# CRISPR/Cas9 & Genomewide CRISPR Screens

Harris Hunter & Franky Shi

# What is gene editing?

Gene editing is the use of technology to change an organism's DNA.

### Why is gene editing important?



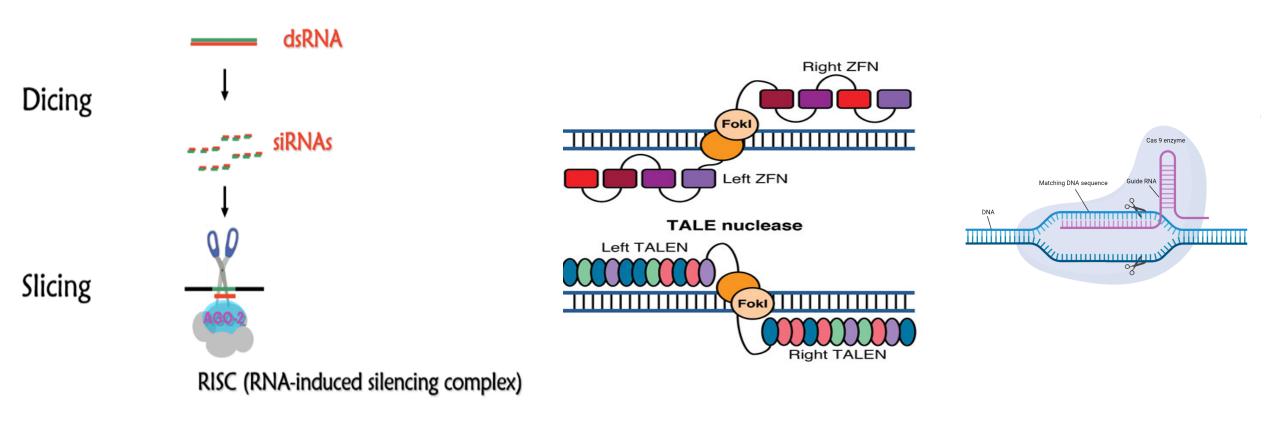
**Cystic Fibrosis** 

Huntington's Disease

Hemophilia

Many researchers are currently using gene editing to cure disorders.

# What are some ways to perform gene editing?

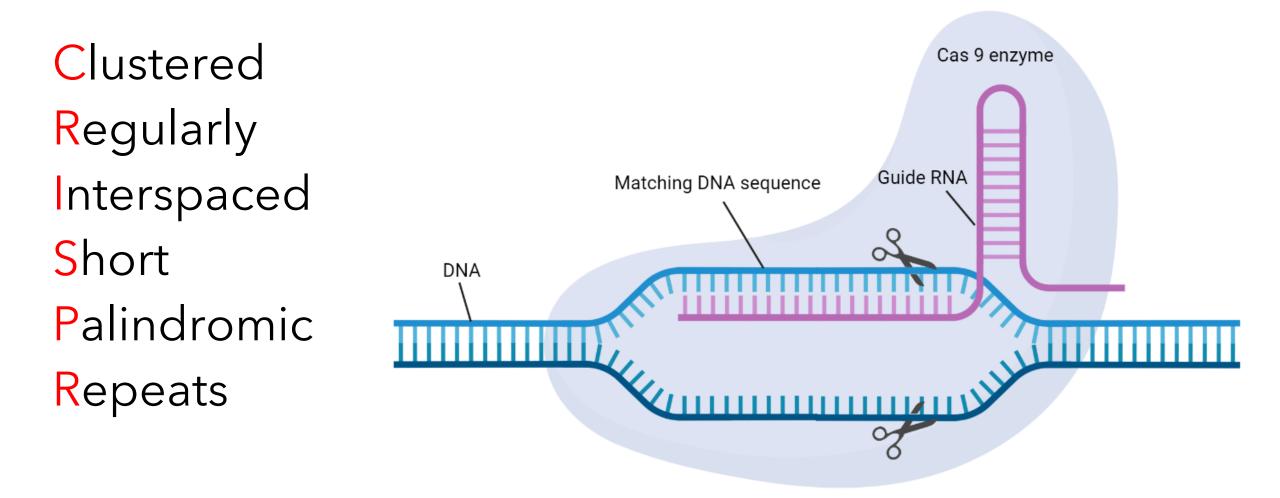


RNAi

**ZFNs/TALENS** 

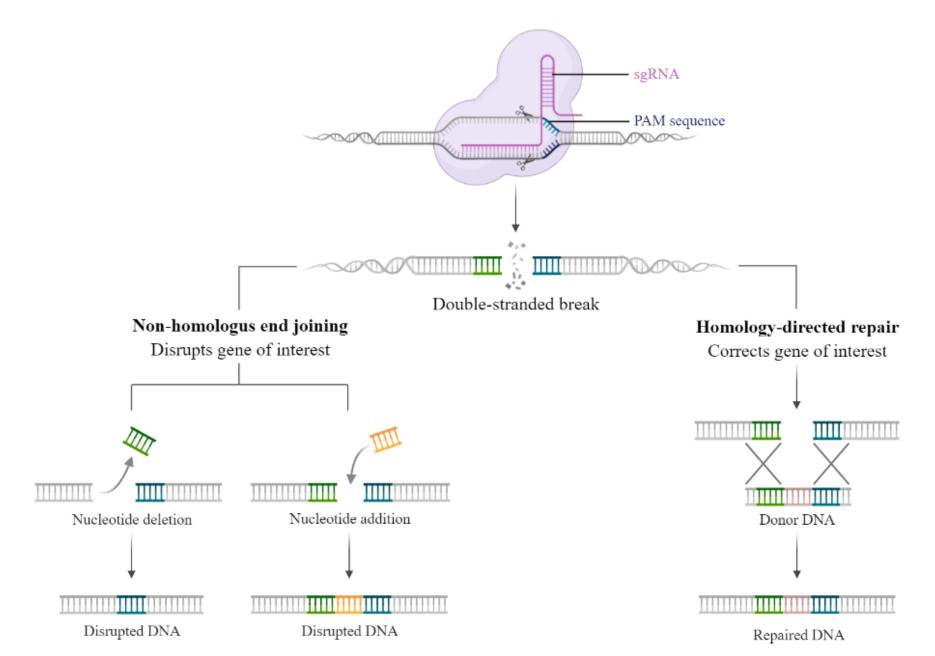
**CRISPR/CAS9** 

# What is CRISPR/Cas9?



This technique uses CRISPR and the Cas 9 protein nuclease to edit the genome.

### How does the CRISPR-CAS9 gene editing system work?



# Who discovered CRISPR/Cas9?

nature

Oct. 2020, Jennifer Doudna and Emmanuelle Charpentier receive Nobel Prize for discovery of CRISPR/Cas 9

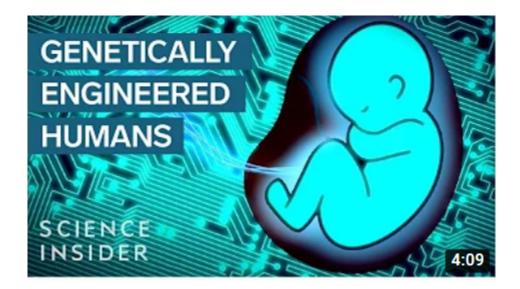
# Why has CRISPR revolutionized science?



It allows us to manipulate genetic sequences in any species

We can perform genome wide editing and screens

# What can CRISPR do in humans? movie



YouTube video: 4min

https://www.youtube.com/watch?v=kkvslrfaCLY

### What are the pros and cons of CRISPR/Cas9?

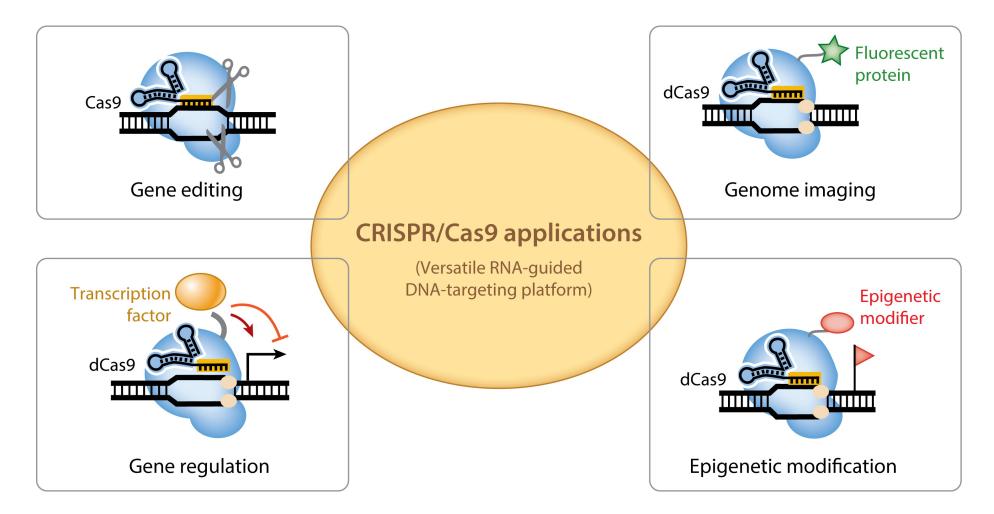


Advantages: Fast Affordable More accurate and efficient

### **Disadvantages**:

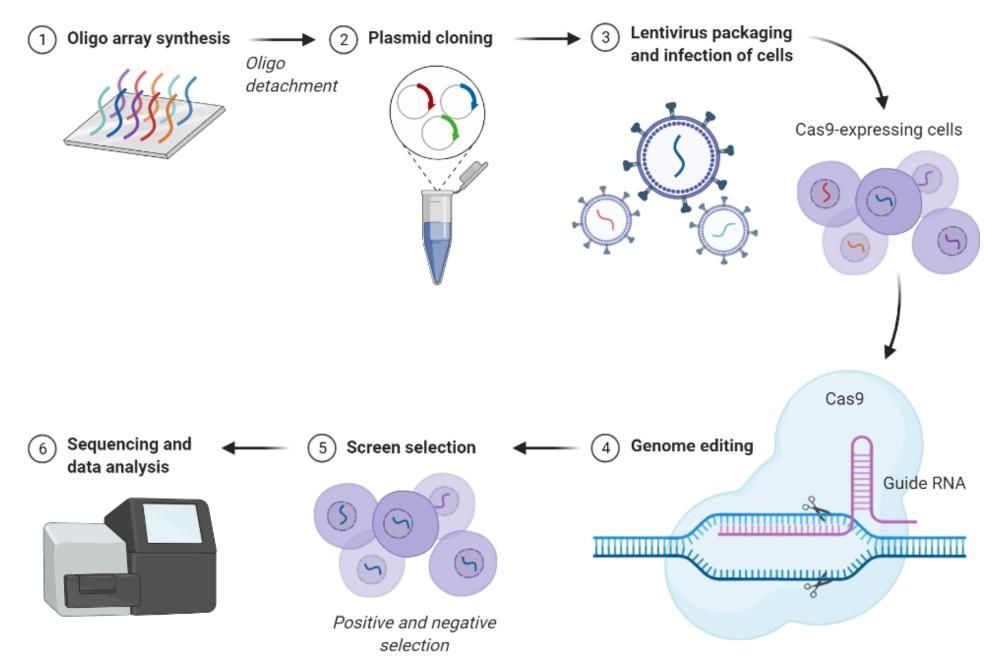
Can cause unintended mutations or loss of genomic sequence Unethical

# What are some other applications of CRISPR/Cas9?



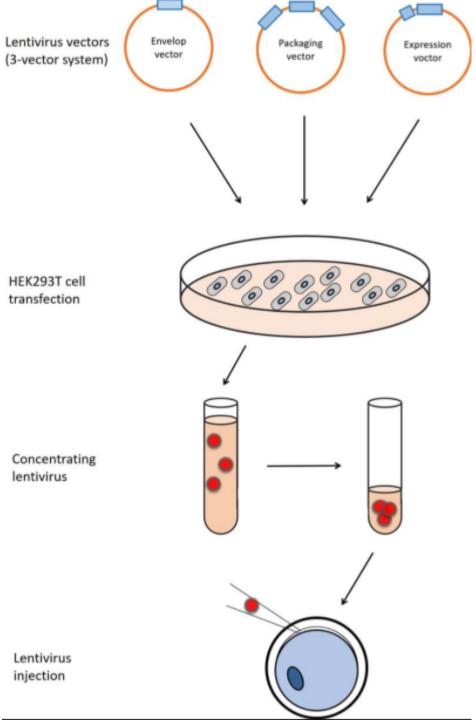
R Xiong X, et al. 2016. Annu. Rev. Genom. Hum. Genet. 17:131–54

## How can CRISPR be used in a genome-wide screen?

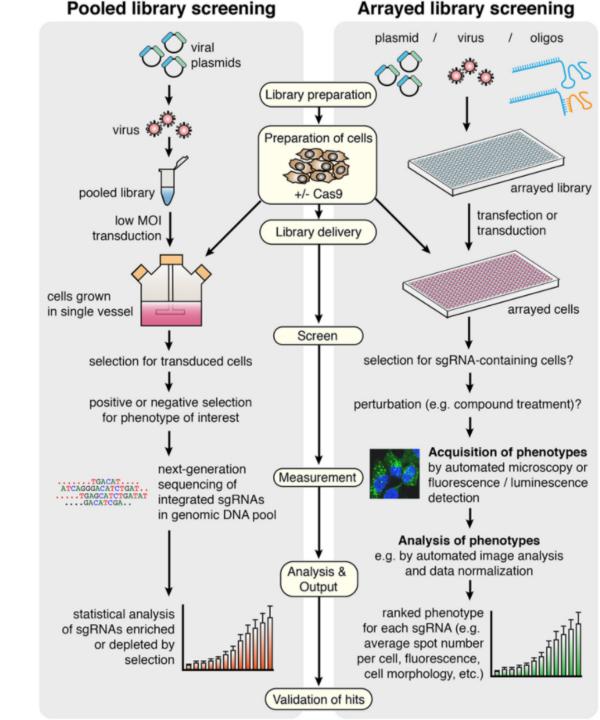


# What are lentiviruses and how are they to moved into the cell?

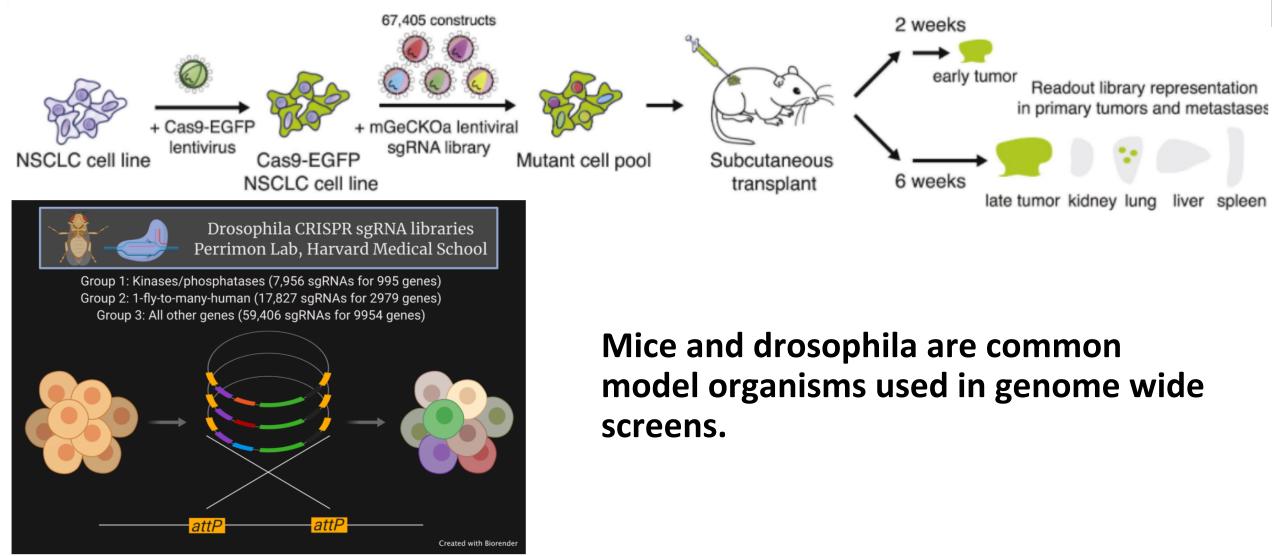




# There are two types of genome-wide CRISPR screens



## What model organisms do genome wide CRISPR screens use?



Perrimon Lab Drosophila CRISPR Pooled Libraries

### How has CRISPR been used to understand SARS-CoV-2 biology?

#### Cell Leading Edge

### A Crisp(r) New Perspective on SARS-CoV-2 Biology

#### Adam L. Bailey<sup>1,4,\*</sup> and Michael S. Diamond<sup>1,2,3,\*</sup>

<sup>1</sup>Department of Pathology & Immunology, Washington University School of Medicine, St. Louis, MO, USA <sup>2</sup>Department of Medicine, Washington University School of Medicine, St. Louis, MO, USA <sup>3</sup>Department of Molecular Microbiology, Washington University School of Medicine, St. Louis, MO, USA <sup>4</sup>Department of Pathology and Laboratory Medicine, University of Wisconsin-Madison, Madison, WI, USA <sup>\*</sup>Correspondence: albailey@wisc.edu (A.L.B.), diamond@wusm.wustl.edu (M.S.D.) https://doi.org/10.1016/j.cell.2020.12.003

Complementary genome-wide CRISPR-Cas9 screens performed by multiple groups reveal new insights into SARS-CoV-2 biology including aspects of viral entry, translation, replication, egress, and the genes regulating these processes. Comparisons with other coronaviruses enhances our understanding of the cellular life cycle of this medically important family of emerging viruses.

JOURNAL OF DRUG TARGETING https://doi.org/10.1080/1061186X.2020.1769637



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### CRISPR-cas systems based molecular diagnostic tool for infectious diseases and emerging 2019 novel coronavirus (COVID-19) pneumonia

Xiaohong Xiang<sup>a</sup>, Keli Qian<sup>b</sup>, Zhen Zhang<sup>c</sup>, Fengyun Lin<sup>a</sup>, Yang Xie<sup>a</sup>, Yang Liu<sup>a</sup> and Zongfa Yang<sup>a</sup>

<sup>a</sup>School of Pharmacy, Chongqing Medical and Pharmaceutical College, Chongqing, China; <sup>b</sup>Department of Infectious Disease, The Fifth People's Hospital of Chongqing, Chongqing, China; <sup>c</sup>Department of Clinical Laboratory, Chongqing General Hospital, Chongqing, China

#### ABSTRACT

Emerging infectious diseases, the persistent potential for destabilising pandemics, remain a global threat leading to excessive morbidity and mortality. The current outbreak of pneumonia caused by 2019 novel coronavirus (COVID-19) illustrated difficulties in lack of effective drugs for treatment. Accurate and rapid diagnostic tools are essential for early recognition and treatment of infectious diseases, allowing timely implementation of infection control, improved clinical care and other public health measures to stop the spread of the disease. CRISPR-Cas technology speed up the development of infectious disease diagnostics with high rapid and accurate. In this review, we summarise current advance regarding diverse CRISPR-Cas systems, including CRISPR-Cas9, CRISPR-Cas12 and CRISPR-Cas13, in the development of fast, accurate and portable diagnostic tests and highlight the potential of CRISPR-Cas13 in COVID-19 Pneumonia and other emerging infectious diseases diagnosis.

#### **ARTICLE HISTORY**

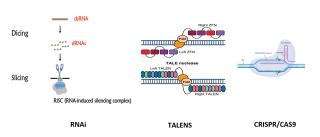
Received 19 March 2020 Revised 15 April 2020 Accepted 12 May 2020

#### KEYWORDS

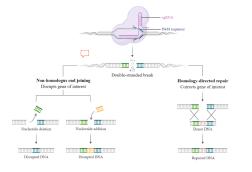
CRISPR-Cas; diagnostics; infectious diseases; COVID-19

# **CRISPR/Cas9 Summary**

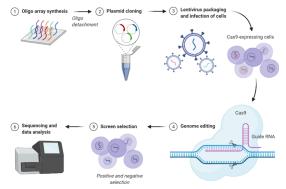
#### What are some ways to perform gene editing?



How does the CRISPR-CAS9 gene editing system work?



How can CRISPR be used in a genome-wide screen?



## Gene editing is used to modify DNA

CRISPR/Cas9 is a gene editing tool that targets and cleaves DNA to edit and regulate genes

CRISPR is used for genome-wide screens to analyze gene functions

# **Recent advancements?**

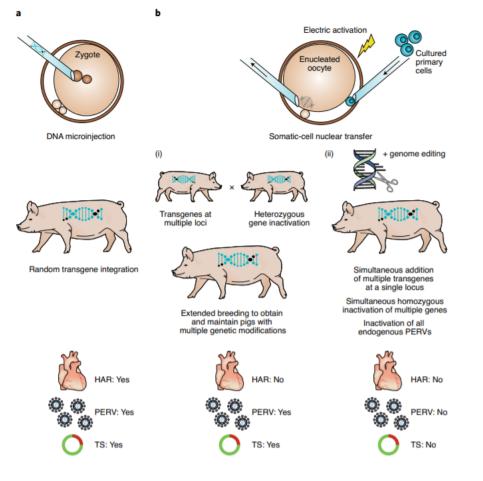
News & Views Published: 12 February 2021

## XENO-ORGAN TRANSPLANTATION **Extensively edited pigs**

Konrad Fischer & Angelika Schnieke 🖂

Nature Biomedical Engineering 5, 128–129(2021) Cite this article 207 Accesses 10 Altmetric Metrics

By using CRISPR and transposon constructs, pigs have been genetically modified to inactivate endogenous retroviruses and to enhance the compatibility of their organs with the human immune and coagulation systems.



CRISPR is being used to modify pigs so that their organs have more compatibility with human organs.

# References

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