Partial Genetic Humanization

Strategy Examples Point mutations, single exon replacement

Advantages ■ Ge mous

Gene retains mouse's wild-type expression pattern

Random insertion is typically the lowestcost option

Humanization

by Transgenic

Knockin

Insertion of

human genomic or

coding sequence into

mouse genome

Drawbacks

Limited design does not model all aspects of homologous human gene

Proper gene regulation may not be maintained

May require knockout of mouse gene and tracking of 2 separate alleles TruHumanization™ Complete Gene Replacement

Entire mouse gene replaced including introns and potentially promoter and UTRs

Gene controlled by human regulatory elements

Most faithful model for study of human gene

Requires high degree of technical skill to create

> ingenious targeting laboratory Custom mouse, rat, and rabbit models