

Socs2 Cas9-CKO Strategy

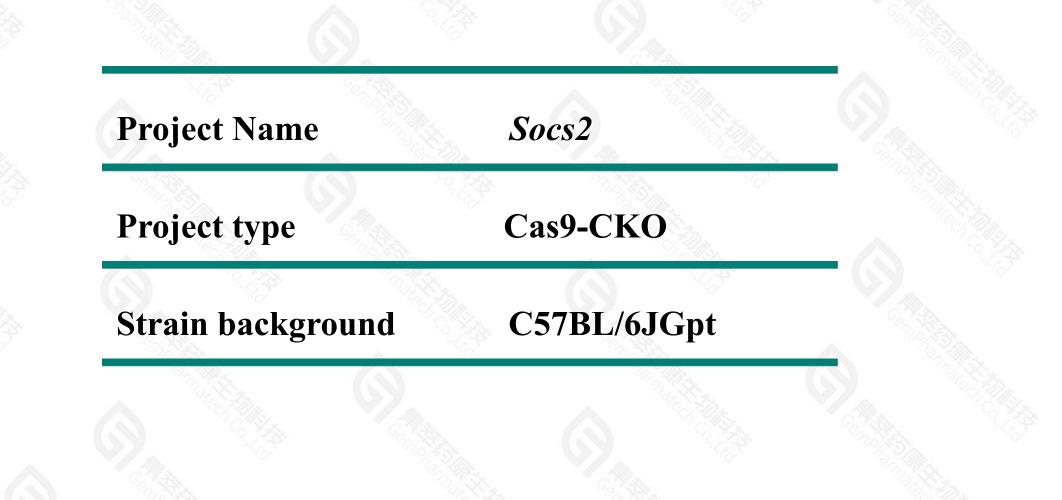
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Reviewer: Daohua Xu

Design Date: 2022-4-22

Project Overview





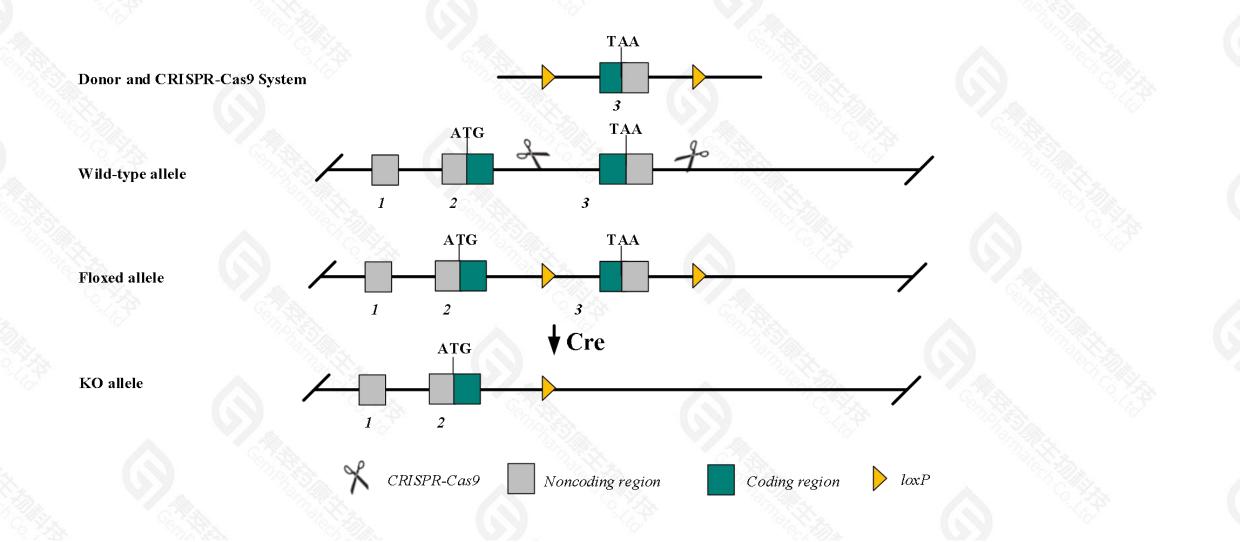
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Conditional Knockout strategy



This model will use CRISPR-Cas9 technology to edit the Socs2 gene. The schematic diagram is as follows:



Technical routes



> The *Socs2* gene has 14 transcripts. According to the structure of *Socs2* gene, exon3 of *Socs2-201*(ENSMUST00000020215.16) transcript is recommended as the knockout region. The region contains most of coding sequence. Knock out the region will result in disruption of protein function.

➤ In this project we use CRISPR-Cas9 technology to modify *Socs2* 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.

 \succ 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.



- > According to the existing MGI data, mutations in this gene cause accelerated postnatal growth. Homozygotes for a targeted mutation also show increased bone growth, enlargement of most organs, collagen deposition in the skin and some ducts and vessels, lower major urinary protein levels, and elevated IGF-I mRNA levels in some tissues.
- The KO region near to the 5730420D15Rik gene. Knockout the region may affect the function of 5730420D15Rik gene.
 The Socs2 gene is located on the Chr10. 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)



Socs2 suppressor of cytokine signaling 2 [Mus musculus (house mouse)]

± Download Datasets

Gene ID: 216233, updated on 7-Mar-2022

Summary

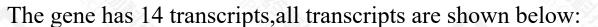
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Official Symbol	Socs2 provided by MGI
Official Full Name	suppressor of cytokine signaling 2 provided by MGI
Primary source	MGI:MGI:1201787
See related	Ensembl:ENSMUSG00000020027 AllianceGenome:MGI:1201787
Gene type	protein coding
RefSeq status	VALIDATED
Organism	Mus musculus
Lineage	Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi; Mammalia; Eutheria; Euarchontoglires; Glires; Rodentia; Myomorpha; Muroidea;
	Muridae; Murinae; Mus; Mus
Also known as	hg; JAB; CIS2; Cish2; SSI-2; SOCS-2; AI527257; AW108012; 8030460M17; D130043N08Rik
Summary	Enables growth hormone receptor binding activity. Acts upstream of or within several processes, including mammary gland development; negative
	regulation of receptor signaling pathway via JAK-STAT; and positive regulation of neuron differentiation. Predicted to be located in cytoplasm.
	Predicted to be part of phosphatidylinositol 3-kinase complex. Is expressed in several structures, including lung; primary sex cord; retina; and
	ureteric tip. Human ortholog(s) of this gene implicated in endometrial cancer and ovarian carcinoma. Orthologous to human SOCS2 (suppressor of
	cytokine signaling 2). [provided by Alliance of Genome Resources, Nov 2021]
Expression	Ubiquitous expression in ovary adult (RPKM 9.8), lung adult (RPKM 5.2) and 24 other tissues See more
Orthologs	human all

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Transcript information (Ensembl)



Transcript ID	Name 🍦	bp 🖕	Protein 🍦	Biotype 🕴	CCDS 🕴	UniProt Match	Flags
ENSMUST0000020215.16	Socs2-201	2222	<u>198aa</u>	Protein coding	CCDS24136	035717@ 054807@	Ensembl Canonical GENCODE basic APPRIS P1 TSL:1
ENSMUST00000170690.8	Socs2-213	2195	<u>198aa</u>	Protein coding	CCDS24136@	<u>035717</u> & <u>054807</u> &	GENCODE basic APPRIS P1 TSL:3
ENSMUST00000172070.8	Socs2-214	2153	<u>198aa</u>	Protein coding	CCDS24136	035717@ 054807@	GENCODE basic APPRIS P1 TSL:2
ENSMUST00000119917.2	Socs2-202	948	<u>198aa</u>	Protein coding	CCDS24136	035717@ 054807@	GENCODE basic APPRIS P1 TSL:1
NSMUST00000129942.2	Socs2-205	623	<u>62aa</u>	Protein coding	분	D3Z441@	TSL:3 CDS 3' incomplete
NSMUST00000150432.8	Socs2-211	605	<u>70aa</u>	Protein coding	×	D3Z355	TSL:2 CDS 3' incomplete
NSMUST00000135822.8	Socs2-208	497	<u>80aa</u>	Protein coding	<i>.</i>	D3YZ26	TSL:3 CDS 3' incomplete
NSMUST00000139210.8	Socs2-209	4242	<u>201aa</u>	Nonsense mediated decay	<u></u>	F6RVG2	TSL:1 CDS 5' incomplete
NSMUST00000155148.2	Socs2-212	893	No protein	Processed transcript	-	-	TSL:3
NSMUST00000134918.2	Socs2-207	874	No protein	Processed transcript	1	1.72	TSL:3
NSMUST00000128363.2	Socs2-203	502	No protein	Processed transcript	-	1	TSL:5
NSMUST00000145847.2	Socs2-210	479	No protein	Processed transcript			TSL:3
NSMUST00000128854.2	Socs2-204	352	No protein	Processed transcript	분	121	TSL:3
ENSMUST00000130784.8	Socs2-206	3285	No protein	Retained intron	*	-	TSL:1

The strategy is based on the design of *Socs2-201* transcript, the transcription is shown below:

< Socs2-201 protein coding

Reverse strand

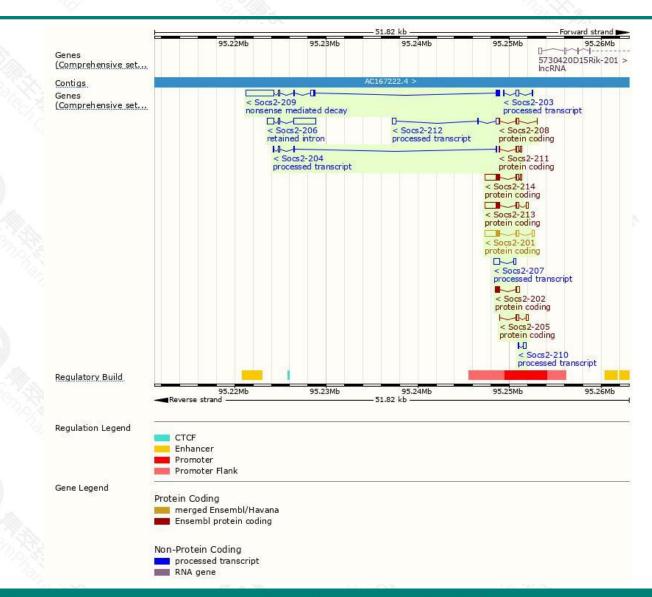
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Genomic location distribution





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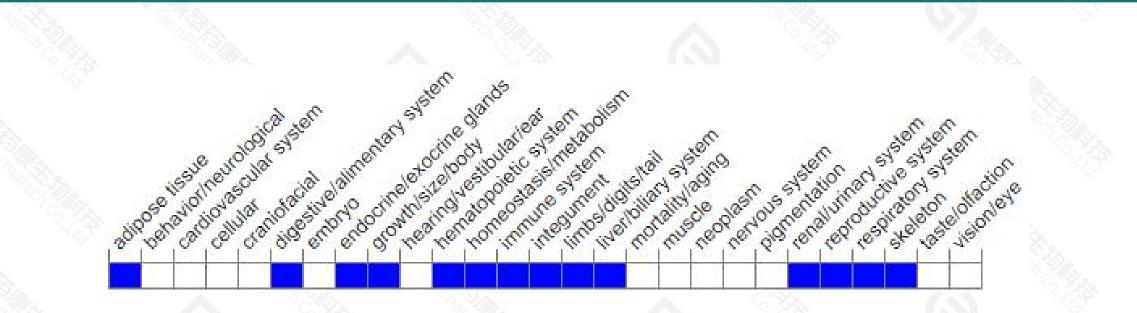
Protein domain



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53.	Scale bar	0	20	40	60	80	100	120	140	160	198	
	Variant Legend	-	synonymous	variant						412.		
	All sequence SNPs/i	Sequ	uence variants	(dbSNP and	d all other so	urces)		•	1	11	20	
	CDD.			SOCS2,	SH2 domain					cd03736	-	
	Gene3D		and contraction of the second s	SH2 domain s	superfamily					1,10,750,20		
	PANTHER		ressor of cytoki 10155	ine signalling	2						_	
	PROSITE profiles			s	12 domain					OCS box domain		
	Pfam.			and the second se	12 domain					SOCS box do	main	
	Prints			SH	12 domain	_				SOCS box dor	main	
	SMART			SHE	2 domain				-	SOCS box domain		
	Low complexity (Seg) Superfamily			SH2 doma	ain superfamil	у.			s	OCS box-like doma	in superfamil	
	ENSMUSP00000020 MobiDB lite	-	-									

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, mutations in this gene cause accelerated postnatal growth. Homozygotes for a targeted mutation also show increased bone growth, enlargement of most organs, collagen deposition in the skin and some ducts and vessels, lower major urinary protein levels, and elevated IGF-I mRNA levels in some tissues.

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



