## Chromosome engineering of a mouse reagent for functional-genomics analyses of human Chromosome

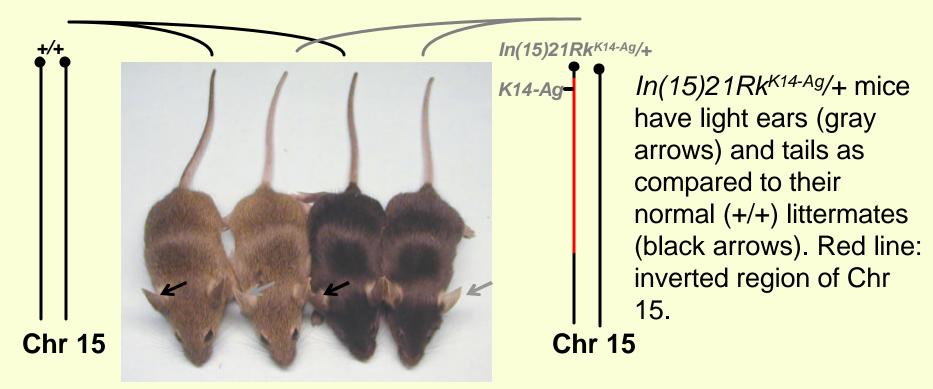
**5p** [Contact: Yun You (youy@ornl.gov); funding source DOE-OBER]

<u>Purpose</u>: To modify existing mouse chromosomal inversions so that they may be used for easy recovery and maintenance of all types of mouse mutations

Accomplishment: Dr. Wallace Chick (Postdoctoral fellow in Dr. Yun You's laboratory at ORNL) successfully constructed a new inversion reagent for the ORNL human Chromosome 5p functional-genomics effort

- Embryonic stem-cell lines were derived from the known In(15)21Rk inversion [In(15)21Rk/+ mice are normal, and In(15)21Rk/In(15)21Rk mice die prenatally].
- Gene-targeting techniques were used to insert the K14-Ag gene, conferring dominant yellow color, into the inverted chromosome in the ES cells.
- In(15)21Rk<sup>K14-Ag</sup>/+ mice generated from the targeted ES cells have lighter ears and tails, now allowing the use of this inversion in mutation recovery and maintenance crosses.





- Mice derived from the targeted ES cells manifest an easily visible phenotype, allowing them to be used as a genetic reagent for the mouse genetics community.
- This modified inversion spans the proximal segment of mouse Chr 15, which is homologous to human Chr 5p, a region of the DOE interest.
- This appropriately marked inversion can now be used for easy, lowcost maintenance of detrimental 5p-homologous mutations derived from either gene- or phenotype-driven mutagenesis screens.

