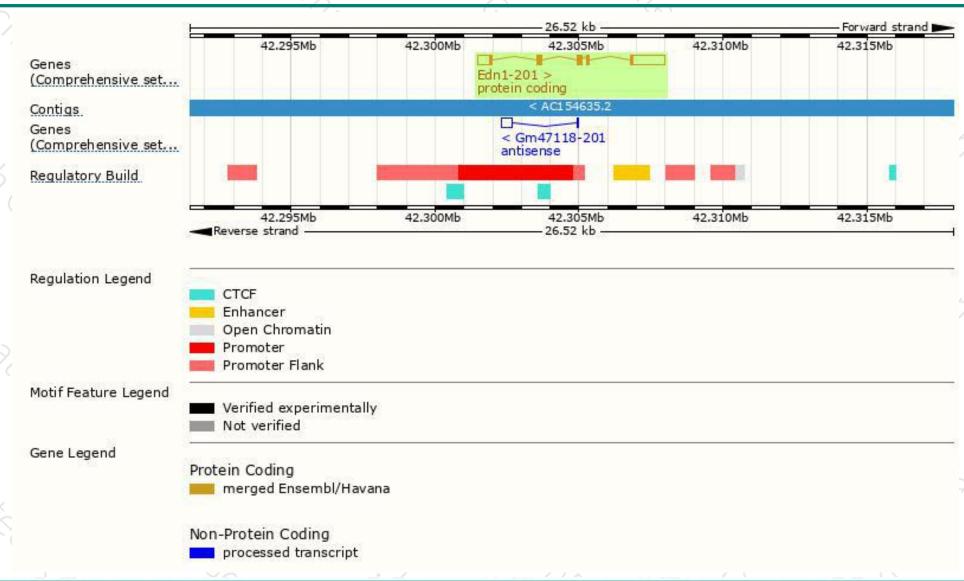
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags	
Edn1-201	ENSMUST00000021796.8	2139	202aa	Protein coding	CCDS26474	P22387 Q544E0	TSL:1 GENCODE basic APPRIS P1	Ľ

The strategy is based on the design of *Edn1-201* transcript, The transcription is shown below



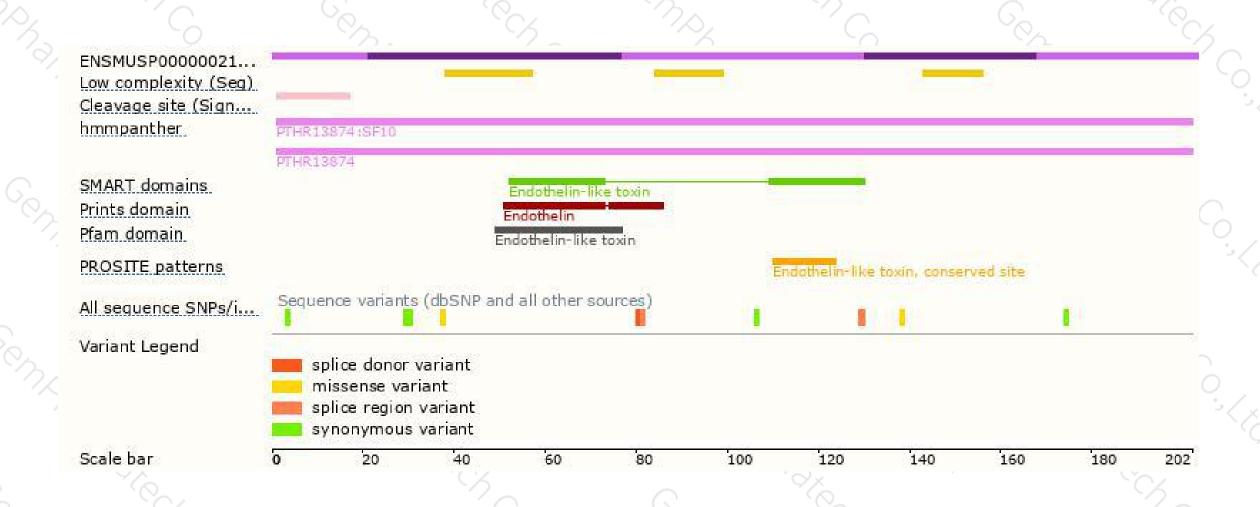
Genomic location distribution





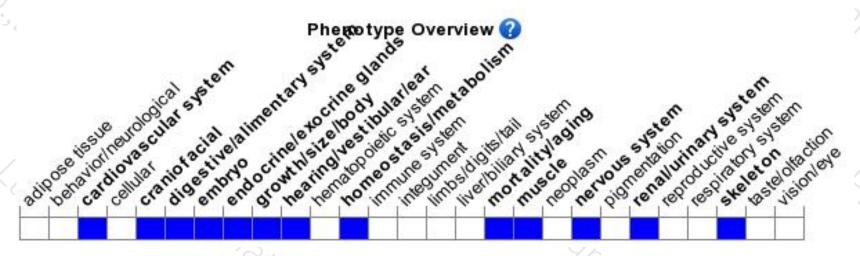
Protein domain





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 cardiovascular malformations, craniofacial abnormalities, and lethality due to respiratory failure at birth. Heterozygotes develop elevated arterial blood pressure.



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





