

Researchers discover unique 'spider web' mechanism that traps, kills viruses

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Researchers Matthew Miller, an associate professor at McMaster's Michael G. DeGroote Institute for Infectious Disease Research and Hannah Stacey, a graduate student and lead author of the study. Credit: Georgia Kirkos, McMaster University

Immunologists at McMaster University have discovered a previously



unknown mechanism which acts like a spider web, trapping and killing pathogens such as influenza or SARS-CoV-2, the virus responsible for COVID-19.

The researchers have found that neutrophils, the most abundant white blood cells in the <u>human body</u>, explode when they bind to such pathogens coated in antibodies and release DNA outside of the cell, creating a sticky tangle which acts as a trap.

The findings, published online in the *Proceedings of the National Academy of Science*, are significant because little is understood about how antibodies neutralize viruses in the respiratory tract.

The discovery has implications for <u>vaccine design</u> and delivery, including aerosol and nasal spray technologies that could help the body head off infections before they have a chance to take hold.

"Vaccines can produce these antibodies that are present in our lungs, which are the first type of antibody to see viruses like flu or COVID-19, which infect our lungs and respiratory tracts," says the study's lead author Matthew Miller, an associate professor at McMaster's Michael G. DeGroote Institute for Infectious Disease Research and Canada's Global Nexus for Pandemics and Biological Threats. "Mechanisms that can stop the infection at the site where it enters our body can prevent the spread and serious complications."

By comparison, injectable vaccines are designed to bolster antibodies in the blood, but those antibodies are not as prevalent at the sites where infection begins.

"We should be thinking carefully about next generation COVID-19 vaccines that could be administered in the respiratory tract to stimulate antibodies. We don't have many candidates right now that are focused on



raising the mucosal response," says Hannah Stacey, a <u>graduate student</u> in the Miller Lab and lead author of the paper, who recently won a major national scholarship from the Canadian Society for Virology for her work on COVID-19.

"If you want a lot of these antibodies that are really abundant in blood, then injections make the most sense, but if you want antibodies that are abundant in the <u>respiratory tract</u>, then a spray or an aerosol makes sense," she says.

Researchers caution that while the body's <u>spider-web</u> mechanism has the potential to be hugely beneficial, it can cause harm too, including inflammation and further illness when the web formation is uncontrollable.

They point to the early waves of the pandemic, prior to vaccinations, when these NETs, or neutrophil extracellular traps, were found in some patients' lungs, and had made their breathing more difficult.

"An <u>immune response</u> that is meant to protect you can end up harming you if it's not properly controlled," says Miller. "It's important to understand the balance of the immune system. If you have a lot of these antibodies before you get infected, they are likely going to protect you, but if the infection itself stimulates a lot of those <u>antibodies</u> it might be harmful."

More information: Hannah D. Stacey et al, IgA potentiates NETosis in response to viral infection, *Proceedings of the National Academy of Sciences* (2021). DOI: 10.1073/pnas.2101497118

Provided by McMaster University



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