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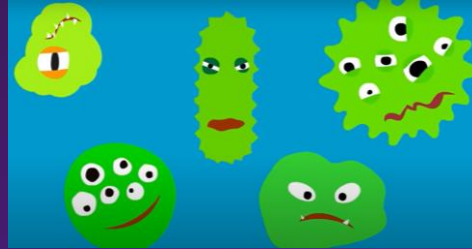
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# HOW DOES THE IMMUNE SYSTEM WORK?



Germs (bacteria/virus)

Our body is exposed to



Food and water  
we take

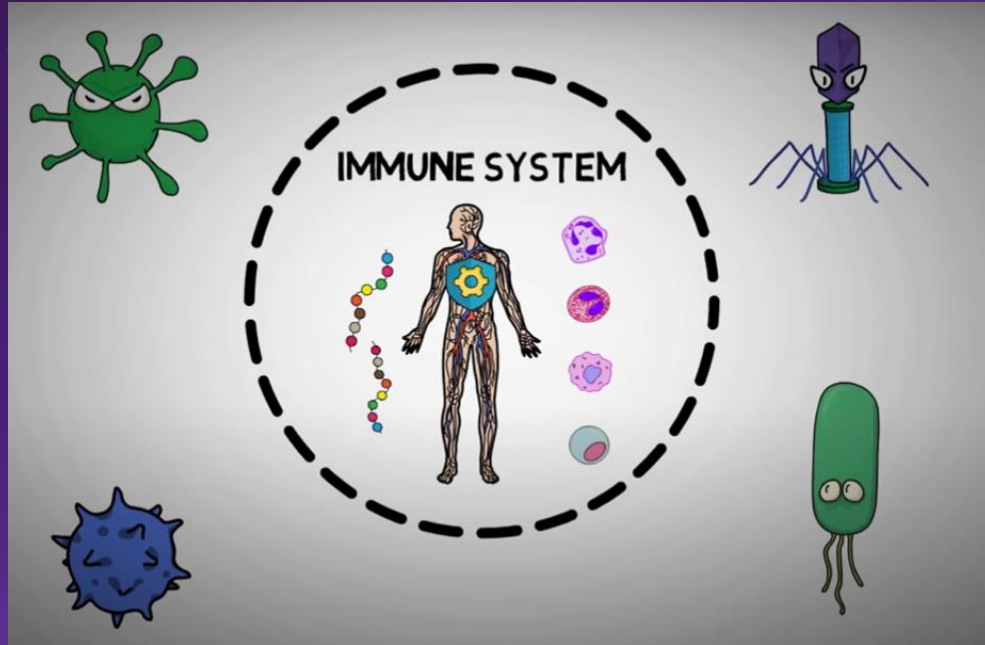


Air we breathe

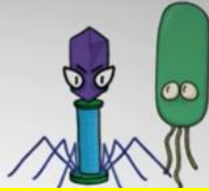


**But we do not fall sick**

# ALL THANKS TO OUR IMMUNE SYSTEM



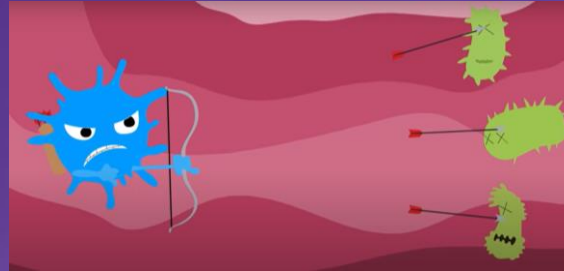
# INNATE IMMUNITY



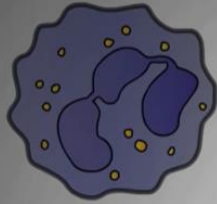
INNATE IMMUNITY

ACQUIRED IMMUNITY

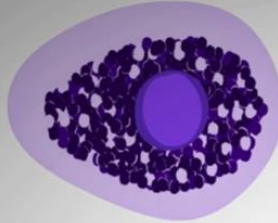
- Immunity we are born with
- Kill bacteria or virus in a non-specific way
- Cells either eat the germs or secrete chemicals to kill them



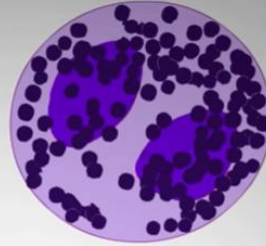
# INNATE IMMUNITY



NEUTROPHILS



MAST CELLS

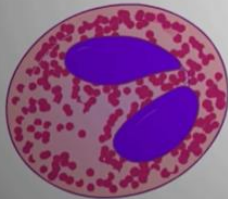


BASOPHILS

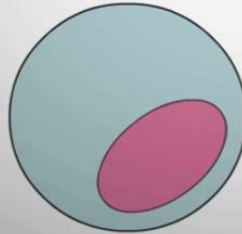


DENDRITIC CELLS

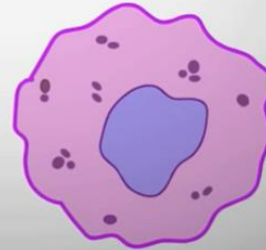
## CELLS OF INNATE IMMUNITY



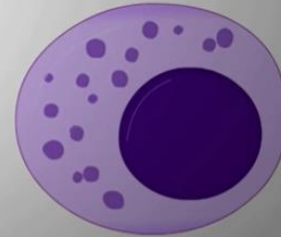
EOSINOPHILS



MONOCYTES



MACROPHAGE



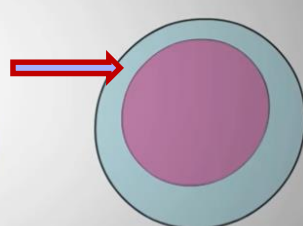
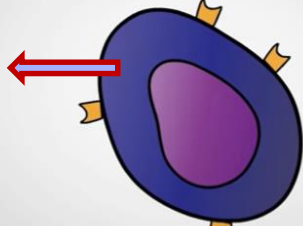
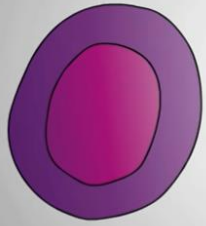
NATURAL  
KILLER CELLS

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# ADAPTIVE IMMUNITY

B LYMPHOCYTES

T LYMPHOCYTES



T-helper

T-Cytotoxic

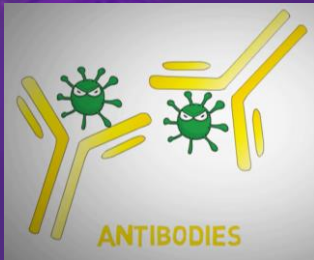
- Immunity we acquire following infection (adapts to the infection)
- Highly specific to the type of bacteria or virus causing infection
- Recognize specific proteins on the surface of germs (antigens)

Help B- and T- cells

Kill infected cells

Produce Antibodies

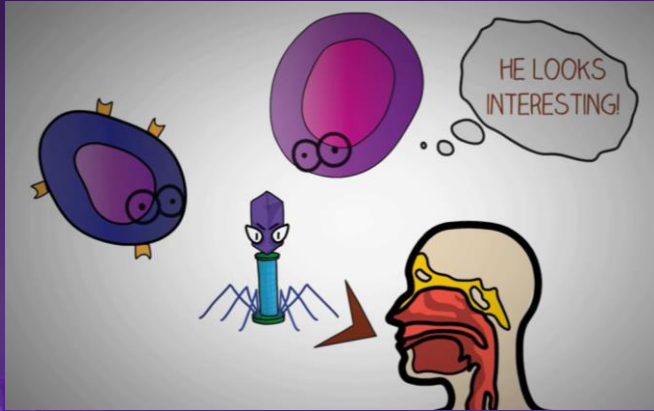
- Bind to the surface of virus/bacteria and blocks it
- Help innate immune cells to find and eat them



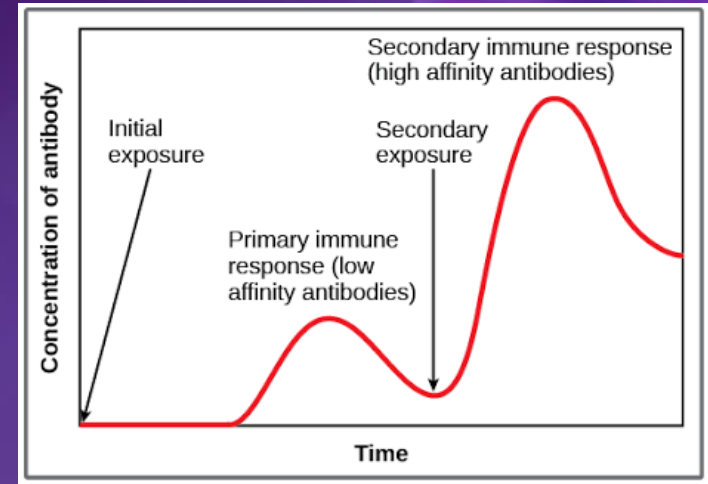


# ADAPTIVE IMMUNITY (RETAIN MEMORY) – THE BASIS OF VACCINATION

## 1<sup>ST</sup> TIME EXPOSURE



- **Body produce B- and T-cell response**
- **But is usually slow as it is still adapting**
- **Also retain memory of the virus/bacteria**
- **When challenged again with same antigen, body mounts a fast and strong immune response**



The secondary (and tertiary and subsequent) immune response is: **FASTER, GREATER and STRONGER**

# PRACTICE MAKES PROGRESS

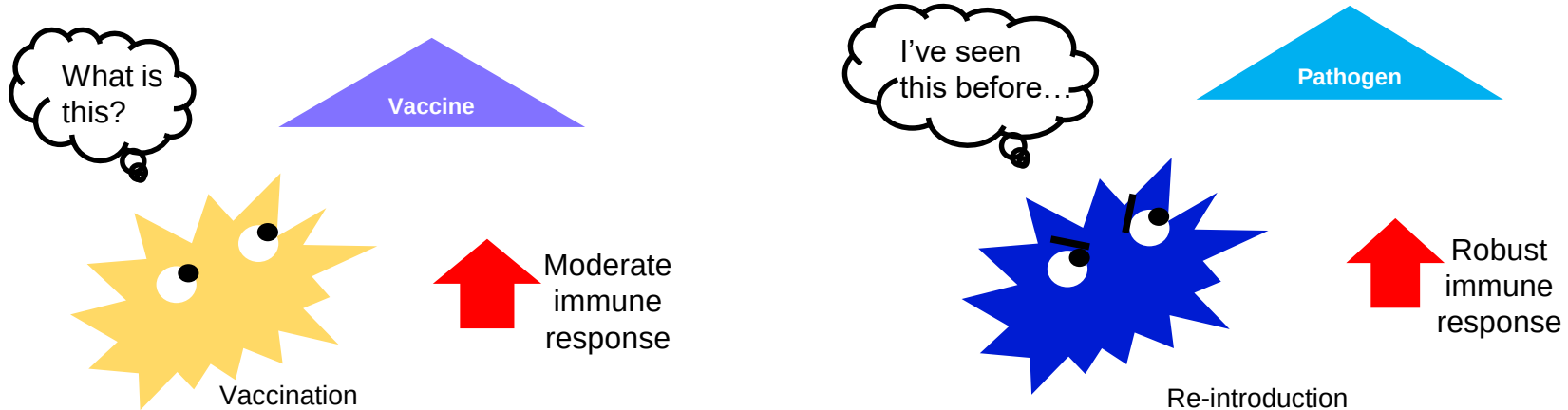
- Practicing is a great way to learn a new skill or reach a new goal
  - Learning an instrument
  - Running a set distance
  - Scoring high marks on a test
  - Landing that job
  - Practice makes perfect





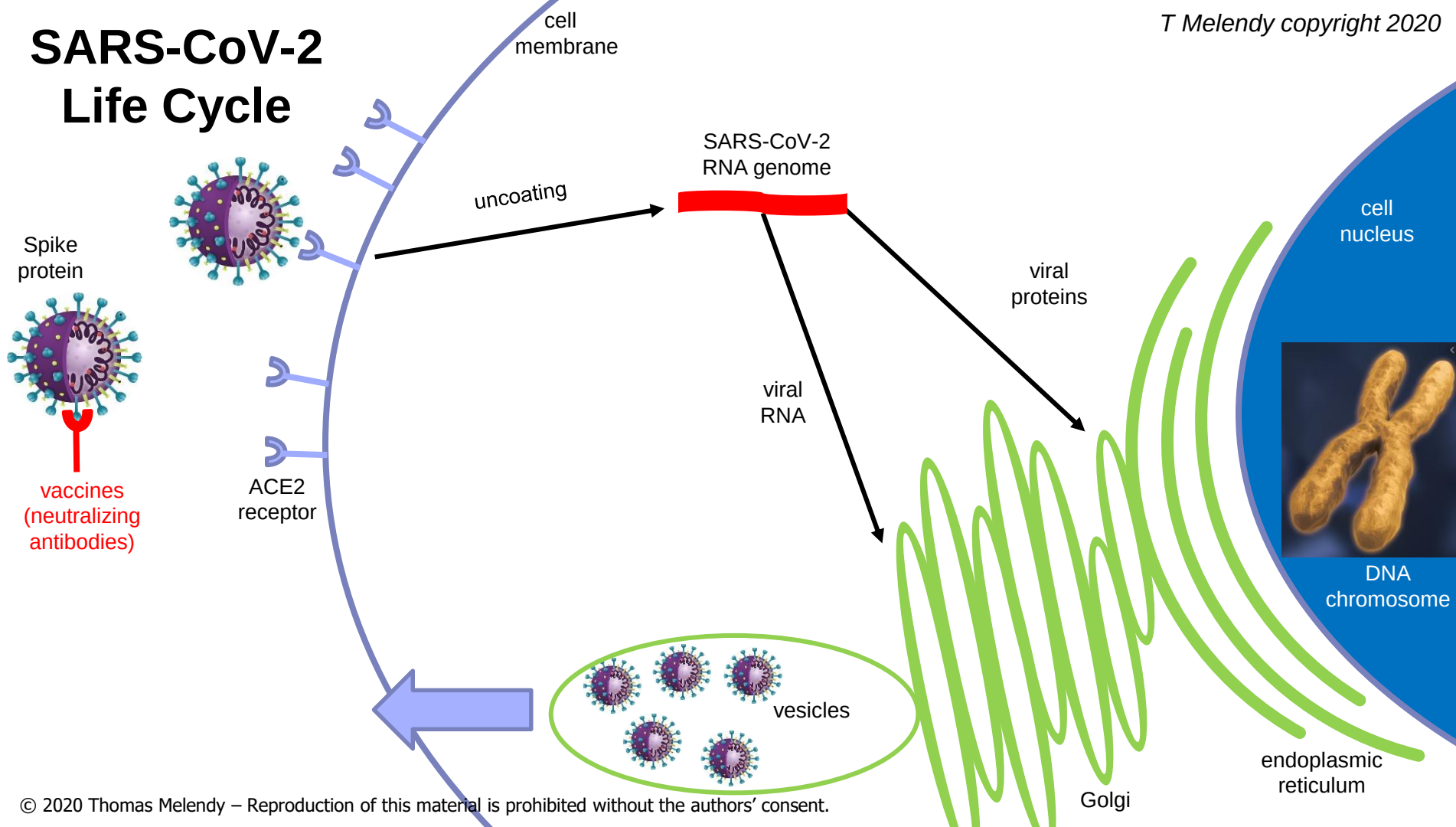
# VACCINATION IS PRACTICE FOR THE IMMUNE SYSTEM – WITHOUT GETTING SICK!

- Controlled exposure to an exogenous molecule to illicit a mild immune response
  - Molecule may be
    - Weakened/inert form of the pathogen
    - Protein expressed by the pathogen
  - Trains the immune system to more rapidly recognize the pathogen in the future



# SARS-CoV-2 Life Cycle

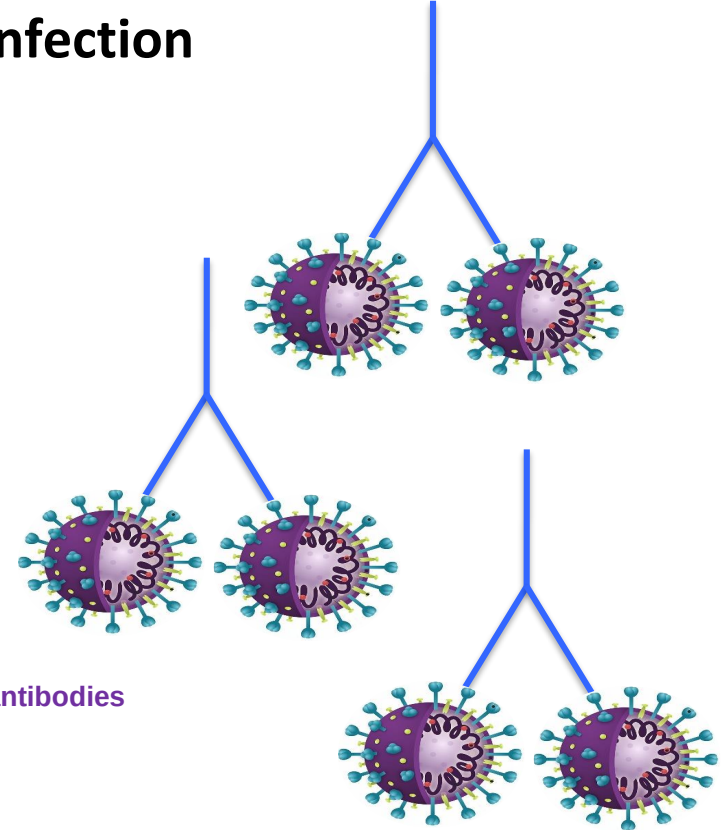
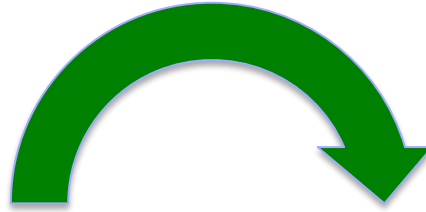
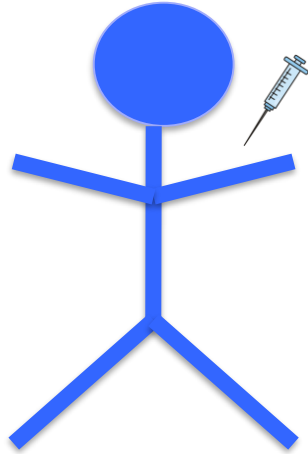
T Melendy copyright 2020



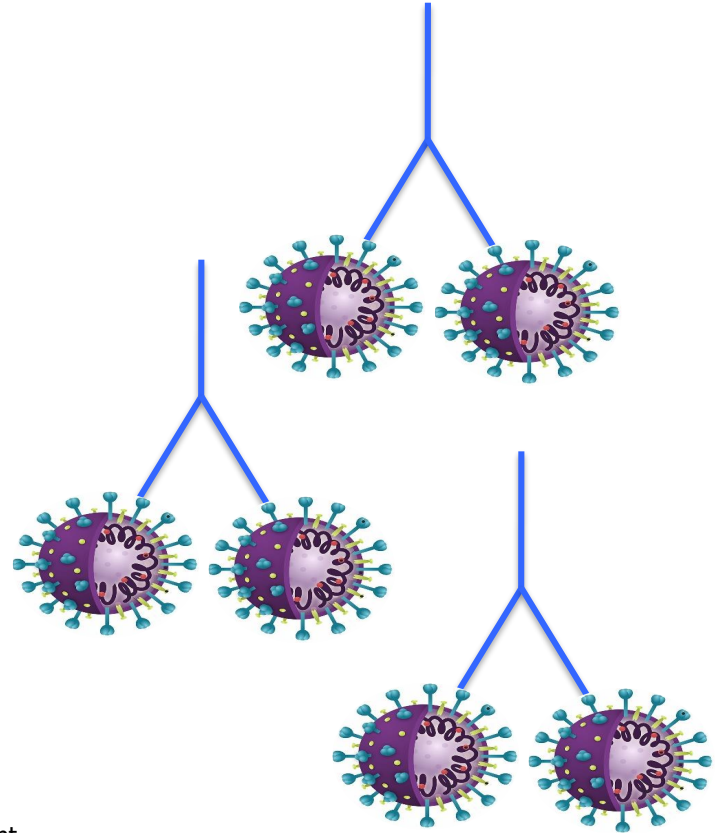
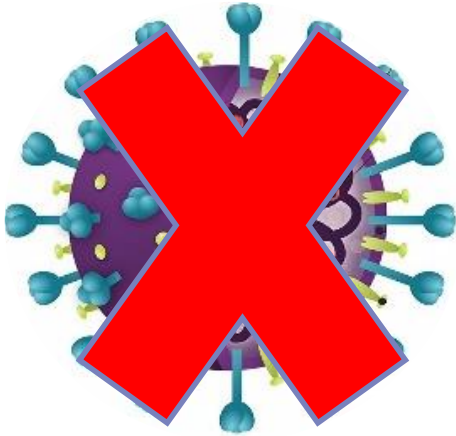
All vaccines work in the same general way – they present a target for the immune system, to “educate” the immune system to recognize a pathogen.

Many different “targets” can be used.

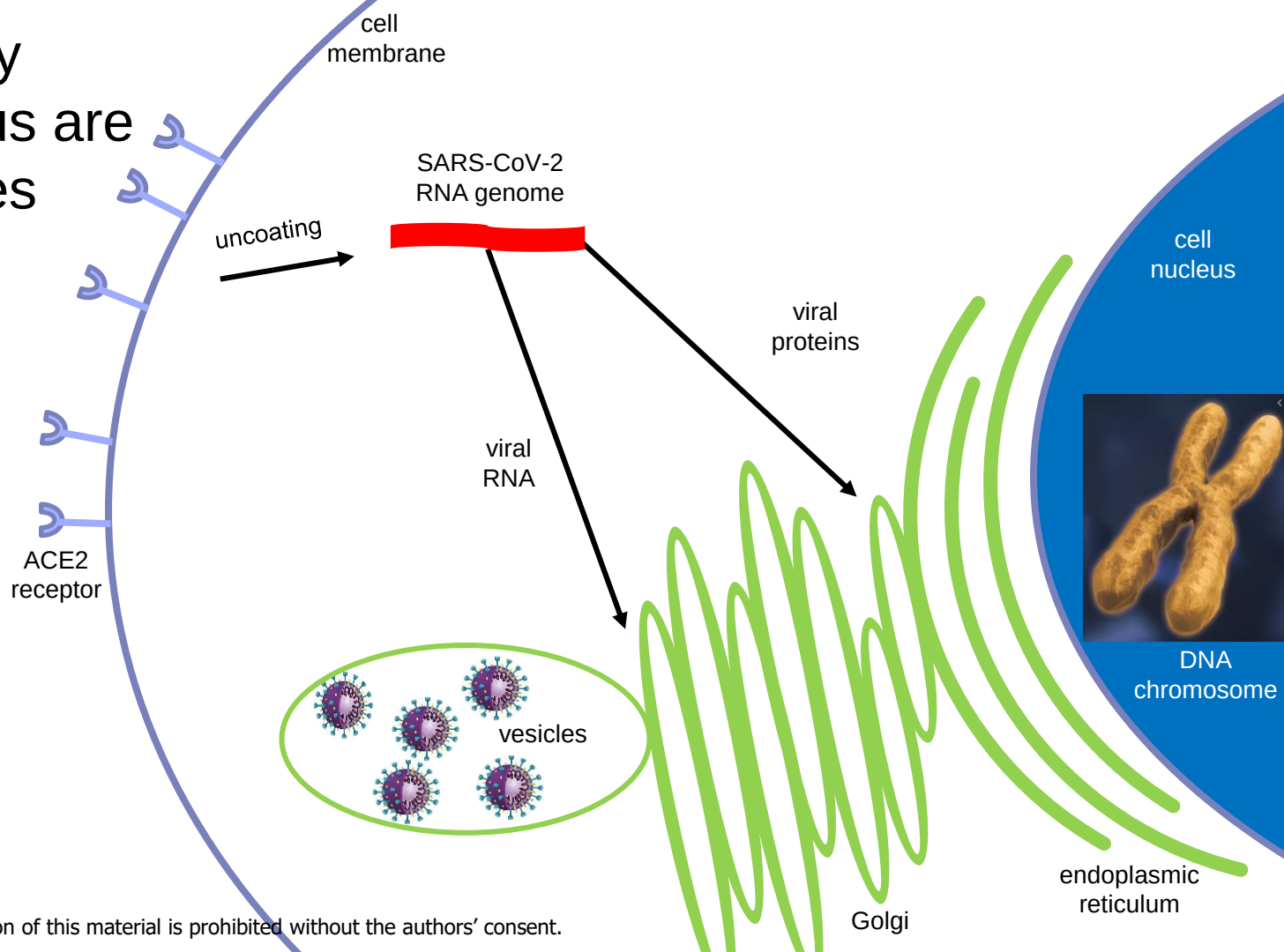
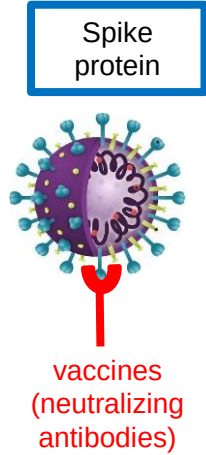
**Vaccines against SARS-CoV-2 train your immune system to recognize the virus, or parts of it, to produce protective antibodies that prevent infection**



**Sometimes inactivated or attenuated virus is used in vaccines to produce an immune response**



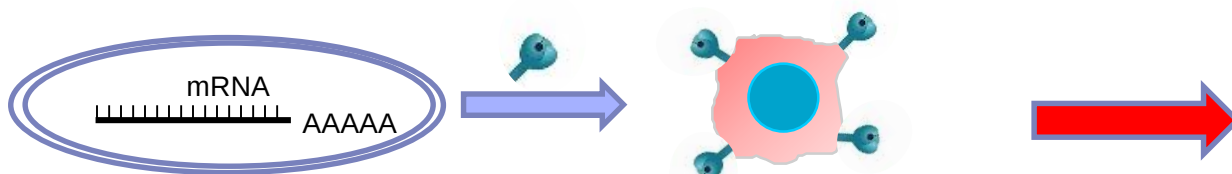
# Sometimes only parts of the virus are used in vaccines



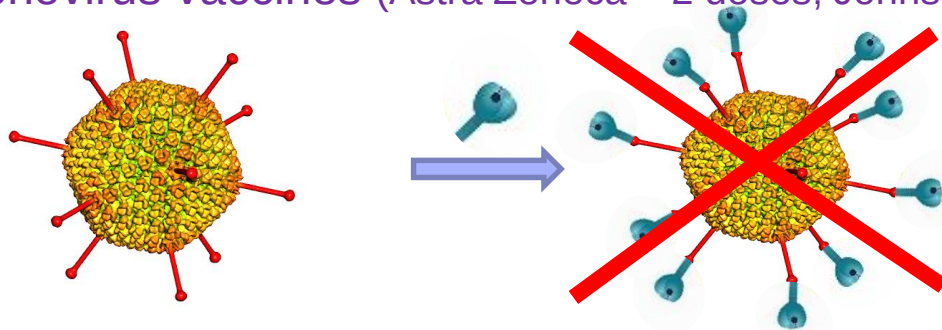


# Different approaches to display the spike protein for the immune system

mRNA vaccines (Pfizer and Moderna – 2 doses)



Adenovirus vaccines (Astra Zeneca – 2 doses; Johnson & Johnson – 1 dose)

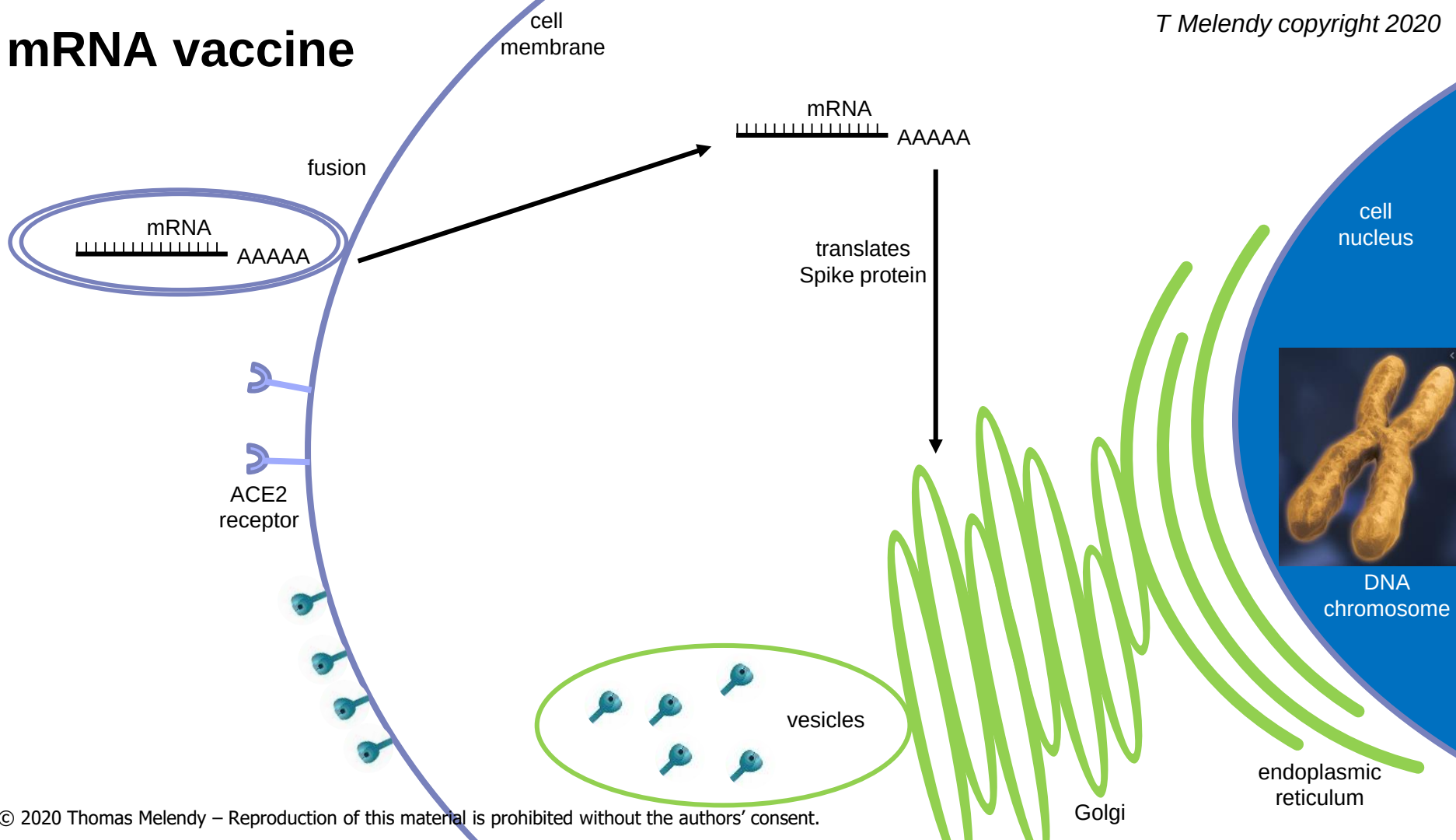


Recombinant protein vaccine (Novavax – 2 doses)



# mRNA vaccine

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# HERD IMMUNITY

Indirect protection of population

- If enough people are immune
- Reduces chance others will get infected
- For COVID-19, estimated:
  - 70-90% of population will need to be vaccinated
  - NY state is at 1% fully vaccinated



Healthy,  
Not immunized



Sick,  
Not immunized

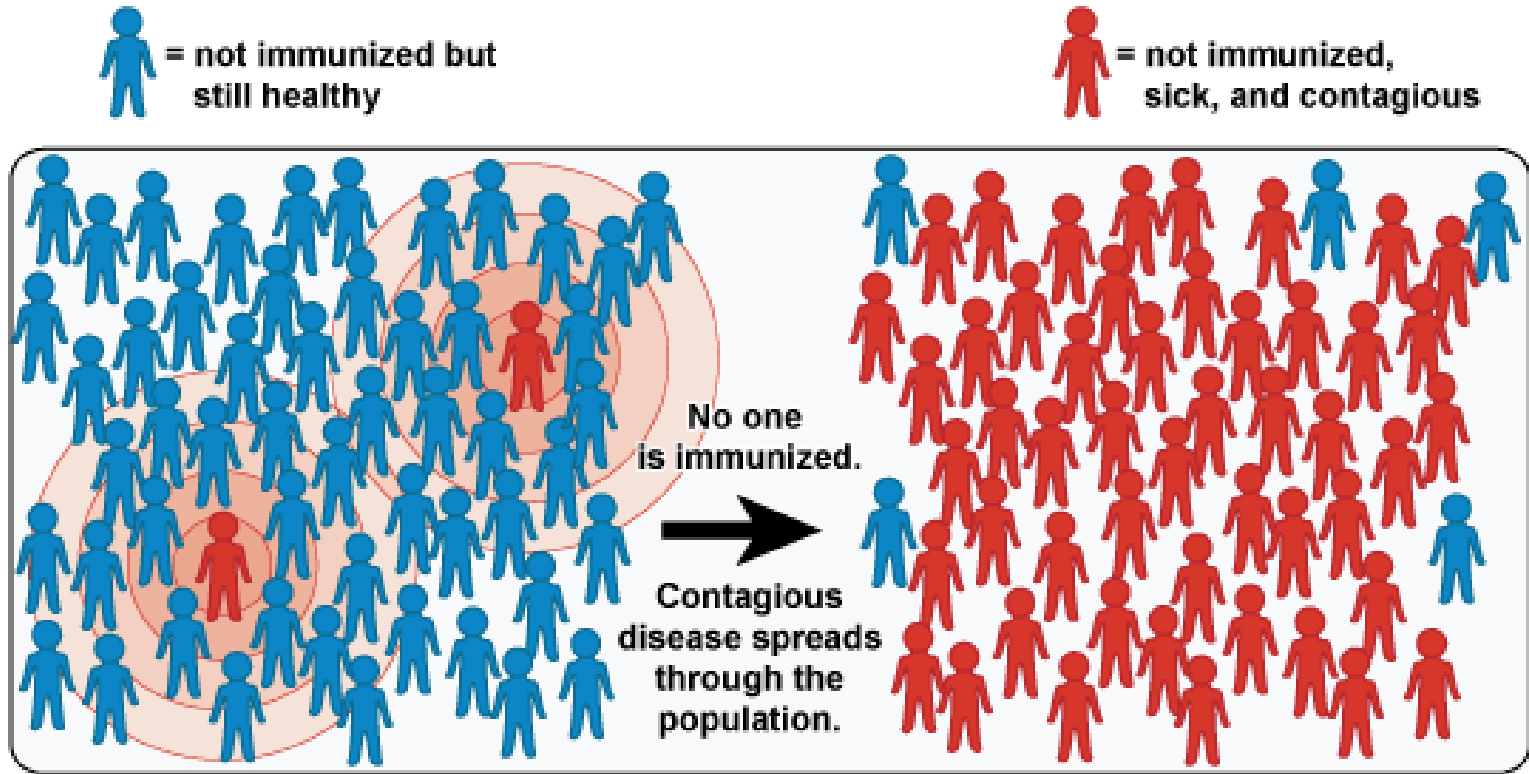


Healthy,  
Immunized

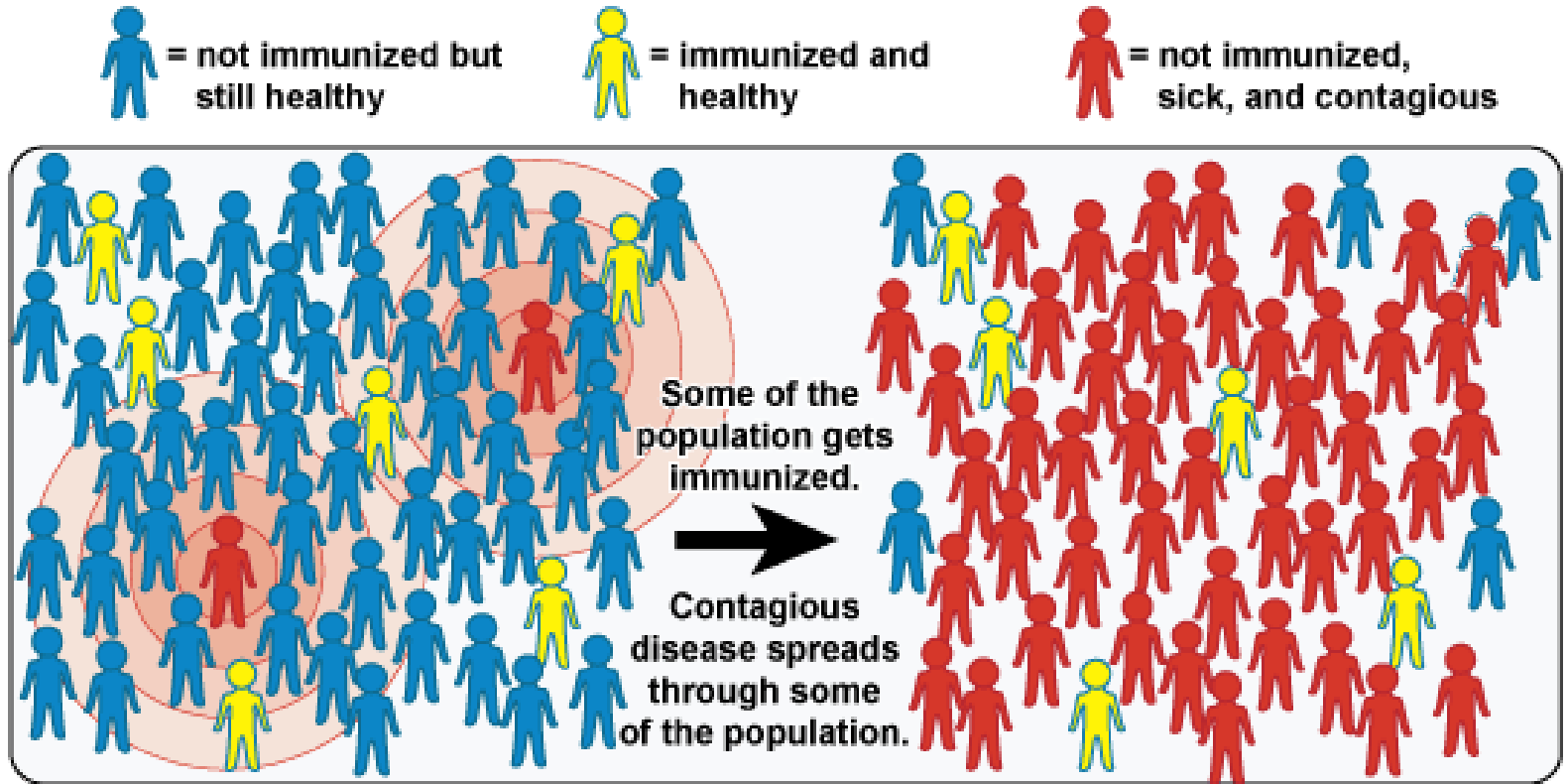
For COVID-19, herd immunity can protect:

- Kids
- Vaccinated, but vaccine didn't work
- Others who didn't get vaccine

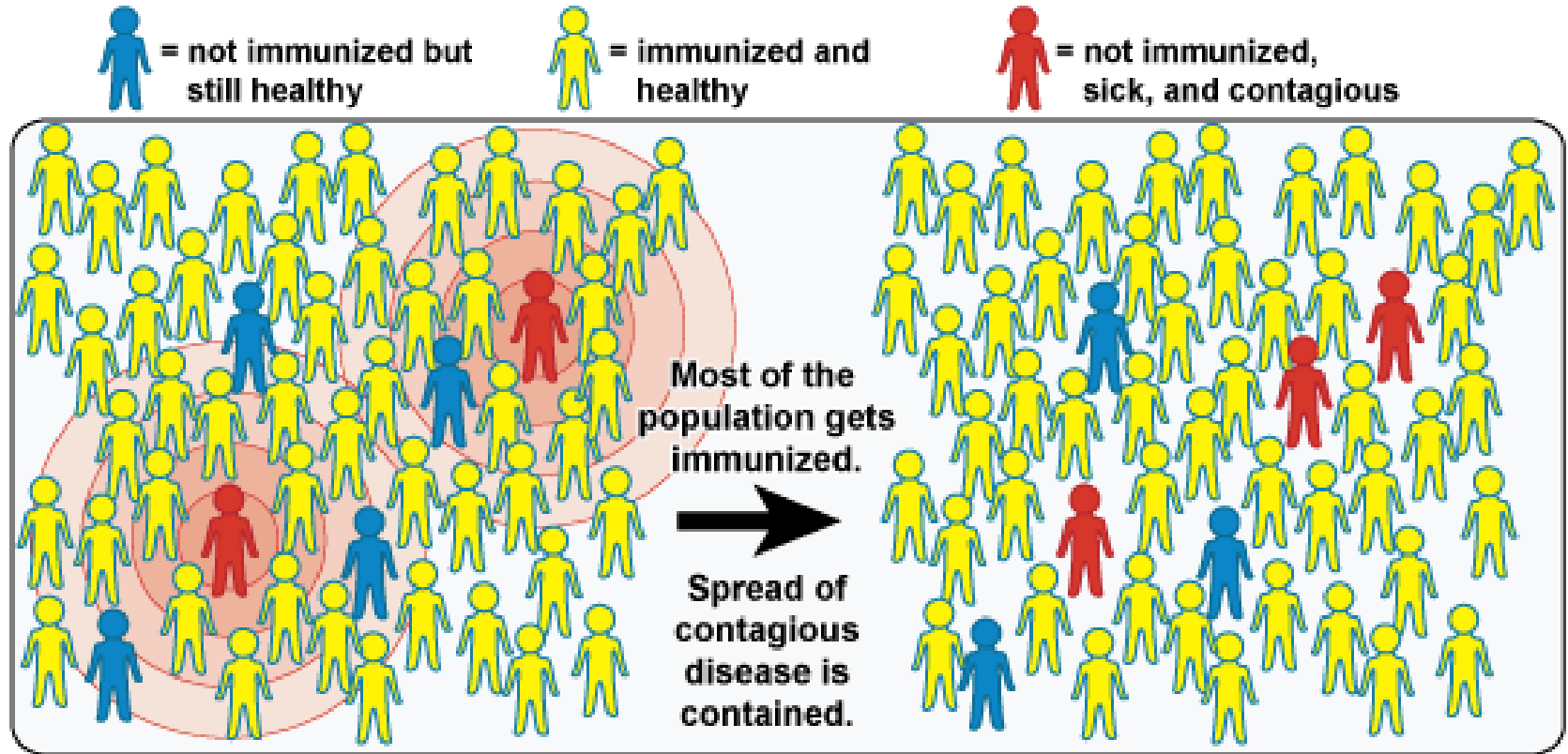
# Unvaccinated population



# Partially vaccinated population



# With widespread immunization, HERD IMMUNITY protects the non-immune!



# CONCERNS ABOUT VACCINES AGAINST SARS-COV-2

- mRNA will integrate into my DNA
- There are fertility concerns with the vaccines
- The side effects aren't worth it
- The vaccines won't work in older people
- The vaccines were rushed – how is it possible to have had this happen so quickly?
- The variants we are hearing about will make the vaccines moot

# POSSIBLE WITH SIGNIFICANT FUNDS TO ALLOW PARALLEL TRACKS OF SCIENCE AND PRODUCTION BACKED BY DECADES OF BASIC SCIENCE RESEARCH

| Date          | Milestone                                                        |
|---------------|------------------------------------------------------------------|
| Dec 1         | Covid-19 illness documented (unpublicized Nov 17 <sup>th</sup> ) |
| Jan 10        | SARS-CoV-2 virus sequenced                                       |
| Jan 15        | NIH designs mRNA vaccine in collaboration with Moderna           |
| Mar 16        | Moderna Phase 1/2 trial begins                                   |
| May 2         | Pfizer/BioNTech Phase 1/2 trial begins                           |
| July 14       | Moderna Phase 1/2 trial published in NEJM                        |
| July 27, 28   | Moderna and Pfizer/BioNTech Phase 3 trial begins                 |
| Aug 12        | Pfizer/BioNTech Phase 1/2 published in Nature                    |
| October 22,27 | Enrollment in both Phase 3 trials complete; >74,000 participants |
| Nov 9         | Pfizer/BioNTech announces interim analysis efficacy > 90%        |
| Nov 16        | Moderna announces interim analysis efficacy 94.5%                |
| Nov 18        | Pfizer/BioNTech announces 95% efficacy as final result           |
| Nov 20        | 1 <sup>st</sup> EUA submitted by Pfizer/BioNTech                 |
| Nov 27        | Distribution of vaccine by UAL charter flights throughout US     |
| Dec 10        | FDA External review of Pfizer/BioNTech EUA                       |
| Dec 11        | Phase 1a Vaccination begins for health care professionals*       |

\*Provisional on positive external review



# Phylogeny

Clade ^

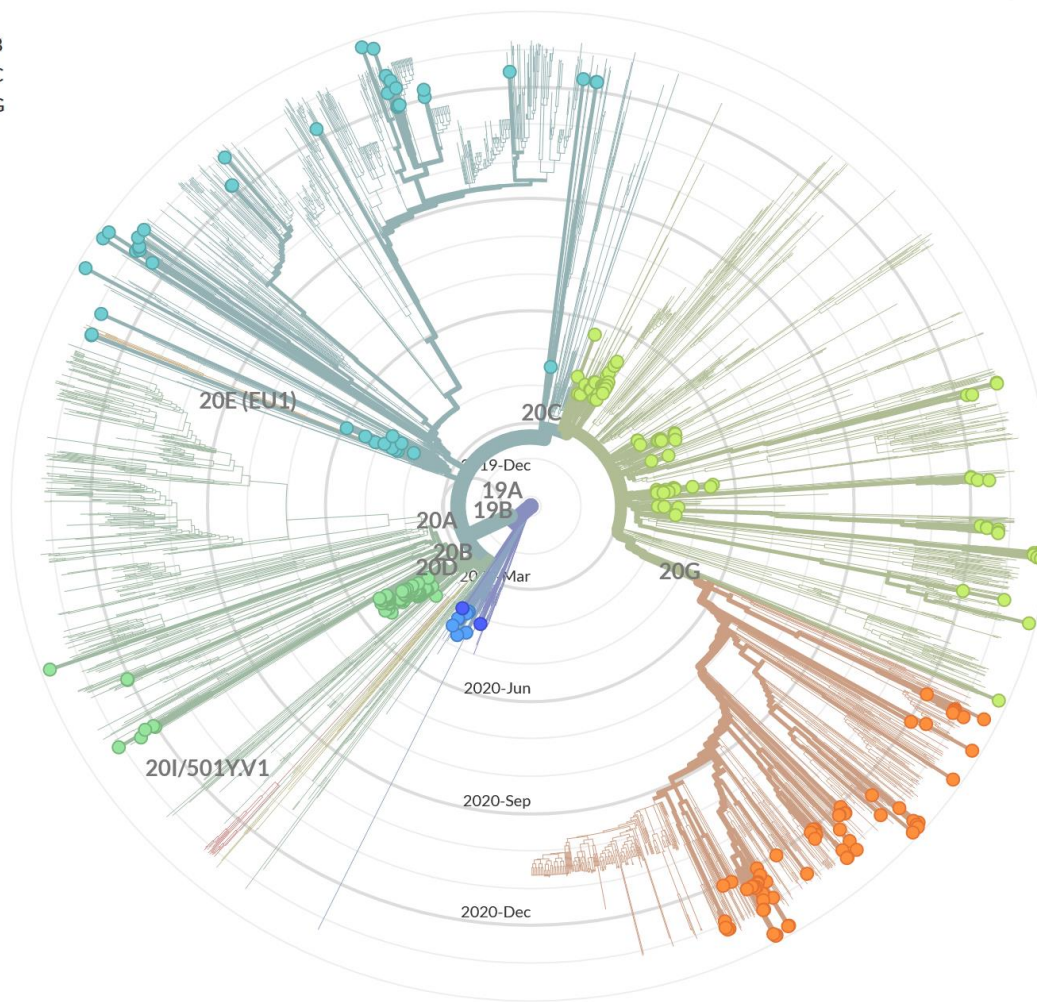
19A  
19B  
20A

20B  
20C  
20G

ZOOM TO SELECTED

RESET LAYOUT

## Variants in NYS



# HOW THE KNOWN VARIANTS OF CONCERN AFFECT COVID-19 AND VACCINES

|                                                                 | B.1.1.7                                                              | B.1.351                                                                                               | P.1                                                   |
|-----------------------------------------------------------------|----------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------|-------------------------------------------------------|
| Alternate name                                                  | 501Y.V1                                                              | 501Y.V2                                                                                               | 501Y.V3                                               |
| Country identified                                              | United Kingdom                                                       | South Africa                                                                                          | Brazil                                                |
| Mutations                                                       | 23                                                                   | 21                                                                                                    | 17                                                    |
| Spike mutations                                                 | 8                                                                    | 9                                                                                                     | 10                                                    |
| Key RBD, spike mutations beyond <b>N501Y in all</b>             | E69/70 deletion, P681H 144Y deletion, A570D                          | <b>E484K, K417N, orf1b deletion</b>                                                                   | <b>E484K, K417T, orf1b deletion</b>                   |
| Other mutations, including N-terminal                           | T7161, S982A, D1118H                                                 | <b>L18F</b> , D80A, D215G, Δ242-244, R264I, A701V                                                     | <b>L18F</b> , T20N, P26S, D138Y, R190S, H655Y, T10271 |
| Transmissibility Δ                                              | <b>&gt;50% increased</b>                                             | No                                                                                                    | Not established                                       |
| Lethality Δ                                                     | <b>Not resolved</b>                                                  | ?                                                                                                     | ?                                                     |
| Immune evasion                                                  | Unclear                                                              | Yes                                                                                                   | Yes, less than B.1.351                                |
| Vaccine efficacy (preserved vs severe infections in all so far) | <b>Modest reduction ~10% point decline in 2 trials (Novavax, AZ)</b> | <b>Yes, reduced in 2 (J&amp;J, Novavax ~20-30% point decline. No efficacy v mild infections w/AZ)</b> | <b>Preserved in J&amp;J trial</b>                     |
| Countries reported                                              | 94                                                                   | 48                                                                                                    | 25                                                    |
| US States reported                                              | 46                                                                   | 17                                                                                                    | 5                                                     |