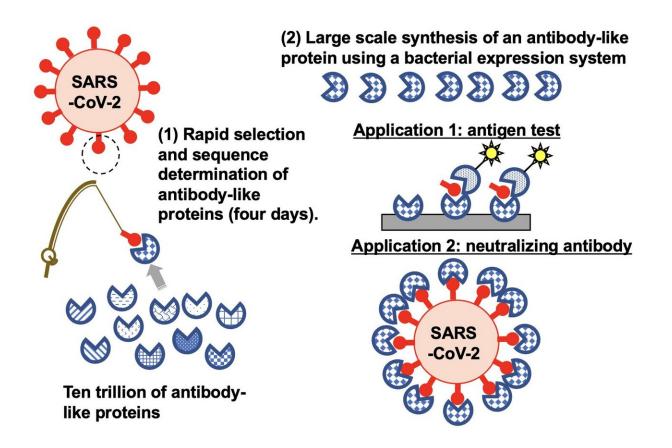


Setting a TRAP for pandemic-causing viruses

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The TRAP display method 'fishes' for synthetic proteins from among a library of trillions for those that can target SARS-CoV-2. The approach was able to identify proteins that can be used for testing for the virus and potentially treating people infected with COVID-19. Credit: Hiroshi Murakami

A research team led by Nagoya University scientists in Japan has



developed an approach that can quickly find synthetic proteins that specifically bind to important targets, such as components of the SARS-CoV-2 virus. The method was published in the journal *Science Advances* and could be used to develop test kits or for finding treatments.

"We developed a laboratory technique for rapid selection of <u>synthetic</u> <u>proteins</u> that strongly bind to SARS-CoV-2," says Nagoya University biomolecular engineer Hiroshi Murakami. "High-affinity synthetic proteins can be used to develop sensitive antigen tests for SARS-CoV-2 and for future use as neutralization antibodies in infected patients."

Murakami and his colleagues had previously developed a protein selection lab test called TRAP display, which stands for "transcriptiontranslation coupled with association of puromycin linker." Their approach skips two time-consuming steps in another commonly used technique for searching through synthetic protein libraries. But their investigations indicated there was a problem with the puromycin linker.

In the current study, the team improved their technique by modifying the puromycin linker. Ultimately, they were able to use their TRAP display to identify nine synthetic proteins that bind to the spike protein on SARS-CoV-2's outer membrane. The approach took only four days compared to the weeks it would take using the commonly used messenger RNA display technology.

TRAP display involves using a large number of DNA templates that code for and synthesize trillions of proteins carrying random peptide sequences. The synthetic proteins are linked to DNA with the help of the modified puromycin linker and then exposed to a <u>target protein</u>. When the whole sample is washed, only the synthetic proteins that bind to the target remain. These are then placed back into the TRAP display for further rounds until only a small number of very specific target-binding synthetic proteins are left.



The researchers investigated the nine synthetic proteins that were found to bind to SARS-CoV-2. Some were specifically able to detect SARS-CoV-2 in nasal swabs from COVID-19 patients, indicating they could be used in test kits. One also attaches to the virus to prevent it from binding to the receptors it uses to gain access to human cells. This suggests this protein could be used as a treatment strategy.

"Our high-speed, improved TRAP <u>display</u> could be useful for implementing rapid responses to subspecies of SARS-CoV-2 and to other potential new viruses causing future pandemics," says Murakami.

This study, "Antibody-like proteins that capture and neutralize SARS-CoV-2," was published online in *Science Advances* on September 18, 2020.

More information: T. Kondo et al. Antibody-like proteins that capture and neutralize SARS-CoV-2, *Science Advances* (2020). <u>DOI:</u> <u>10.1126/sciadv.abd3916</u>

Provided by Nagoya University

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