**TREATMENT**: The National Institutes of Health (NIH) recommend the following therapies for non-hospitalized patients with confirmed COVID-19 and mild or moderate symptoms at high risk for progression to hospitalization or death. For a list of risk factors, see the CDC webpage Underlying Medical Conditions Associated with High Risk for Severe COVID-19.

The NIH recommends preference for these therapies in order listed in the table below.<sup>1</sup>

Therapeutic Agent		Dose	Но	Prevention of spitalizations or Death over 28 days		Clinical Considerations
1.	Ritonavir- boosted Nirmatrelvir (PAXLOVID) <sup>2</sup>	Nirmatrelvir 300 mg plus ritonavir 100 mg, orally twice daily for 5 days, within 5 days of symptom onset	•	ARR: 6.3% → 0.8% RRR: 88% NNT: 19	•	Significant drug-drug interactions  No known risk of ritonavir in pregnancy  Decrease nirmatrelvir to 150 mg if eGFR ≥30  to <60 mL/min  Avoid in severe hepatic impairment
2.	Sotrovimab <sup>3</sup>	500 mg IV once, within 10 days of symptom onset	•	ARR: 7.2% → 1.0% RRR: 85% NNT: 17	•	No drug-drug interactions Only mAb on this list; safer option in pregnancy
3.	Remdesivir (VEKLURY) <sup>4</sup>	200 mg IV day 1, within 7 days of symptom onset, then 100 mg IV on Days 2 and 3	•	ARR: 5.3% → 0.7% RRR: 87% NNT: 22	•	Option if IV services easily accessible Commercially available; not available through federal distribution
4.	Molnupiravir <sup>5</sup>	800 mg, orally twice daily for 5 days, within 5 days of symptom onset	ed alo	ARR: 9.7% → 6.8% RRR: 30% NNT: 35 merular filtration rate: IV = int	• • •	Option if other therapies unavailable <sup>1</sup> Concern for mutagenicity (theoretical) Avoid in pregnancy, growing children hous; mAb = monoclonal antibody; NNT = number needed-

Abbreviations: ARR = absolute risk reduction; eGFR = estimated glomerular filtration rate; IV = intravenous; mAb = monoclonal antibody; NNT = number needed-to-treat to prevent one hospitalization or death over 28 days; RRR = relative risk reduction.

<u>PRE-EXPOSURE PROPHYLAXIS</u>: Pre-exposure prophylaxis is an option for individuals who are moderately or severely immunocompromised and who are not expected to mount an adequate response to COVID-19 vaccination. It may also be used for individuals who have medical contraindications to COVID-19 vaccination.

Therapeutic Agent	Dose	Prevention of symptomatic COVID-19 infection over 6 months	Clinical Considerations			
Tixagevimab and cilgavimab (EVUSHELD) <sup>6</sup>	Tixagevimab 150 mg and cilgavimab 150 mg, IM once each	<ul> <li>1.0% → 0.2%</li> <li>RRR: 77%</li> <li>NNT: 125</li> </ul>	<ul> <li>No dose adjustments needed</li> <li>Caution with use in patients with CVD</li> <li>Avoid within 2 weeks of COVID-19 vaccine</li> <li>May repeat dosing every 6 months</li> </ul>			
Abbreviations: CVD = cardiovascular disease; IM = intramuscular; NNT = number needed-to-treat to prevent one case of symptomatic COVID-19 infection over 6 months; RRR = relative risk reduction.						

<u>POST-EXPOSURE PROPHYLAXIS</u>: No current therapies known to be effective against the Omicron variant have received EUA for post-exposure prophylaxis. Although REGEN-COV (casirivimab plus imdevimab) and bamlanivimab plus etesevimab have received EUA for post-exposure prophylaxis, they are not recommended for the Omicron variant.<sup>1</sup>

THERAPEUTICS ALLOCATION & REQUESTS: Please visit the OHA therapeutics landing page for information and resources on oral antivirals and monoclonal antibodies at: <a href="https://www.oregon.gov/oha/covid19/Pages/therapeutics.aspx">https://www.oregon.gov/oha/covid19/Pages/therapeutics.aspx</a>. Please also note guidance to providers for prescribing of therapeutics in times of constrained supply in this slide deck from U.S. Department of Health and Human Services: <a href="https://www.oregon.gov/oha/covid19/Documents/HHS-tiering-recommendation.pdf">https://www.oregon.gov/oha/covid19/Documents/HHS-tiering-recommendation.pdf</a>. Please pay special attention to slides 6-9 as it pertains to prioritization of patients.

## THERAPEUTIC OPTIONS FOR NONHOSPITALIZED PATIENTS WITH OMICRON COVID-19 VARIANT (1-18-22)

## References:

- Statement on Therapies for High-Risk, Nonhospitalized Patients. COVID-19 Treatment Guidelines, National Institutes of Health (last updated: December 30, 2021). Available at: <a href="https://www.covid19treatmentguidelines.nih.gov/therapies/statement-on-therapies-for-high-risk-nonhospitalized-patients/">https://www.covid19treatmentguidelines.nih.gov/therapies/statement-on-therapies-for-high-risk-nonhospitalized-patients/</a>. Accessed 18 January 2022.
- 2. US Food and Drug Administration. Fact sheet for healthcare providers: emergency use authorization for Paxlovid (nirmatrelvir tablets; ritonavir tablets). Available at: <a href="https://www.fda.gov/media/155050/download">https://www.fda.gov/media/155050/download</a>. Accessed January 18, 2022.
- 3. Gupta A, Gonzalez-Rojas Y, Juarez E, Casal MC, et al. Early treatment of Covid-19 with SARS-CoV-2 neutralizing antibody sotrovimab. *N Engl J Med*. 2021; 385(21): 1941-50. doi: 10.1056/NEJMoa2107934.
- 4. Gottlieb RL, Vaca CE, Paredes R, Mera J, Webb BJ, et al. Early remdesivir to prevent progression to severe Covid-19 in outpatients. *N Engl J Med*. 2021 Dec 22. doi: 10.1056/NEJMoa2116846. Online ahead of print.
- 5. Bernal AJ, Gomes da Silva MM, Musungaie DB, Kovalchuk E, Gonzalez A, et al. Molnupiravir for oral treatment of Covid-19 in nonhospitalized patients. *N Engl J Med*. 2021 Dec 16. doi: 10.1056/NEJMoa2116044. Online ahead of print.
- US Food and Drug Administration. Fact sheet for healthcare providers: emergency use authorization for Evusheld (tixagevimab co-packaged with cilgavimab). Available at: <a href="https://www.fda.gov/media/154701/download">https://www.fda.gov/media/154701/download</a>. Accessed January 18, 2022.