

Wdr77 Cas9-CKO Strategy

Designer: JiaYu

Reviewer: Xiaojing Li

Design Date: 2020-7-21

Project Overview



Project Name

Wdr77

Project type

Cas9-CKO

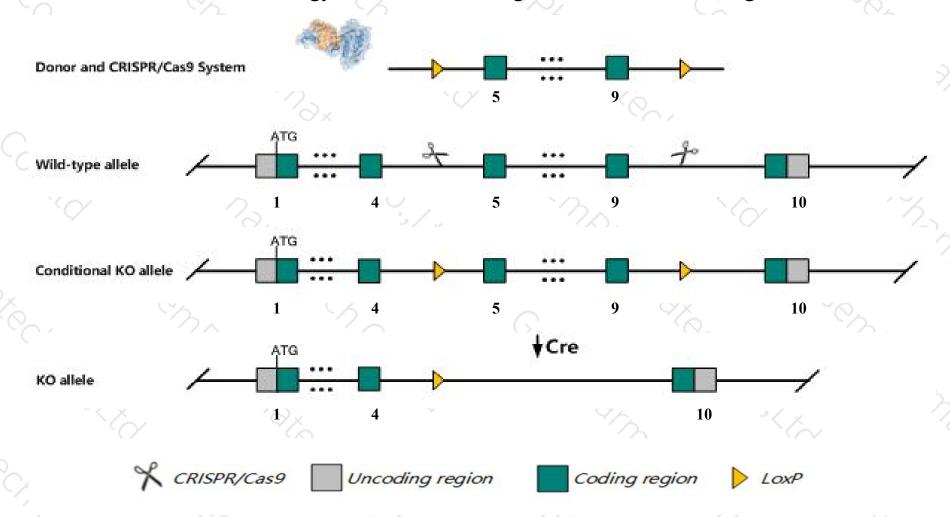
Strain background

C57BL/6JGpt

Conditional Knockout strategy



This model will use CRISPR/Cas9 technology to edit the *Wdr77* gene. The schematic diagram is as follows:



Technical routes



- The *Wdr77* gene has 5 transcripts. According to the structure of *Wdr77* gene, exon5-exon9 of *Wdr77*-201(ENSMUST00000010278.11) transcript is recommended as the knockout region. The region contains 376bp coding sequence. Knock out the region will result in disruption of protein function.
- ➤ In this project we use CRISPR/Cas9 technology to modify *Wdr77* 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, mice homozygous for a knock-out allele die prior to E8.5 for unknown reasons. Heterozygotes develop multifocal hyperplasia in the dorsal prostate; however, no prostate tumors are detected up to 12 months of age.
- > Some amino acids will remain at the N-terminus and some functions may be retained.
- > The *Wdr77* gene is located on the Chr3. 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)



Wdr77 WD repeat domain 77 [Mus musculus (house mouse)]

Gene ID: 70465, updated on 13-Mar-2020





Official Symbol Wdr77 provided by MGI

Official Full Name WD repeat domain 77 provided by MGI

Primary source MGI:MGI:1917715

See related Ensembl: ENSMUSG000000000561

Gene type protein coding
RefSeq status PROVISIONAL
Organism <u>Mus musculus</u>

Lineage Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha;

Muroidea; Muridae; Murinae; Mus; Mus

Also known as 2610003I18Rik, 2610312E17Rik, C79984, p44/MEP50

Expression Ubiquitous expression in duodenum adult (RPKM 34.7), liver adult (RPKM 22.0) and 28 other tissuesSee more

Orthologs <u>human</u> <u>all</u>

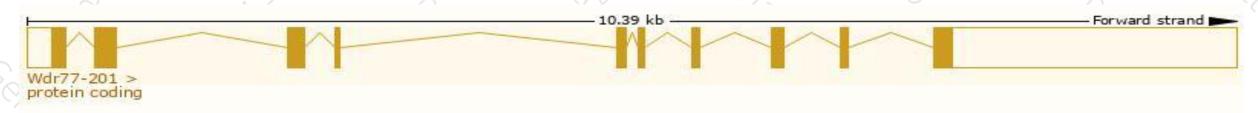
Transcript information (Ensembl)



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

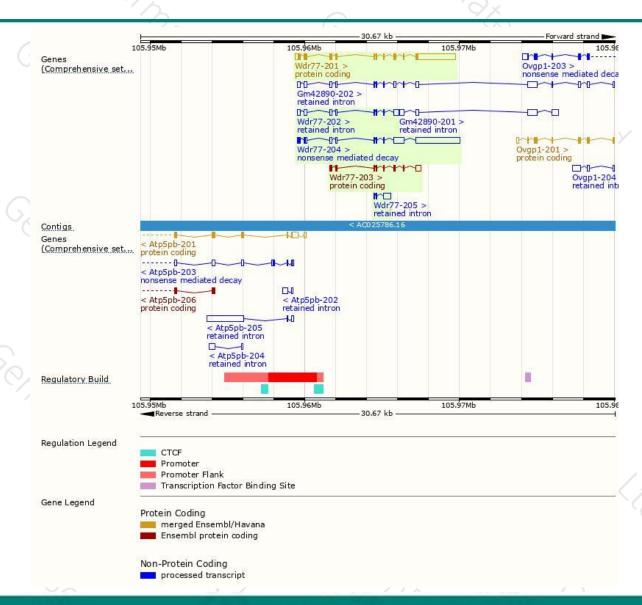
						1 / km	
Name	Transcript ID	bp	Protein	Biotype	CCDS	UniProt	Flags
Wdr77-201	ENSMUST00000010278.11	3682	<u>342aa</u>	Protein coding	CCDS17715	Q99J09	TSL:1 GENCODE basic APPRIS P1
Wdr77-203	ENSMUST00000128005.1	911	<u>167aa</u>	Protein coding	-	<u>F7D5L2</u>	CDS 5' incomplete TSL:3
Wdr77-204	ENSMUST00000130994.7	4286	<u>60aa</u>	Nonsense mediated decay	858	D6RIL4	TSL:1
Wdr77-202	ENSMUST00000127464.7	1095	No protein	Retained intron	100	-	TSL:1
Wdr77-205	ENSMUST00000151263.2	590	No protein	Retained intron	125	Ψ.	TSL:2

The strategy is based on the design of Wdr77-201 transcript, the transcription is shown below:



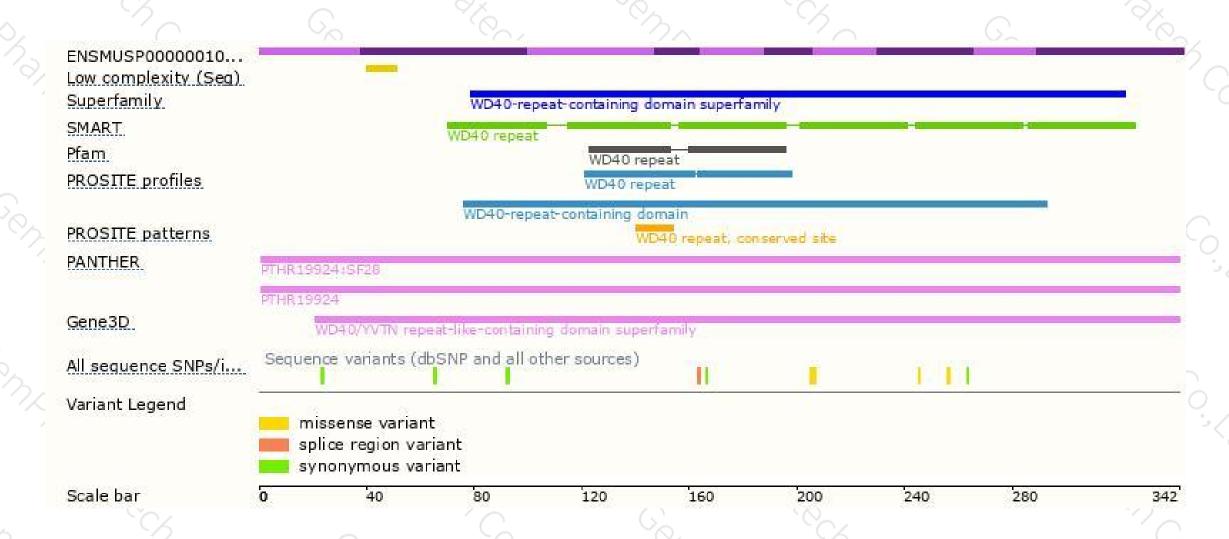
Genomic location distribution





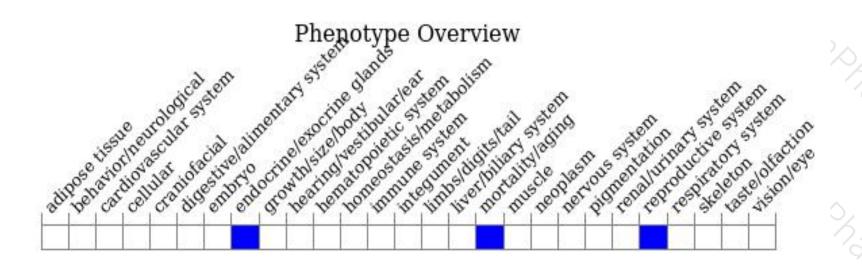
Protein domain





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,mice homozygous for a knock-out allele die prior to E8.5 for unknown reasons. Heterozygotes develop multifocal hyperplasia in the dorsal prostate; however, no prostate tumors are detected up to 12 months of age.



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





