

# How the omicron subvariant BA.5 became a master of disguise, and what it means for the current COVID-19 surge

July 27 2022, by Suresh V. Kuchipudi

---



Credit: Pixabay/CC0 Public Domain

The omicron subvariant known as BA.5 was [first detected in South Africa](#) in February 2022 and spread rapidly throughout the world. As of the second week of July 2022, BA.5 constituted [nearly 80% of COVID-19 variants in the United States](#).

Soon after researchers in South Africa reported the [original version of the omicron variant \(B.1.1.529\)](#) on Nov. 24, 2021, many scientists—including me—speculated that if omicron's numerous mutations made it either more transmissible or better at immune evasion than the preceding [delta variant](#), omicron could become the dominant [variant](#) around the world.

The omicron variant did indeed become [dominant early in 2022](#), and several sublineages, or subvariants, of omicron have since emerged: BA.1, BA.2, BA.4 and BA.5, among others. With the continued appearance of such highly transmissible variants, it is evident that SARS-CoV-2, the virus that causes COVID-19, is effectively using classic techniques that viruses use to escape the immune system. These escape strategies range from [changing the shape of key proteins](#) recognized by your immune system's protective antibodies to [camouflaging its genetic material](#) to fool [human cells](#) into considering it a part of themselves instead of an invader to attack.

I am a [virologist](#) who studies emerging viruses and viruses that jumped from animals to humans, [such as SARS-CoV-2](#). My research group has been tracking the transmission and evolution of SARS-CoV-2, evaluating changes in how well the omicron subvariants evade the [immune system](#) and the severity of disease they cause after infection.

## How is virus transmissibility in a population measured?

The basic reproduction number,  $R_0$ —pronounced "R-naught"—[measures the transmissibility of a virus](#) in a yet-uninfected population.

Once a proportion of individuals in a population become immune due to

prior infection or vaccination, epidemiologists use the term [effective reproduction number](#), called  $R_e$  or  $R_t$ , to measure the transmissibility of the virus. The  $R_e$  of the omicron variant has been estimated to be [2.5 times higher](#) than the delta variant. This increased transmissibility most likely helped omicron out-compete delta to become the dominant variant.

The larger question, then, is what is driving the evolution of omicron sublineages? The answer to that is a well-known process called [natural selection](#). Natural selection is an evolutionary process where traits that give a species a reproductive advantage continue to be passed down to the next generation, while traits that don't are phased out through competition. As SARS-CoV-2 continues to circulate, [natural selection](#) will favor mutations that give the virus the greatest survival advantage.

### **What makes omicron and its offshoots so stealthy at spreading?**

Several mechanisms contribute to the increased transmissibility of SARS-CoV-2 variants. One is the ability to [bind more strongly to the ACE2 receptor](#), a protein in the body that primarily helps regulate blood pressure but can also help SARS-CoV-2 enter cells. The more recent omicron sublineages have mutations that make them better at escaping antibody protection while retaining their ability to effectively bind to ACE2 receptors. The BA.5 sublineage can [evade antibodies](#) from both vaccination and prior infection.

Omicron sublineages BA.4 and BA.5 share several mutations with earlier omicron sublineages, but also have three unique mutations: [L452R](#), [F486V](#) and reversion (or the lack of mutation) of [R493Q](#). L452R and F486V in the spike protein help BA.5 escape antibodies. In addition, the L452R mutation helps the virus [bind more effectively](#) to the membrane of its host cell, a crucial feature associated with COVID-19 disease severity.

While the other mutation in BA.5, F486V, may help the sublineage escape from certain types of antibodies, it could decrease its ability to bind to ACE2. Strikingly, BA.5 appears to compensate for decreased ACE2 binding strength through another mutation, R493Q reversion, that is thought to [restore its lost affinity for ACE2](#). The ability to successfully escape immunity while maintaining its ability to bind to ACE2 may have potentially contributed to the rapid global spread of BA.5.

In addition to these immune-evading [mutations](#), SARS-CoV-2 has been evolving to [suppress its hosts'—in this case, humans'—innate immunity](#). Innate immunity is the body's first line of defense against invading pathogens, comprised of antiviral proteins that help fight viruses. SARS-CoV-2 has the ability to suppress the activation of some of these key antiviral proteins, meaning it's able to effectively get past many of the body's defenses. This explains the spread of infections among vaccinated or previously infected people.

Innate immunity exerts a strong selective pressure on SARS-CoV-2. Delta and [omicron](#), the two most recent and highly successful SARS-CoV-2 variants, [share several mutations](#) that could be key in helping the virus breach innate immunity. However, scientists do not yet fully understand what changes in BA.5 might allow it to do so.

## What's next?

BA.5 will not be the end game. As the virus continues to circulate, this evolutionary trend will likely lead to the emergence of more transmissible variants that are capable of immune escape.

While it is difficult to predict what variants will arrive next, we researchers cannot rule out the possibility that some of these variants could lead to increased disease severity and higher hospitalization rates. As the virus continues to evolve, most people will [get COVID-19](#)

[multiple times](#) despite vaccination status. This could be confusing and frustrating for some, and may contribute to [vaccine hesitancy](#). Therefore, it is essential to recognize that vaccines [protect you from severe disease and death](#), not necessarily from getting infected.

Research over the past two and a half years has helped scientists like me learn a lot about this new virus. However, many unanswered questions remain because the virus constantly evolves, and we are left trying to target a constantly moving goal post. While [updating vaccines](#) to match circulating variants is an option, it may not be practical in the short term because the [virus](#) evolves too quickly. Vaccines that [generate antibodies against a broad range of SARS-CoV-2 variants](#) and a cocktail of [broad-ranging treatments](#), including monoclonal antibodies and antiviral drugs, will be critical in the fight against COVID-19.

This article is republished from [The Conversation](#) under a Creative Commons license. Read the [original article](#).

Provided by The Conversation

Citation: How the omicron subvariant BA.5 became a master of disguise, and what it means for the current COVID-19 surge (2022, July 27) retrieved 10 April 2024 from <https://medicalxpress.com/news/2022-07-omicron-subvariant-ba5-master-disguise.html>

<p>This document is subject to copyright. Apart from any fair dealing for the purpose of private study or research, no part may be reproduced without the written permission. The content is provided for information purposes only.</p>
--