## Vaccine Chat or: How I Learned to Stop Worrying and Love Adaptive Immunity

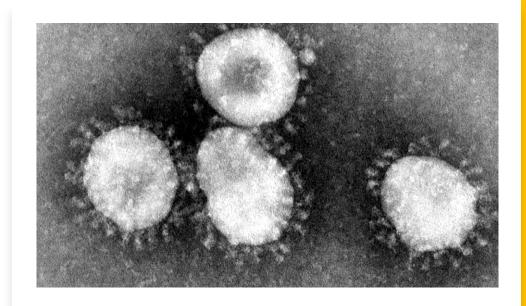
#### Talk Overview

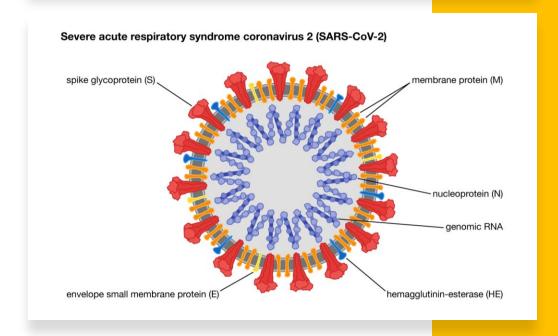
- Background on COVID-19
- COVID-19 Mitigations
- How Vaccines Work
- Types of Vaccines
- COVID-19-Specific Vaccines
- COVID-19 Vaccine Side Effects and a Word on Variants

# COVID-19 Background & Mitigations

#### What is COVID-19?

- SARS-CoV-2 (Severe Acute Respiratory Syndrome Coronavirus 2) is the causative agent of COVID-19 (COronaVIrus Disease 2019)
- Coronaviruses were discovered in the 1960s.
  - Named for appearance under microscope
  - Make up 30% of viruses that cause the common cold
  - Usually present very mild symptoms, resolve in days





### Serious Coronavirus Outbreaks in History

#### SARS-CoV (Severe Acute Respiratory Syndrome)

- Outbreak occurred Nov.
   2002 July 2003,
   primarily in China
- ~8000 cases, 774 deaths
   (9.6% mortality rate)
- No new cases since 2004

#### MERS-CoV (Middle East Respiratory Syndrome)

- Outbreak began June
   2012 in Saudi Arabia
- ~2500 cases worldwide,
   862 deaths (34%
   mortality rate)
- 200-300 new cases every year

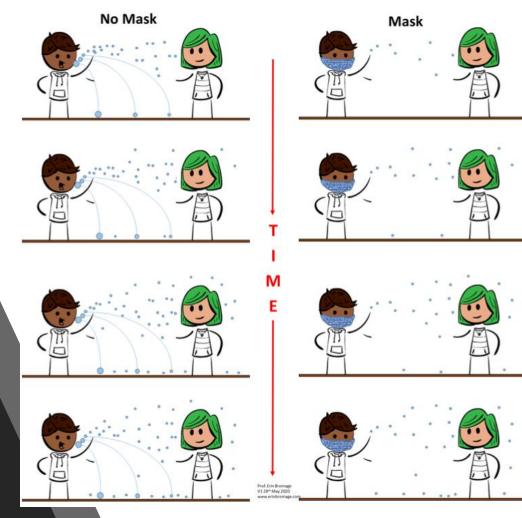
### SARS-CoV-2 (COVID-19)

- Outbreak began
   December 2019 in China
- Millions of worldwide cases, hundreds of thousands of deaths (true mortality rate unknown)
- Thousands of new cases and deaths every day

### COVID-19 Mitigations

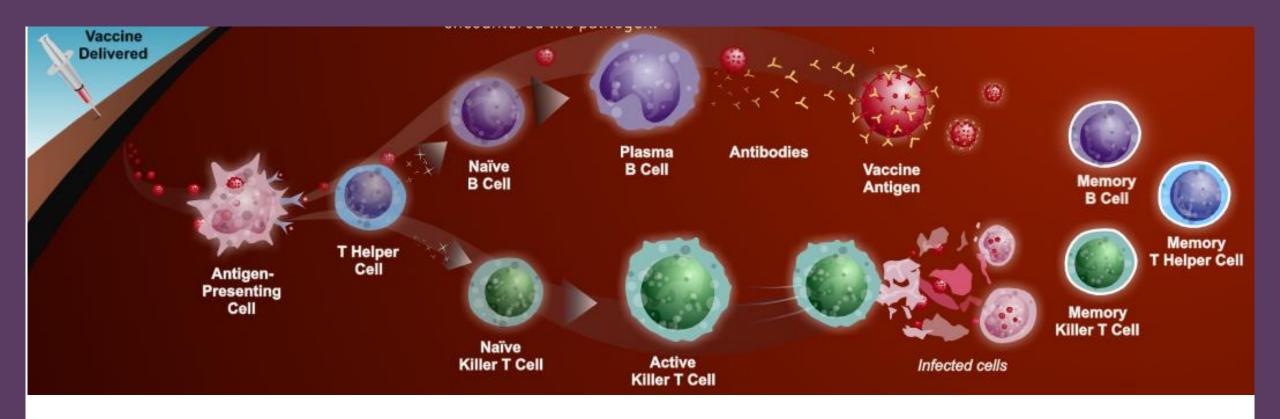
#### Successful infection = Virus Exposure x Time

- Masks
  - Masks can catch virtually all droplets and ~50% of droplet nuclei
  - Masking is a "social contract"
  - Simulations show high rate of mask usage could significantly decrease infections and deaths
- Shared Spaces
  - Minimize time in shared spaces -> minimize chances of viral infection
  - Minimize interactions with large groups
- Hygiene
  - Wash/sanitize hands frequently
  - Do not touch your face with unclean hands!





### Vaccine Background



### How Do Vaccines Work?

### Types of Vaccines

Different **types of vaccines** introduce people to harmless parts of the virus in different ways

some vaccines introduce "pre-made" viral pieces killed virus weakened viral protein virus-like

eakened viral protein virus-like virus (premade) particles





**mRNA** 





aka inactivated aka live,
virus attenuated virus

other vaccines introduce genetic instructions for making viral proteins (viral genes) but rely on the person's cells to do the actual viral protein making - but differ in how they deliver the gene

protein

#### viral vector

stick a coronavirus gene in a different, harmless, virus



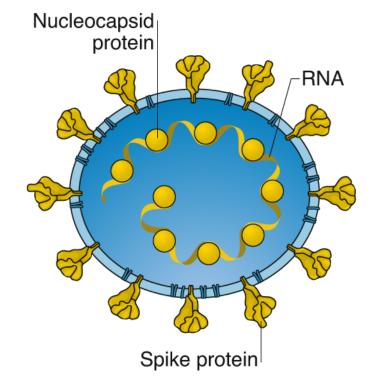
stick a coronavirus gene in a small circular piece of DNA called a plasmid & deliver it to cells (e.g. by electroporation (using electricity))

the cells then makes messenger RNA copies of this recipe from which ribosomes can make protein

or you can give the mRNA directly (e.g. enclosed in oily membranes (Lipid NanoParticles)



#### SARS-CoV-2



COVID-19 Vaccines – How They Work, Side Effects, and Efficacy

#### Moderna's mRNA Vaccine Approach Closely mimics a native viral infection leading to B and T cell responses Muscle cell Formulation Viral antigens MHCI ncoded mRNA Muscle cell Nucleus Specialized antigen presenting cell (APC) in lymph node Exogenous viral antigen Muscle cell CD4' T cell moderna Nucleus APC cell © 2017 Moderna Therapeutics

Specific
Example –
mRNA-1273
(Moderna –
Pfizer's
vaccine works
the same way)

# COVID-19 Vaccines Myths

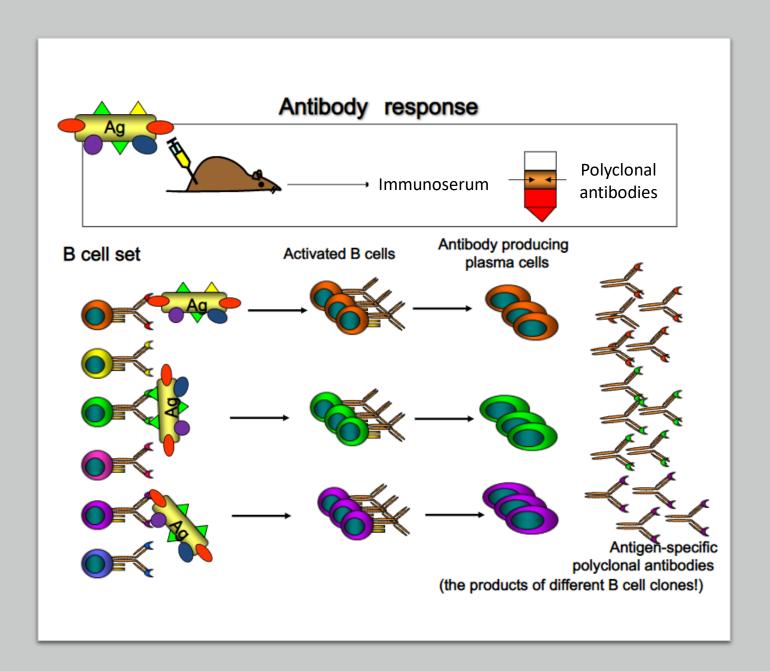
- COVID-19 vaccines can give a person COVID-19
  - Myth: These vaccines contain only a piece of the viral particle, not the full virus; infection with SARS-CoV-2 from the vaccine is not possible
- mRNA vaccine technology is brand-new and was rushed out in an unsafe manner
  - Myth: mRNA vaccines have been under development for ~30 years, and they have been trialed in humans for diseases like rabies, Zika, influenza, and cytomegalovirus
- There is a risk of the genetic material contained within mRNA vaccines of interfering with our DNA/proteins/etc. long-term
  - Myth: Without highly specialized enzymes, RNA cannot interfere with DNA; further, mRNA is transient, and both it and the protein it encodes are degraded by our cells in a number of days
- It's a bad thing when vaccine trials are paused to examine adverse events
  - Myth: All clinical trials have the potential to produce adverse events; it would be worrying if the trials were NOT paused to examine them

### Vaccine Side Effects?

- Both Pfizer and Moderna have released safety and efficacy data to the FDA, which released it to the public
- Each vaccine will require two doses spaced roughly 3-4 weeks apart
  - Booster is required to see ~10-fold increase in antibodies against SARS-CoV-2
  - 95% efficacy for protection at Day 28 post-first dose (assuming two doses given)

- Those who are older (>55) tend to experience less/more mild side effects than those who are younger
- SEVERE side effects are vanishingly rare, like other vaccines (e.g. flu)
- Side effects typically last 12-24 hours and resolve rapidly

Side Effect Comparison	Pfizer (BNT162b2)	Moderna (mRNA-1273)
Side Effect #1 (Dose 1/Dose 2/Placebo 2)	Fatigue ( <b>47.4%/59.4%/22.8%</b> )	Fatigue (38.5%/67.6%/24.5%)
Side Effect #2 (Dose 1/Dose 2/Placebo 2)	Headache ( <b>41.9%/51.7%/24.1%</b> )	Headache (35.4%/62.8%/25.4%)
Side Effect #3 (Dose 1/Dose 2/Placebo 2)	Muscle Pain (21.3%/37.3%/8.2%)	Chills (9.2%/48.3%/5.9%)
Side Effect #4 (Dose 1/Dose 2/Placebo 2)	Chills (14.0%/35.1%/3.8%)	Joint Pain (16.6%/45.2%/10.5%)
Side Effect #5 (Dose 1/Dose 2/Placebo 2)	Joint Pain (11.0%/21.9%/5.2%)	Nausea ( <b>9.4%/21.3%/7.3%)</b>



### SARS-CoV-2 Variants and the Vaccines

### Questions?

