

# Immune responses to COVID-19 vaccines in Hong Kong

Ben Cowling

Co-Director, WHO Collaborating Centre for Infectious Disease Epidemiology and Control  
School of Public Health, The University of Hong Kong, Hong Kong

23 November 2021

# COVID-19 vaccinations

- **Immunity can be acquired by infection or vaccination.** Infections tend to spread until population immunity reaches a high level, but surges in COVID-19 transmission would threaten to overwhelm our healthcare system and cause many thousands of local deaths (infection in unvaccinated individuals is 10-20 times more serious than influenza, and no pre-existing population immunity to COVID-19 prior to 2020)
- Increasing our population immunity through vaccines provides a pathway back to “normal life”, eventually with no further need for face masks, social distancing policies, and other public health measures

# Overarching aims

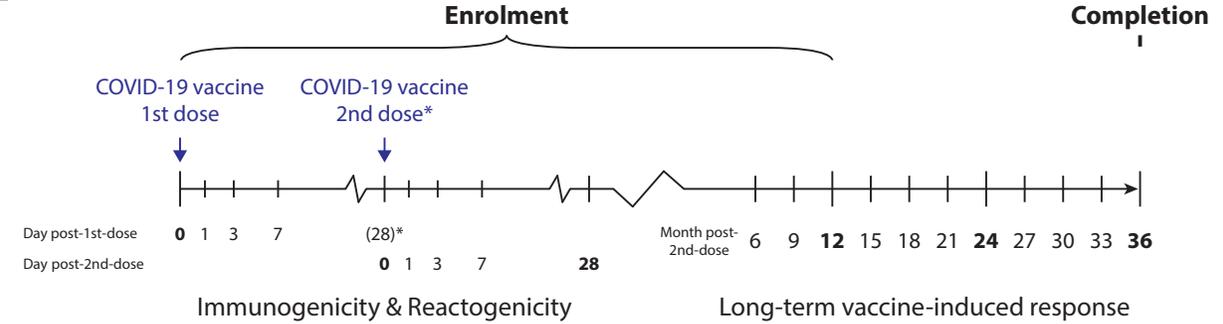
- Our long-term aim is to improve our assessment of population immunity against COVID-19 and inform policy accordingly
- Short term aims:
  - Measure population immunity from natural SARS-CoV-2 infections (EPI-HK study and other studies, presented by Prof Peiris this afternoon)
  - Measure population immunity over time since receipt of COVID-19 vaccine doses
  - Identify “correlates of protection”<sup>1</sup> i.e. translate information on immune markers into actual immunity – only possible once infections are prevalent in community
- Same approaches could also be applied to influenza

# COVAR study

3-year study of immunity in 1500 persons following vaccination, includes:

- Humoral and cellular immune responses to initial vaccine doses and subsequent boosters
- Waning over time between vaccinations
- Biological and epidemiological correlates of immune responses

## Longitudinal follow-up:

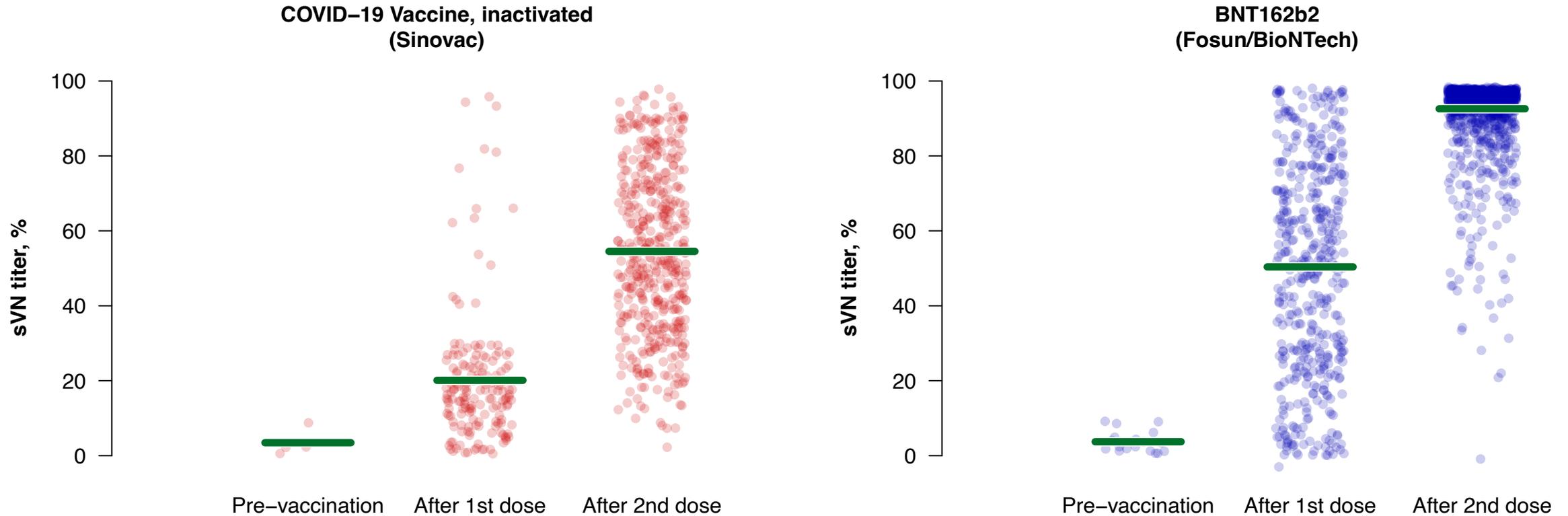


### **Specimen collection**

Response Type	Day 0	Day 1	Day 3	Day 7	Day (28)*	Day 0	Day 1	Day 3	Day 7	Day 28	Month 6	Month 9	Month 12	Month 15	Month 18	Month 21	Month 24	Month 27	Month 30	Month 33	Month 36	
<b>Antibody response-1y (40% participants)</b>																						
Clotted blood	●				●					●												
Whole blood																						
Nasosorption <sup>^</sup>	○				○					○												
Stool <sup>^</sup>	●				●					●												
<b>Antibody response-3m (20% participants)</b>																						
Clotted blood	●				●					●												
Whole blood																						
Nasosorption <sup>^</sup>	○				○					○												
Stool <sup>^</sup>	●				●					●												
<b>Innate response (20% participants)</b>																						
Clotted blood	●				●					●												
Whole blood	●	●			●	●				●												
Nasosorption <sup>^</sup>	○	○			○	○				○												
Stool <sup>^</sup>	●				●					●												
<b>Cellular response (20% participants)</b>																						
Clotted blood	●				●					●												
Whole blood	●		●		●		●			●												
Nasosorption <sup>^</sup>	○		○		○		○			○												
Stool <sup>^</sup>	●				●					●												

\*For individuals who do not plan to receive the second dose of COVID-19 vaccine, or plan to receive >35 days after the first dose, we will arrange blood collection 28 days after the first dose of vaccination. <sup>^</sup>optional

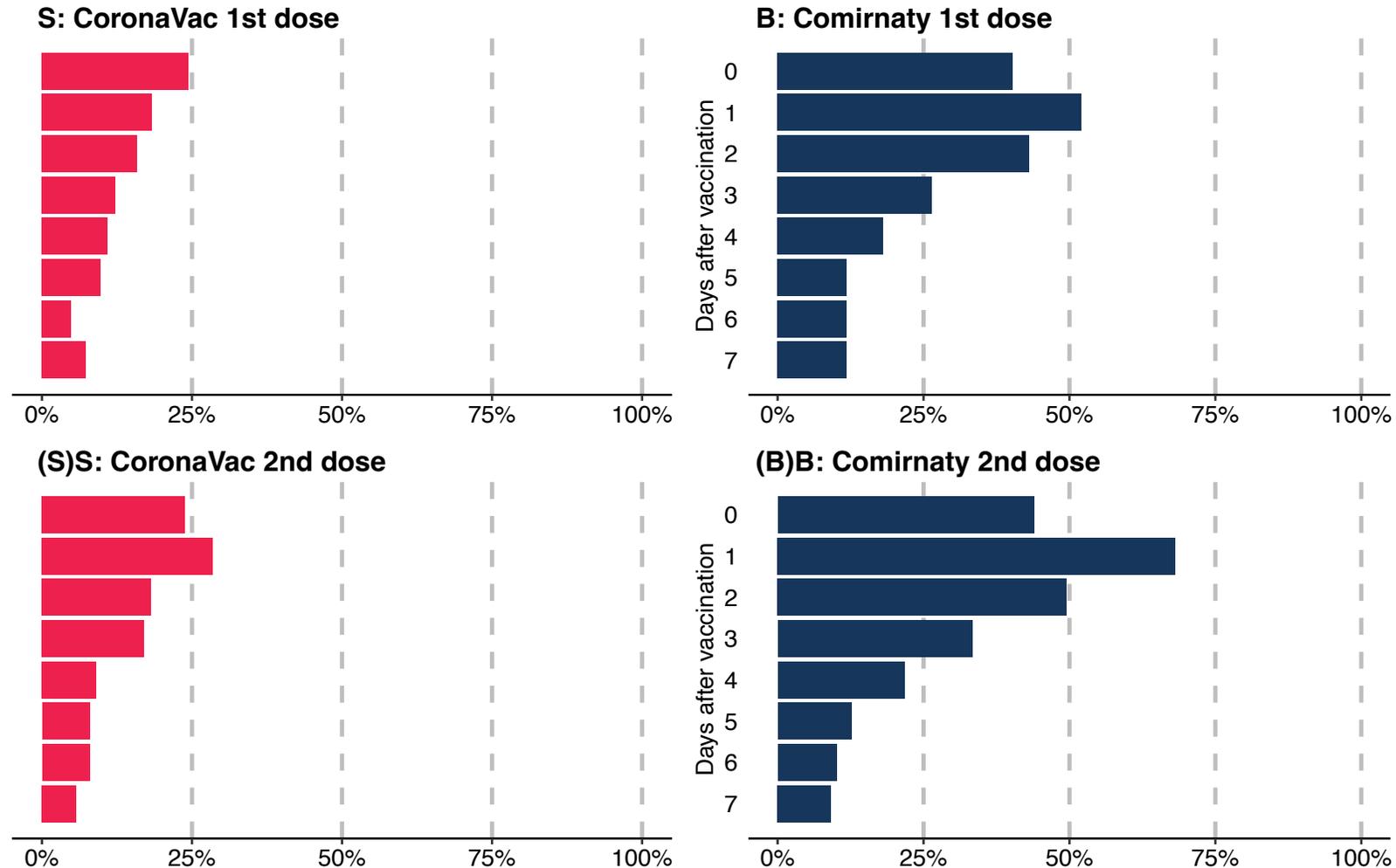
# COVAR study – rises in antibody titers



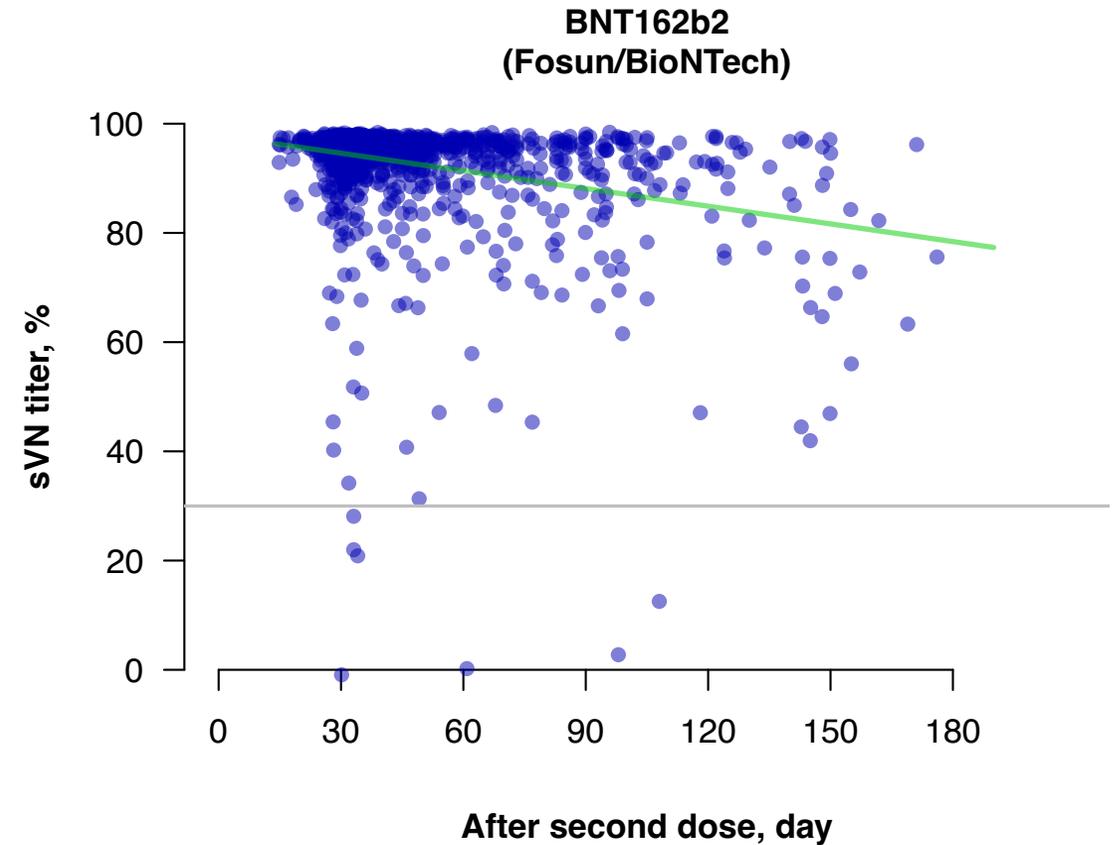
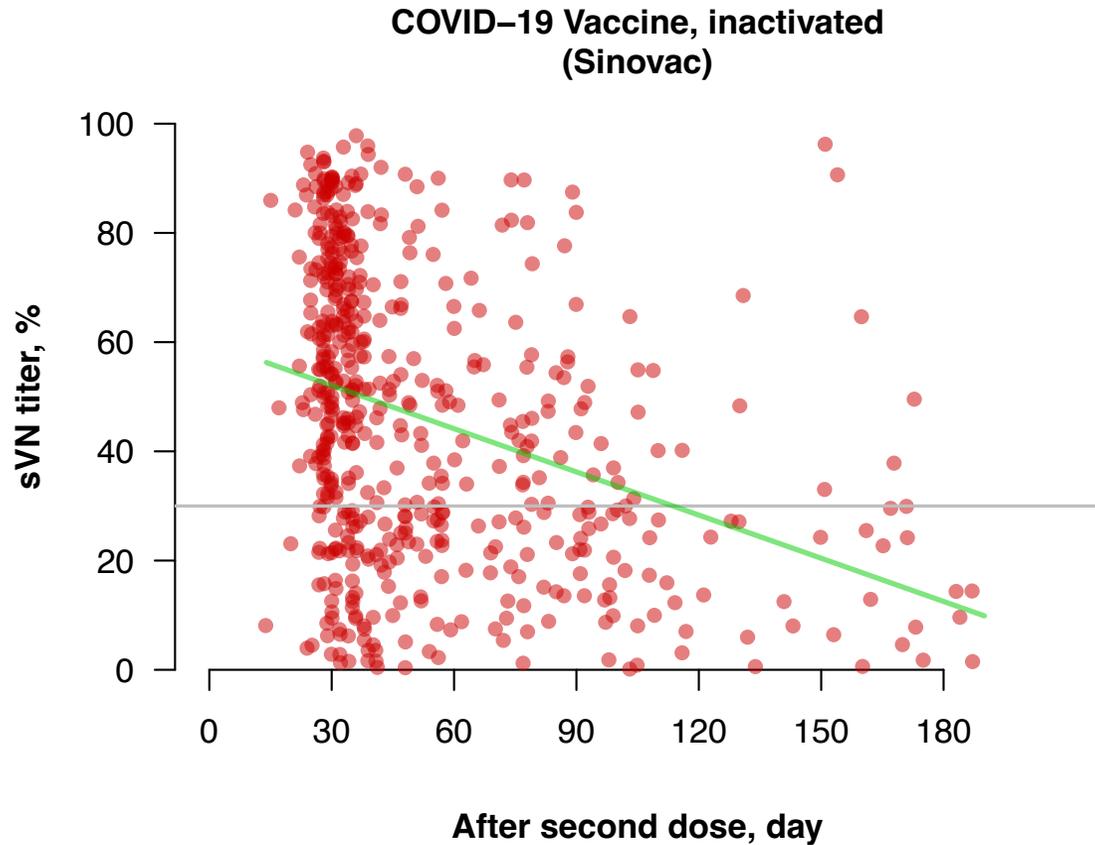
Data from 929 participants (262 S + 667 B)

# COVAR study – signs/symptoms after vaccination

## Feeling unwell in the last 24 hours



# COVAR study – declines in titers up to 6m after 2<sup>nd</sup> dose

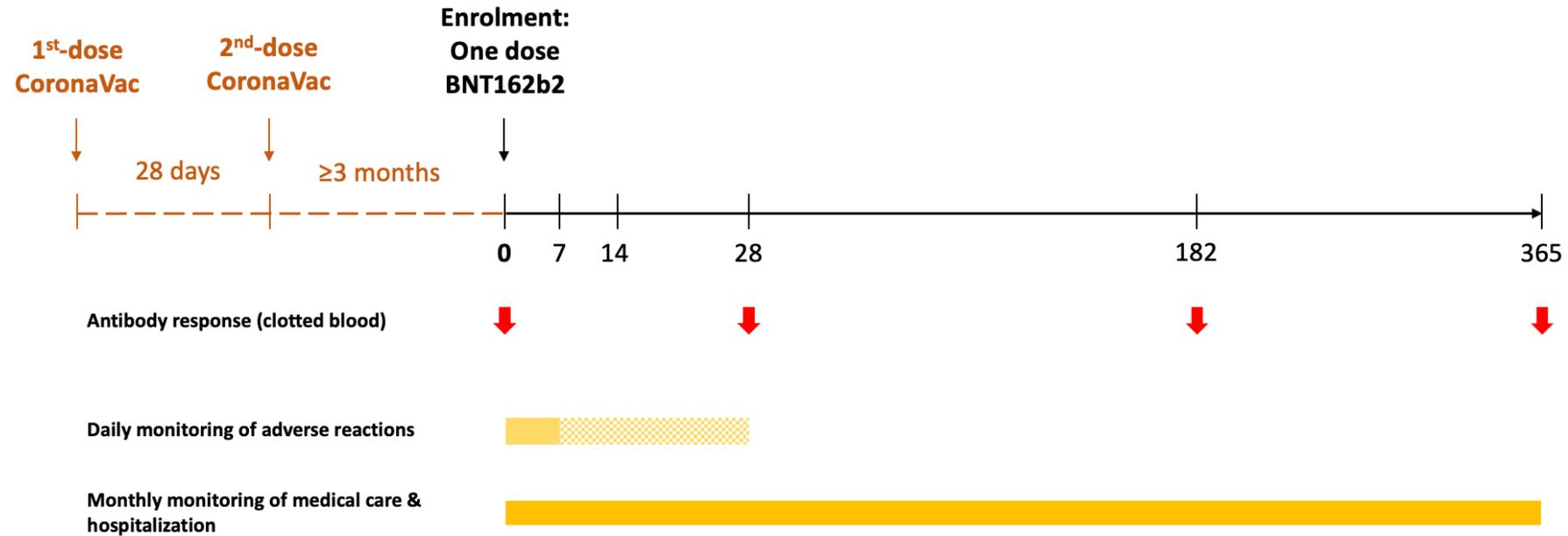


# Potential need for third dose?

- Vaccines maintaining high level of protection against severe disease, although there is evidence of reductions in efficacy against symptomatic infections with Delta variant for two reasons – antigenic mismatch, and immune decay over time (waning)
- Third doses could boost antibody titers and improve protection

# mBoost trial

(\*not supported by HMRF)

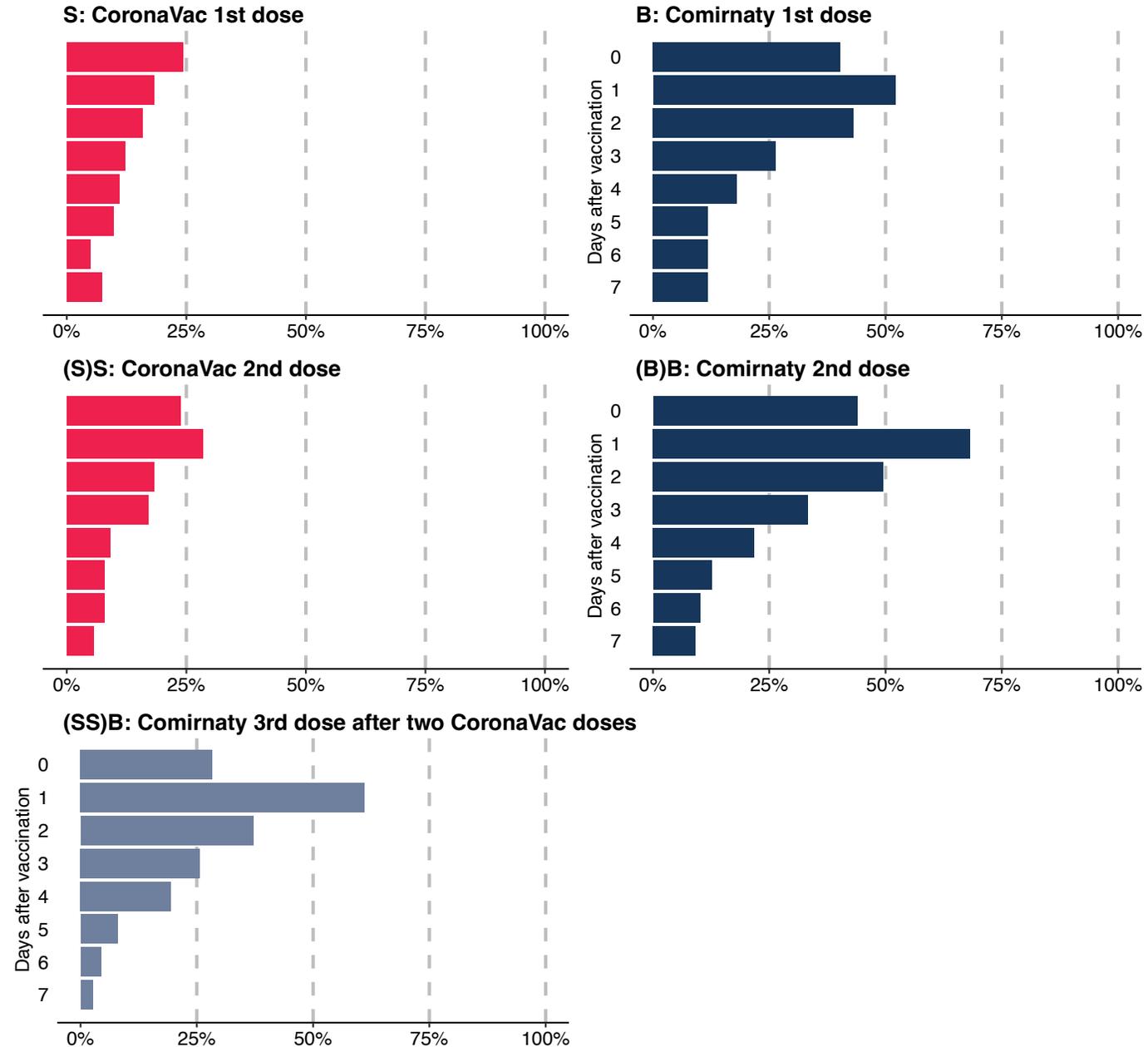


We have enrolled and vaccinated 350 adults age  $\geq 30$ ,  
collection of day 28 samples is ongoing

# mBoost vs COVAR

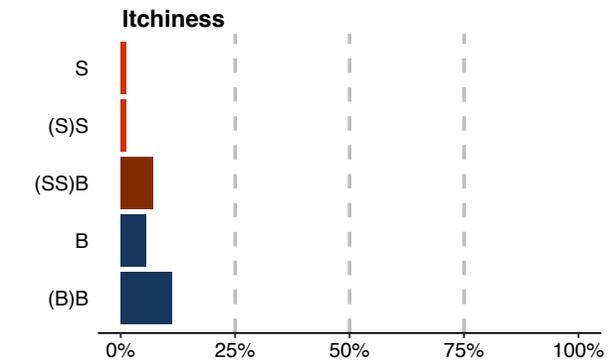
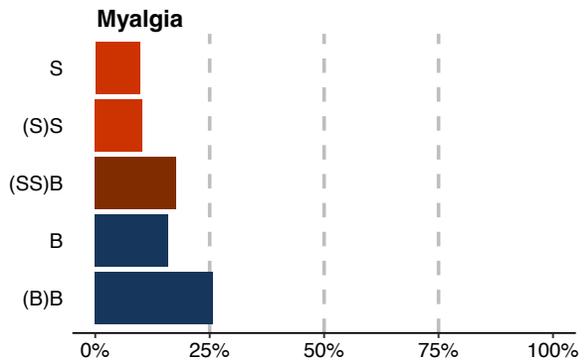
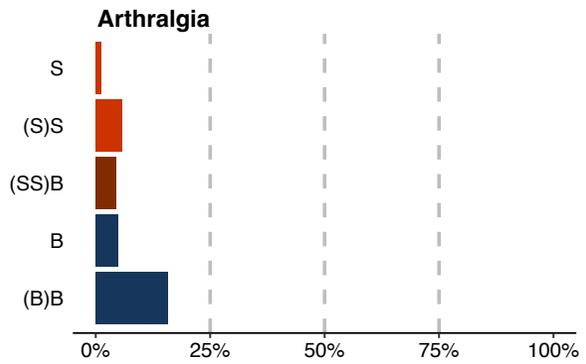
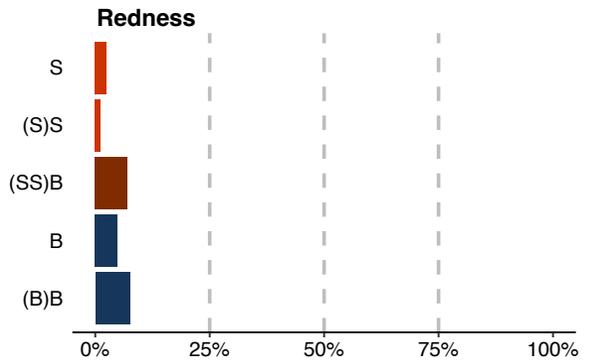
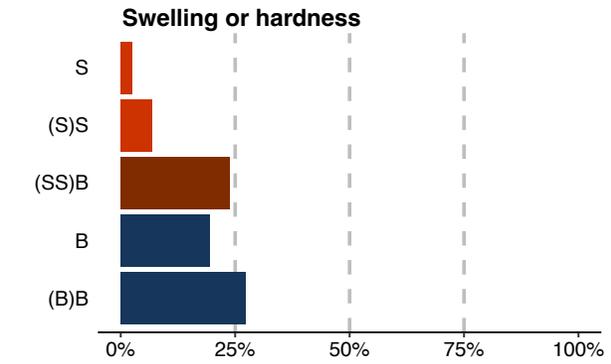
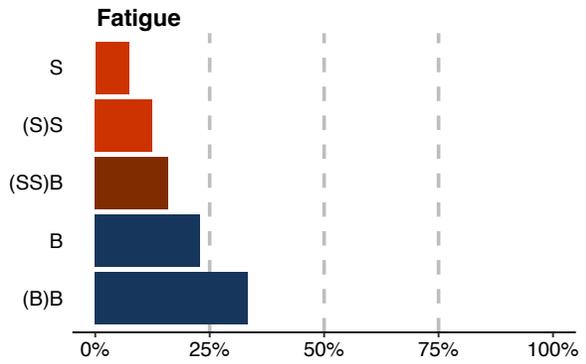
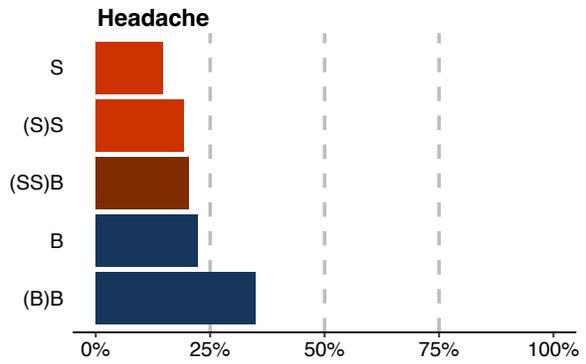
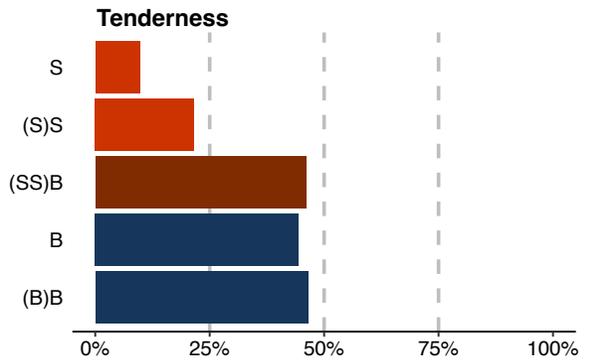
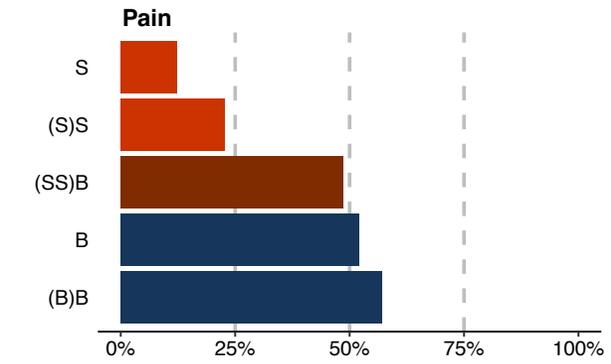
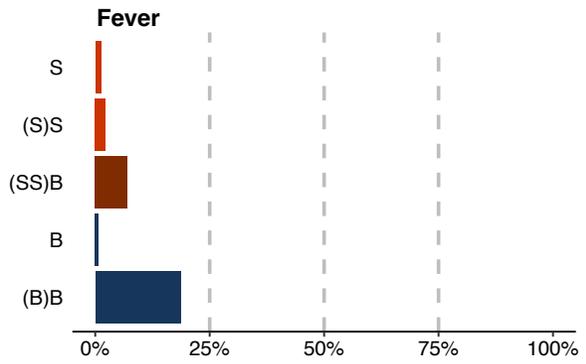
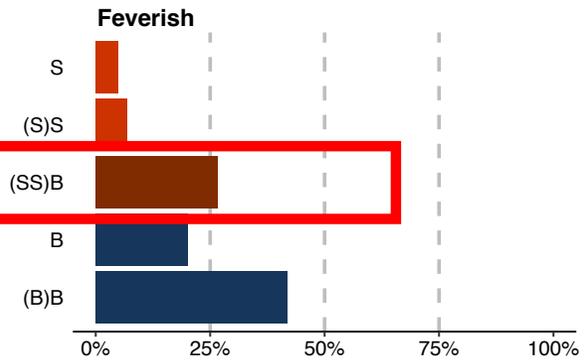
Immediate reactions after BioNTech 3<sup>rd</sup> dose in those who previously received two doses of CoronaVac are quite comparable to the reactions after first BioNTech dose

## Feeling unwell in the last 24 hours

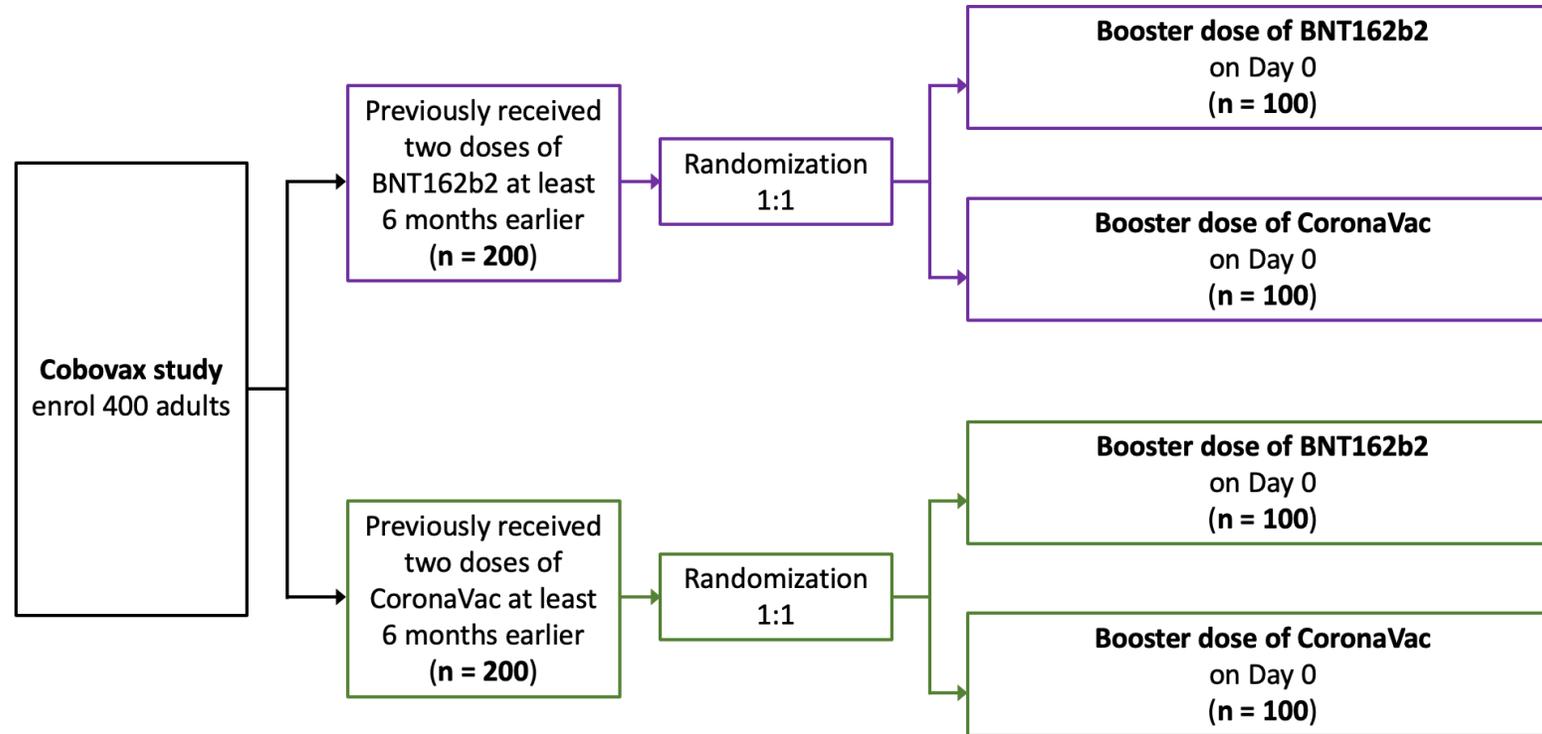


Relevant systemic symptoms up to 7 days after vaccination by vaccine type and dose

Local reactions up to 7 days after vaccination by vaccine type and dose



# Cobovax trial



Currently enrolling 400 adults age  $\geq 18$

Similar timing of blood draws as the mBoost study (previous slide), plus CMI analysis

# PIVOTe cohort study

- We complement the data from the COVAR study with an elderly cohort
- This spring we collected 1290 blood samples from 1299 cohort participants all  $\geq 69$  years of age
  - Only 153 reported receipt of a COVID-19 vaccine between February and the date of the spring draw up to June 2021 (49 BioNTech, 104 Sinovac). Among the remaining participants, only 40 planned to get COVID-19 vaccination.
- Now collecting autumn blood draws and updating information on receipt of COVID-19 vaccine doses, will have laboratory results from spring and autumn draws soon

# Final comments

- Stronger antibody responses to BioNTech vaccine compared to Sinovac vaccine, but similar T cell responses (not shown today)
- mBoost trial will provide data specifically on 3rd doses of BioNTech following two CoronaVac
- Cobovax trial will soon provide data on immune responses to third doses with the same or different platform from the first two doses
- COVAR and PIVOTe studies will provide continuous observational data on immunity through time

# Acknowledgments

- Financial support from HMRF:
  - COVID1903001 (COVAR and PIVOTe)
  - COVID19F01 (Cobovax)
- Nancy Leung
- Malik Peiris, Sophie Valkenburg
- Gabriel Leung, Mario Martín-Sánchez, Irene Wong
- Staff at Ap Lei Chau, Central Library, Gleneagles CVCs
- Participants!!