

Potential game changers: a spotlight on selected emerging technologies.

Gene drives and RNAi applications for rodent eradication.

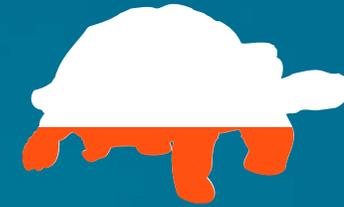


We can do this...

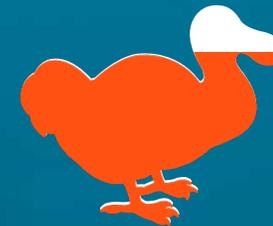
# Islands Represent



Less than **5%** of land mass



**40%** of endangered species

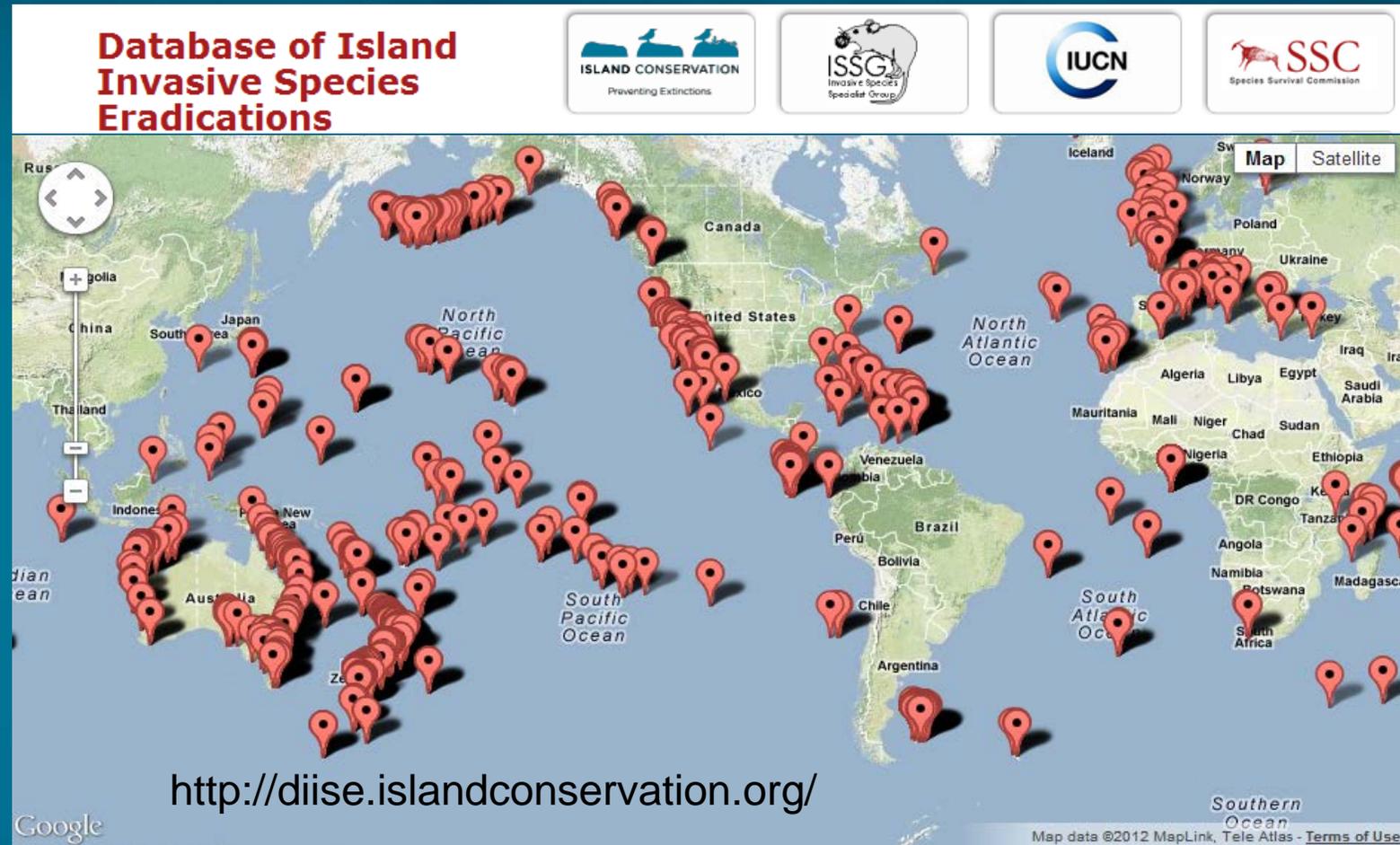


**80%** of extinctions since 1500

We can do this...

# Invasive alien mammal eradications

- 1100 successful
- Result in native species recovery



## Invasive mammal eradication on islands results in substantial conservation gains

Holly P. Jones<sup>a,b,1</sup>, Nick D. Holmes<sup>c</sup>, Stuart H. M. Butchart<sup>d</sup>, Bernie R. Tershy<sup>e</sup>, Peter J. Kappes<sup>f</sup>, Ilse Corkery<sup>g</sup>, Alfonso Aguirre-Muñoz<sup>h</sup>, Doug P. Armstrong<sup>i</sup>, Elsa Bonnaud<sup>j</sup>, Andrew A. Burbidge<sup>k</sup>, Karl Campbell<sup>c,l</sup>, Franck Courchamp<sup>j</sup>, Philip E. Cowan<sup>m</sup>, Richard J. Cuthbert<sup>n,o</sup>, Steve Ebbert<sup>p</sup>, Piero Genovesi<sup>q,r</sup>, Gregg R. Howald<sup>c</sup>, Bradford S. Keitt<sup>c</sup>, Stephen W. Kress<sup>s</sup>, Colin M. Miskelly<sup>t</sup>, Steffen Oppel<sup>n</sup>, Sally Poncet<sup>u</sup>, Mark J. Rauzon<sup>v</sup>, Gérard Rocamora<sup>w,x</sup>, James C. Russell<sup>y,z</sup>, Araceli Samaniego-Herrera<sup>h</sup>, Philip J. Seddon<sup>aa</sup>, Dena R. Spatz<sup>ce</sup>, David R. Towns<sup>bb,cc</sup>, and Donald A. Croll<sup>e</sup>

We can do this...



ISLAND CONSERVATION



**Pinzon Giant Tortoise**

**Pinzon Island, Galapagos**

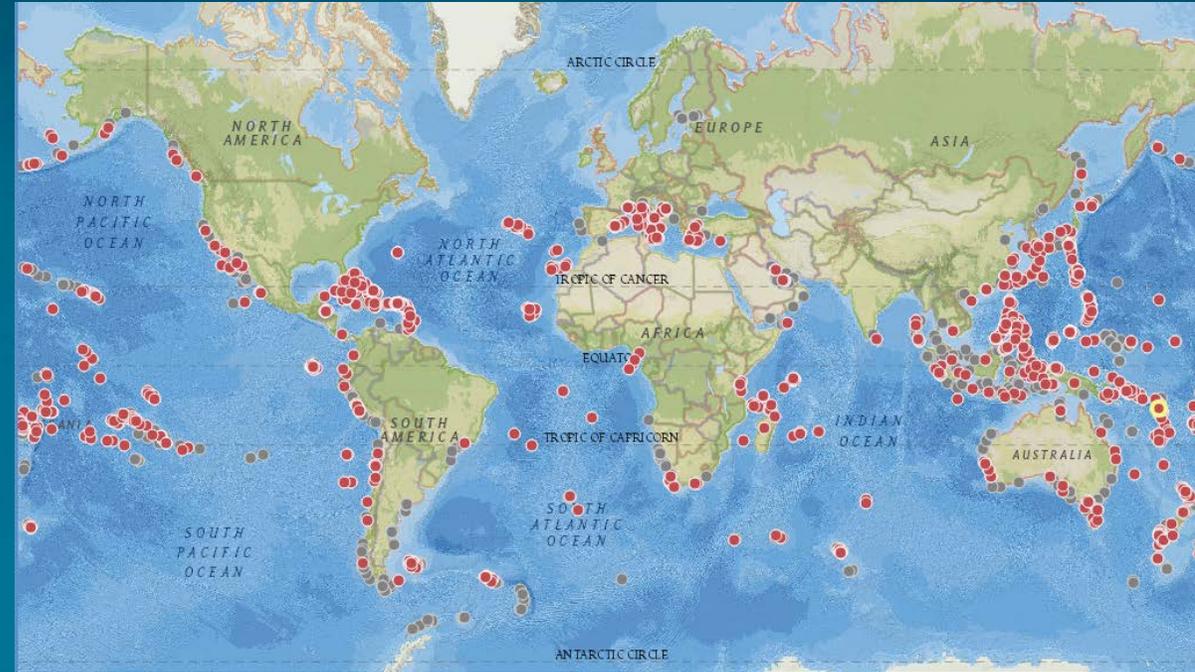
We can do this...

**Four spp. invasive rodents impact 88% of CR/EN vertebrate spp. on islands**

**Current methods feasible for islands holding 15% of those CR/EN spp.**

We can do this...

<http://tib.islandconservation.org>



# Innovation Strategy

Identify point of greatest impact

- Invasive rodents

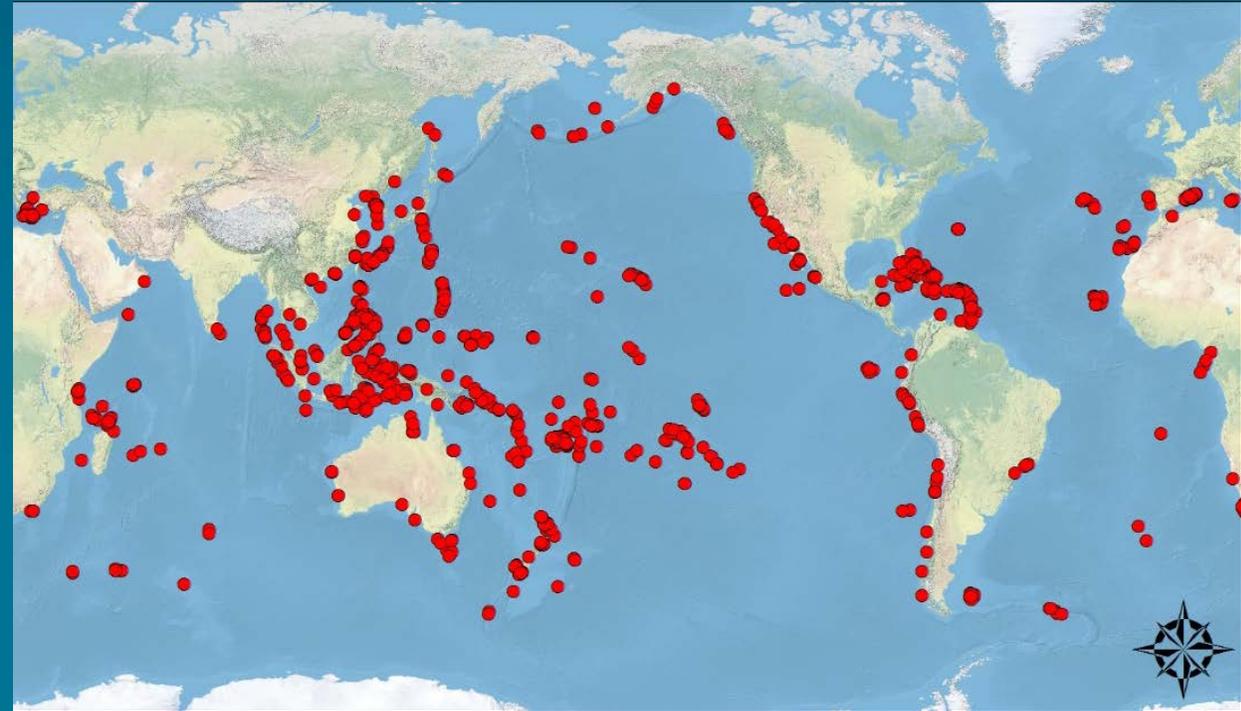
Match technology to the need

- Horizon scanning

Select investments

- Incremental
- Transformative

We can do this...



Contents lists available at ScienceDirect

Biological Conservation

journal homepage: [www.elsevier.com/locate/biocon](http://www.elsevier.com/locate/biocon)



Special Issue Article: Tropical rat eradication

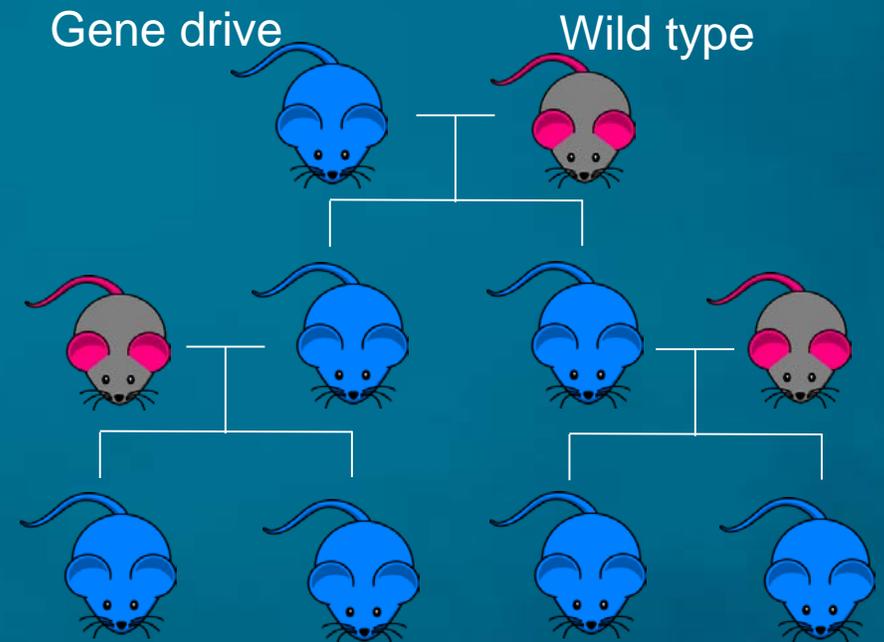
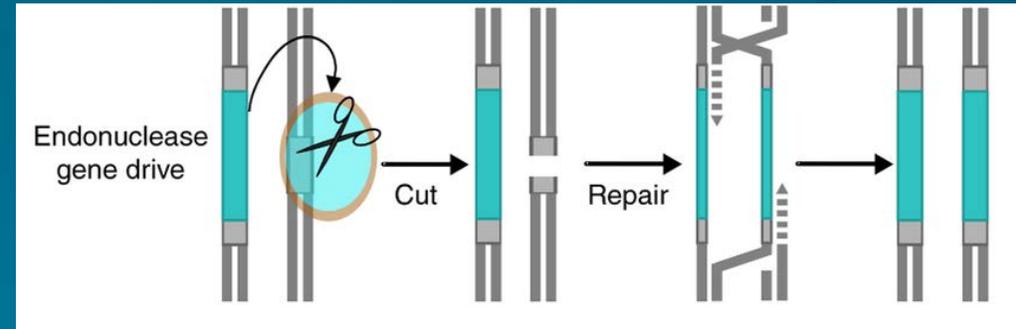
The next generation of rodent eradications: Innovative technologies and tools to improve species specificity and increase their feasibility on islands



Karl J. Campbell<sup>a,h,\*</sup>, Joe Beek<sup>a</sup>, Charles T. Eason<sup>c,d</sup>, Alistair S. Glen<sup>e</sup>, John Godwin<sup>f</sup>, Fred Gould<sup>g</sup>, Nick D. Holmes<sup>a</sup>, Gregg R. Howald<sup>a</sup>, Francine M. Madden<sup>h</sup>, Julia B. Ponder<sup>i</sup>, David W. Threadgill<sup>j,k</sup>, Alexander S. Wegmann<sup>a</sup>, Greg S. Baxter<sup>b</sup>

# Gene drives

- Cause genes to be inherited more frequently than normal – up to 100%
- Sexual reproduction
- Ability to modify wild populations *by design*
  - Insert new, modify or delete genes
- CRISPR-Cas9

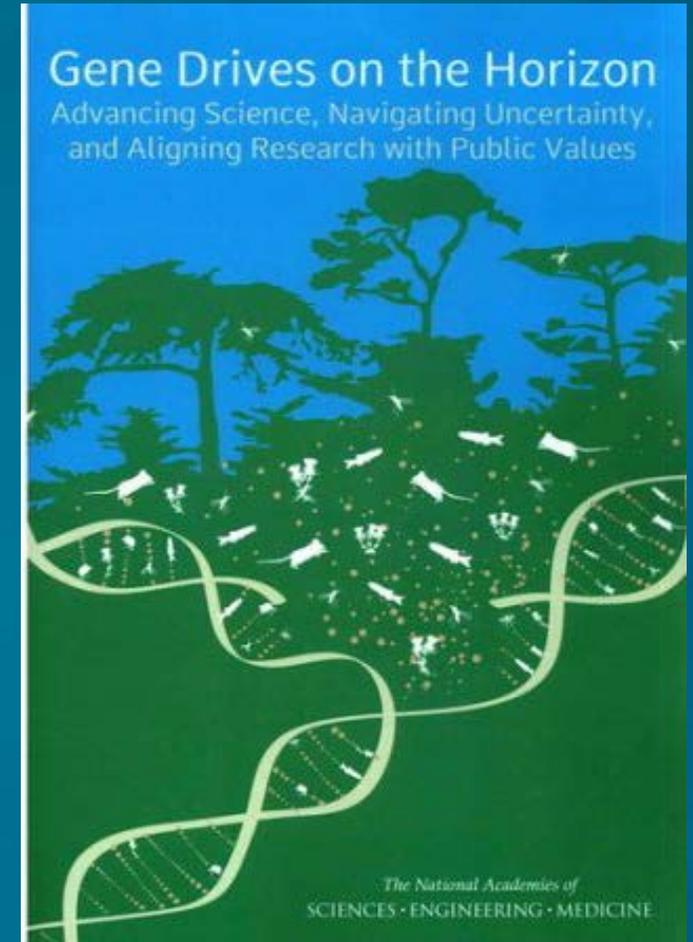


We can do this...

# Gene drives

- Cause genes to be inherited more frequently than normal – up to 100%
- Sexual reproduction
- Ability to modify wild populations *by design*
  - Insert new, modify or delete genes
- CRISPR-Cas9
- Nascent fast-moving field
  - Pre-caution & phased approaches needed

We can do this...



NASEM 2016; Kaebnick et al. 2016

# Genetic Biocontrol of Invasive Rodents program

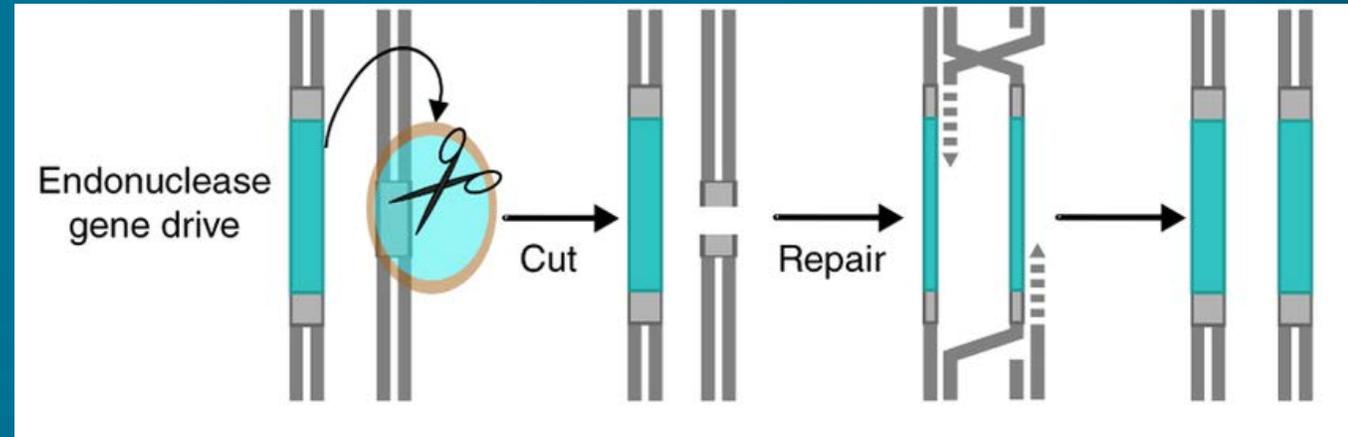
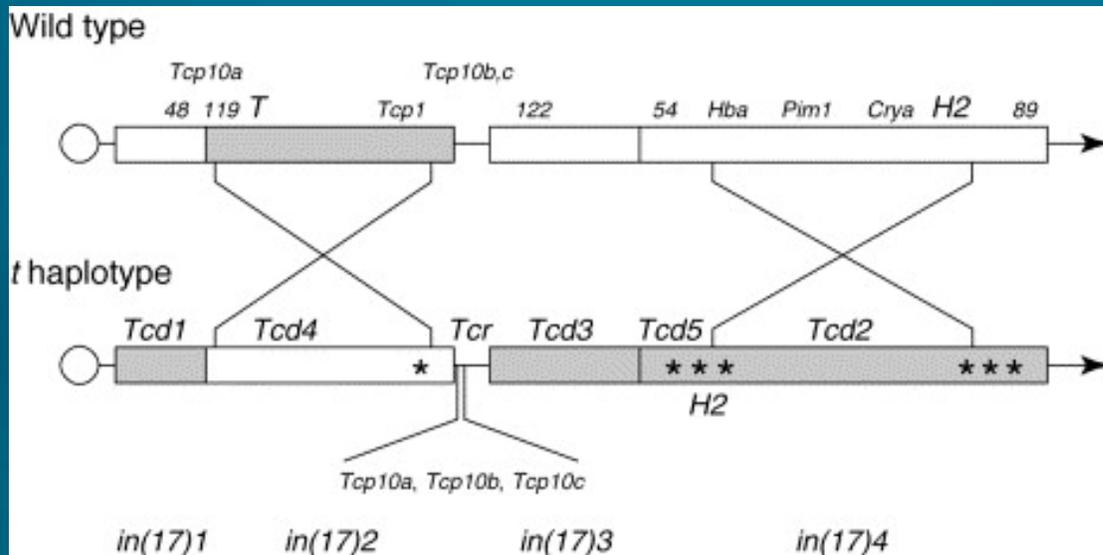


We can do this...

# Gene Drive

T-Complex – natural

CRISPR/Cas9 - synthetic



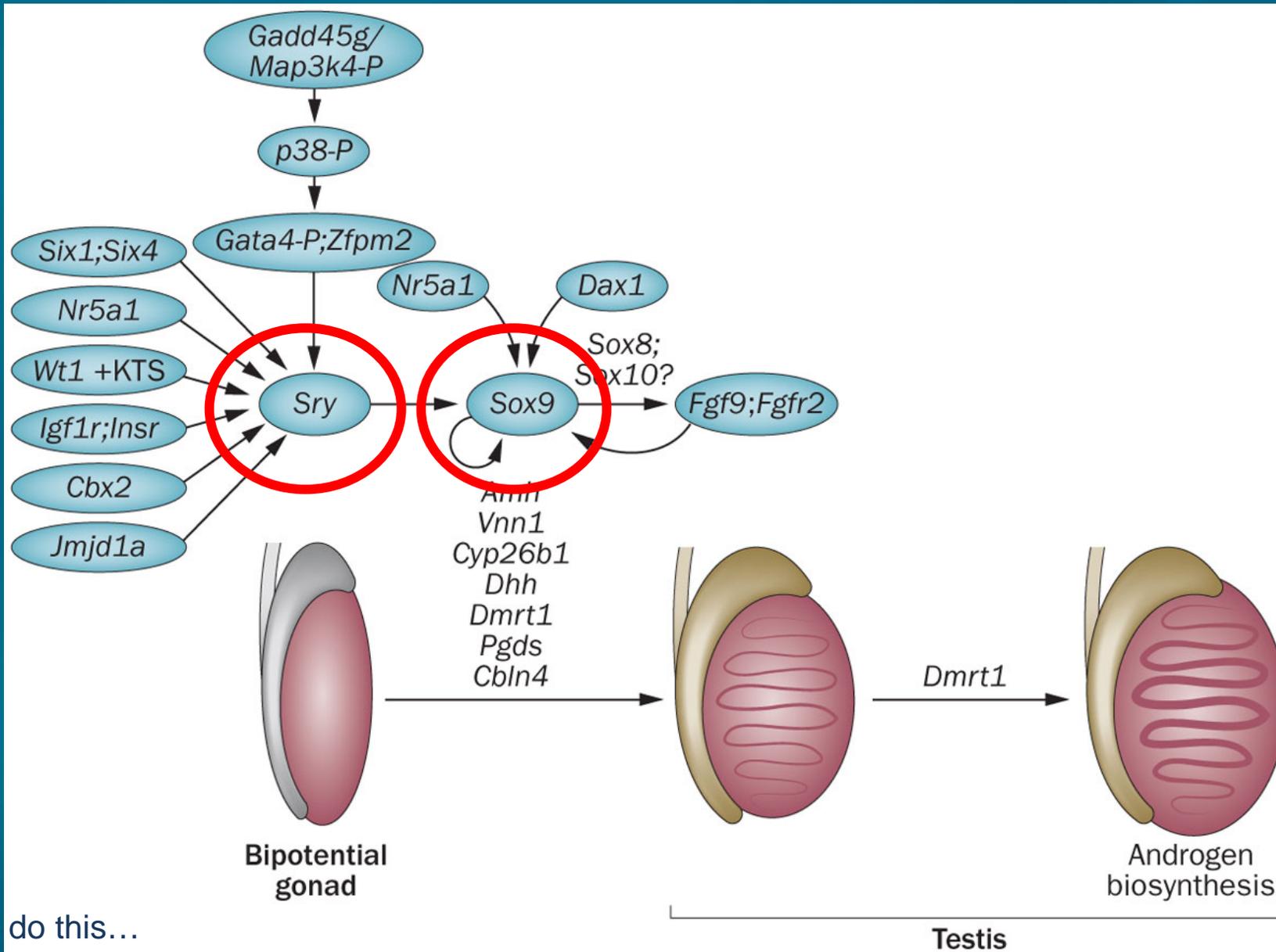
92% n = 691



THE UNIVERSITY  
of ADELAIDE

We can do this...

# Mammalian Sex Determination



Bias sex ratio:

- Sry
- Sox9

We can do this...

Eggers *et al.* 2014

# Genetic Biocontrol of Invasive Rodents

- *T-Complex/Sry*
- *CRISPR-Cas9/Sox9*
- Control mechanisms
- Risk assessment
- Mathematical models
- Breeding behavior
- Biocontainment
- Communications
- Island selection
- Genetic characterization of mouse populations & monitoring
- Regulatory (multi-national)
- Stakeholder/comm/public engagement
- Intellectual property
- Funding
- Program management
- Governance
  - Ethics committee
  - Advisory committee

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**NC STATE**  
UNIVERSITY



# Leverage biology

- Population ~1000 house mice
- Released 42 male & 35 female mice from Eday Island
- After 18 months, all mice trapped were hybrids (n=70)
- Males were disproportionately responsible for 'invasion'
- Leverage biology and promiscuity

Isle of May (57ha)



# RNA interference (RNAi)

## Ribonucleic acid (RNA)

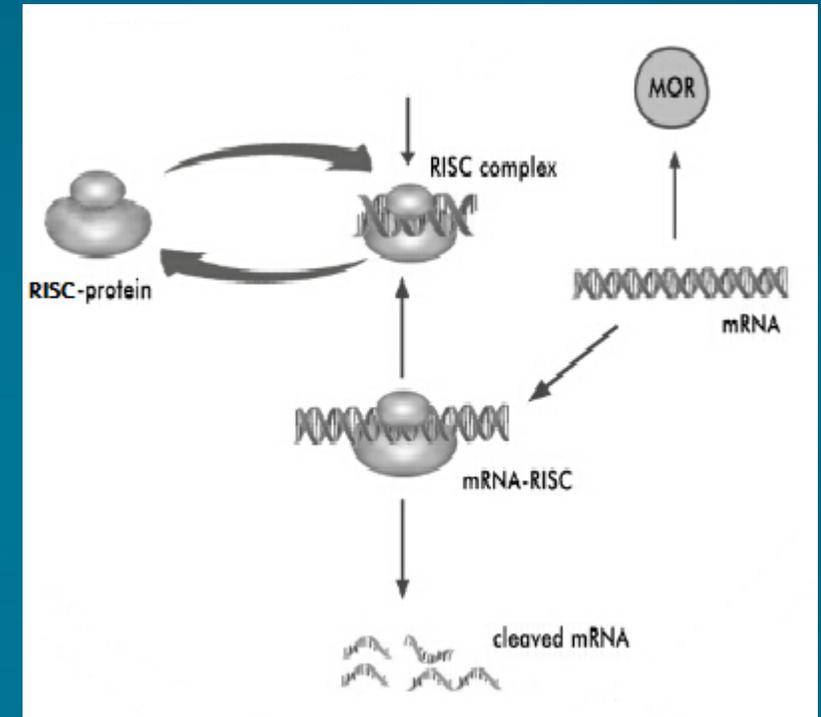
- Normally single stranded

## Double-stranded RNA

- Elicits immune response
  - Double-stranded RNA and matching single-stranded RNA destroyed

## Focus of a significant body of research:

- Invertebrate pest control
- Animal models (e.g. mice) as a potential cure for cancer and other diseases



# RNA interference (RNAi)

Double-stranded RNA can be synthesized chemically

- Use as taxa-specific bio-pesticides

Can be synthesized biologically (species can produce it)

- Plants & animals can be vectors

Vertebrates digest RNA

- Nanoparticles as carriers



# Gene drives & RNAi: Power-tools in the tool box?

Cautious investigation of opportunities  
required

Big picture

- Reduce risks
- Increase impact

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