

# Dolar Day Co. Nedd41 Cas9-KO Strategy Rohalmakech Co.

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## **Project Overview**



**Project Name** 

Nedd4l

**Project type** 

Cas9-KO

Strain background

C57BL/6JGpt

# **Knockout strategy**



This model will use CRISPR/Cas9 technology to edit the Nedd4l gene. The schematic diagram is as follows:



#### **Technical routes**



- The *Nedd4l* gene has 34 transcripts. According to the structure of *Nedd4l* gene, exon2-exon9 of *Nedd4l-215*(ENSMUST00000235343.1) transcript is recommended as the knockout region. The region contains 632bp coding sequence.

  Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Nedd4l* gene. The brief process is as follows: CRISPR/Cas9 system

#### **Notice**



- ➤ According to the existing MGI data, Mice homozygous for a null mutation display salt sensitive hypertension and high salt diet induced cardiac hypertrophy. A spontaneous mutation results in overt diabetes insipidus. Mice homozygous for a knock-out allele exhibit neonatal lethality with primary atelectasis.
- > The *Nedd4l* gene is located on the Chr18. 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)



#### Nedd4l neural precursor cell expressed, developmentally down-regulated gene 4-like [Mus musculus (house mouse)]

Gene ID: 83814, updated on 12-Mar-2019

#### Summary



Official Symbol Nedd4l provided by MGI

Official Full Name neural precursor cell expressed, developmentally down-regulated gene 4-like provided by MGI

Primary source MGI:MGI:1933754

See related Ensembl:ENSMUSG00000024589

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 1300012C07Rik, Nedd4-2, Nedd4b

Expression Ubiquitous expression in whole brain E14.5 (RPKM 4.8), CNS E18 (RPKM 4.7) and 26 other tissuesSee more

Orthologs <u>human</u> all

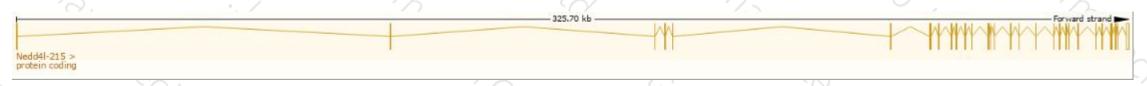
# Transcript information (Ensembl)



The gene has 34 transcripts, all transcripts are shown below:

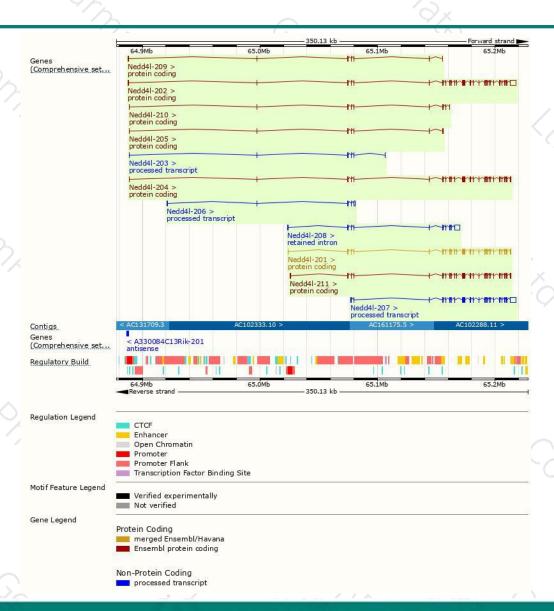
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Nedd4I-215	ENSMUST00000235343.1	3792	976aa	Protein coding	CCDS50309		GENCODE basic APPRIS P2
ledd4I-201	ENSMUST00000080418.6	3671	855aa	Protein coding	CCDS29305	G3X9H8	TSL:1 GENCODE basic
ledd41-228	ENSMUST00000237384.1	3592	855aa	Protein coding	CCDS29305		GENCODE basic
ledd4I-211	ENSMUST00000226058.1	3393	855aa	Protein coding	CCDS29305	<u> G3Х9Н8</u>	GENCODE basic
ledd41-204	ENSMUST00000224347.2	3295	855aa	Protein coding	CCDS29305	A0A286YCM8	GENCODE basic
ledd41-202	ENSMUST00000163516.8	7787	835aa	Protein coding		E9PXB7	TSL:5 GENCODE basic
ledd41-233	ENSMUST00000237854.1	4845	835aa	Protein coding	-		GENCODE basic
edd41-223	ENSMUST00000236736.1	4802	956aa	Protein coding	- 4		GENCODE basic APPRIS ALT2
edd4I-221	ENSMUST00000236209.1	3254	912aa	Protein coding			GENCODE basic
edd41-220	ENSMUST00000236103.1	3182	751aa	Protein coding			GENCODE basic
edd41-225	ENSMUST00000236898.1	3034	1004aa	Protein coding	-		GENCODE basic
edd41-229	ENSMUST00000237410.1	2974	984aa	Protein coding	- 4		GENCODE basic
edd4I-218	ENSMUST00000235743.1	2917	872aa	Protein coding			GENCODE basic
edd4I-231	ENSMUST00000237644.1	2018	527aa	Protein coding			CDS 5' incomplete
edd4I-217	ENSMUST00000235577.1	770	256aa	Protein coding	-		5' and 3' truncations in transcript evidence prevent annotation of the start and the end of the CDS, CDS 5' and 3' incompl
edd4l-210	ENSMUST00000225261.2	747	224aa	Protein coding	- 1	A0A286YCR1	CDS 3' incomplete
edd41-205	ENSMUST00000224385.2	735	113aa	Protein coding		A0A286YDV4	CDS 3' incomplete
edd4I-213	ENSMUST00000235310.1	586	94aa	Protein coding			CDS 3' incomplete
edd4I-224	ENSMUST00000236764.1	548	37aa	Protein coding			CDS 3' incomplete
edd4I-207	ENSMUST00000224663.2	4875	No protein	Processed transcript	10		
edd4l-226	ENSMUST00000236916.1	792	No protein	Processed transcript			
edd4I-212	ENSMUST00000235204.1	776	No protein	Processed transcript			
edd4I-222	ENSMUST00000236595.1	717	No protein	Processed transcript	-		
edd41-203	ENSMUST00000223959.1	414	No protein	Processed transcript			
edd41-209	ENSMUST00000225057.2	381	No protein	Processed transcript			
edd41-234	ENSMUST00000238122.1	373	No protein	Processed transcript			
edd4I-219	ENSMUST00000236100.1	348	No protein	Processed transcript	-		
edd4I-227	ENSMUST00000236929.1	240	No protein	Processed transcript			
edd4I-208	ENSMUST00000224890.1	5814	No protein	Retained intron			
edd4I-216	ENSMUST00000235571.1	1116	No protein	Retained intron			
edd4I-206	ENSMUST00000224516.1	1045	No protein	Retained intron			
edd4I-214	ENSMUST00000235339.1	772	No protein	Retained intron			
edd41-232	ENSMUST00000237818.1	550	No protein	Retained intron			
ledd4I-230	ENSMUST00000237492.1	437	No protein	Retained intron			

The strategy is based on the design of Nedd4l-215 transcript, The transcription is shown below



### Genomic location distribution





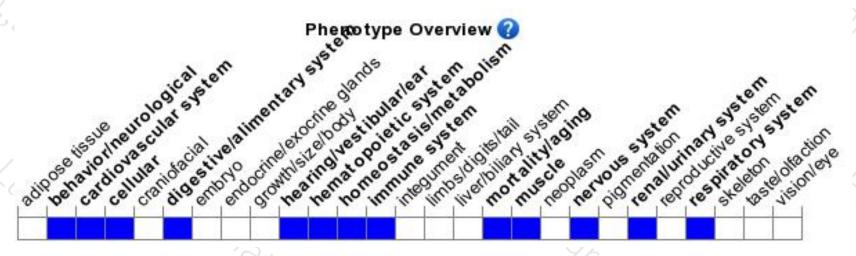
#### Protein domain





# 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, Mice homozygous for a null mutation display salt sensitive hypertension and high salt diet induced cardiac hypertrophy. A spontaneous mutation results in overt diabetes insipidus. Mice homozygous for a knock-out allele exhibit neonatal lethality with primary atelectasis.



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





