

# Atxn7 Cas9-CKO Strategy

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## **Project Overview**



**Project Name** 

Atxn7

**Project type** 

Cas9-CKO

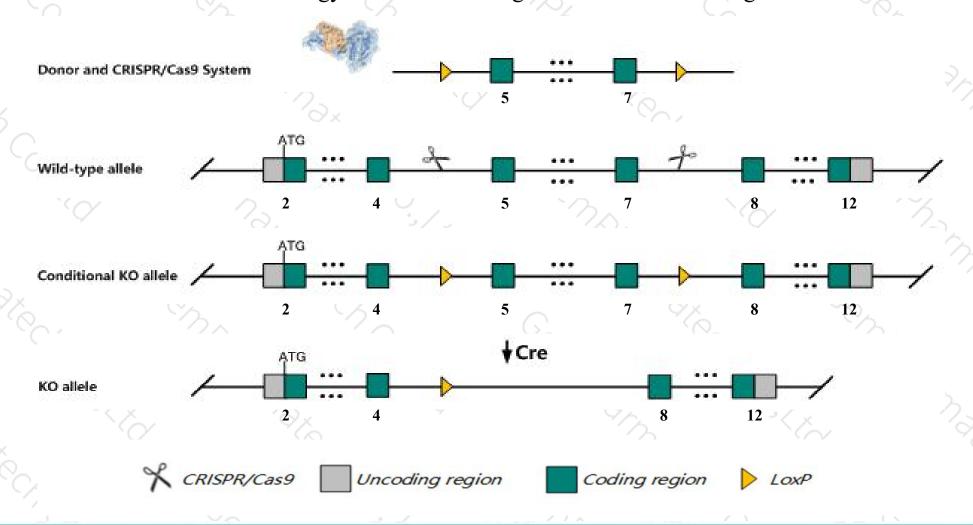
Strain background

C57BL/6JGpt

## Conditional Knockout strategy



This model will use CRISPR/Cas9 technology to edit the Atxn7 gene. The schematic diagram is as follows:



### Technical routes



- The Atxn7 gene has 9 transcripts. According to the structure of Atxn7 gene, exon5-exon7 of Atxn7-201 (ENSMUST00000022257.3) transcript is recommended as the knockout region. The region contains 593bp coding sequence. Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Atxn7* 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, heterozygotes for a targeted mutation with an expanded polyglutamine tract exhibit impaired coordination, ataxia, reduced growth, kyphosis, eye defects, poor reproduction, and high mortality at around 4 months. homozygotes die at 7-8 weeks of age.
- $\rightarrow$  Transcript Atxn7-205 is incomplete, so the effect on it is unknown.
- > The Atxn7 gene is located on the Chr14. 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)



#### Atxn7 ataxin 7 [ Mus musculus (house mouse) ]

Gene ID: 246103, updated on 12-May-2020

#### Summary

△ ?

Official Symbol Atxn7 provided by MGI

Official Full Name ataxin 7 provided by MGI

Primary source MGI:MGI:2179277

See related Ensembl: ENSMUSG00000021738

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 Sca7; Al627028; ataxin-7; A430107N12Rik

Expression Ubiquitous expression in lung adult (RPKM 4.5), bladder adult (RPKM 4.4) and 28 other tissues See more

Orthologs human all

## Transcript information (Ensembl)



The gene has 9 transcripts, all transcripts are shown below:

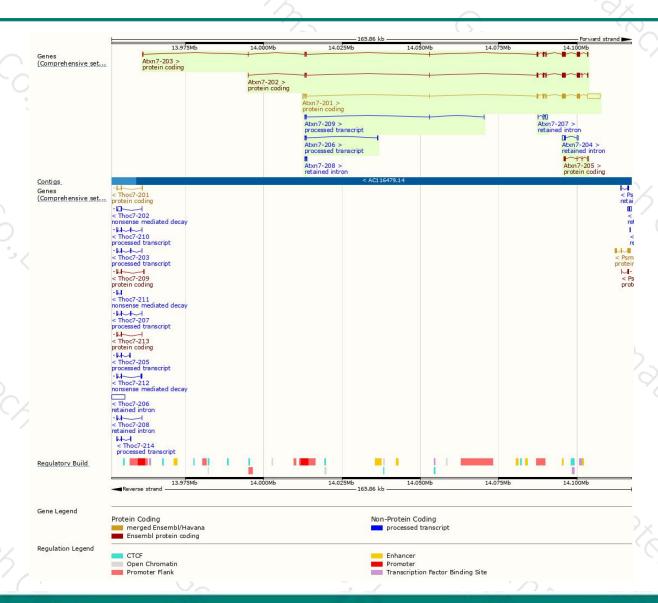
Name	Transcript ID	bp 🛊	Protein	Biotype	CCDS	UniProt	Flags
Atxn7-201	ENSMUST00000022257.3	6853	867aa	Protein coding	CCDS26823 €	Q8R4I1₽	TSL:1 GENCODE basic APPRIS P2
Atxn7-203	ENSMUST00000223880.1	2946	867aa	Protein coding	CCDS26823 €	Q8R4I1₽	GENCODE basic   APPRIS P2
Atxn7-202	ENSMUST00000223714.1	2954	920aa	Protein coding	-	<u>A0A286YDW9</u> ₽	GENCODE basic APPRIS ALT2
Atxn7-205	ENSMUST00000224315.1	779	<u>106aa</u>	Protein coding	-	A0A286YCL5₽	CDS 5' incomplete
Atxn7-209	ENSMUST00000226073.1	492	No protein	Processed transcript	323	92	<u> </u>
Atxn7-206	ENSMUST00000224370.1	392	No protein	Processed transcript	5 <del>.</del> 5		: 5.
Atxn7-204	ENSMUST00000223932.1	1081	No protein	Retained intron	-	5	5.
Atxn7-207	ENSMUST00000224616.1	766	No protein	Retained intron	9 <del>-</del> 2		. 5
Atxn7-208	ENSMUST00000225164.1	534	No protein	Retained intron	(-)	-	

The strategy is based on the design of *Atxn7-201* transcript, the transcription is shown below:



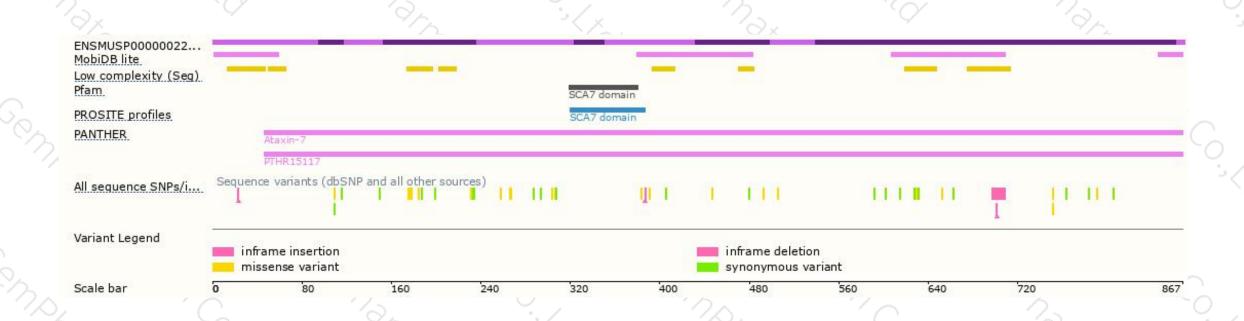
### Genomic location distribution





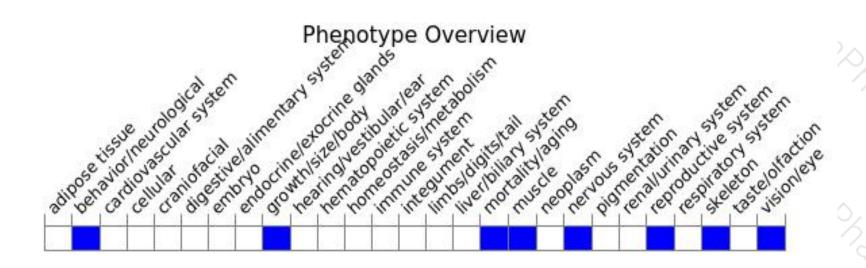
## Protein domain





## 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, heterozygotes for a targeted mutation with an expanded polyglutamine tract exhibit impaired coordination, ataxia, reduced growth, kyphosis, eye defects, poor reproduction, and high mortality at around 4 months. Homozygotes die at 7-8 weeks of age.



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





