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Novel Mouse Lines for Study of Cancer, Wound Healing and Stem Cell Biology

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BACKGROUND

The Dact2 protein may play an important role in oncogenesis, metastasis, wound healing and stem cell biology. Molecular signaling pathways potentially involving Dact2 include the Wnt/β-catenin pathway, the Wnt/PCP non-β-catenin-dependent pathway, regulation of small GTPases of the Rho family, and pathways involving the Dishevelled signal transduction molecule, such as p120-catenin signaling. There is evidence that Dact2 also regulates the TGF-β pathway. In vitro and in vivo knockdown models would be very useful for studying Dact2 function in disease, but have not been reported to date.

TECHNOLOGY DESCRIPTION

UCSF investigators have generated Dact2 conditional, constitutive and partial knockout mouse lines by making a targeted mutation in the genetic locus corresponding to Dact2, a conserved homolog of the Dapper/Frodo genes originally described in *Xenopus laevis*. This mutation was designed such that the essential exon 2 of the gene is flanked by loxP sites, which allows for its excision by the Cre recombinase protein to render the allele non-functional. ES cells carrying this mutation were made by homologous recombination, and a mouse line derived from the correctly targeted ES cells was created using embryo manipulation techniques. Subsequently, through genetic crosses to a few transgenic lines of mice, several different mouse lines have been created and characterized, including a neo- version of the conditional allele, both neo+ and neo- versions of a constitutive knock-out allele, and a hypomorphic (partial knockout) allele. This invention is a valuable model that enables study of Dact2 gene's roles during development as well as postnatally by making use of Cre and Flipase to partially or fully eliminate the gene in primary tissue culture cell lines, immortalized cell lines or in mouse organs and tissues at specific developmental times by crossing to Cre and/or Flipase-expressing transgenic animals.

APPLICATIONS

- Study Dact2's role in cancer etiology, pathogenesis, metastasis and invasiveness
- Study Dact2's role in planar cell polarity pathways involved in wound healing and stem cell biology

CONTACT Sunita R. Rajdev sunita.rajdev@ucs

sunita.rajdev@ucsf.edu tel: 415-340-2476.



OTHER INFORMATION

KEYWORDS

Dact2

CATEGORIZED AS

Medical

- Disease: Cancer
- Research Tools
- Stem Cell

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ADDRESS

UCSF

Innovation Ventures

600 16th St, Genentech Hall, S-272,

San Francisco,CA 94158

CONTACT

Tel:

innovation@ucsf.edu

https://innovation.ucsf.edu

Fax:

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