

Updated Information on Availability and Use of Treatments for Outpatients with Mild to Moderate COVID-19 Who are at Increased Risk for Severe Outcomes of COVID-19






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Summary


The Centers for Disease Control and Prevention (CDC) is issuing this Health Alert Network (HAN) Health Advisory to update healthcare providers, public health departments, and the public about the availability and use of recommended therapies for COVID-19 and to advise against using unproven treatments that have known or potential harms for outpatients with mild to moderate COVID-19. For patients with mild to moderate COVID-19 who are not hospitalized and who are at [increased risk](#) for severe COVID-19 outcomes, several [treatment options](#)  are now widely available and accessible.


Systemic corticosteroids are [not recommended](#)  to treat patients with mild to moderate COVID-19 who do not require supplemental oxygen; patients who are receiving dexamethasone or another corticosteroid for other indications should continue therapy for their underlying conditions as directed by their healthcare providers. Antibacterial therapy is [not recommended](#)  for the treatment of COVID-19 in the absence of another indication.

Staying [up to date](#) with COVID-19 vaccination is still the best way to prevent serious outcomes of COVID-19, including severe disease, hospitalization, and death.

Background

Early outpatient treatment of COVID-19 can avert serious, potentially life-threatening illness and reduce burden on the healthcare system. CDC issued a [HAN Health Advisory on December 31, 2021](#) to address using therapeutics in the outpatient setting for people with COVID-19. At that time, Omicron cases were increasing rapidly in the United States and some COVID-19 therapeutics were in short supply. Now antivirals for COVID-19 are widely available and can be accessed with a provider prescription at pharmacies nationwide and at [Test to Treat](#)  locations.

Data from CDC (1, 2) (highlighted in a [February 13, 2021 CDC/](#)  Infectious Diseases Society of America COVID-19 Clinical Call) and the Food and Drug Administration (3) suggest that there has been increasing use of systemic corticosteroids and antibiotics to treat outpatients with COVID-19. However, these drugs can cause harm and provide no demonstrated benefit in patients with COVID-19 with no supplemental oxygen requirement or bacterial coinfection. Short courses of systemic corticosteroids have been associated with adverse events such as hyperglycemia, gastrointestinal bleeding, psychosis, infections, and longer-term effects (4–7).

The National Institutes of Health (NIH) provides [COVID-19 Treatment Guidelines](#) . The guidelines panel provides treatment options and recommends against using systemic corticosteroids to treat patients with mild to moderate COVID-19 who do not require supplemental oxygen (**Figure**). Patients who are receiving dexamethasone or another corticosteroid for other indications should continue therapy for their underlying conditions as directed by their healthcare providers. Systemic corticosteroids are recommended for hospitalized patients with COVID-19 who require supplemental oxygen or higher-level respiratory support.

The guidelines panel also recommends against using antibacterial therapy for COVID-19 in the absence of another indication. Antibacterial drugs have no benefit in treating viral infections and can cause harm.

Figure. Therapeutic Management of Nonhospitalized Adults with COVID-19 (from [NIH COVID-19 Treatment Guidelines](#) , last updated: April 8, 2022)

PATIENT DISPOSITION	PANEL'S RECOMMENDATIONS
Does Not Require Hospitalization or Supplemental Oxygen	<p>All patients should be offered symptomatic management (AIII).</p> <p>For patients who are at high risk of progressing to severe COVID-19,^a use 1 of the following treatment options:</p> <p>Preferred Therapies <i>Listed in order of preference:</i></p> <ul style="list-style-type: none">• Ritonavir-boosted nirmatrelvir (Paxlovid)^{b,c} (AIIa)• Remdesivir^{c,d} (BIIa) <p>Alternative Therapies <i>For use <u>ONLY</u> when neither of the preferred therapies are available, feasible to use, or clinically appropriate. Listed in alphabetical order:</i></p> <ul style="list-style-type: none">• Bebtelovimab^e (CIII)• Molnupiravir^{c,f} (CIIa) <p>The Panel recommends against the use of dexamethasone^g or other systemic corticosteroids in the absence of another indication (AIII).</p>

Rating of Recommendations: A = Strong; B = Moderate; C = Weak
Rating of Evidence: I = One or more randomized trials without major limitations; IIa = Other randomized trials or subgroup analyses of randomized trials; IIb = Nonrandomized trials or observational cohort studies; III = Expert opinion

^a For a list of risk factors, see the CDC webpage [Underlying Medical Conditions Associated With Higher Risk for Severe COVID-19](#).

^b Ritonavir-boosted nirmatrelvir has significant drug-drug interactions. Clinicians should carefully review a patient's concomitant medications and evaluate potential drug-drug interactions.

^c If a patient requires hospitalization after starting treatment, the full treatment course can be completed at the healthcare provider's discretion.





^d Administration of remdesivir requires 3 consecutive days of IV infusion.



^e Bebtelovimab is active in vitro against all circulating Omicron subvariants, but there are no clinical efficacy data from placebo-controlled trials that evaluated the use of bebtelovimab in patients who are at high risk of progressing to severe COVID-19. Therefore, bebtelovimab should be used only when the preferred treatment options are not available, feasible to use, or clinically appropriate.

^f Molnupiravir has lower efficacy than the preferred treatment options. Therefore, it should be used only when the preferred options are not available, feasible to use, or clinically appropriate.



^g There is currently a lack of safety and efficacy data on the use of this agent in outpatients with COVID-19; using systemic glucocorticoids in this setting may cause harm.

Recommendations for Healthcare Providers



1. Obtain updated information on appropriate use of clinically indicated therapeutics through [NIH's COVID-19 Treatment Guidelines](#) .
2. Prescribe **COVID-19 therapeutics** for patients when clinically indicated.
 - There are considerable differences in efficacy, risk profiles, and use restrictions between the two oral antivirals. Healthcare providers need to be familiar with these distinctions to make clinical decisions and inform patients. In addition, initiating treatment with these oral antivirals must begin within five days of symptom onset to maintain product efficacy.
 - Please see [NIH's COVID-19 Treatment Guidelines](#)  for important therapeutic considerations, such as the potential for significant drug-drug interactions with ritonavir-boosted nirmatrelvir ([Paxlovid](#) ) and dosing regimens for patients with renal impairment.
3. Obtain information on **access to outpatient COVID-19 treatments**, including pharmacies where antivirals for COVID-19 are distributed and [Test to Treat](#)  locations.

4. Do not use **dexamethasone and other systemic corticosteroids** to treat patients with mild to moderate COVID-19 who do not require hospitalization or supplemental oxygen; these drugs have no proven benefit in these patients and can cause harm.
5. Do not use **antibacterial therapy** to treat COVID-19 in the absence of another indication; these drugs have no benefit for treating viral infections and can cause harm.
6. To prevent serious outcomes of COVID-19, including severe disease, hospitalization, and death, encourage all patients to remain **up to date with COVID-19 vaccination**.
 - People who are immunocompromised or severely allergic to COVID-19 vaccines may receive tixagevimab co-packaged with cilgavimab ([Evusheld](#) ) , a **long-acting combination monoclonal antibody therapy** given by intramuscular injection for pre-exposure prophylaxis of COVID-19. To find Evusheld distribution locations, providers can go to the [COVID-19 Therapeutics Locator](#)  , call the support line at **1-800-232-0233** (TTY **888-720-7489**), or contact their individual state or territorial health planners.






Recommendations for Public Health Departments and Public Health Jurisdictions

1. Maintain awareness of **locations of available therapeutics** within your jurisdictions, including pharmacies where antivirals for COVID-19 are distributed and [Test to Treat](#)  locations.
2. Communicate ongoing and up-to-date information on therapeutics for COVID-19 and their availability to healthcare providers within your jurisdiction.
3. Disseminate information for the Test to Treat call center at **1-800-232-0233** (TTY **1-888-720-7489**) which provides information in more than 150 languages, and for the [Disability Information and Access Line](#)  at 1-888-677-1199.



Recommendations for the Public






1. If you test positive and are an older adult or someone who is at **increased risk** of getting very sick from COVID-19, **treatment is available**. **Contact a healthcare provider right away after a positive test to determine if you are eligible for treatment**, even if your symptoms are mild. You can also visit a [Test to Treat](#)  location and, if eligible, receive a prescription from a provider at that location.
2. Follow [CDC guidance on testing for COVID-19](#) and use the [Test to Treat locator](#)  or call **1-800-232-0233** (TTY **1-888-720-7489**) to find a testing location that can provide treatment if you test positive.
3. **Don't delay**: Treatment must be started within the first few days of when your symptoms started to be effective.
4. Staying **up to date with COVID-19 vaccination** is still the best way to prevent serious outcomes of COVID-19, including severe disease, hospitalization, and death.

For More Information

- [CDC COVID-19 Treatment website](#)
- [NIH COVID-19 Treatment Guidelines](#) 
- [NIH COVID-19 Treatment Guidelines: Therapeutic Management of Nonhospitalized Adults with COVID-19](#) 
- [Interim Clinical Considerations for Use of COVID-19 Vaccines | CDC](#)
- [NIH COVID-19 Treatment Guidelines: Prevention of SARS-CoV-2 Infection](#) 
- [Office of the Assistant Secretary for Preparedness & Response \(ASPR\) Test to Treat website](#) 
- [U.S. Food and Drug Administration COVID-19 Therapeutic Product Emergency Use Authorizations](#) 
- [CDC COVID Data Tracker](#)

References

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2. Tsay SV, Bartoces M, Goulin K, Kabbani S, Hicks, LA. Antibiotic prescriptions associated with COVID-19 visits among Medicare beneficiaries, April 2020 to April 2021. JAMA 2022. <https://jamanetwork.com/journals/jama/fullarticle/2791077> 

3. Bradley MC, Perez-Vilar S, Chillarige Y, Dong D, Martinez AI, Weckstein AR, Dal Pan GJ. Systemic corticosteroid use for COVID-19 in U.S. outpatient settings from April 2020 to August 2021. JAMA 2022. <https://jamanetwork.com/journals/jama/fullarticle/2791078> 
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5. The RECOVERY Collaborative Group. Dexamethasone in hospitalized patients with COVID-19. N Engl J Med 2021;384:693-704. <https://www.nejm.org/doi/full/10.1056/nejmoa2021436> 
6. Crothers K, DeFaccio R, Tate J, et al. Dexamethasone in hospitalised coronavirus-19 patients not on intensive respiratory support. Eur Resp J 2021. <https://erj.ersjournals.com/content/early/2021/11/18/13993003.02532-2021> 
7. Li Q, Li W, Jin Y, et al. Efficacy evaluation of early, low-dose, short-term corticosteroids in adults hospitalized with non-severe COVID-19 pneumonia: a retrospective cohort study. Infect Dis Ther 2020;9:823-36. <https://pubmed.ncbi.nlm.nih.gov/32880102/> 

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DEPARTMENT OF HEALTH AND HUMAN SERVICES

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- **Health Advisory:** Provides important information for a specific incident or situation; may not require immediate action.
- **Health Update:** Provides updated information regarding an incident or situation; unlikely to require immediate action.
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This message was distributed to state and local health officers, state and local epidemiologists, state and local laboratory directors, public information officers, HAN coordinators, and clinician organizations.

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