

Biodefense solutions to protect our nation

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## Army Study: DNA Vaccine and Duck Eggs Protect Against Hantavirus Disease

Army scientists and industry collaborators have successfully protected laboratory animals from lethal hantavirus disease using a novel approach that combines DNA vaccines and duck eggs. The work appears in a recent edition of the online scientific journal PLoS ONE, published by the Public Library of Science.

According to first author Jay W. Hooper of the U.S. Army Medical Research Institute of Infectious Diseases (USAMRIID), this is the first time that the DNA vaccine/duck egg system has been shown to produce an antiviral product capable of protecting against hantavirus disease.

Hantavirus causes a condition known as hantavirus pulmonary syndrome (HPS), which has a case fatality rate of 35-40 percent. Currently there are no vaccines, prophylactics, or therapeutics to prevent or treat this highly pathogenic disease.

In this study, the research team used a hamster model of Andes virus, which is the predominant cause of HPS in South America and the only hantavirus known to be transmitted person-to-person. Infection of Syrian hamsters with Andes virus, as demonstrated in earlier studies at USAMRIID, results in a disease that closely mimics human HPS in incubation time, symptoms of respiratory distress, and disease pathology. This makes it an ideal system for evaluating the feasibility of postexposure protection strategies.

Collaborating with Aldevron of Fargo, N.D. and the Universidad del Desarrollo in Santiago, Chile, Hooper and his team first evaluated a natural product, human polyclonal antibody, which was obtained as fresh frozen plasma (FFP) from a patient who survived HPS. Their results indicate that FFP shows promise as a post-exposure preventive treatment for HPS.

The team then vaccinated ducks with a DNA vaccine against Andes virus. This vaccine, initially developed and tested at USAMRIID, uses genetic material, or DNA, that encodes a specific hantavirus gene to elicit an immune response in the recipient.

Next, they purified an antibody called IgY from the yolks of the duck eggs. This purified IgY, as well as a similar version produced in duck eggs, was capable of neutralizing Andes virus when tested in cell culture. More importantly, it also protected Syrian hamsters from lethal HPS—even when administered as a single injection several days after the hamsters had been exposed to a lethal dose of virus.

The work demonstrates the feasibility of using DNA vaccine technology, coupled with the duck/egg system, to manufacture a product that could supplement or replace FFP. Furthermore, the new approach can be scaled as needed and eliminates the necessity of using blood products from HPS survivors, which may be in limited supply.

According to Hooper, another advantage of this technique is that duck IgY naturally loses a part of the antibody that has been associated with "serum sickness" when animal antibodies have been used in humans, making the product potentially less reactogenic.

"This antiviral product, if fully developed and manufactured, has the potential to be used in future outbreak situations," Hooper said. "It also could be used to treat health care workers and others who have close contact with HPS patients."

In addition, the authors suggest, the flexibility of the DNA vaccine/duck egg system could be applied to the production of antibodies against other infectious agents and toxins.

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Reference: Brocato R, Josleyn M, Ballantyne J, Vial P, Hooper JW (2012). DNA Vaccine-Generated Duck Polyclonal Antibodies as a Postexposure Prophylactic to Prevent Hantavirus Pulmonary Syndrome (HPS). PLoS One 7(4): e35996. doi: 10.1371/journal.pone.0035996

About USAMRIID: USAMRIID's mission is to protect the warfighter from biological threats and to be prepared to investigate disease outbreaks or threats to public health. Research conducted at USAMRIID leads to medical solutions—vaccines, drugs, diagnostics, and information—that benefit both military personnel and civilians. The Institute plays a key role as the lead military medical research laboratory for the Defense Threat Reduction Agency's Joint Science and Technology Office for Chemical and Biological Defense. USAMRIID is a subordinate laboratory of the U.S. Army Medical Research and Materiel Command.

About Aldevron: Aldevron manufactures plasmid DNA, proteins, and antibodies for many applications. Aldevron's genetic immunization technology is used to generate valuable antibodies directly from a DNA sequence without the use of conventional antigens. The Company works with all sectors of the biotech industry and has many clients in government, academia, and industry. Aldevron is based in Fargo, ND and has divisions in Wisconsin and Germany.

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