

C57BL/6N-Tg(Ins2-Chrm3*)SJwe/J

Stock No: **028966** | β -GsD Tg (or β -R-s Tg)

 Coisogenic, Transgenic

Typically mice are recovered in 10-14 weeks. Contact Customer Service to place an order or for more information.

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following administration of the pharmacologically inert molecule clozapine-*N*-oxide (CNO). These mice may be useful for chemogenetic/pharmacogenetic applications to express an activating DREADD for studying G protein regulation of β -cell function.

Donating Investigator

Jürgen Wess, National Institute of Diabetes and Digestive and Kidney Diseases (NIH-NIDDK)

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GENETIC OVERVIEW

Genetic Background

Generation

Tg(Ins2-Chrm3*)SJwe

Alele Type

Transgenic (Inserted expressed sequence)

VIEW GENETICS

RESEARCH APPLICATIONS

Neurobiology Research

Research Tools

Immunology, Inflammation and Autoimmunity Research

Diabetes and Obesity Research

Cancer Research

Cell Biology Research

BASE PRICE

Starting at:

\$2,854.50 Domestic price Cryo Recovery

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Details

Detailed Description

The β -GsD (or β -R-s) transgene has the rat insulin 2 promoter directing expression of a hemagglutinin epitope-tagged, mutant G protein-coupled receptor (GsD [also called rM3Ds or R-s]) to pancreatic β -cells. The GsD is a G_s -coupled rat muscarinic 3 DREADD modified to have its second and third intracellular loops replaced with the corresponding turkey β 1-adrenergic receptor sequences. Two amino acid substitutions (Y148C and A238G) have been added that abolish receptor affinity for the native ligand, acetylcholine (ACh), but allow receptor binding and subsequent activation by the small pharmacologically inert molecule clozapine-*N*-oxide (CNO). CNO binding to GsD induces the selective activation of the canonical $G\alpha_s$ signaling pathway, effectively increasing intracellular cAMP levels.

Mice that are hemizygous for the transgene (β -GsD Tg or β -R-s Tg) are viable and fertile. The donating investigator reports they have not tried to generate homozygous mice (May 2016). The phenotype described below compares β -GsD Tg with a similarly created transgenic mouse that expresses a rat insulin 2 promoter-driven rhM3Dq (β -rhM3Dq Tg ; Stock No. [028964](#)).

The transgenic mouse lines β -rhM3Dq Tg (Stock No. 028964) and β -GsD Tg (Stock No. 028966) express similar numbers of rhM3Dq and GsD in their pancreatic islets, respectively. RT-PCR studies confirmed that rhM3Dq and GsD transcripts were not detectable in tissues other than pancreatic islets.

In the absence of CNO, both transgenic lines have normal growth, insulin sensitivity, plasma insulin levels and glucagon levels. While freely-fed β -rhM3Dq Tg have normal blood glucose levels, freely-fed β -GsD Tg have decreased blood glucose levels. Taken together, this indicates the rhM3Dq receptor is devoid of ligand-independent signaling *in vivo*, whereas the GsD receptor retains some degree of agonist-independent signaling *in vivo*.

CNO treatment of transgenic mice leads to the activation of β -cell $G_{q/11}$ or $G\alpha_s$ signaling pathways, respectively, in a conditional and β -cell-selective fashion. Specifically, following CNO administration in β -rhM3Dq Tg, the canonical $G_{q/11}$ pathway is activated in rhM3Dq-expressing pancreatic β -cells. This leads to significant increases in both first- and second-phase insulin release, greatly improved glucose tolerance in obese, insulin-resistant mice, and elevated β -cell mass, associated with pathway-specific alterations in islet gene expression levels. For β -GsD Tg, CNO-induced activation of the canonical $G\alpha_s$ pathway in GsD-expressing pancreatic β -cells results in qualitatively similar *in vivo* metabolic effects.

Development

Expression Data

[+ Control Suggestions](#)

[+ Selected References](#)

[- Genetics](#)

[+ Tg\(Ins2-Chrm3*\)SJwe](#)

[- Disease/Phenotype](#)

[+ Disease Terms](#)

[+ Research Areas By Phenotype](#)

[+ Mammalian Phenotype Terms by Genotype](#)

[+ References](#)

[- Technical Support](#)

C O N T A C T T E C H N I C A L S U P P O R T

Genotyping Protocols

Standard PCR: [Tg\(Ins2-Chrm3*\)SJwe](#)

[Genotyping resources and troubleshooting](#)

Breeding Considerations

When maintaining a live colony, hemizygous mice may be bred together, to wildtype (noncarrier) mice from the colony, or to C57BL/6NJ inbred mice (Stock No. [005304](#)). To date (May 2016), it has not been attempted to make this strain homozygous.

[Additional Breeding and Husbandry Support](#)

Mating System

Noncarrier x Hemizygote

Hemizygote x Noncarrier

Citation

When using the β -GSD Tg (or β -R-s Tg) mouse strain in a publication, please [cite the originating article\(s\)](#) and include JAX stock #028966 in your Materials and Methods section.

Animal Health Reports

[Facility Barrier Level Descriptions](#)

Production of mice from cryopreserved embryos or sperm occurs in a maximum barrier room, [G200](#)

🔻 Pricing & Availability



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Cryorecovery - Domestic Pricing

SERVICE/PRODUCT	DESCRIPTION	PRICE
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Related Products and Services

Frozen Mouse Embryo	C57BL/6N-Tg(Ins2-Chrm3*)SJwe/J Frozen Embryo	\$2595.00
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