

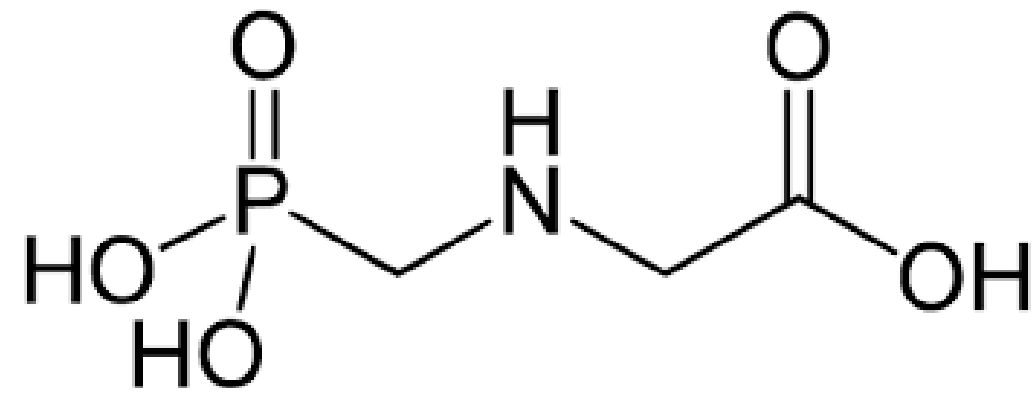
Living Technologies for Ecological Problems

Kevin M. Esvelt

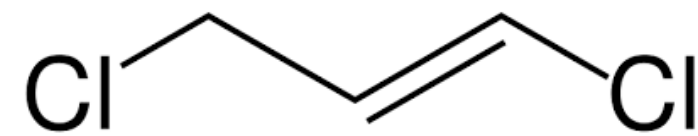
Assistant Professor, MIT Media Lab

National Academy of Sciences
16 November 2016

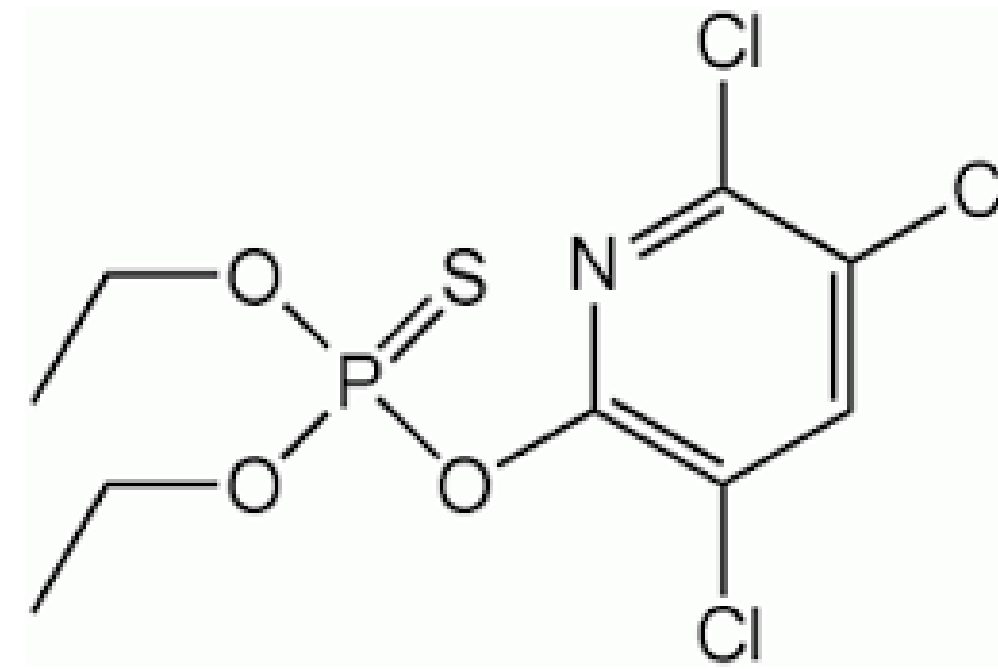




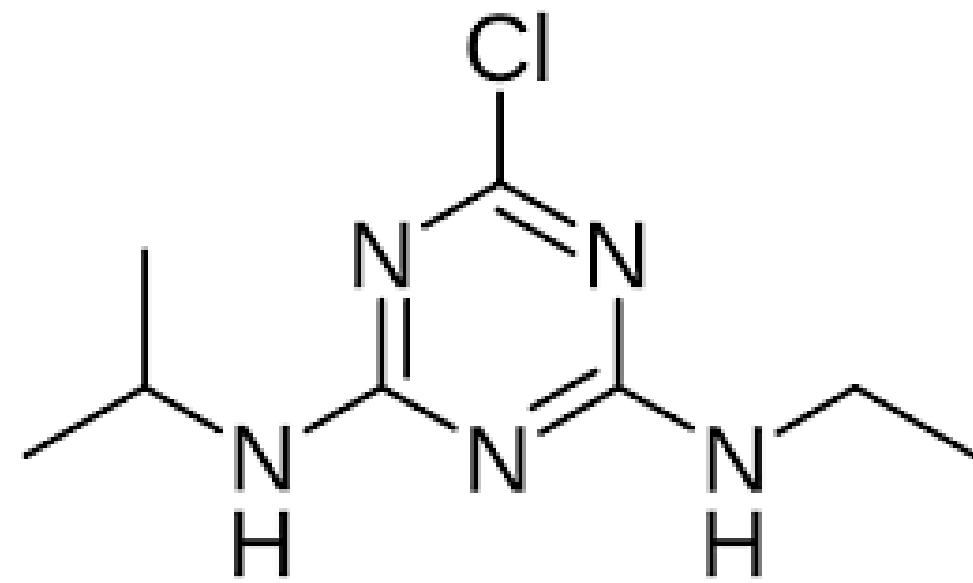
Glyphosate



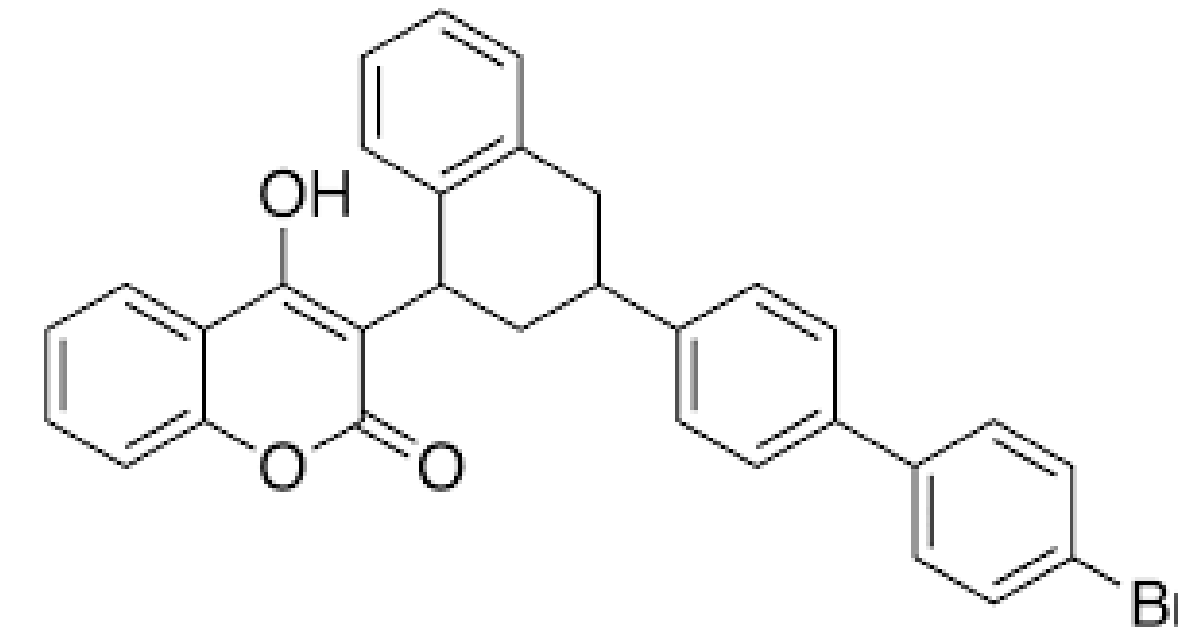
Dichloropropene



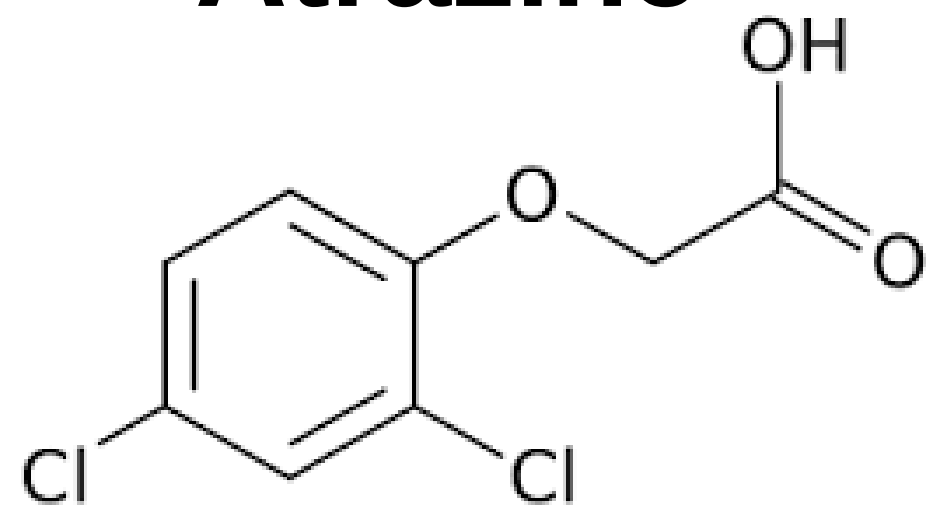
Chlorpyrifos



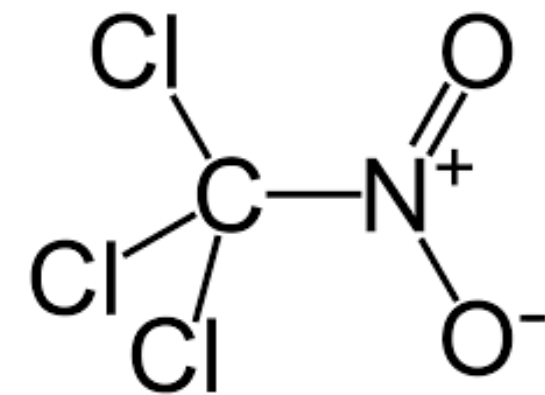
Atrazine



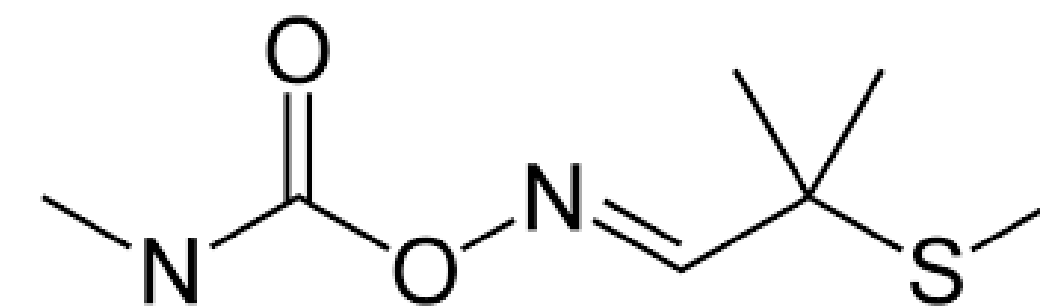
Brodifacoum



2,4-D



Chloropicrin



Aldicarb



Imagine a world that solves ecological problems using biology...

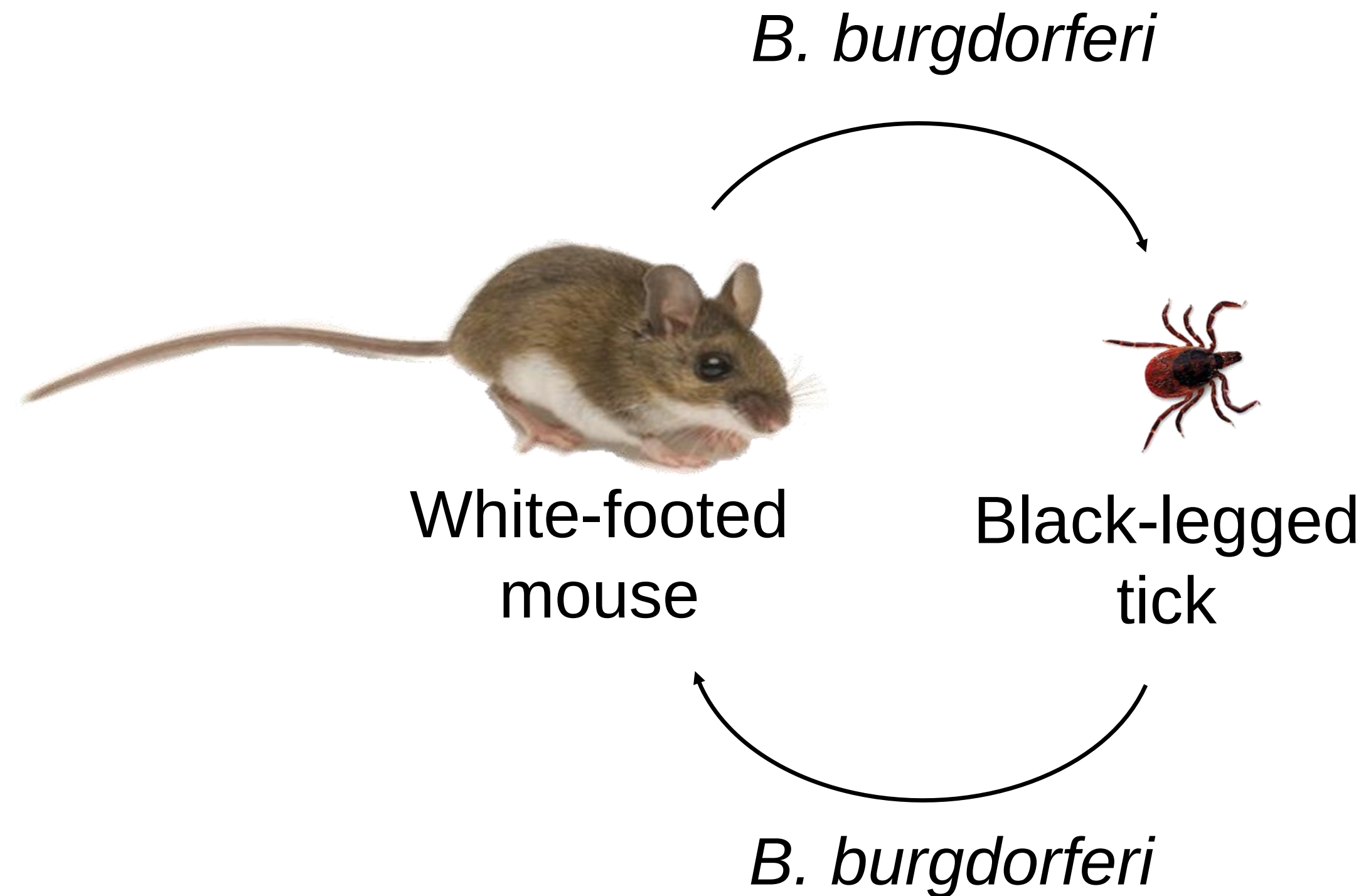


Preventing Tick-Borne Disease



photo: Yousur Al-Hilou/
The New York Times

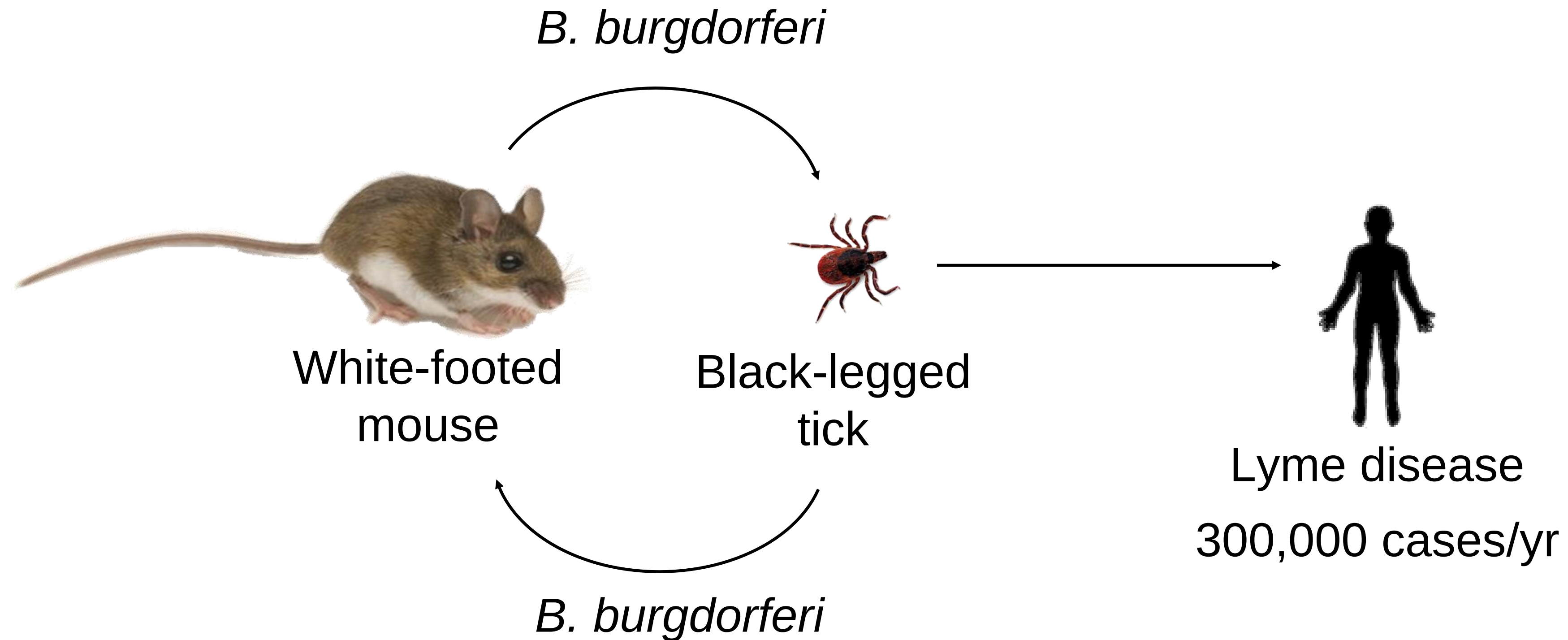
Tick-borne disease is an ecological problem



The spirochetes that cause Lyme disease persist by moving between mice and ticks



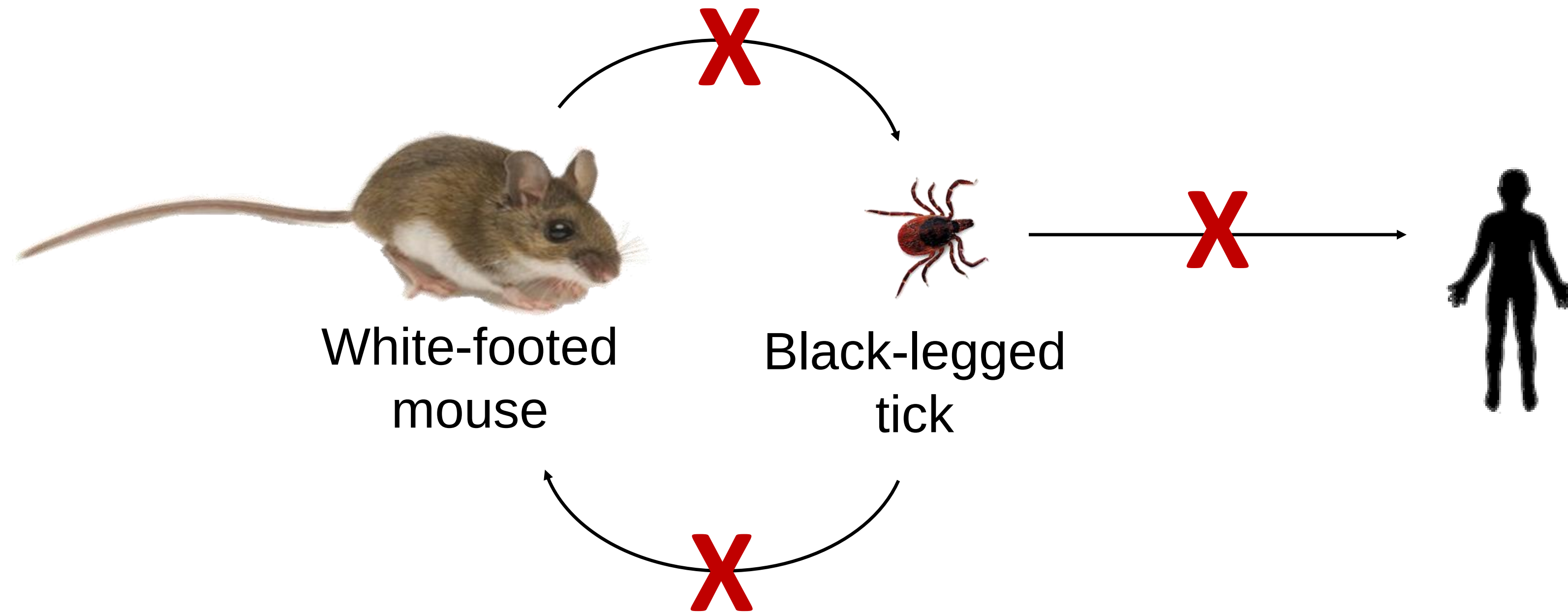
Tick-borne disease is an ecological problem



Ticks pass the spirochetes to humans, causing Lyme disease
(a similar cycle holds for other tick-borne diseases)



Tick-borne disease is an ecological problem



What if white-footed mice couldn't become infected?



Ecological Vaccination



White-footed mice

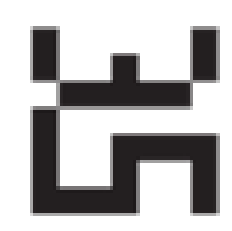
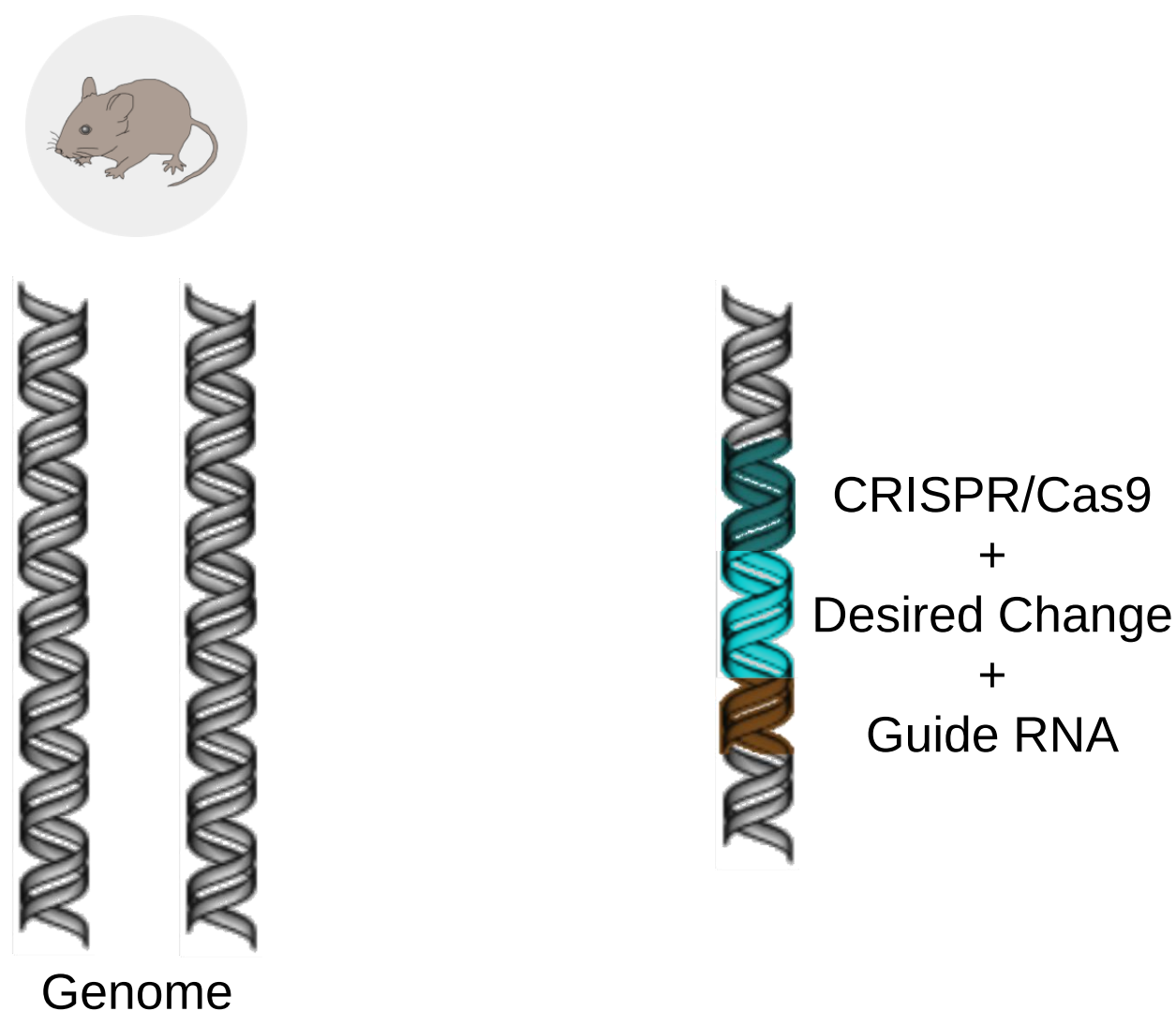
**What is the smallest possible change
that can solve the problem?**

Start local

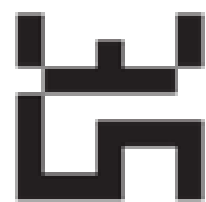
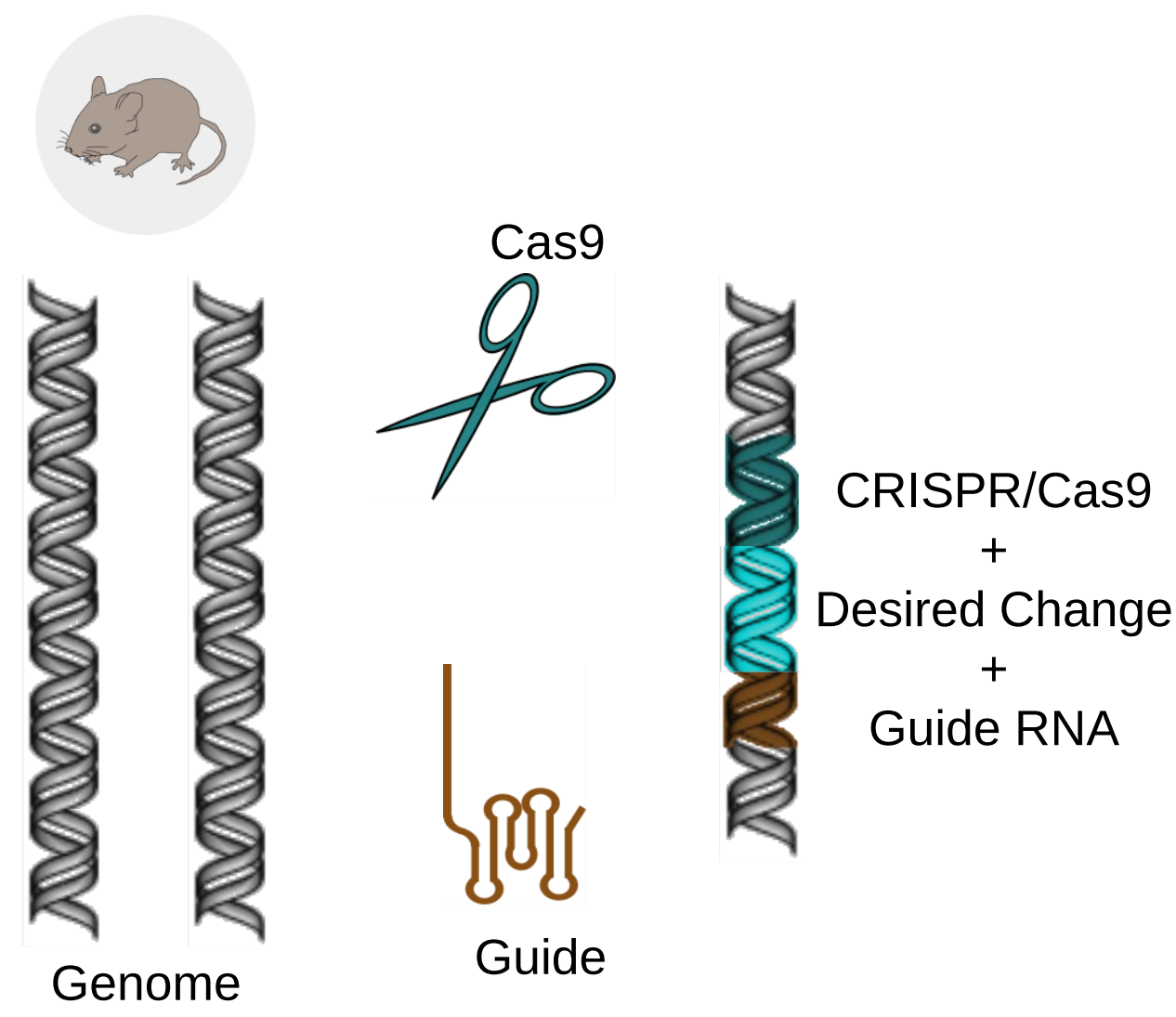
Scale up if it works



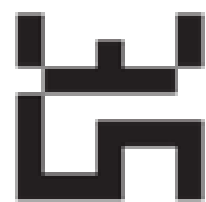
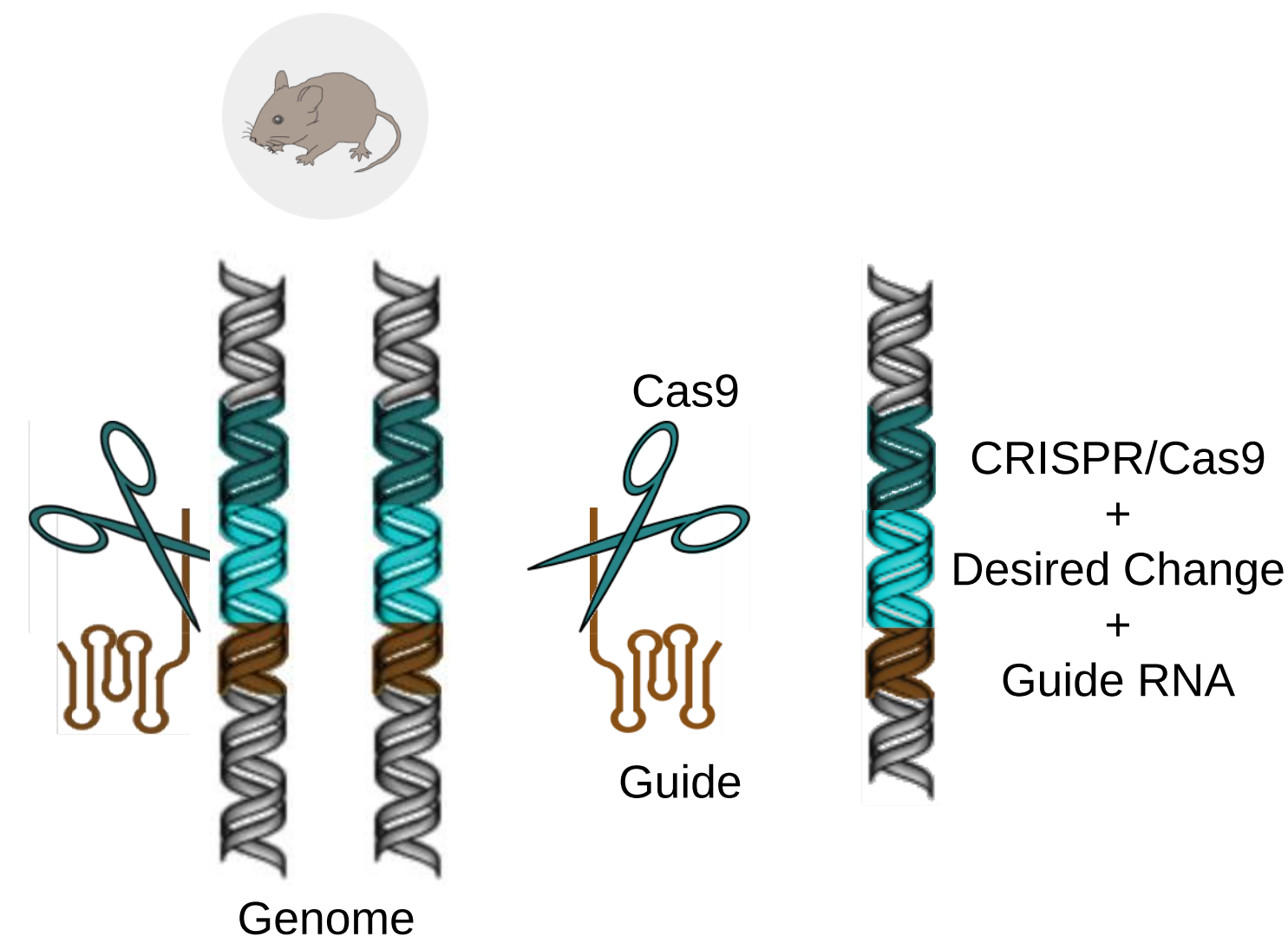
Global Gene Drive



Global Gene Drive

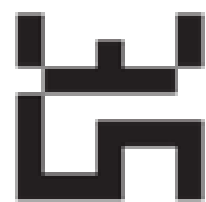
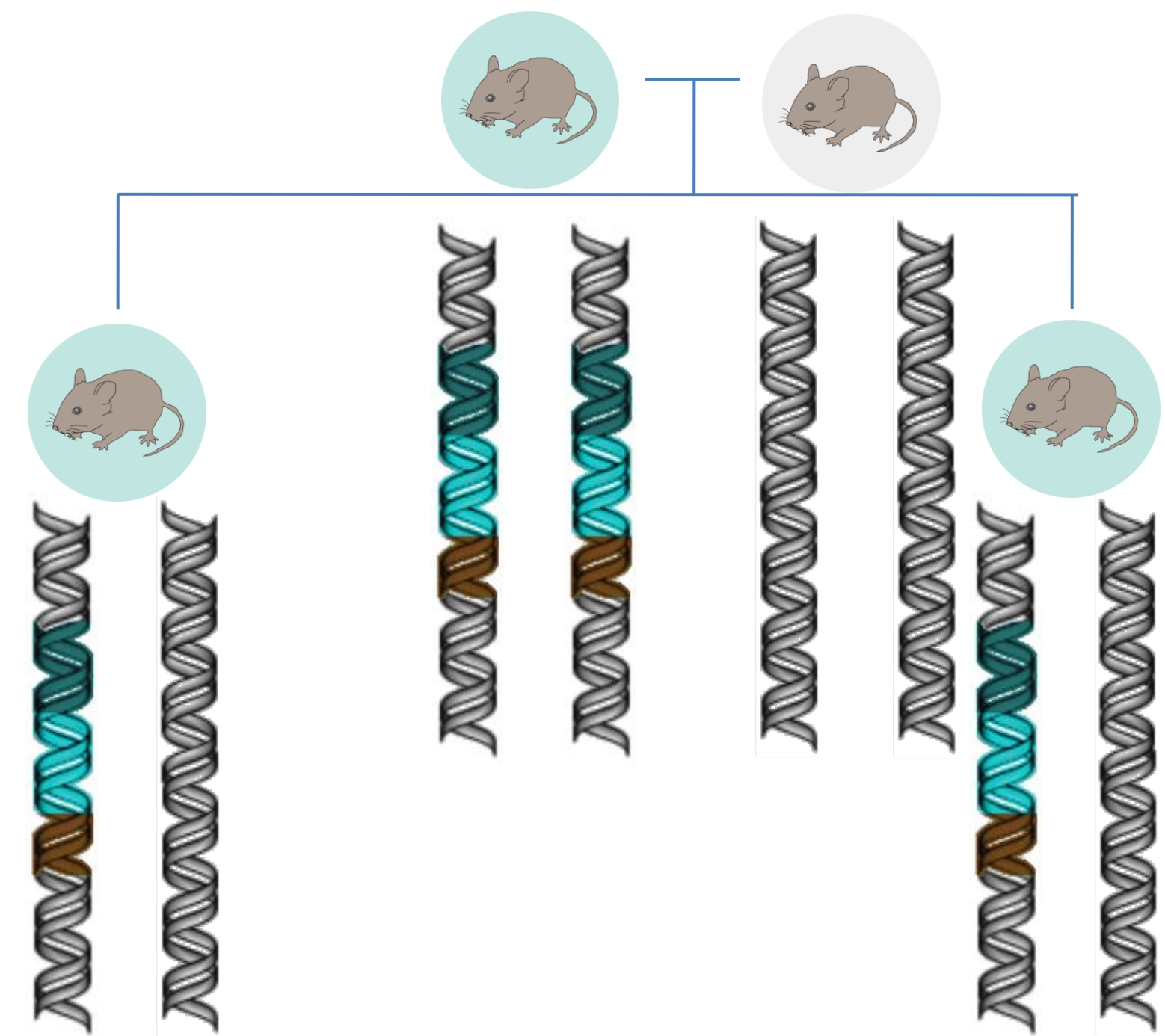


Global Gene Drive



Esvelt KM, Smidler AL, Catteruccia F, Church GM (2014) *eLife*
Oye K, Esvelt K et al. (2014) *Science*

Global Gene Drive

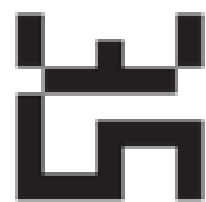
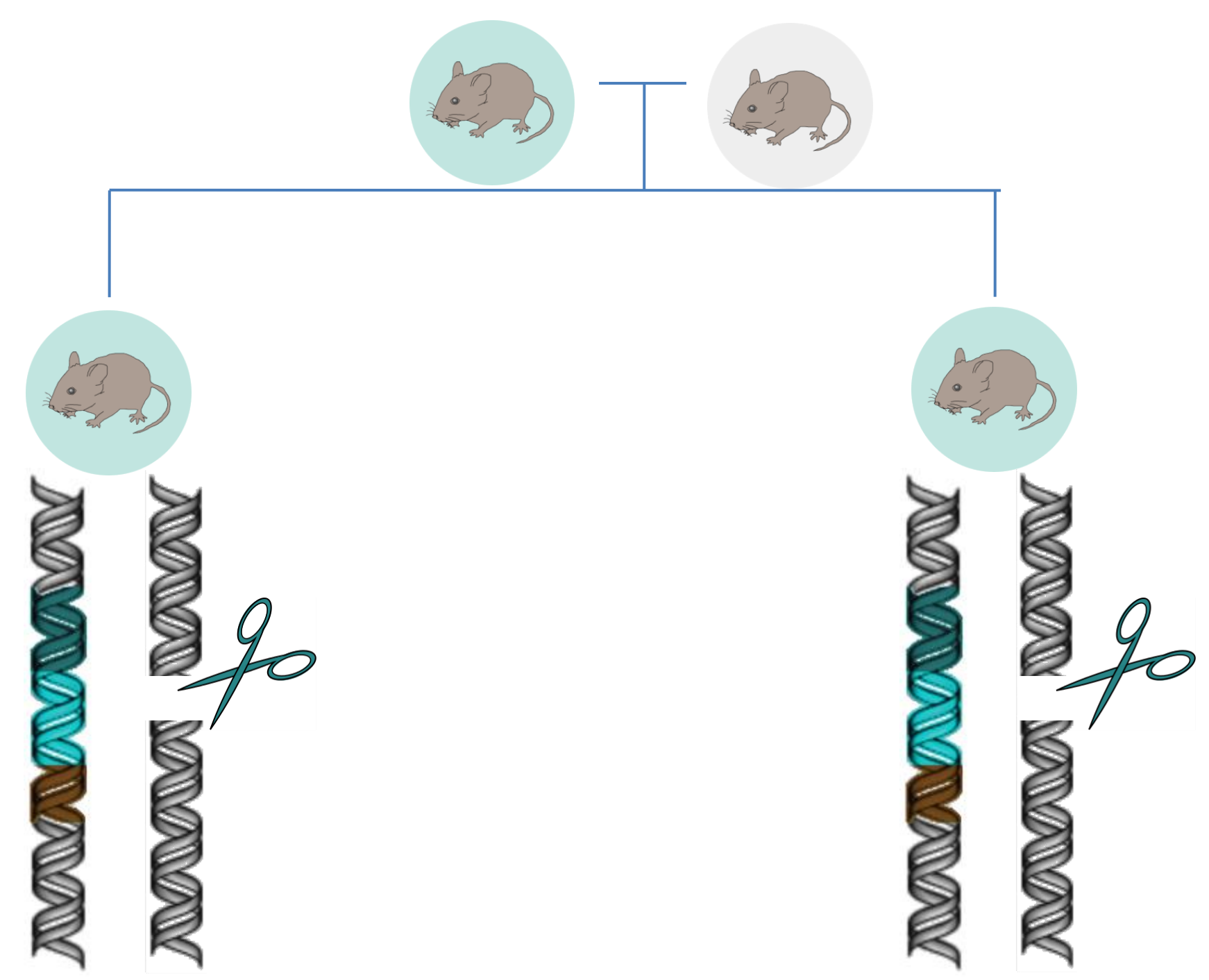


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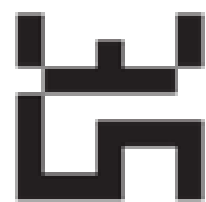
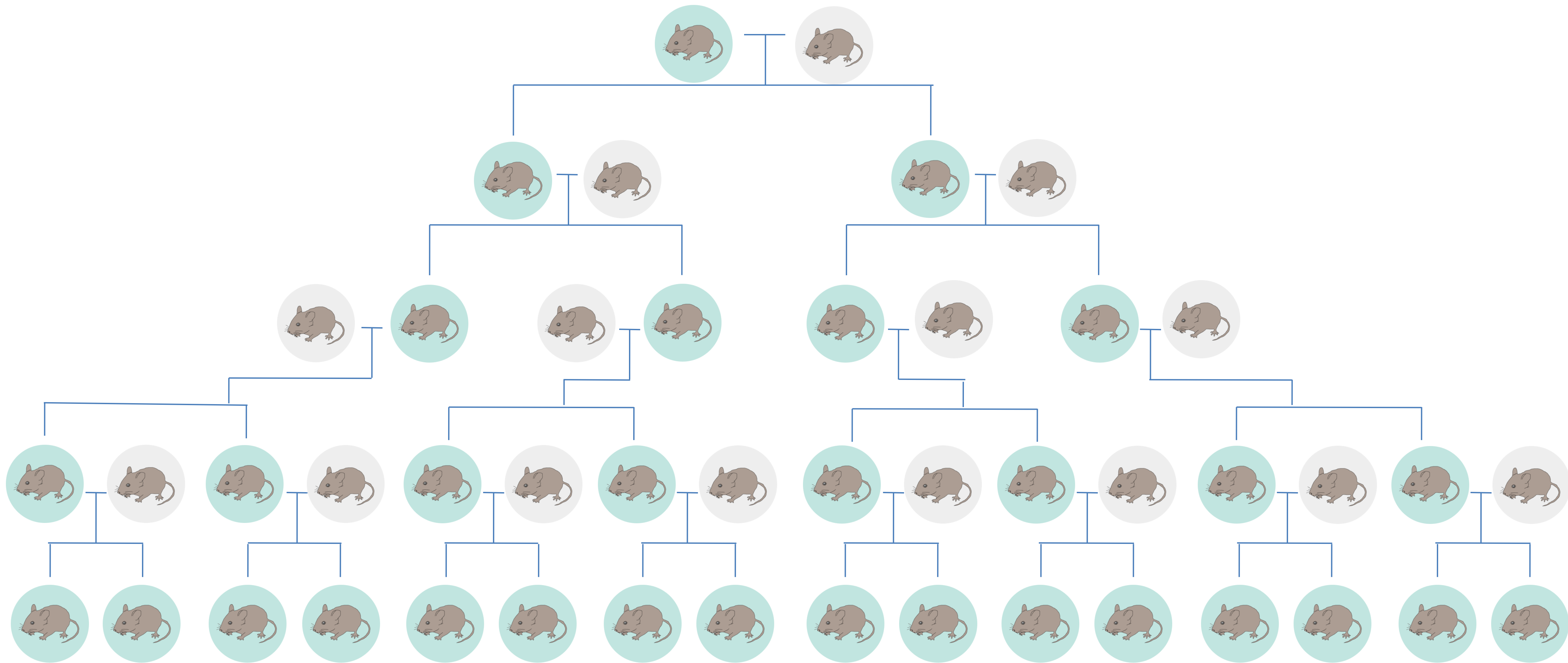
Living Technologies for
Ecological Problems
NAS

Kevin M. Esvelt
Sculpting Evolution Group
MIT Media Lab

Global Gene Drive



Global Gene Drive



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**What is the smallest possible change
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Start local

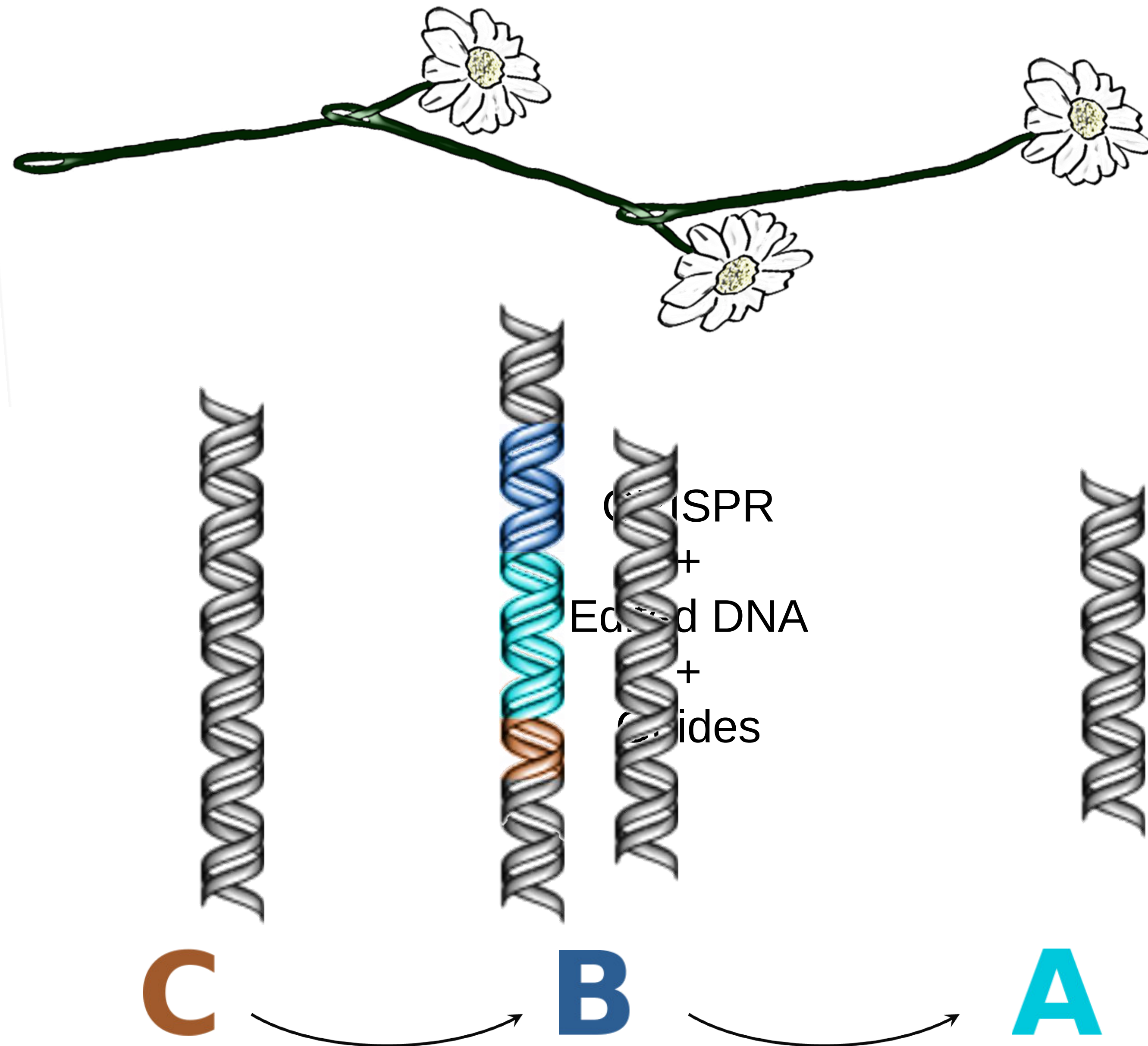
Scale up if it works



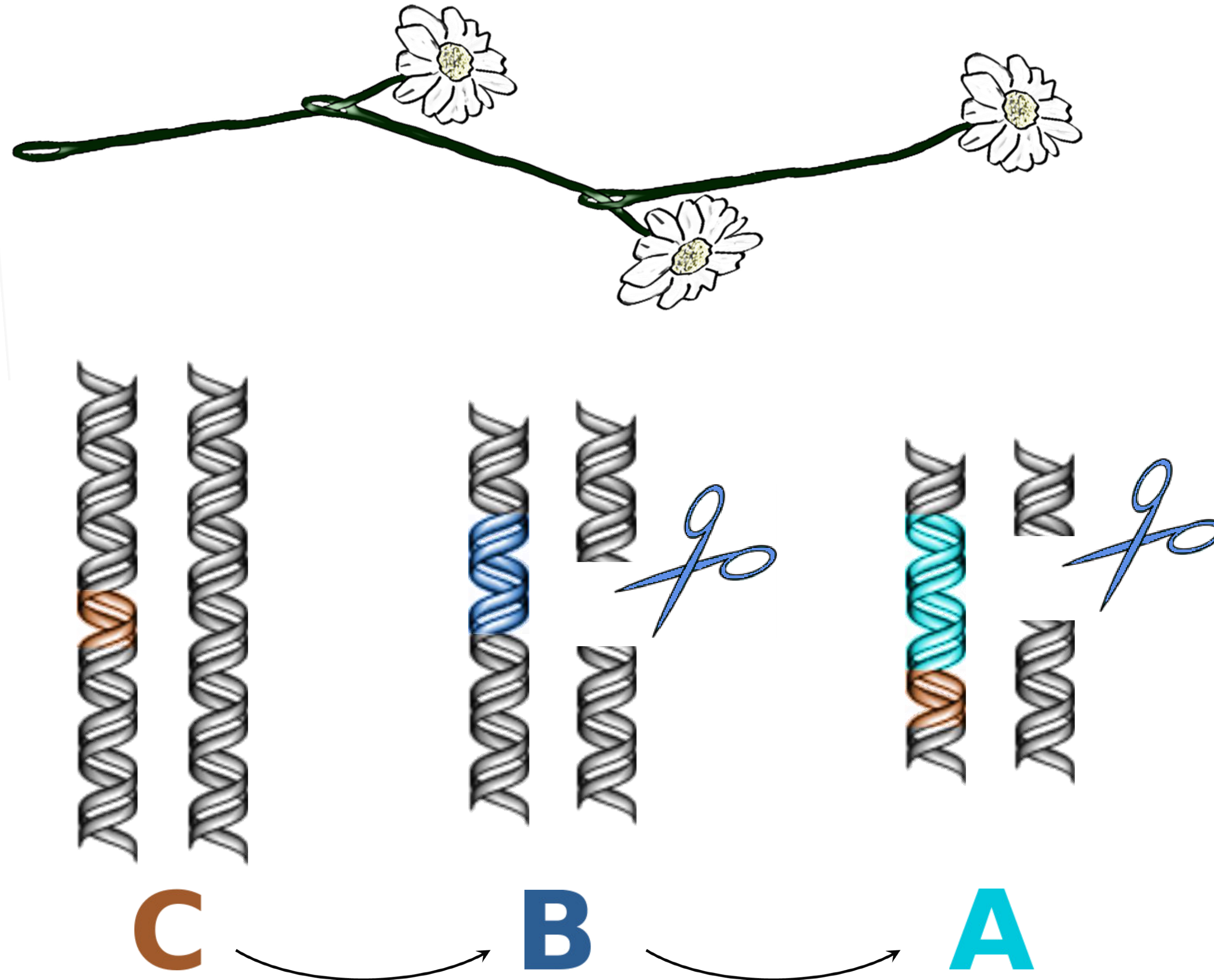
How do you make a local CRISPR-based drive system?



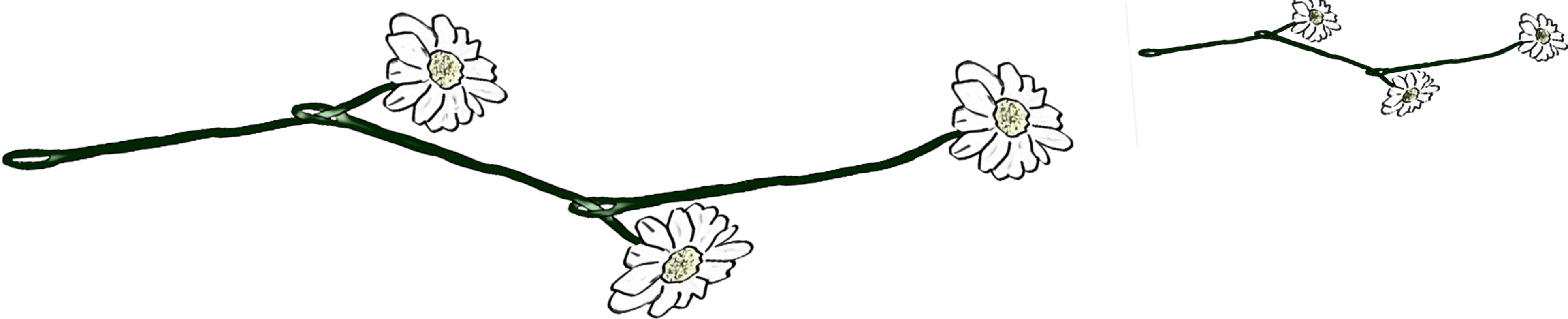
Daisy Drive



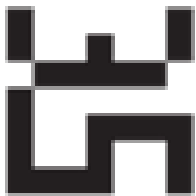
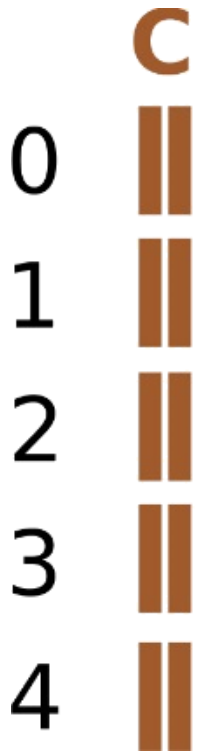
Daisy Drive



Daisy Drive



Generations



Daisy Drive

Frequency
in Local
Population

A

B

C



CBA

0

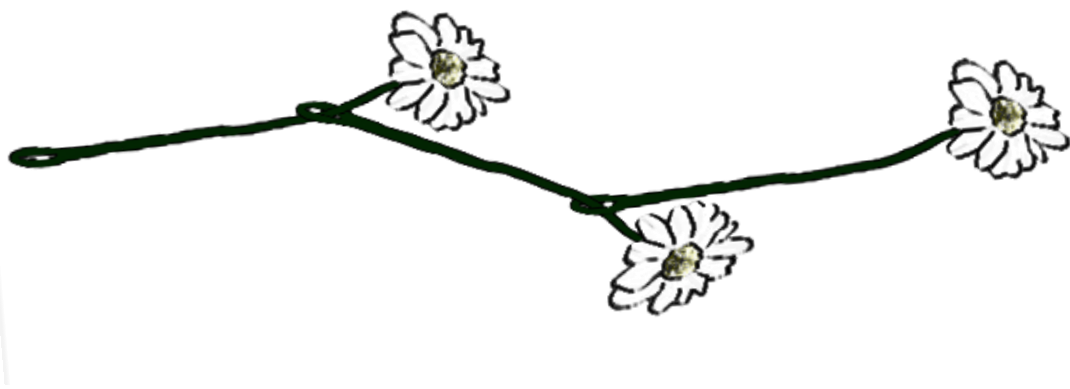
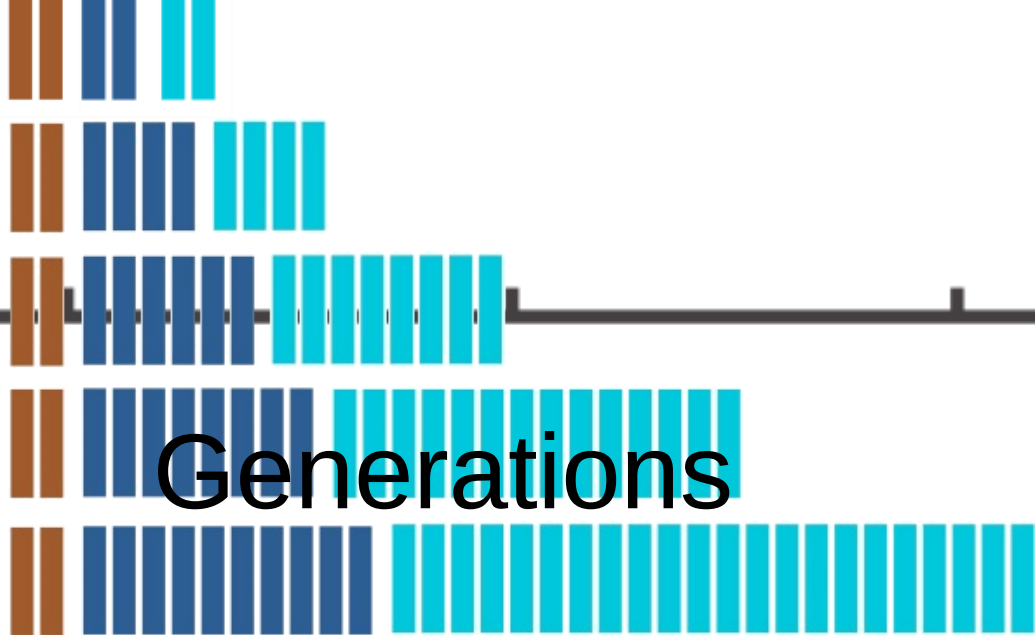
1

2

3

4

Generations



Daisy Drive

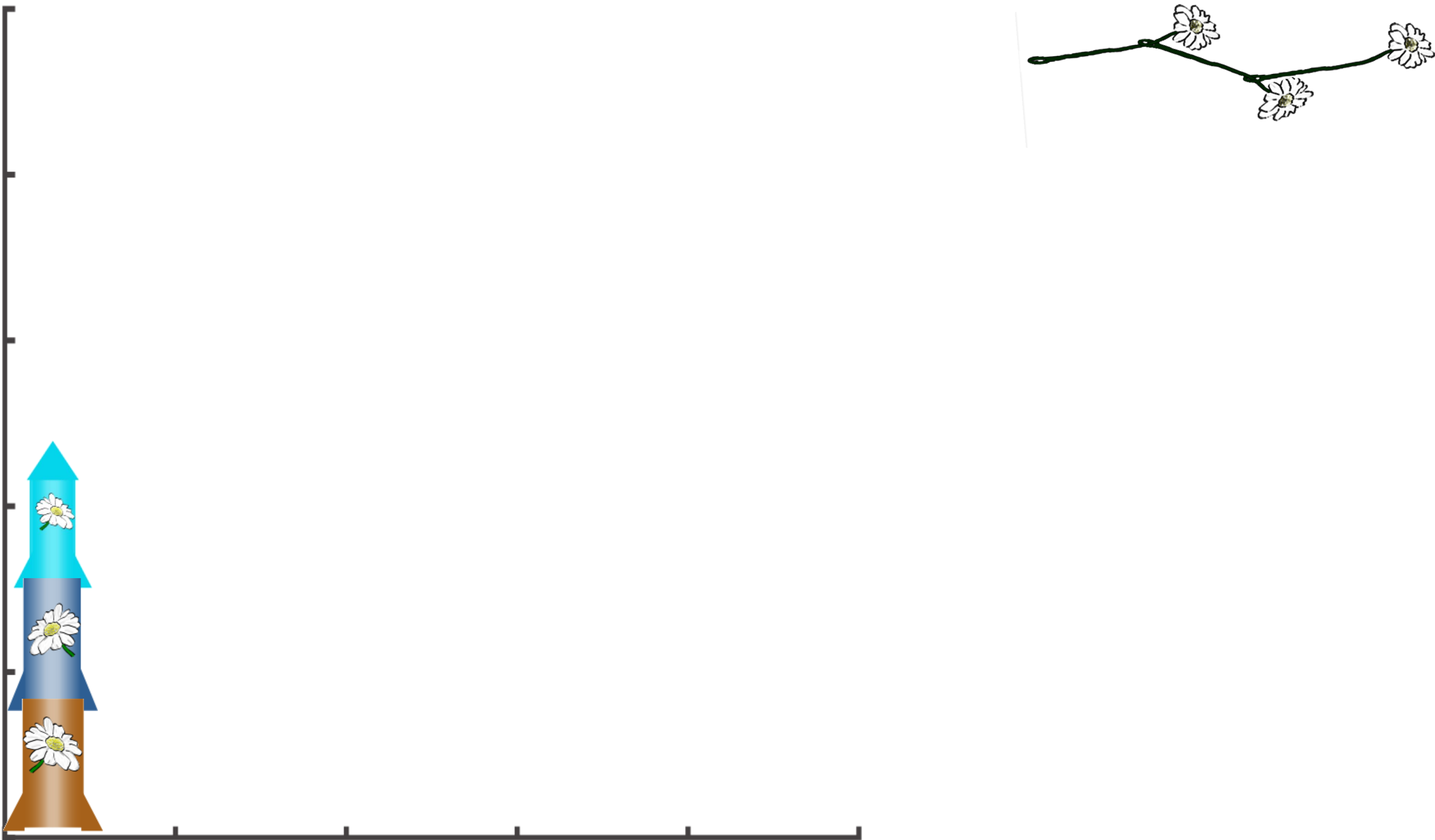
Frequency
in Local
Population

A

B

C

Generations



Daisy Drive

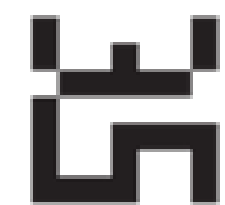
Frequency in Local Population

Generations



B

C



Noble C*, Min J*, Olejarz J, Buchthal J, Chavez A, Smidler AL, DeBenedictis EA, Church GM, Nowak M, Esvelt KM (2016) *bioRxiv*

Living Technologies for Ecological Problems

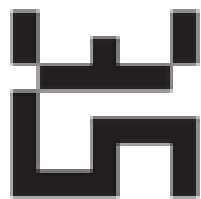
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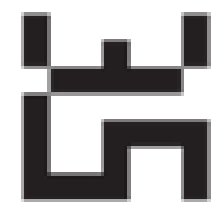
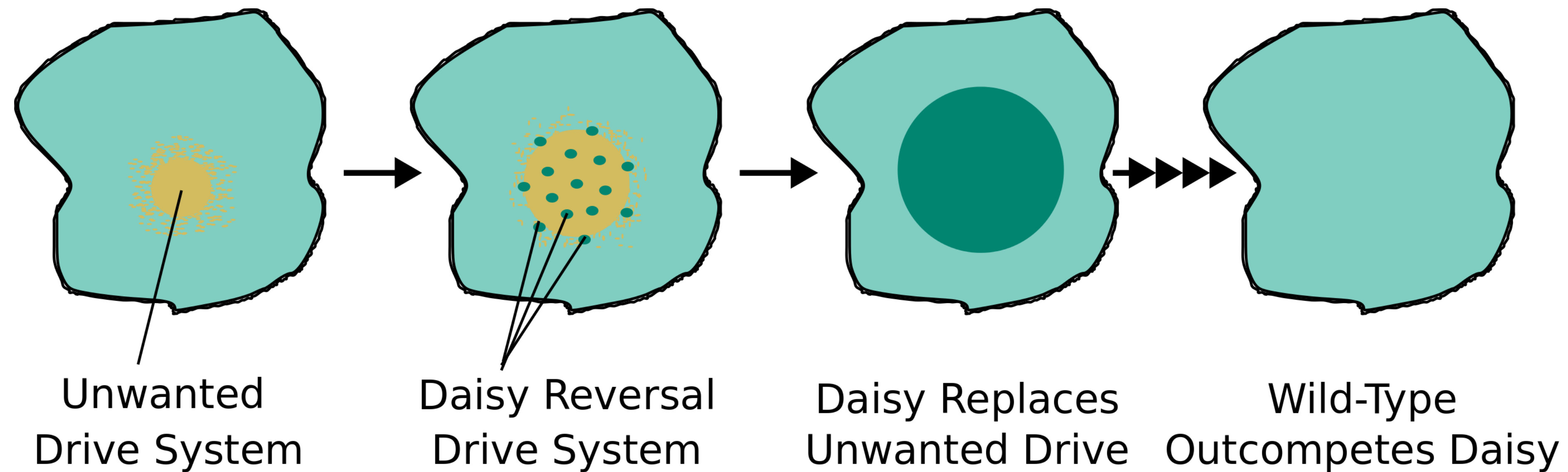
Daisy Drive

Maximum
Frequency

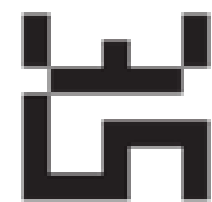
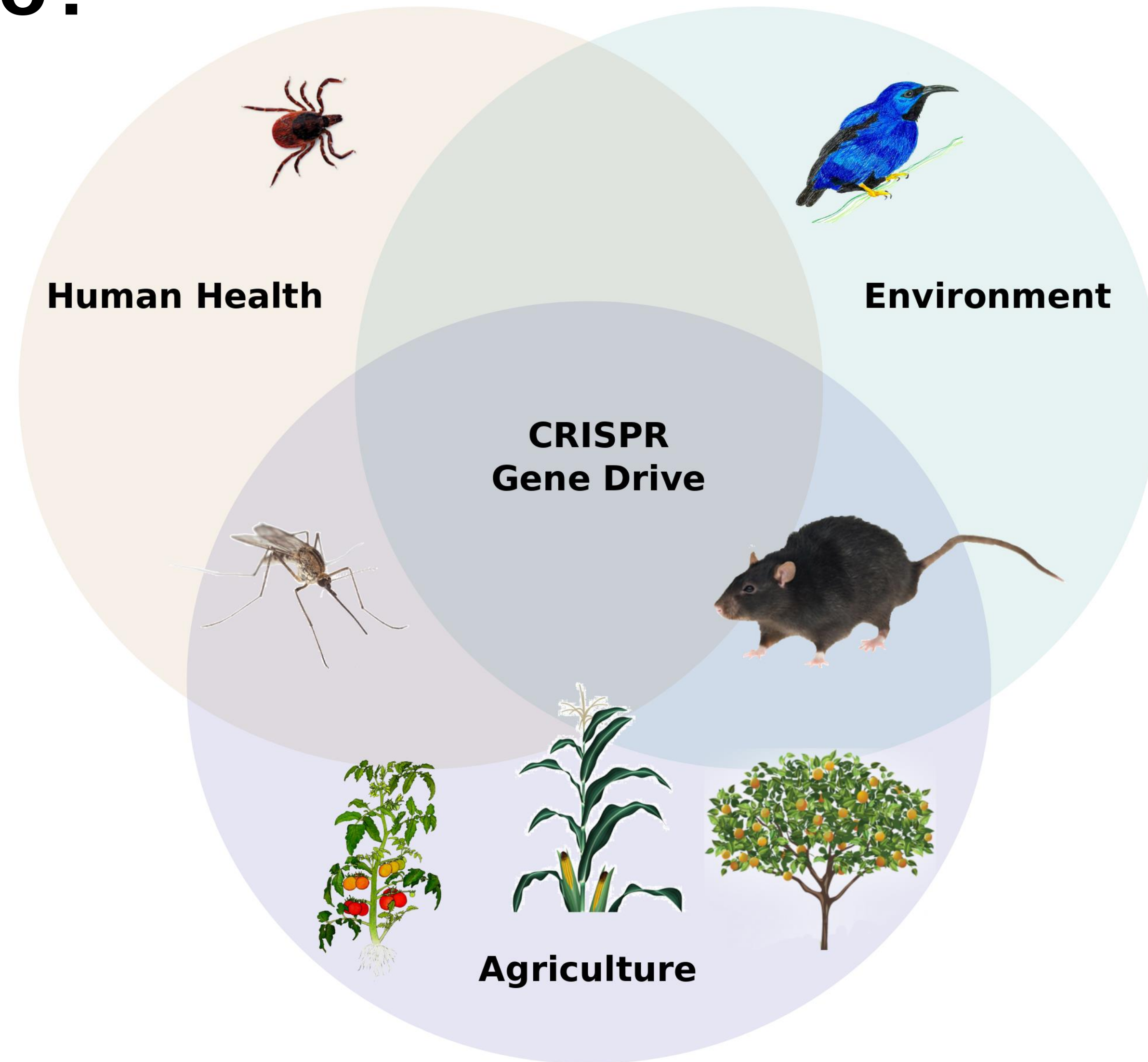
of Daisy Elements



Daisy Drive – Perfect Restoration to Wild-Type



What can we do?



“The best course of action is to ensure that the people who could be affected by a proposed project or policy have an opportunity to have a voice in decisions about it.”

Gene drive technology cannot be developed in secret

“Experts acting alone will not be able to identify or weigh the true costs and benefits of gene drives.”

- The U.S. National Academies report

We are working with funders, journals, and IP experts to ensure openness



Open science is faster, safer, and more trustworthy



Local, Open, Responsive Science

- Aim to make the smallest possible change
- Openly share proposals before experiments begin
- Actively invite concerns
- Request community guidance
- Arrange for independent assessment
- Start local and scale up



Acknowledgements

Ecological Immunization

<i>Joanna Buchthal</i>	Sam Telford	Ryan Kelly
<i>Devora Najjar</i>	Linden Hu	Jeantine Lunshof
<i>Avery Normandin</i>	Aurelie Kerns	

CRISPR Gene Drive (Esvelt et al, Oye et al 2014)

Andrea Smidler	Kenneth Oye	Tod Kuiken
Flaminia Catteruccia	Shlomiya Lightfoot	Evan Appleton
George Church	James P. Collins	Julie McNamara

Daisy Drive (Noble, Min et al 2016)

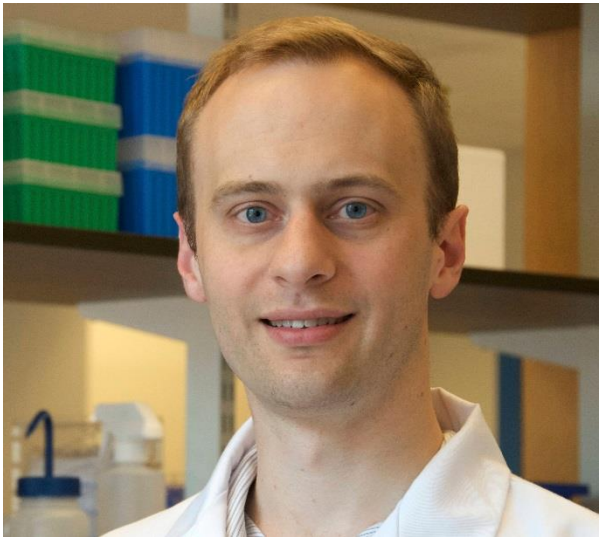
<i>John Min</i>	<i>Joanna Buchthal</i>	<i>Erika DeBenedictis</i>
Charleston Noble	George Church	Andrea Smidler
Alex Chavez	Martin Nowak	Jason Olejarz

Responsive Science

<i>Joanna Buchthal</i>	Sam Weiss Evans
Shlomiya Lightfoot	Travis Rich
Ariel Conn	Jeantine Lunshof



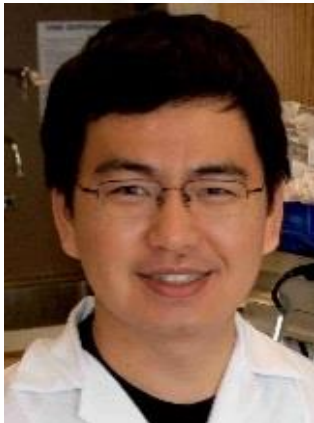
exploring evolutionary and ecological engineering



Kevin Esvelt



Georgiana Ujomu



John Min



Erika DeBenedictis



Joanna Buchthal



Cody Gilleland



Devora Najjar