

Long distance signals protect brain from viral infections

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The brain contains a defense system that prevents at least two unrelated viruses—and possibly many more—from invading the brain at large. The research is published online ahead of print in the *Journal of Virology*.

"Our work points to the remarkable ability of the immune system, even within the <u>brain</u>, to protect us against opportunistic <u>viruses</u>," says Anthony van den Pol of Yale University, an author on the study.

The research explains a long-standing mystery. The olfactory mucosa in the nose can serve as a conduit for a number of viruses to enter the brain including rabies, polio and <u>influenza viruses</u>. Yet infections in the central nervous system rarely occur. The mechanism responsible for protecting the brain from viruses that successfully invade the <u>olfactory bulb</u> (OB), the first site of infection in nasal mucosa, remains elusive.

Van den Pol and his colleagues have discovered that, in response to viral infection, cells in the olfactory bulb release long-distance signaling molecules that tell cells in uninfected parts of the brain to produce antiviral interferon, a first line of defense against invading viruses.

In the study, normal mice expunged the infection, while mice lacking receptors for the initial signaling molecules succumbed as the viruses spread throughout the brain, proving the critical role of these molecules.

These signaling molecules are different than regular neurotransmitters. Van den Pol notes that during neuronal signaling, neurotransmitters



released by one cell travel across a mere 20 nanometers of synapse to the next nerve cell. Yet, the long-distance signaling molecules diffuse up to 15 millimeters.

"That distance is almost a million times greater than the distance across a synapse," he says.

"The success of the immune system in blocking two unrelated viruses, vesicular stomatitis virus and cytomegalovirus, suggests that our results may generalize to many other viruses that can enter the brain through the olfactory nerve," says van den Pol.

More information: jvi.asm.org/content/early/2014 ... 509-13.full.pdf+html

The final version of the article is scheduled for the April 2014 issue of the *Journal of Virology*.

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