Ai79(TITL-Jaws)-D (also called Ai79D) mice are a Cre/Tet-dependent, optogenetic line, created by targeted insertion at the Igs7 locus (TIGRE; an intergenic region on mouse chromosome 9 that allows reporter expression to be tightly regulated). Following Cre-mediated removal of the STOP cassette, they may be used to generate Tet-Off/Tet-On mutant animals with conditional (inducible/reversible) expression of an improved Halorhodopsin/EGFP fusion protein (Halo57*K200R*W214F/EGFP or Jaws). Subsequent illumination of Jaws-expressing (EGFP fluorescent) cells with yellow-to-red light leads to reversible photoinhibition of action potential firing/neural activity in these cells.

Donating Investigator

Hongkui Zeng, Allen Institute for Brain Science
Ai79(TITL-Jaws)-D mice are a Cre/Tet-dependent, optogenetic line, created by targeted insertion at the Igs7 locus (TIGRE; an intergenic region on mouse chromosome 9 that allows reporter expression to be tightly regulated). Ai79D mice harbor the TIGRE-Ins-TRE-LSL-Jaws conditional allele, designed with a modified Tet response element (TRE or tetO) and loxP-flanked STOP cassette upstream of an improved Halorhodopsin/EGFP fusion protein (Jaws; see detailed description below). When bred with other mice expressing Cre recombinase, tetracycline-controlled transactivator protein (tTA) and/or reverse tetracycline-controlled transactivator protein (rtTA), Jaws expression in cells/tissues where the expression patterns of the individual promoters driving Cre and tTA/rtTA overlap can be regulated with tetracycline or its analog doxycycline (dox). Following induction of Jaws expression (EGFP fluorescence), illuminating Jaws-expressing neurons with yellow-to-red light leads to reversible photoinhibition of action potential firing/neural activity in these cells.

Specifically, the donating investigator reports Ai79D mice have no reported levels of EGFP fluorescence in absence of Cre and tTA. When Ai79D mice are bred with tTA-driver and Cre-driver lines to create triple transgenic animals (Jaws+/tTA+/Cre+), cells expressing both tTA and Cre exhibit robust fluorescence. Light-induced expression of the inhibitory opsin is expected to occur at levels sufficient to effectively silence the activity of cortical neurons. Jaws expression/function in tissues other than brain has not yet been evaluated by the donating investigator (March 2014).

Heterozygous Ai79D mice are viable and fertile with no reported gross physical or behavioral abnormalities. As of April 2017, it is the experience of The Jackson Laboratory that homozygous Ai79D mice are viable and fertile.

The bacterial opsins are retinal-binding proteins that combine a light-sensitive domain with an ion channel or pump; providing light-dependent ion transport, membrane potential alteration, and sensory functions to bacteria. The Halobacterium halobium (strain shark)-derived halorhodopsin Halo57 is an inward chloride ion pump that causes hyperpolarization and prevents action potentials. Halo57 allows robust photocurrents/action potential inhibition when exposed to light ranging from yellow to red (~540-640 nm, peak ~600 nm), with function out to the hemoglobin absorption trough range (~660 nm). Halo57 is significantly red-light shifted relative to other opsins; displaying larger red-light induced photocurrents and less aggregation than NpHR.

The Jaws fusion protein (Halo57*K200R*W214F/EGFP) is composed of a modified Halo57 fused in-frame to EGFP. The Halo57 sequence has K200R and W214F mutations to substantially boost photocurrents. An endoplasmic reticulum exporting sequence (ER2) from the Kir2.1 inward rectifier potassium channel gene at the C-terminal of EGFP improves membrane trafficking/prevents aggregation. The Halo57*K200R*W214F variant in Jaws exhibits photocurrents driven by red light that are over three times greater than those of previous neural silencing opsins, and illumination of Jaws-expressing neurons leads to reversible photoinhibition (silencing) of action potential firing/neural activity in these cells in the awake mammalian brain with millisecond latency.
Genotyping Protocols
Standard PCR: Igs7
Genotyping resources and troubleshooting

Breeding Considerations
When maintaining a live colony, heterozygous mice may be bred to wildtype mice from the colony or to C57BL/6J inbred mice (Stock No. 000664). As of April 2017, it is the experience of The Jackson Laboratory that homozygous Ai79D mice are viable and fertile. Therefore, this colony may also be maintained by breeding homozygous mice together.

Additional Breeding and Husbandry Support
Citation
When using the Ai79(TITL-Jaws)-D or Ai79D mouse strain in a publication, please cite the originating article(s) and include JAX stock #023529 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

מערכת העדacles בבריעה

Pricing & Availability

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

CRYORECOVERY - DOMESTIC NOT-FOR-PROFIT & ACADEMIC PRICING

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<th>SERVICE/PRODUCT</th>
<th>DESCRIPTION</th>
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RELATED PRODUCTS AND SERVICES

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<td>Frozen Mouse Embryo</td>
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PAYMENT TERMS AND CONDITIONS

Terms are granted by individual review and stated on the customer invoice(s) and account statement. These transactions are payable in U.S. currency within the granted terms. Payment for services, products, shipping containers, and shipping costs that are rendered are expected within the payment terms indicated on the invoice or stated by contract. Invoices and account balances in arrears of stated terms may result in The Jackson Laboratory pursuing collection activities including but not limited to outside agencies and court filings.

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The Jackson Laboratory has rigorous genetic quality control and mutant gene genotyping programs to ensure the genetic background of JAX® Mice strains as well as the genotypes of strains with identified molecular mutations. JAX® Mice strains are only made available to researchers after meeting our standards. However, the phenotype of each strain may not be fully characterized and/or captured in the strain data sheets. Therefore, we cannot guarantee a strain’s phenotype will meet all expectations. To ensure that JAX® Mice will meet the needs of individual research projects or when requesting a strain that is new to your research, we suggest ordering and performing tests on a small number of mice to determine suitability for your particular project. We do not guarantee breeding performance and therefore suggest that investigators order more than one breeding pair to avoid delays in their research.
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Related Strains

All

By Allele

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Leading the search for
TOMORROW'S CURES

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