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Manuscript Type: *Basic Study*

Homologous recombination mediates stable Fah gene integration and phenotypic correction in tyrosinaemia mouse-model

Junge N *et al* Phenotypic correction in tyrosinaemia mouse-model

Norman Junge, Qinggong Yuan, Thu Huong Vu, Simon Krooss, Christien Bednarski, Asha Balakrishnan, Toni Cathomen, Michael P. Manns, Ulrich Baumann, Amar Deep Sharma and Michael Ott

Abstract

AIM

To stably correct tyrosinaemia in proliferating livers of fumarylacetoacetate-hydrolase knockout (*Fah*^{-/-}) mice by homologous-recombination-mediated targeted

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So far there has been no report of any clinical or preclinical evidence for chromosomal vector integration following adenovirus (Ad) vector-mediated gene transfer in vivo. ... We used the FAH^{Δexon5} mouse model (8) of tyrosinemia type I (MIM 27670) to analyze potential homologous and heterologous recombination events ...

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PK Held, EC Olivares, CP Aguilar, M Finegold... - Molecular Therapy, 2005 - Elsevier

... integration of plasmid DNA into host cell genomes, including mammalian cells [8]. In nature, it mediates a unidirectional ... the middle of the site, then the pseudo attP site near p21 has 37% homology to the ... In addition, the recombination is unidirectional and integration is stable ...

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We chose to study DNA transposition in a mouse model of hereditary tyrosinemia type 1 (HT1) [19], because of its unique advantages for the characterization of stable integration events in hepatocytes in vivo. HT1 is a lethal recessive liver disease caused by deficiency of fumarylacetoacetate hydrolase (FAH) [19], the ...

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