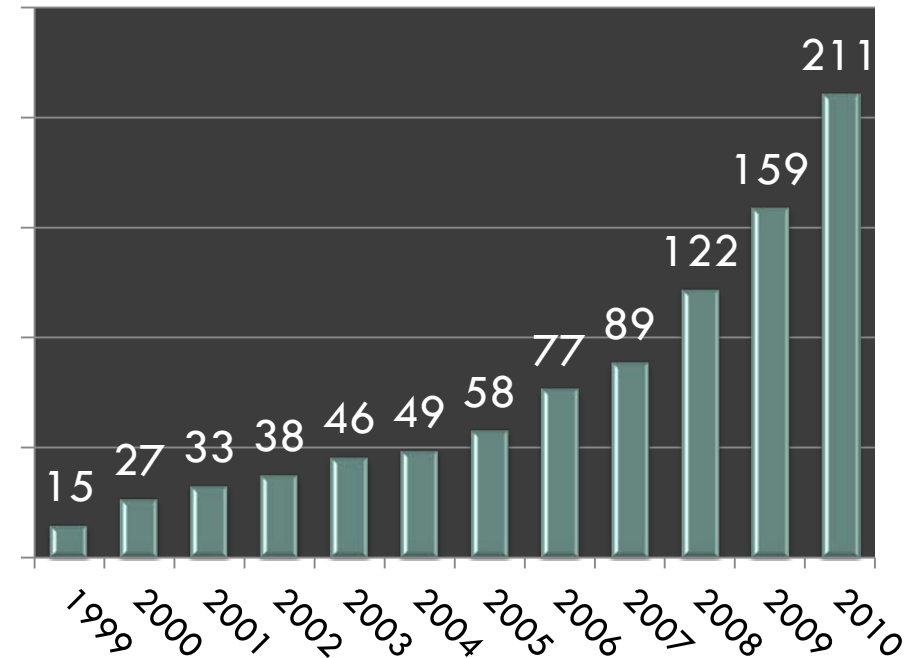
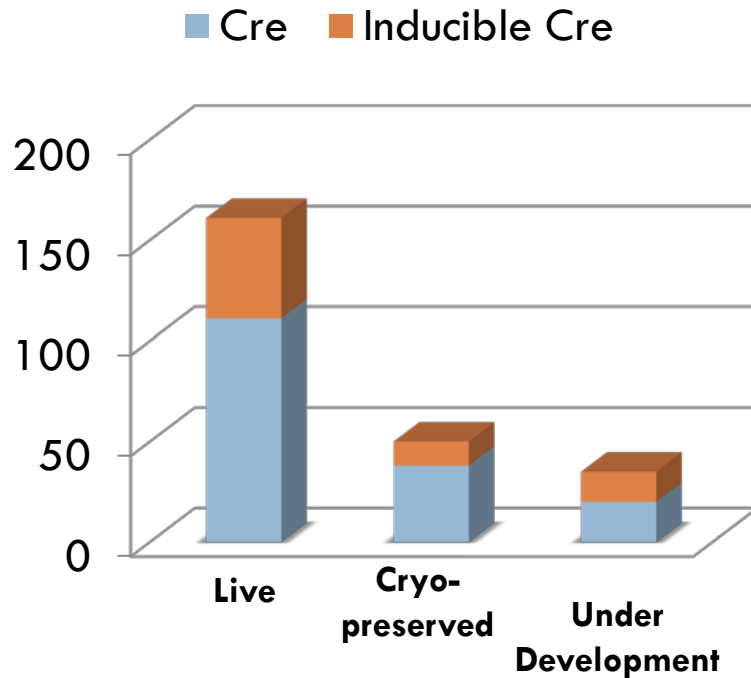


# Leveraging conditional IKMC alleles: Cre driver strain resources

- Enabling access: critical role for Repositories
  - ▣ Live versus cryo
  - ▣ Quality control: transfer of responsibility
  - ▣ Directed versus “laissez faire” promotion strategies
- Detailed characterization
  - ▣ Add to and complement published data
  - ▣ Particularly important for Cres developed in large sets
  - ▣ Positive AND negative data
- Data dissemination
  - ▣ Centralized sources of Cre functional data
  - ▣ CREATE BioMART and CrePortal

# Cre strain distribution

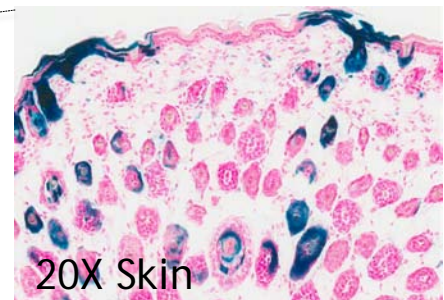
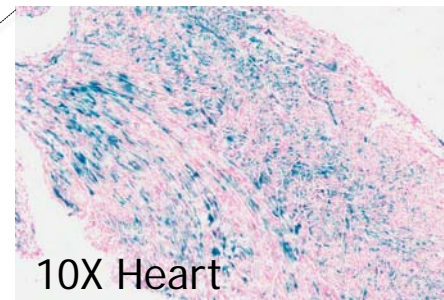
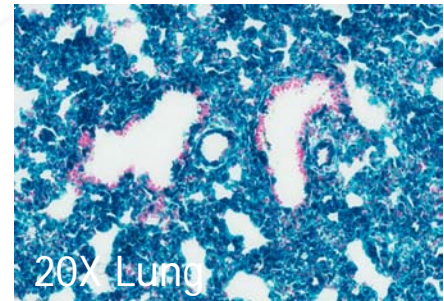
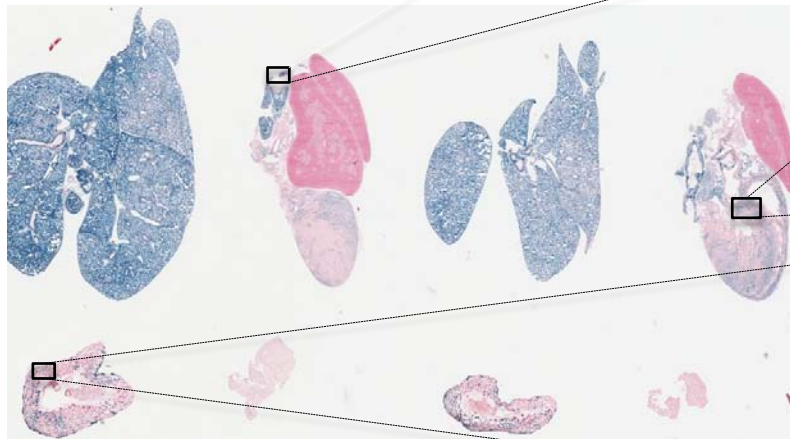


Growth increasingly driven by large sets (e.g. Neuroscience Blueprint) that have limited community use prior to distribution.

Open question: how do we best manage our Cre strain distribution to best serve the community?

# Comprehensive characterization of Cre driver pipeline (cre.jax.org)

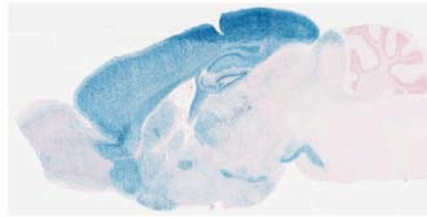
- R26R-LacZ Reporter strain
- Four time points: E10.5, E15.5, P7 and P56 (adult)
- Whole-mount (E10.5) or frozen sections
- Full necropsy at P7 and P56 (11 organ systems, 30 organs/structures, 89 substructures )
- NanoZoomer slide scanner to capture data at 20X
- Data collected for 45 strains, 44 in progress
- Data delivered to creportal.org



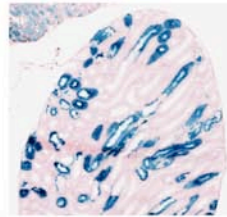
**129(Cg)-Foxg1<sup>tm1(cre)Skh</sup>/J** Characterized with R26R LacZ Reporter

# Off-target Cre activity: a surprisingly common problem

*Camk2a-cre*

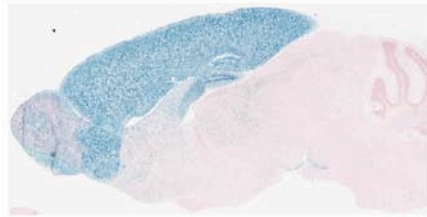


Brain

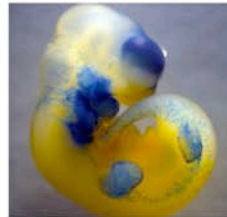


Testis

*Emx1-cre*



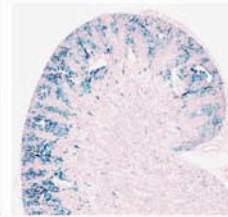
Brain



Embryo (E10.5)

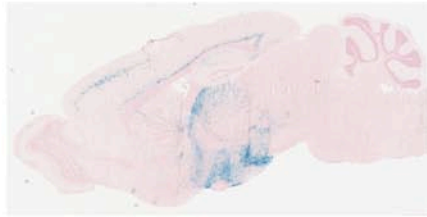


Embryo (E15.5)

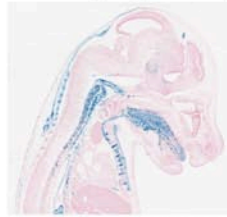


Kidney

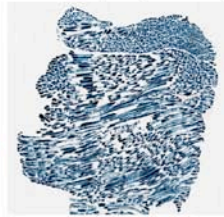
*Sim1-cre*



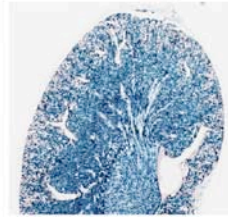
Brain



Embryo (E15.5)

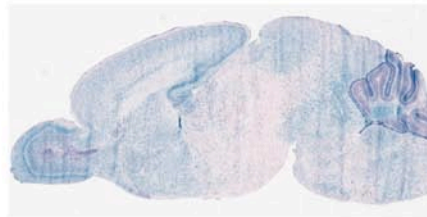


Muscle

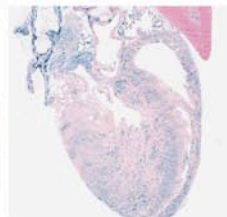


Kidney

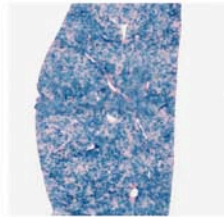
*Foxg1-cre*



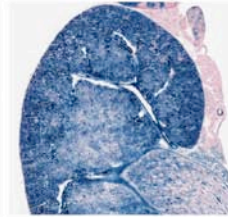
Brain



Heart



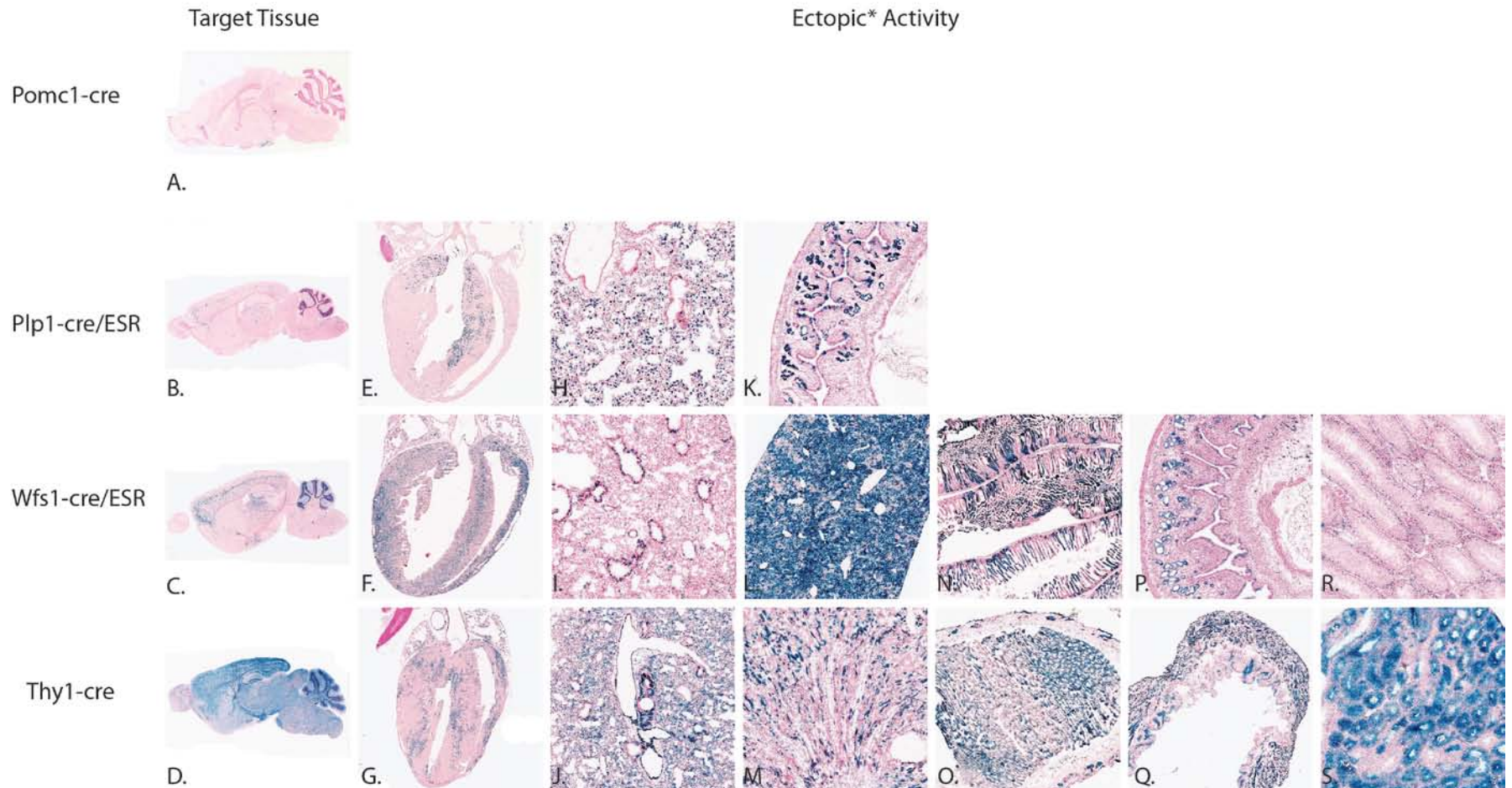
Liver



Kidney



# Ectopic\* activity continued

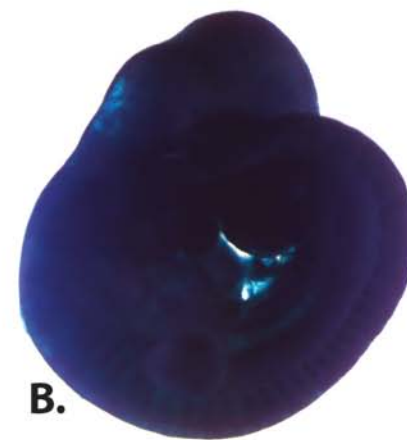
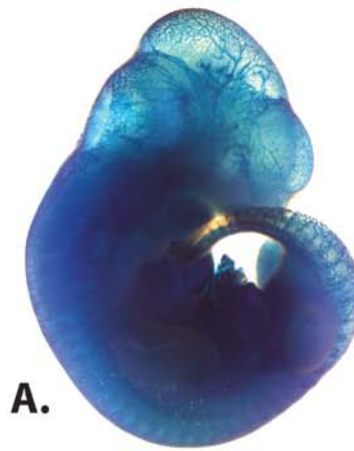


# Inconsistent activity

Regionally Restricted

Widespread

B6.Cg-Tg(Tek-cre)12Flv/J



B6.Cg-Tg(Vav1-cre)A2Kio/J



# Recombinase (cre) Home (www.creportal.org)

## Recombinase (cre) Specificity

MGI collects and annotates expression and activity data for recombinase-containing transgenes and knock-in alleles.

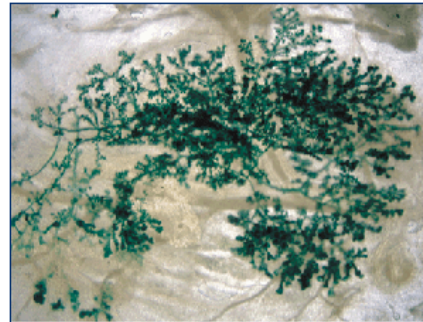


Image reproduced with permission of the Journal of Cell Science.  
Rijnkels M and Rosen, JM, J Cell Sci 2001, 114(17): 3147-3153.

### Access Data

#### FIND RECOMBINASE-CARRYING ALLELES

Search for alleles assayed for specificity/expression in a particular anatomical system.

Recombinase specificity in: (choose one)

Search for alleles by promoter/driver specificity.

Recombinase driven by: (choose one)

#### RETRIEVE ALL ALLELES

Retrieve a list of all recombinase-containing transgenes and knock-in alleles.

[MGI Recombinase Alleles Report](#)

[Tab-delimited version](#)

#### RELATED

Search  
Alleles

Search

Search for cre activity by anatomical system or by driver/promoter

Get a report of all recombinase knock-ins and transgenes

1,516 total Cre driver strains

~50% fully curated for Cre activity

### FAQs

How do I...

.. find existing recombinase-expressing transgenes and knock-ins that have a given promoter (driver)? [FAQ](#)

ie? [FAQ](#)

12 Mar 2010

October 29, 2009

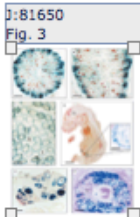
curated from the scientific literature and integrated from projects underway at collaborating organizations. (See the **Collaborators** tab, below).  
Alleles) consortium.



# Tg(AMH-cre)1Flor - Reproductive System

Recombinase Specificity Detail

[Allele Information](#) | [Tissue Information](#) | [Images](#) | [Recombinase Specificity](#) | [References](#)

Allele Information	Allele: <b>Tg(AMH-cre)1Flor</b> transgene insertion 1, Florian Guillou  Synonym: AMH-  Molecular description: The t show  Find mice (IMSR): Mous  Additional Tissues: Tg(A alim	Driver: AMH Type: Transgenic (Cre/Flp)	<ul style="list-style-type: none"><li>• Molecular description of the transgene or knock-in</li><li>• Links to mice in IMSR</li><li>• Other tissues with cre activity</li></ul>															
Tissue Information	Reproductive System	Other ▶ Alp	em tissues: mi(cre)Lowl ... (more)															
Images	Drag images to compare to others or J:81650 Fig. 3 																	
Recombinase Specificity	Click heading to resort table. ⓘ <table><thead><tr><th>Structure</th><th>Assayed</th></tr></thead><tbody><tr><td>testis</td><td>embryon 15</td></tr><tr><td>seminiferous tubule</td><td>postnatal week 9</td></tr></tbody></table> <table><thead><tr><th>Assays</th><th>Genotypic Background</th><th>Result Notes</th></tr></thead><tbody><tr><td>Recombinase reporter</td><td>lacZ</td><td>Direct Detection</td></tr><tr><td>Recombinase reporter</td><td>lacZ</td><td>Direct Detection</td></tr></tbody></table>			Structure	Assayed	testis	embryon 15	seminiferous tubule	postnatal week 9	Assays	Genotypic Background	Result Notes	Recombinase reporter	lacZ	Direct Detection	Recombinase reporter	lacZ	Direct Detection
Structure	Assayed																	
testis	embryon 15																	
seminiferous tubule	postnatal week 9																	
Assays	Genotypic Background	Result Notes																
Recombinase reporter	lacZ	Direct Detection																
Recombinase reporter	lacZ	Direct Detection																
References	All for this allele: 34 reference(s)																	

Recombinase activity data

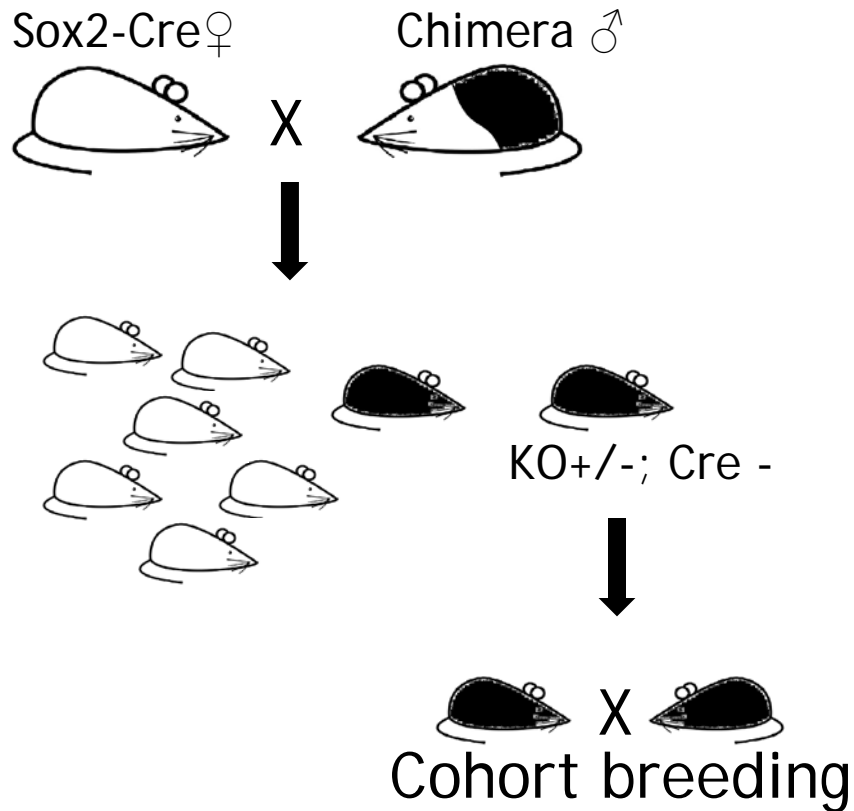
Link to References



# Cre strains for KOMP2, IMPC

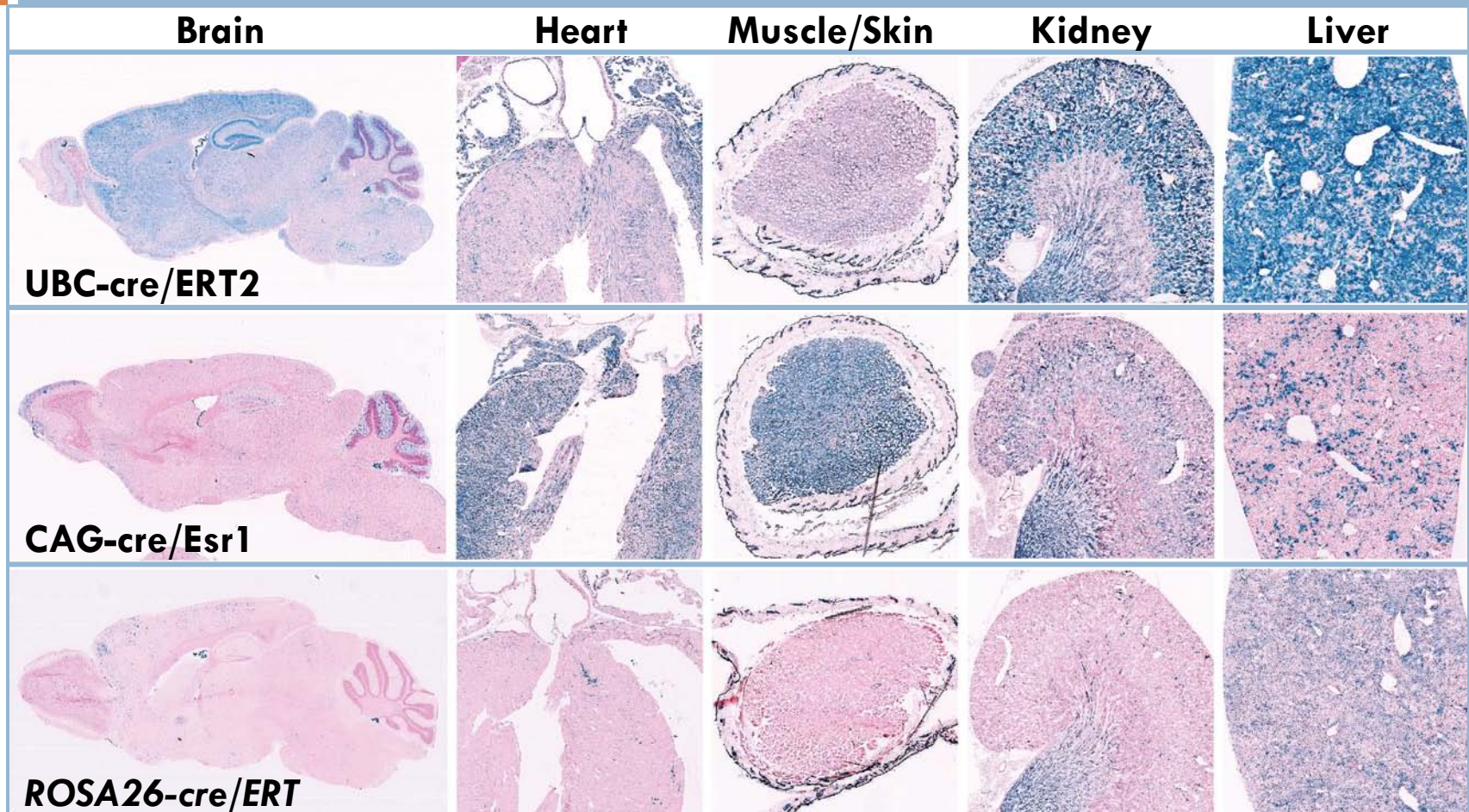
- Cre deleter
  - ▣ C57BL/6N background
  - ▣ Highly efficient
  - ▣ Maternal deletion 'feature' (Sox2-Cre; ICS Cre-GFP)
- FLPe germline deleter strains on B6N needed to convert to conditional alleles
- Inducible Cre (ERT2) and tissue specific Cre strains for use with lethals
- Need to move well-characterized, well-published Cres to B6N

# JAX Cre deleter choice: Sox2-Cre



- Deletion activity independent of Cre allele segregation
- ▣ Speed congenic to C57BL/6N using new N/J SNP panel

# Comparison of ubiquitous CreER lines



Thanks to...

## Cre Characterization Pipeline

Caleb Heffner

Herb Pratt

## JAX Repository

Leah Rae Donahue

Cat Lutz

Steve Rockwood

Mike Sasner

Supported by ARRA RR026117

RR032656

## CrePortal

Randal Babiuk

Kim Forthofer

Peter Frost

Martin Ringwald

James Kadin

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and NIH grants HD062499 and RR032656