

Growing Capability - Virus Protein Production for Vaccine Research



Ferrier Research Institute
Te Kāuru

Kerstin Thornton 300375672
Faculty of Science



Introduction

Pandemics have focused global attention on gaining the capability to develop, test, and then locally produce vaccines at pace. The Vaccine Alliance Aotearoa New Zealand (VAANZ) is actively working toward this goal. As part of this endeavor, this project was to develop a protein subunit vaccine candidate against COVID-19.

A protein subunit vaccine uses portions of the virus to create a lasting protective immune response. A key benefit of this approach is that it avoids the challenges of working with active virus.

Method

The receptor binding domain (RBD) of the SARS-CoV-2 virus spike protein made an ideal target as the virus uses it to enter host cells. The production of antibodies recognising this site will prevent viral entry and replication.

The encoding sequence of the RBD was introduced into a commercial yeast strain using targeted integration. After testing RBD production was successful, a high yield clone was selected and grown at lab scale. The protein was then harvested and purified.

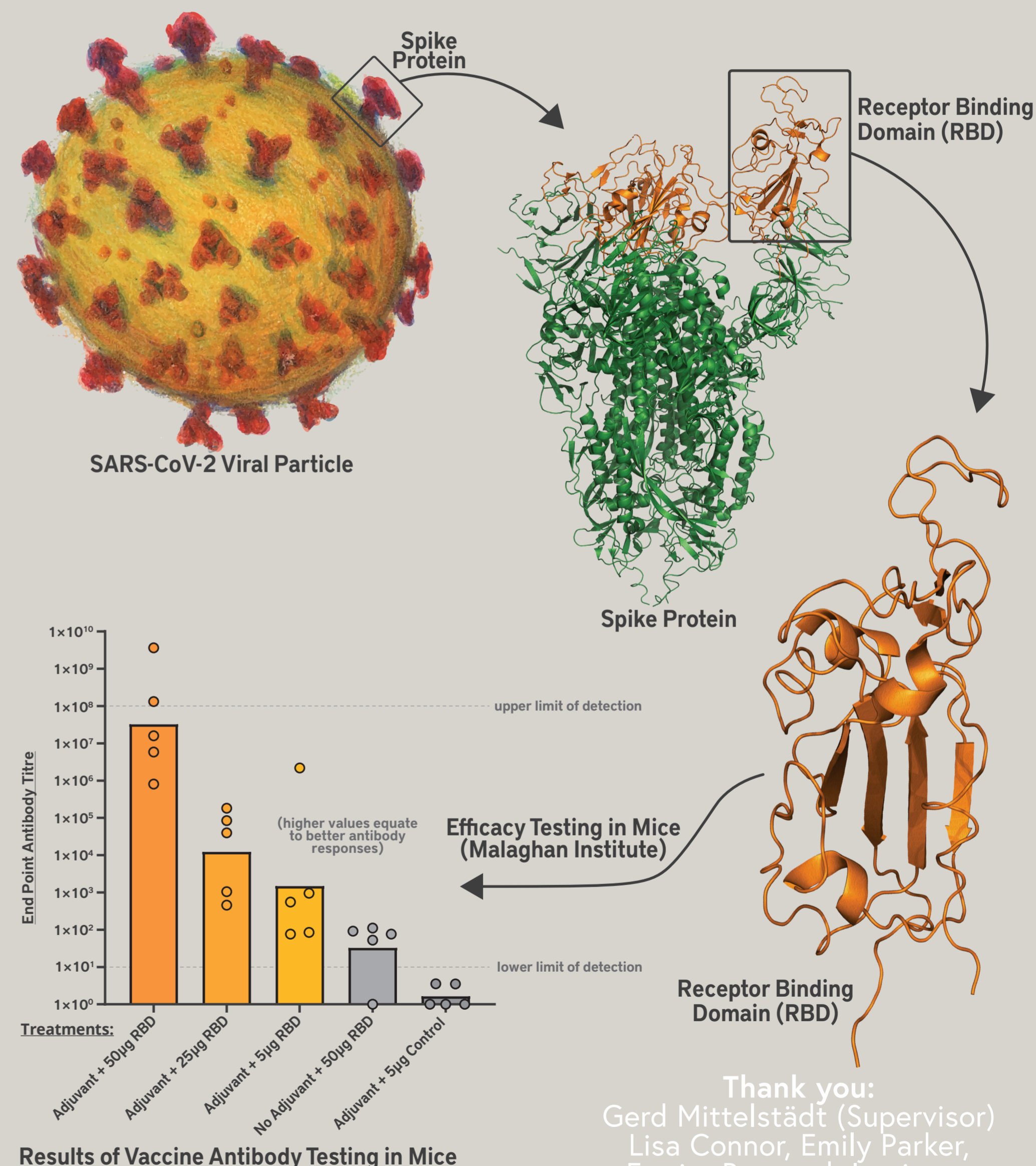
Results and Conclusion

Production of SARS-CoV-2 RBD in yeast at lab scale was successfully demonstrated. Vaccine testing in mice was undertaken to ascertain the effectiveness of the resultant RBD as a vaccine candidate. The candidate vaccine was shown to stimulate the generation of neutralising antibodies. Further development is now being undertaken to optimise the production process and efficacy.

This research project was a successful incremental step toward providing increased disease response capabilities here in New Zealand.

Sources

- [1] Zhou *et al.*, 2020. Cryo-EM Structures Delineate a pH-Dependent Switch that Mediates Endosomal Positioning of SARS-CoV-2 Spike Receptor-Binding Domains. SSRN Electronic Journal.
- [2] Chen *et al.*, 2013. Yeast-expressed recombinant protein of the receptor-binding domain in SARS-CoV spike protein with deglycosylated forms as a SARS vaccine candidate. *Human Vaccines & Immunotherapeutics*, 10(3), pp.648-658.
- [3] Dai, *et al.*, 2020. A Universal Design of Betacoronavirus Vaccines against COVID-19, MERS, and SARS. *Cell*, 182(3), pp.722-733.e11.
- [4] Eckert and Higgins., 2020. Details - Public Health Image Library (PHIL) #23312 Ultrastructural Morphology Exhibited by Coronaviruses. [online] <https://phil.cdc.gov/Details.aspxpid=23312> [Accessed 2 February 2021].



Results of Vaccine Antibody Testing in Mice

Thank you:
Gerd Mittelstädt (Supervisor)
Lisa Connor, Emily Parker,
Ferrier Research Institute,
VUW Faculty of Science, VAANZ, &
Malaghan Institute of Medical Research