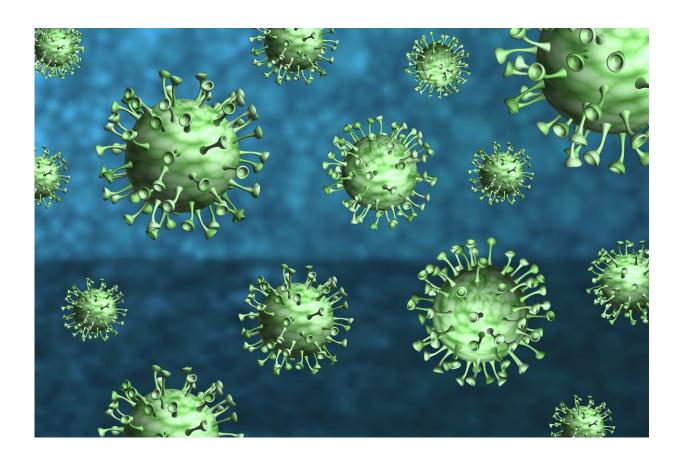


## Immune cell responses in COVID-19 patients far from optimal

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Credit: Thiago Lazarino from Pixabay

Melbourne researchers have discovered that killer T cells, key immune cells in fighting viral infections, are present at much lower levels in people with COVID-19, compared to influenza or glandular fever.



Publishing their findings today in *PNAS*, the team from the Peter Doherty Institute for *Infection and Immunity* (Doherty Institute) looked at 22 COVID-19 samples from patients who experienced asymptomatic, mild or moderate illness.

First author, University of Melbourne's Jennifer Habel, a Ph.D. student at the Doherty Institute, said their research concentrated on CD8 T cells (killer T cells), which are integral to mounting an effective and rapid recovery from viruses such as influenza.

The team looked at T cells in people who express a protein called human antigen leucocyte (HLA) serotype HLA-A2. HLA proteins are important for T cell recognition and vary across individuals.

"What we found is those key immune cells weren't stimulated optimally for rapid proliferation and expansion to fight SAR-CoV-2," Ms Habel said.

"The magnitude of the killer T cells was only five times higher than those of the naïve immune cells. To give that perspective, it's 10 times lower than what we see during an influenza or glandular fever response."

In addition to magnitude, the team also looked at the activation profile of these immune cells and found that not only was activation poor, but in some cases these cells remained largely naïve, as if they hadn't been exposed to the virus at all.

University of Melbourne Professor Katherine Kedzierska, a laboratory head at the Doherty Institute and a world-leading influenza immunology researcher who led this study, said these striking findings support the 'prime and boost' vaccine model.

"Knowing the specific T cells and proteins to target will inform the



design of an effective vaccine," Professor Kedzierska said. "This research shows that potentially, if a vaccine was able to prime the immune system of killer T cells and then boost them a short time later, the immune response is likely to be much more robust to fight SARS-CoV2."

Co-author, Doherty Institute Patron Laureate Professor Peter Doherty said: "If the SARS-CoV-2 virus is indeed compromising the killer T cell response that can be important for recovery, the best bet may be to get the right vaccine before the virus gets to us."

**More information:** Jennifer R. Habel et al. Suboptimal SARS-CoV-2–specific CD8+ T cell response associated with the prominent HLA-A\*02:01 phenotype, *Proceedings of the National Academy of Sciences* (2020). DOI: 10.1073/pnas.2015486117

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