

**B6;129-Raf1<sup>tm2Ara</sup>/J**

Stock No: **021233**

 Targeted Mutation

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

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disorder Noonan syndrome (NS).

### Donating Investigator

Benjamin Neel, NYU School of Medicine

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## GENETIC OVERVIEW

Genetic Background

Generation

*Raf1<sup>tm2Ara</sup>*

**Alele Type**

Targeted

**Gene Symbol**

*Raf1*

**Gene Name**

v-raf-leukemia viral oncogene 1

VIEW GENETICS

## RESEARCH APPLICATIONS

Developmental Biology Research

Cardiovascular Research

Research Tools

VIEW ALL RESEARCH APPLICATIONS

## BASE PRICE

Starting at:

\$2,854.50 Domestic price Cryo Recovery

VIEW PRICE LIST

### Details

#### Detailed Description

Mice homozygous for the *Raf1*<sup>loxP/loxP</sup> allele are viable and fertile. This allele contains a floxed sequence containing a cDNA fragment encoding v-raf-leukemia viral oncogene 1 (*Raf1*) wild-type exons 13 to 16 and a neomycin (neo) resistance cassette downstream of exon 12 of the *Raf1* gene. A point mutation was introduced downstream of the floxed sequence, resulting in the missense mutation D486N, associated with the autosomal dominant disorder Noonan syndrome (NS). *Raf1* is a MAP kinase kinase kinase (MAP3K), which is activated by Ras GTPases, which in turn activates MEK1/2 and extracellular signal-regulated kinase (ERK)1/2 pathways. Activated ERKs are involved in the cell division cycle, apoptosis, cell differentiation and cell migration. The D486N, kinase-impaired, mutation is associated with the autosomal dominant disorder Noonan syndrome (NS) characterized by short stature, facial dysmorphia, cardiovascular abnormalities, and myeloproliferative disease (MPD). Homozygous mice are viable and fertile. When these mutant mice are bred to mice that express Cre recombinase, resulting offspring will have the floxed sequence deleted in cre-expressing tissues.

For example, when bred to mice expressing germ line *cre* (see B6.FVB-Tg(Ella-cre)C5379Lmgd/J Stock No. [003724](#)), removal of the floxed sequence permits expression of mutant *Raf1*<sup>D486N</sup>. Heterozygous D486N/+ females exhibit a mild growth defect. One-third of homozygous mice had body length and weight only 50% that of littermate controls. A majority of these smaller mice (s-D486N/D486N) die shortly after weaning, while survivors only live 4 and 8 months. The longer-lived s-D486N/D486N mice exhibit reduced body size, a hunched appearance with ruffled fur, and frequent tremors. These mice have a slight decrease in skull width resulting in a "triangular" facial appearance. The mice with normal length and weight (n-D486N/D486N) display an increase in heart weight/body weight ratio with an increase also seen in the left ventricular diastolic posterior wall thickness. They exhibit a decrease in left ventricular internal endsystolic dimension, and an increase in stroke volume, fractional shortening, cardiac output, and ejection fraction. *Raf1*<sup>D486N</sup>-expressing cells show enhanced ERK/MAPK pathway activation and increased heterodimerization with BRAF.

#### Development

#### Control Suggestions

#### Selected References

### Genetics

#### *Raf1*<sup>tm2Ara</sup>

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## – Disease/Phenotype

+ Disease Terms

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+ Research Areas By Phenotype

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+ Mammalian Phenotype Terms by Genotype

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+ References

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## – 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

End Point Analysis:[Raf1 End Point](#)

[Genotyping resources and troubleshooting](#)

### Breeding Considerations

When maintaining a live colony, homozygous mice may be bred together.

[Additional Breeding and Husbandry Support](#)

### Citation

When using the B6;129-Raf1<sup>tm2Ara</sup>/J mouse strain in a publication, please [cite the originating article\(s\)](#) and include JAX stock #021233 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](#)*

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## – Pricing & Availability



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## Domestic International

Pricing effective for USA, Canada and Mexico shipping destinations

### CRYORECOVERY - DOMESTIC PRICING

SERVICE/PRODUCT	DESCRIPTION	PRICE
<a href="#">Cryo Recovery</a>	Heterozygous or wildtype for Raf1<tm2Ara>	\$2,854.50

### RELATED PRODUCTS AND SERVICES

<a href="#">Frozen Mouse Embryo</a>	B6;129-Raf1<tm2Ara>/J	\$2595.00
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## PAYMENT TERMS AND CONDITIONS

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