

SARS-CoV-2 mutations come and go, and physicians' advice remains the same: Please vaccinate

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This is an illustration of a SARS-CoV-2 particle, top left. To the right of the particle, the viral genome is stretched out. Genes are represented by boxes of different colors. This single strand RNA is made up of 29,903 nucleotides (adenine, cytosine, guanine and uracil). These nucleotides are read in sets (genes) and provide instruction on how to make proteins, an example of which is shown for the box labeled "spike," which are instructions for the spike proteins. When



nucleotides are deleted, inserted, substituted, or switched (recombination), changes in the resulting protein and the eventual structure of the viral particle can lead to changes that make the virus more effective at infection, less adapted to infect humans, or provide no harm or benefit at all. Credit: Mayo Clinic

Mayo Clinic experts say that regardless of the variant, prevention of infection works. The vaccines reduce and prevent hospitalization and death based on current knowledge. If a person can get vaccinated or is eligible for a booster, they should do it now.

But what about these variants? Why do they show up, and how? Here is what Mayo Clinic experts are saying:

Two years ago, on Dec. 12, a group of people were infected and sickened from a virus that had never been known to infect humans. SARS-CoV-2, the virus that causes COVID-19, has spread around the globe, causing more than 267 million cases of infection in these two years of the pandemic, including 49 million cases in the U.S. Within each host as the virus spreads, the viral genome changes or mutates. This genetic alteration, called mutagenicity, can affect how easily the virus spreads, the severity of COVID-19 once infected, or the level of immunity from a vaccine or prior infection. The most concerning changes are tracked as they become more prominent in different geographic regions of the world. Researchers and clinicians keep an eye on these variants—like delta or omicron—so they can figure out how the mutations in these variants affect testing or patient care.

A virus reproduces by copying its genetic code and forming new virus particles to infect as many host cells as possible. In the case of SARS-CoV-2, the virus stores its <u>genetic code</u> as RNA. It prioritizes the speed of copying over accuracy, and mutations can show up in each round of



replication. From the time of infection through the first few days, SARS-CoV-2 particles may double in number about every six hours.

As that RNA replication happens, errors are made. These errors are called mutations, and if they change the virus in ways that help it spread or infect cells, it can lead to a new strain of virus, called a variant. Mutations in the viral genome, especially to the instructions for spike proteins that help SARS-CoV-2 infect cells, can result in a viral particle that is better at avoiding the immune system, resistant to previous drug treatments, or one that can infect cells more efficiently. But other copying errors may hurt the virus's reproduction or have no effect at all.

Eventually, unless precautions are taken, an assortment of viral particles spreads to the next host, where the virus gets to try again.





Diagnostic samples from people with a SARS-CoV-2 infection are sequenced globally. From the unique mutations that appear in specific regions or at certain times, scientists can create a "family tree" of what mutations are helping a particular variant spread. As a lineage diverges, it gets a letter name (A, B) or if it's a mutation of a mutation, a number (B, B.1, B.1.1.1). This naming system is called a PANGO lineage. Credit: Mayo Clinic

"You get infected by a population of viruses," says Richard Kennedy, Ph.D., an immunologist and co-director of Mayo Clinic's Vaccine Research Group. "Each individual virus could be different from the others because mutations happen at random locations."

With millions of SARS-CoV-2 infections, scientists will likely see most of the options for mutations, writes Andrew Badley, M.D., Mayo Clinic infectious diseases specialist, in a recent publication. He is chair of the Mayo Clinic SARS-CoV-2 COVID-19 Research Task Force, which oversees Mayo Clinic's COVID-19-related research activities. In the commentary, Dr. Badley notes that it essential to screen for and collect information about these mutations, and evaluate their impact on how efficiently the virus spreads, how sick they make people, and whether the new variants resist current treatments and experimental therapeutics for COVID-19.

Not all mutations worry scientists—remember, some show up and then die out if they don't help the virus spread. But some help it spread directly or indirectly. As part of the immune response, the body makes a variety of antibodies, all of which bind to a small piece of viral protein. If the virus mutates, the protein made from those instructions might change enough that the antibody can't bind to it or binds less tightly. And that, writes Dr. Badley, could reduce or nullify beneficial effects of



therapies that rely heavily on antibodies: monoclonal antibodies therapies and, to a certain extent, vaccination.

Equally important are the mutations at sites used in molecular diagnostic tests, writes Joseph Yao, M.D., Mayo Clinic microbiologist and expert in diagnostic testing. Those changes could decrease the accuracy or precision of these tests, making it harder to detect these infections. And that will indirectly lead to the spread of SARS-CoV-2.

The known effects of variants are a potential increase in transmissibility of the virus, even to the point of reinfection of those who are vaccinated or have recovered from a previous SARS-CoV-2 infection. New variants may increase hospitalization and death or infect either younger or older people than previous variants. They also may require additional tests, writes Dr. Yao, including sequencing of the viral genome more often to maintain surveillance on the variants and ensure the current diagnostic tests remain accurate.

Regardless, Mayo's experts agree: Prevention is worth a pound of cure.

"Masking, social distancing and hand-washing will all decrease our chance of exposure, no matter what variants are out there," Dr. Kennedy says. "If we follow all the recommendations for masking, distancing and hand-washing, and get a COVID-19 <u>vaccine</u>, we will have multiple layers of protection in place to keep us safe."

Provided by Mayo Clinic

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