## **Coronavirus** Singapore

## S'pore study shows length of immunity varies greatly

## Depending on patient, it ranges from 35 days to 41 years, so Covid-19 reinfections a possibility

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Scientists in Singapore, using an algorithm to predict the longevity of immunity to Covid-19, have found that it can last just 35 days in some people and up to four decades in others.

The implications are that some people who have recovered from Covid-19 may still get reinfected, said Professor Wang Linfa, a corresponding author of the study.

He added that this would limit the viability of a natural herd immunity strategy.

Prof Wang, who is from Duke-NUS Medical School's Emerging Infectious Diseases Programme, also said that assuming the immunity provided by vaccinations wanes like naturally produced antibodies, yearly Covid-19 vaccination may be needed. This is to prevent future outbreaks of the disease.

But in the study published in The Lancet Microbe on Tuesday, he and the other scientists also pointed out that waning antibodies do not always mean waning immunity.

People with low levels of neutralising antibodies may still be protected from Covid-19 if they have a robust T-cell immunity, they noted.

In the Singapore study, scientists from Duke-NUS Medical School, the National Centre for Infectious Diseases (NCID) and the Agency for Science, Technology and Research Infectious Diseases Labs followed 164 Covid-19 patients in Singapore for at least six months after their infection.

During that time, the scientists analysed the patients' blood samples for neutralising antibodies against the coronavirus, as well as T-cells and immune system signalling molecules.

They found that some people had neutralising antibodies – a key arm of the immune system - that waned quickly.

In others, they decreased slowly. For some, the neutralising antibodies persisted.

Generally, the more ill a person was with Covid-19, the more likely his neutralising antibody levels would persist. Those who were mildly ill saw their neutralising antibodies wane more rapidly.

Using a machine learning algorithm, the scientists projected that the longevity of coronavirus neutralising antibodies can range from 35 days to 41 years.

Prof Wang said the prediction of immunity longevity can be accurately determined only at the individual level. To do this, three data

points are needed. "We need to take blood samples during the infection, three months after the infection and six months after that, and we can predict if you are immune to Covid for how

many months or years," he said. In the study, the scientists sorted the patients into five groups.

The "negative group" (11.6 per cent of the patients in the study) were people who never developed detectable neutralising antibodies.

The "rapid waning" group (slightly less than 27 per cent) had varying early levels of antibodies that waned quickly.

The "slow waning" group (29 per cent) tested mostly positive for antibodies at six months.

Nearly one-third – the "persistent" group - showed little change in their antibody levels for up to six months after their infection.

Finally, there was an unusual "delayed response" group of three people who showed a marked rise in neutralising antibodies months af-

ter recovering from severe illness. Prof Wang said: "The key message from this study is that the longevity of functional neutralising antibodies against Sars-CoV-2 can vary greatly, and it is important to monitor this at an individual level.

"This work may have implications for immunity longevity after vaccination, which will be part of our follow-up studies.'

The study had limitations, including the fact that the patients had a median age of 44. Older adults or children might have different immune profiles, the scientists said.

The study also found that patients from all groups, including

those who did not have detectable neutralising antibodies, had sustained T-cell immunity six months

Neutralising antibodies are important in protecting against Covid-19, but they are just a part of parts include T-cells.

Scientists are starting to see the importance of T-cells in the fight

"maintain the memory" of a pathogen for decades and will be rapidly recruited upon new infection, said Professor Antonio Bertoletti of Duke-NUS' Emerging Infectious Diseases Programme.

Prof Bertoletti was also involved

In earlier research, he and a team of scientists found that 23 individuals who had severe acute respiratory syndrome (Sars) in 2003 were recently all found to still have T-cells that recognise the virus that causes Sars - 17 years after infection.

Unlike antibodies, T-cells are harder to detect and study, and some classical vaccines, such as the seasonal influenza vaccine,

tion, said Prof Bertoletti.

Associate Professor David Lye said the knowledge that antibodies against Sars-CoV-2 wane in different people at different rates shows how important public health and social measures are in the ongoing pandemic outbreak response.

Prof Lye, who is the director of the Infectious Disease Research and Training Office at NCID, is also a corresponding author of the latest study.

"However, the presence of T-cell immunity provides hope of longerterm protection which will require more studies and time for epidemiological and clinical evidence to confirm," he added.







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