

## Study shows laboratory-developed protein spikes consistent with COVID-19 virus

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A new international study has found that the key properties of the spikes of SARS-CoV-2 virus which causes COVID-19 are consistent with those of several laboratory-developed protein spikes, designed to mimic the infectious virus.

A central component in designing serological tests and vaccines to protect against COVID-19 is the manufacture of protein "spikes". These recombinant spikes closely mimic those sticking out of surface of the infectious virus and trigger the body's immune system into action.

Laboratory manufactured spikes are also used for serological testing (also referred to as antibody testing) and as research reagents. The findings show how that viral spike manufactured through different methods in laboratories across the globe are highly similar and provide reassurance that the spike can be robustly manufactured with minimal variations between laboratories.

The spikes on the SARS-CoV-2 virus are coated in sugars, known as glycans, which they use to disguise themselves from the human immune system. The abundance of these glycans has the potential to create significant discrepancies between studies that use different recombinant spikes.

In this new study, published in the journal *Biochemistry*, the research team studied the <u>glycan</u> coatings on recombinant spikes developed in five laboratories around the world and compared them to those on the



spikes of the infectious virus.

"The speed at which <u>scientific community</u> has moved to tackle the COVID-19 pandemic has put considerable pressure on laboratories around the world to validate their findings quickly," Explained Max Crispin, Professor of Glycobiology at the University of Southampton, who led the study. "Over the last year we have seen vaccines developed around the world at an unprecedented rate and the rapid development, and validation, of recombinant proteins have been fundamental to that success story," he continued.

In April 2020, Professor Crispin and his team from the University of Southampton mapped the glycan coating of the SARS-CoV-2 spike for the first time. In the present study, they extend their analysis to examine recombinant spike developed in laboratories at the Amsterdam University Medical Centre, Harvard Medical School, the University of Oxford, and the Swiss company ExcellGene. All the different batches of spike protein were shown to mimic key features of the glycosylation of inactivated virions analysed at Tsinghua University, China.

The study also used <u>computational methods</u> to examine the protein features that were shaping some of the glycosylation features that were seen across all the samples. Dr. Peter Bond, Senior Principal Investigator at the Bioinformatics Institute of the Agency for Science, Technology and Research (A\*STAR), Singapore, who led the computational work said, "Our modelling enabled us to shed light on how the protein influences the structure of the glycans and why the glycosylation was so consistent. This predictive approach could also be of potential value in therapeutics development against new variants or other emerging viruses."

"The ability to produce mimics of the SARS-CoV-2 spike <u>protein</u> with <u>high fidelity</u> at many different laboratories, all of which recapitulate the



glycan signatures of the authentic <u>virus</u>, is of significant benefit for vaccine design, antibody testing and drug discovery," concluded Professor Crispin.

The study, "Site-specific steric control of SARS-CoV-2 spike glycosylation," has been published in *Biochemistry*.

**More information:** Joel D. Allen et al, Site-Specific Steric Control of SARS-CoV-2 Spike Glycosylation, *Biochemistry* (2021). <u>DOI:</u> <u>10.1021/acs.biochem.1c00279</u>

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