

Ghr Cas9-KO Strategy

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Reviewer: Yanhua Shen

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Overview

Target Gene Name

• Ghr

Project Type

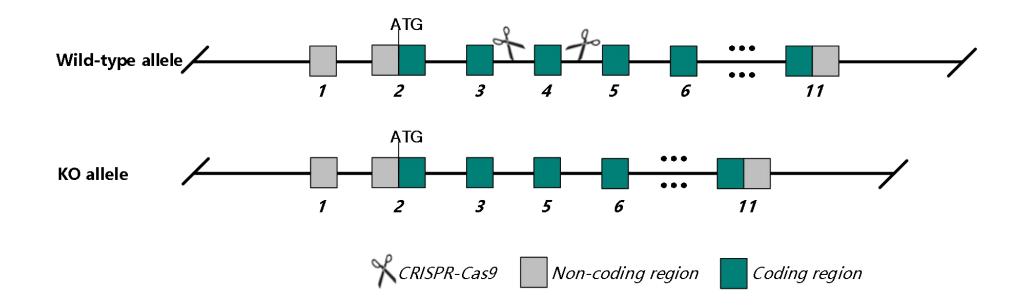
• Cas9-KO

Genetic Background

• C57BL/6JGpt



Strain Strategy



Schematic representation of CRISPR-Cas9 engineering used to edit the Ghr gene.

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Technical Information

- The *Ghr* gene has 10 transcripts. According to the structure of *Ghr* gene, exon4 of *Ghr*-201 (ENSMUST0000069451.11) transcript is recommended as the knockout region. The region contains 130bp of coding sequences. Knocking out the region will result in disruption of protein function.
- In this project we use CRISPR-Cas9 technology to modify *Ghr* gene. The brief process is as follows: gRNAs were transcribed in vitro. Cas9 and gRNAs 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 on-target amplicon sequencing. A stable F1-generation mouse strain was obtained by mating positive F0-generation mice with C57BL/6JGpt mice and confirmation of the desired mutant allele was carried out by PCR and on-target amplicon sequencing.

Gene Information

Ghr growth hormone receptor [Mus musculus (house mouse)]

Gene ID: 14600, updated on 10-Jan-2023

🗄 Download Datasets

Official Symbol	Ghr provided by MGI	
Official Full Name	growth hormone receptor provided by MGI	
Primary source	MGI:MGI:95708	
See related	Ensembl:ENSMUSC00000055737 AllianceGenome:MGI:95708	
Gene type	protein coding	
RefSeq status	VALIDATED	
Organism	Mus musculus	
Lineage	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomor	pha; Muroidea;
	Muridae; Murinae; Mus	
	GHBP; GHR/BP	
Summary	Enables growth hormone receptor activity and peptide hormone binding activity. Involved in growth hormone receptor signaling pathw of peptidyl-tyrosine phosphorylation; and regulation of growth. Acts upstream of or within taurine metabolic process. Located in nucleu membrane. Is extrinsic component of membrane. Is expressed in several structures, including alimentary system; brain; early concep system; and integumental system. Used to study Laron syndrome. Human ortholog(s) of this gene implicated in several diseases, inc syndrome; familial hypercholesterolemia; isolated growth hormone deficiency; osteoarthritis; and type 2 diabetes mellitus. Orthologou (growth hormone receptor). [provided by Alliance of Genome Resources, Apr 2022]	us and plasma tus; genitourinary luding Laron
Expression Orthologs		
NEW	Try the new <u>Gene table</u> Try the new <u>Transcript table</u>	
enomic context		

Source: https://www.ncbi.nlm.nih.gov/

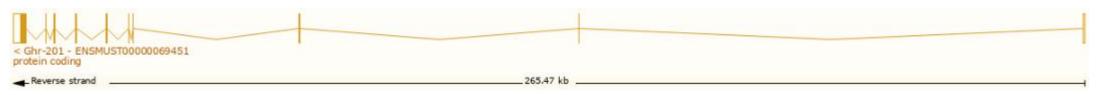


Transcript Information

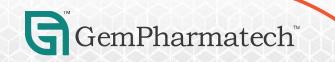
The gene has 10 transcripts, all transcripts are shown below:

Transcript ID	Name 🍦	bp 💧	Protein	Biotype	CCDS 🍵	UniProt Match	Flags			
ENSMUST0000069451.11	Ghr-201	4175	<u>650aa</u>	Protein coding	CCDS27358	Q3UP14@	Ensembl Canonical GENCODE basic APPRIS P1 TSL:1			
ENSMUST00000161561.8	Ghr-208	2749	<u>650aa</u>	Protein coding	<u>CCDS27358</u> മ	Q3UP14@	GENCODE basic APPRIS P1 TSL:1			
ENSMUST00000110697.9	Ghr-202	1213	<u>297aa</u>	Protein coding	CCDS37024@	P16882-2@	GENCODE basic TSL:5			
ENSMUST00000110698.9	Ghr-203	1171	<u>297aa</u>	Protein coding	CCDS37024@	P16882-2@	GENCODE basic TSL1			
ENSMUST00000161770.2	Ghr-209	898	<u>183aa</u>	Protein coding		E0CXS6译	TSL:3 CDS 3' incomplete			
ENSMUST00000159912.2	Ghr-205	738	No protein	Protein coding CDS not defined		-	TSL:3			
ENSMUST00000159508.8	Ghr-204	598	No protein	Protein coding CDS not defined		844).	TSL:5			
ENSMUST00000160343.8	Ghr-206	472	No protein	Protein coding CDS not defined		100	TSL:3			
ENSMUST00000161180.2	Ghr-207	2481	No protein	Retained intron		64-5	TSL:1			
ENSMUST00000162993.8	Ghr-210	1794	No protein	Retained intron		(T .)	TSL:1			

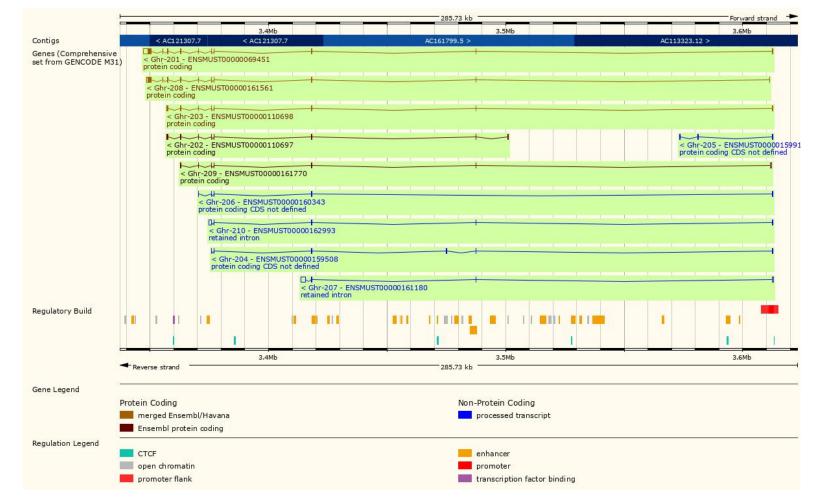
The strategy is based on the design of *Ghr*-201 transcript, the transcription is shown below:



Source: https://www.ensembl.org



Genomic Information



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Source: : https://www.ensembl.org

Protein Information

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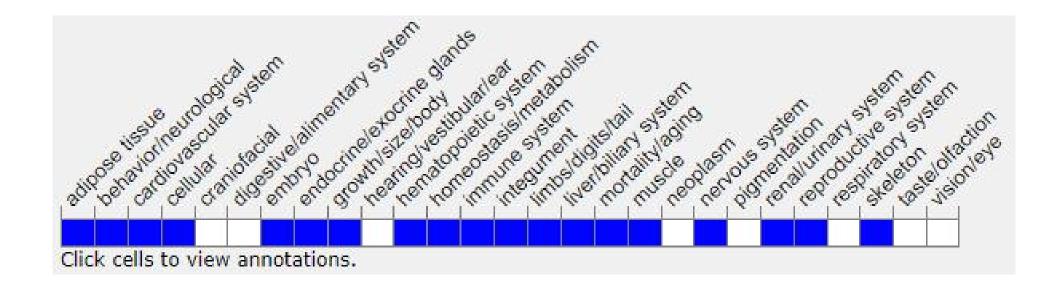
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ENSMUSP00000069453 Transmembrane helice Cleavage site (Sig Superfamily Pfam		ceptor, ligand binding		Growth hormone-binding pr	otein				
PROSITE profiles		Fibronectin type 111							
PROSITE patterns									
PANTHER	PTHR23036								
	PTHR23036:5F108								
Gene3D CDD	Immunoglobulin-like fold	Fibronectin type III	-						
All sequence SNPs/	Sequence variants (dbSNP and all other sound and all other sound and all other sound all other sound all other sound and all other sound all o	ces)	I.	ana apara		THE OFTEN	(1,1),(1,-1)	${}^{\rm T}{\cal H}$	• 11
Variant Legend									73
	frameshift variant			stop lost					
	missense variant			synonymous varian	t				
Scale bar	b 60 120	180	240 300	360	420	480	540		650

Source: : https://www.ensembl.org

Mouse Phenotype Information (MGI)

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• Homozygotes for targeted null mutations exhibit retarded postnatal growth, proportionate dwarfism, decreased plasma insulin-like growth factor I levels, small pituitaries, reduced fecundity in females, and extended life-span.

Source: https://www.informatics.jax.org

Important Information

- The effect of *Ghr*-205 and *Ghr*-207 gene is unknown.
- *Ghr* is located on Chr15. If the knockout mice are crossed with other mouse strains to obtain double homozygous mutant offspring, please avoid the situation that the second gene is on the same chromosome.
- This Strategy is designed based on genetic information in existing databases. Due to the complexity of biological processes, all risks of the mutation on gene transcription, RNA splicing and protein translation cannot be predicted at the existing technology level.

