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How the Novavax Vaccine Works

6-7 minutes

The Maryland-based company Novavax has developed a protein-based coronavirus vaccine called **NVX-CoV2373**. In March the company announced an efficacy rate of 96 percent against the original coronavirus, 86 percent against the [B.1.1.7 variant](#) and 49 percent against the [B.1.351 variant](#).

Coronavirus Proteins

The SARS-CoV-2 virus is [studied with proteins](#) that it uses to enter human cells. These so-called spike proteins make a tempting target for potential [vaccines](#) and [treatments](#).

The Novavax vaccine works by teaching the immune system to make antibodies to the spike protein.

Growing Spike Proteins

To create their vaccine, Novavax researchers started with a modified spike gene. They inserted the gene into a different virus, called a [baculovirus](#), and allowed it to infect

moth cells. The infected cells produced spike proteins that spontaneously joined together to form spikes, as they do on the surface of the coronavirus.

A similar method of growing and harvesting virus proteins is [already used](#) to make licensed vaccines for diseases including influenza and HPV.

Building Nanoparticles

The researchers harvested the spike proteins from the moth cells and assembled them into nanoparticles.

While the nanoparticles mimicked the molecular structure of the coronavirus, they could not replicate or cause Covid-19.

Presenting the Spike

The vaccine is injected into the muscles of the arm. Each injection includes many spike nanoparticles, along with a compound extracted from the soapbark tree. The compound attracts immune cells to the site of the injection and causes them to respond more strongly to the nanoparticles.

Spotting the Intruder

Immune cells called antigen-presenting cells encounter the vaccine nanoparticles and take them up.

An antigen-presenting cell tears apart the spike proteins and displays some

of their fragments on its surface. A so-called helper T cell may detect the fragments. If a fragment fits into one of its surface proteins, the T cell becomes activated. Now it can recruit other immune cells to respond to the vaccine.

Making Antibodies

Another type of immune cell, called a B cell, may also encounter the vaccine nanoparticles. B cells have surface proteins in a huge variety of shapes, and a few might have the right shape to latch onto a spike protein. If a B cell does latch on, it can pull the vaccine particle inside and present spike

protein fragments on its surface.

If a helper T cell activated against the spike protein latches onto one of these fragments, it activates the B cell. Now the B cell proliferates and pours out antibodies that have the same shape as its surface proteins.

Stopping the Coronavirus

If vaccinated people are later exposed to the coronavirus, their antibodies can lock onto the spike proteins. The coronavirus cannot enter cells, and the infection is blocked.

Killing Infected Cells

The Novavax vaccine can also trigger

another kind of protection by destroying infected cells. When a coronavirus invades, infected cells put fragments of its spike protein on their surface. Antigen-presenting cells can activate a type of immune cell called a killer T cell. It can recognize [coronavirus-infected cells](#) and destroy them before they have a chance to produce new viruses.

Remembering the Virus

Novavax's vaccine would be easier to distribute and store than the vaccines from [Pfizer-BioNTech](#) and [Moderna](#).

While those vaccines have to be kept frozen, NVX-CoV2373 can stay stable

for up to three months in a refrigerator. But if the vaccine does turn out to be effective, scientists won't know for sure how long it provides protection.

If it works like protein-based vaccines for other diseases, it may create a group of special cells called memory B cells and memory T cells. These cells will retain information about the coronavirus for years or even decades, enabling a quick counterattack in response to a new infection.

Vaccine Timeline

January, 2020 Novavax begins work on a coronavirus vaccine.

A screen showing protein structures at

a Novavax lab in Maryland. Andrew Caballero-Reynolds/Agence France-
Presse

May Novavax launches clinical trials for their vaccine.

July The U.S. government awards Novavax [\\$1.6 billion](#) to support the vaccine's clinical trials and manufacturing.

August Novavax launched a [Phase 2 trial](#) on 2,900 people in South Africa.

Preparing an injection in Johannesburg, South Africa. Joao Silva/The New York Times

September Novavax [launches](#) a [Phase 3 trial](#) with up to 15,000

volunteers in the United Kingdom. The trial is expected to [deliver results](#) in early 2021.

Dec. 28 Novavax launches a [Phase 3 trial](#) with 30,000 people in the United States. The trial had been delayed because of problems with manufacturing the doses required for the study.

March 11, 2021 Novavax [reported](#) that their United Kingdom trial determined an efficacy rate of 96 percent against the original coronavirus. But in South Africa, where volunteers were exposed to the variant B.1.351, the efficacy was only 49 percent. The company is developing a new version of the

vaccine that is tailored to that variant.

April 13 Novavax [says](#) it could reach its production goal of 150 million doses per month by the third quarter of 2021.

May 3 Novavax expands its U.S. Phase 3 trial to include [volunteers as young as 12](#).

Sources: National Center for Biotechnology Information; Nature Reviews Immunology; Science; Maria Elena Bottazzi, Baylor College of Medicine; Matthew Frieman, University of Maryland School of Medicine.

Tracking the Coronavirus