



# **Regional SARS-CoV-2 Variants and their Impact on Inpatient Treatment**

*Supported by an educational grant from Gilead Sciences, Inc.*

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## LEARNING OBJECTIVE 1

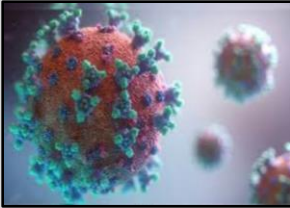
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*Analyze regional differences in SARS-CoV-2 variants and their influence on the effectiveness of inpatient COVID-19 treatment strategies*

# SARS-CoV-2 Misconceptions between Patients and Clinicians



Natural immunity provides superior and efficient protection without vaccination

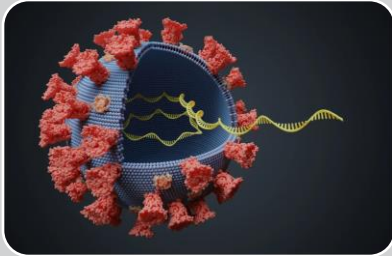


Emerging variants are more severe and current vaccines are ineffective

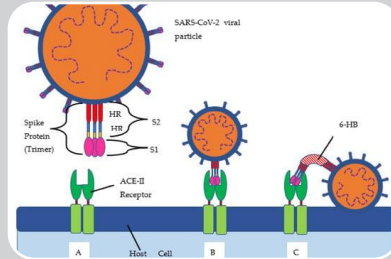


Long COVID is not a real or serious medical condition

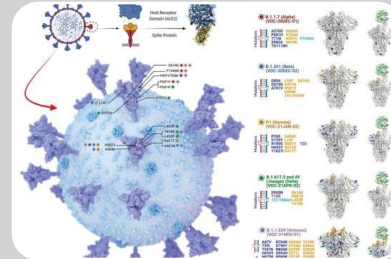
# Evolutionary Trajectory: Wuhan to Stratus/Nimbus



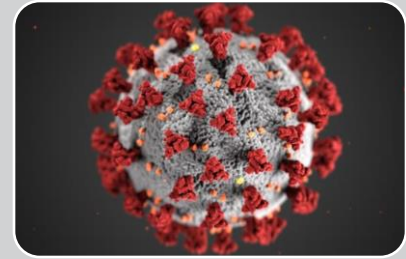
**Wuhan**  
(2019)



**Delta**  
(2021)



**Omicron**  
(2022)



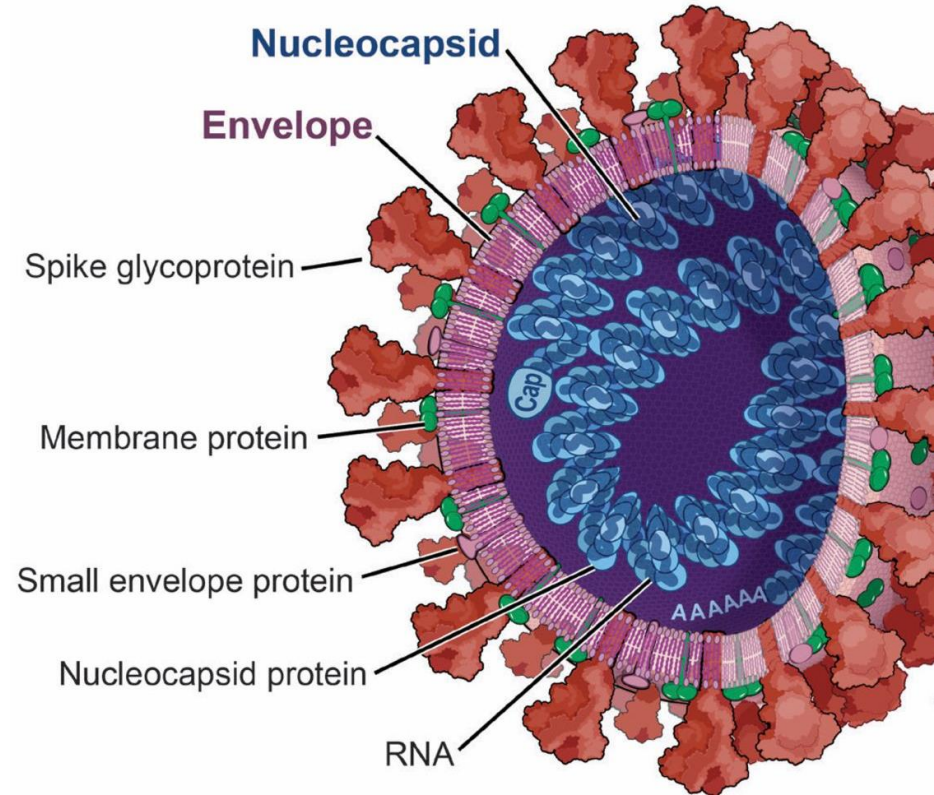
**Stratus/  
Nimbus**  
(2025)



# Biological Mechanism of SARS-CoV-2

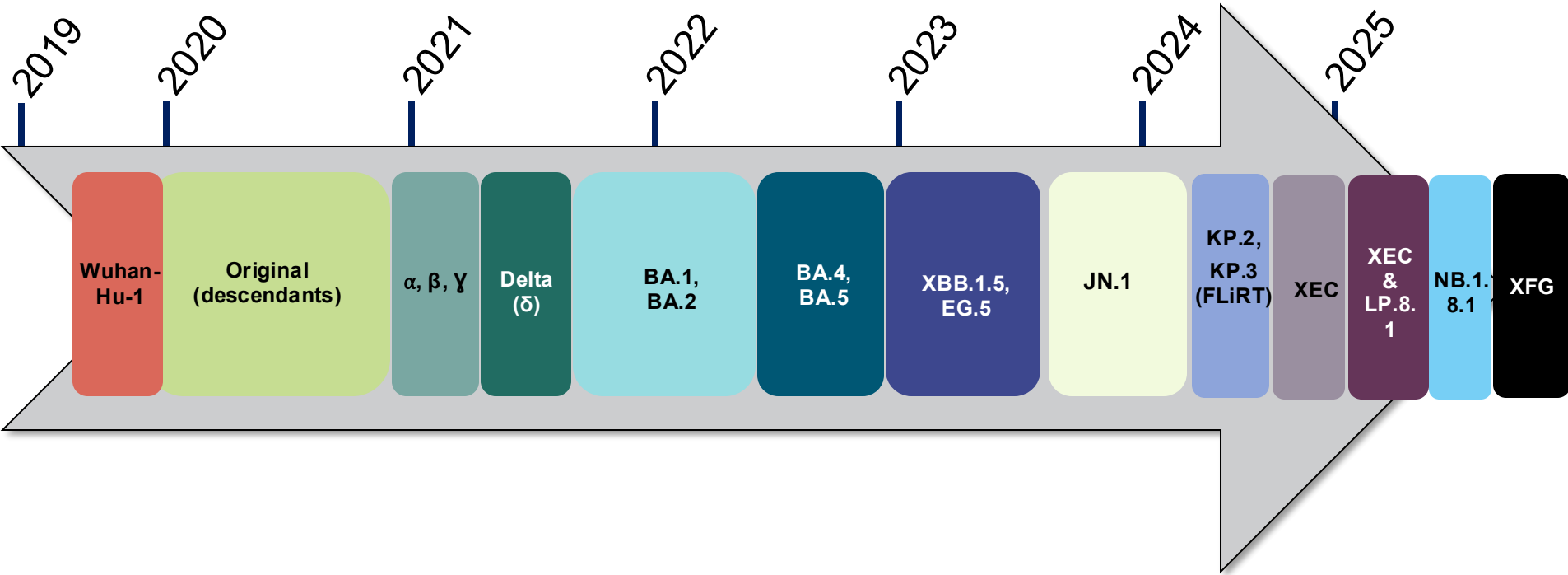
- Enveloped +ssRNA virus
- Uses **ACE2** as the obligate entry receptor
- ACE2 expressed in:
  - Nasal epithelium
  - Bronchial epithelium
  - Type II alveolar cells
  - Small intestine
  - Heart and kidney

Structural Protein	Function
<b>Nucleocapsid (N)</b>	Encapsulates the RNA genome into a helical ribonucleocapsid complex
<b>Spike (S)</b>	Viral attachment and entry into host cells
<b>Envelope (E)</b>	Virion assembly, budding, and release
<b>Membrane (M)</b>	Virus assembly and morphogenesis



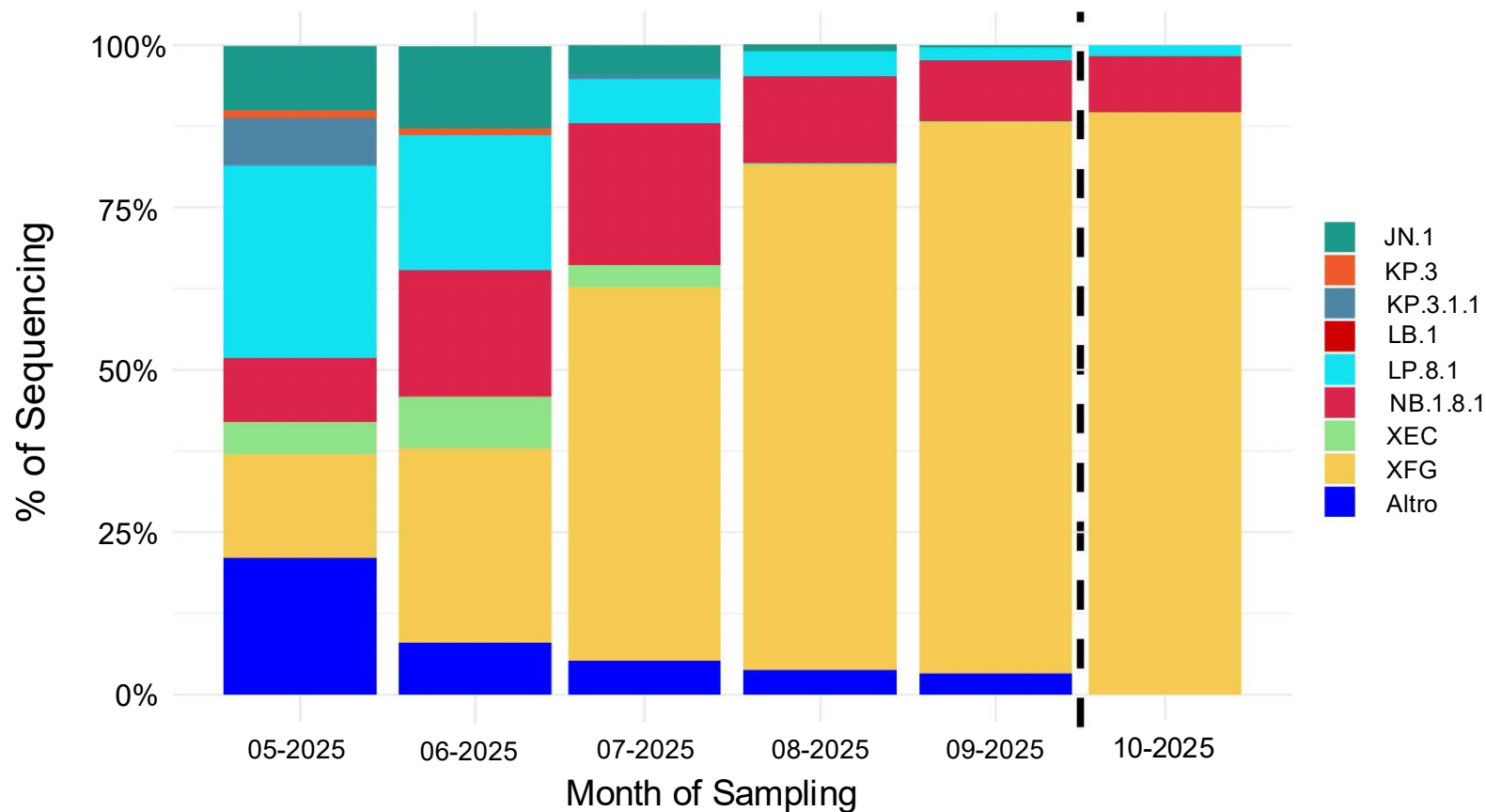
ACE2 = angiotensin-converting enzyme 2; +ssRNA = positive single-stranded ribonucleic acid

# SARS-CoV-2 Variant Timeline Progression

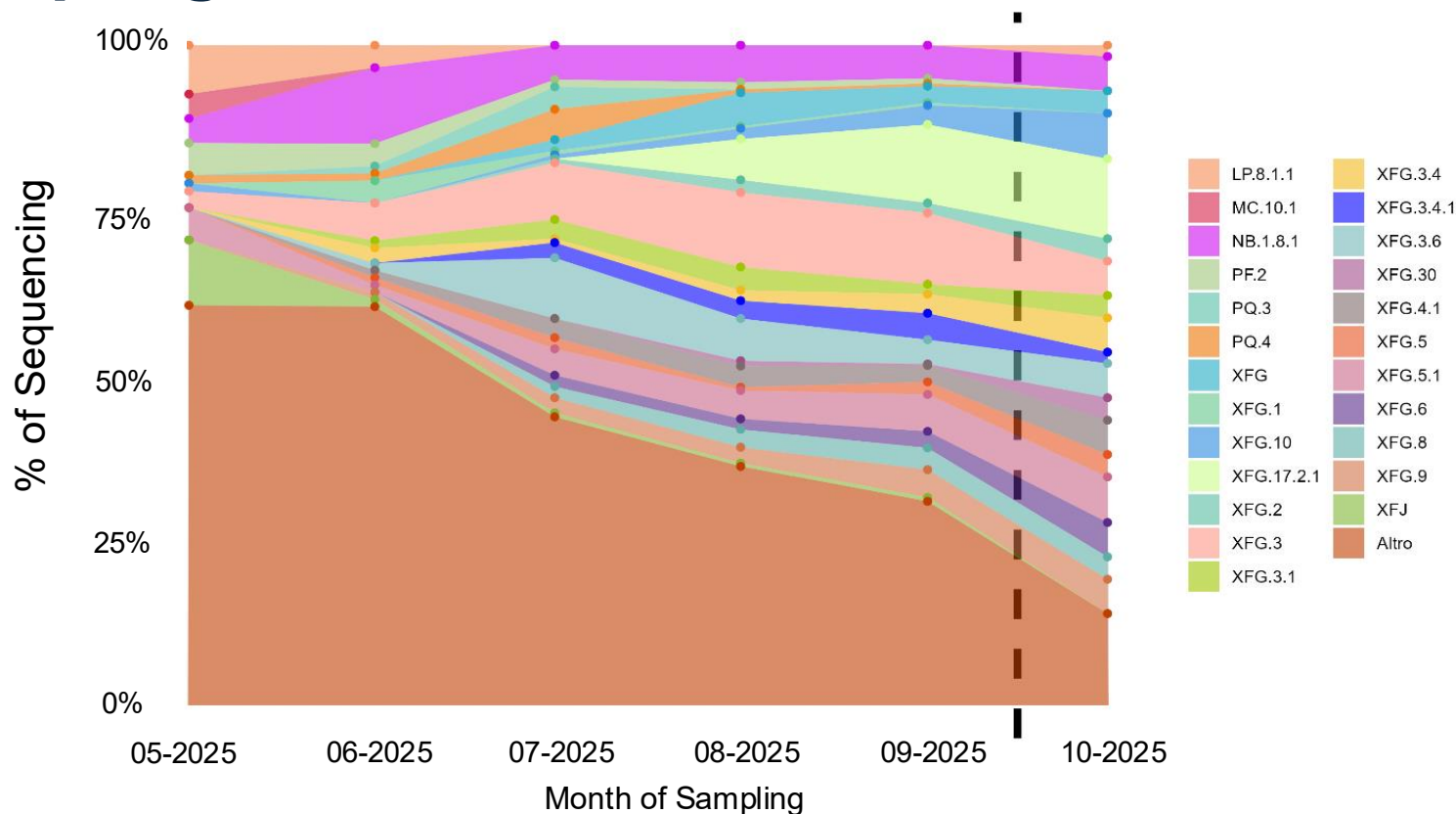




# Percentage (%) of Major SARS-CoV-2 Lineages by Sampling Month



# Percentage (%) of Major SARS-CoV-2 Sub-lineages by Sampling Month



# Regional SARS-CoV-2 Variant Circulation Overview (October 5 – November 5, 2025)



Variants of interest	Variants under monitoring	Variants of concern
<ul style="list-style-type: none"><li>• JN.1</li></ul>	<ul style="list-style-type: none"><li>• XFG</li><li>• NB.1.8.1</li><li>• LP.8.1</li><li>• KP.3.1.1</li><li>• XEC</li></ul>	<ul style="list-style-type: none"><li>• None</li></ul>

## **Dominant Circulating Variants**

XFG – Europe and Americas

NB.1.8.1 – Western Pacific

# Patient Case: PW



- PW is a 29-yr old female with rheumatoid arthritis (RA) on maintenance therapy of prednisone (10mg/day)



## Initial Presentation (Day 0)

- Reports 2 days of fever, cough, and chills

### Vitals in ED

- Temp: 38.8°C (101.8°F)
- HR: 104 bpm
- BP: 104/55 mmHg
- RR: 18 breaths/min
- SpO2 97% on RA

### Diagnostics

- CBC with diff – WBC 11,000/ $\mu$ L; neutrophils 70%, lymphocytes 12% (absolute 0.9K), eosinophils 0%
- CMP normal
- CRP mild elevation
- CXR - unremarkable

### Management

- Admitted for monitoring due to immunosuppression

# Patient Case: PW

## Hospital Course

### Day 2 (am)

Increasing dyspnea  
SpO2 92% RA  
CXR reveals bilateral infiltrates

**Started on O2 2-3L via nasal cannula; IV dexamethasone (6mg/day)**

### Day 2 (pm)

- SpO2 drops to 85%
- CRP rises to 160 mg/L - tocilizumab initiated (single IV dose)
- NIV attempted (low tidal volume, high PEEP, proning cycles initiated)

### Days 5-7

- Lung mechanics improve; FiO2 requirements fall

### Days 7-9

- Successfully extubated to nasal cannula

### Days 9-12

- Stable on room air, transferred to ward

### Vitals Chart

Day	HR	BP	Temp	RR	SpO2
Day 1	98	108/60	100.4°F	20	95% RA
Day 2 AM	112	102/58	100°F	24	92% RA
Day 2 PM	118	98/54	100.2°F	32	85% despite O2
Day 3	114	100/58	99°F	Vent-controlled	SpO2 94% FiO2 60%
Day 5	112	102/60	98.6°F	Vent-controlled	SpO2 96% FiO2 40%
Day 7	106	104/58	98°F	Vent-controlled	SpO2 97% FiO2 30%
Day 9	92	110/65	98.2°F	18	Extubated → 94% on 3 L NC
Day 12	86	112/68	98.0°F	16	97% RA



# XFG vs. NB.1.8.1 Variant Comparison



## Key Findings:

- ✓ Enhanced transmissibility, no increased severity
- ✓ Public health Risk: LOW for both variants
- ✓ Vaccines remain effective
- ✓ Consistent symptom profiles with prior omicron variants

Parameter	XFG	NB.1.8.1
Global Prevalence (Oct-Nov 2025)	71% of sequences	15% of sequences
Transmissibility	Moderate growth advantage; low ACE2 binding affinity highest relative growth rate	High ACE2 binding: 2.5-fold higher infectivity than LP.8.1
Severity	No increase vs. prior variants	No increase vs. prior variants
Immune Evasion	2-fold antibody reduction in antibody neutralization	1.6-fold antibody reduction in antibody neutralization
WHO Public Health Risk Level	Low	Low
Vaccine Effectiveness	Expected to remain effective	Expected to remain effective

World Health Organization (WHO). 2025. <https://data.who.int/dashboards/covid19/summary>; Geddes, et al. Gavi. 2025. <https://www.gavi.org/vaccineswork/eight-things-you-need-know-about-new-nimbus-and-stratus-covid-variants>; World Health Organization (WHO). *WHO TAG-VE Risk Evaluation for SARS-CoV-2 Variant Under Monitoring: XFG2025*. [https://www.who.int/docs/default-source/coronaviruse/25062025\\_xfg\\_ire.pdf](https://www.who.int/docs/default-source/coronaviruse/25062025_xfg_ire.pdf); World Health Organization (WHO). *WHO TAG-VE Risk Evaluation for SARS-CoV-2 Variant Under Monitoring: NB.1.8.1*. 2025. [https://cdn.who.int/media/docs/default-source/documents/epp/tracking-sars-cov-2/23052025\\_nb.1.8.1\\_ire.pdf](https://cdn.who.int/media/docs/default-source/documents/epp/tracking-sars-cov-2/23052025_nb.1.8.1_ire.pdf); Kavanagh K. Nature Website. 2025. <https://www.nature.com/articles/d41586-025-03412-x>; Guo C, et al. *Lancet Infect Dis*. 2025;25(7):374-377; Abir HP, et al. *Health Sci Rep*. 2025;8(11):e71453.

# Inpatient Treatment: Regional Variations

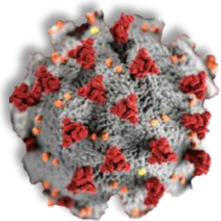
Organization	Severity Classifications	First-Line Treatments	Second-Line Treatments	VTE Prophylaxis
<b>WHO</b>	<ul style="list-style-type: none"> <li>• Non-severe</li> <li>• Severe (SpO<sub>2</sub> &lt; 90%)</li> <li>• Critical (ARDS/sepsis/shock)</li> </ul>	<ul style="list-style-type: none"> <li>• Corticosteroids (dexamethasone)</li> <li>• Antivirals</li> <li>• Supportive care</li> </ul>	<ul style="list-style-type: none"> <li>• Tocilizumab/baricitinib for severe cases</li> </ul>	<ul style="list-style-type: none"> <li>• Standard prophylactic doses LMWH</li> </ul>
<b>NICE (UK)</b>	<ul style="list-style-type: none"> <li>• Similar to WHO classifications</li> </ul>	<ul style="list-style-type: none"> <li>• Corticosteroids</li> <li>• Antivirals</li> <li>• Oxygen therapy (CPAP/HFNC)</li> </ul>	<ul style="list-style-type: none"> <li>• Tocilizumab for severe cases</li> </ul>	<ul style="list-style-type: none"> <li>• Risk-based approach</li> </ul>
<b>IDSA (USA)</b>	<ul style="list-style-type: none"> <li>• Mild-Moderate (SpO<sub>2</sub> &gt; 94%)</li> <li>• Severe (SpO<sub>2</sub> ≤ 94%)</li> <li>• Critical (mechanical ventilation/ECMO)</li> </ul>	<ul style="list-style-type: none"> <li>• Nirmatrelvir/ritonavir</li> <li>• Remdesivir</li> <li>• Corticosteroids</li> </ul>	<ul style="list-style-type: none"> <li>• Tocilizumab/baricitinib for severe cases</li> </ul>	<ul style="list-style-type: none"> <li>• ASH recommends prophylactic over intermediate doses</li> </ul>
<b>ICMR (India)</b>	<ul style="list-style-type: none"> <li>• Mild: URTI, no hypoxia</li> <li>• Moderate: RR ≥ 24/min, SpO<sub>2</sub> 90–93% on room air</li> <li>• Severe: RR &gt; 30/min, SpO<sub>2</sub> &lt; 90% on room air</li> </ul>	<ul style="list-style-type: none"> <li>• Awake proning for all hypoxic patients</li> <li>• Corticosteroids: (moderate/severe)</li> <li>• Antivirals: Restricted emergency use for moderate cases</li> </ul>	<ul style="list-style-type: none"> <li>• Tocilizumab (off-label): Considered only for rapidly progressive severe disease with raised inflammatory markers within 24-48h of steroid use</li> </ul>	<ul style="list-style-type: none"> <li>• Prophylactic anticoagulation: UFH or LMWH</li> </ul>

ASH = American Society of hematology; ARDS = acute respiratory distress syndrome; CPAP = continuous positive airway pressure; ECMO = extracorporeal membrane oxygenation; HFNC = high-flow nasal cannula; ICMR = Indian Council of Medical Research; IDSA = Infectious Diseases Society of America; LMWH = low molecular weight heparin; NICE = National Institute for Health and Care Excellence; UFH = unfractionated heparin; URTI = upper respiratory tract infection; VTE = venous thromboembolism

Infectious Diseases Society of America (IDSA). 2025. <https://www.idsociety.org/practice-guideline/SARS-CoV-2-guideline-treatment-and-management/>.  
 National Institute for Health and Care Excellence (NICE). 2021. <https://www.nice.org.uk/guidance/ng191/resources/covid19-rapid-guideline-managing-covid19-pdf-66142077109189>. World Health Organization (WHO). 2025. <https://iris.who.int/server/api/core/bitstreams/d1021eff-f570-4c22-b630-a44b4267a6c/content>. Siegal et al. *Blood Advances*. 2025; 9(6):1247–1260;  
 Sharma S, et al. *Sens Int*. 2020; 1:100013; Indian Council for Medical Research. *Clinical Guidance for Management of Adult COVID-19 Patients*. 2023. <https://covid19dashboard.mohfw.gov.in/pdf/ClinicalGuidanceforManagementofAdultCOVID19Patientsupdatetason05thjan2023.pdf>.



# Key Takeaways for Global HealthCare



## Variants do not dictate treatment

- Apply the same evidence-based standard of care across variants according more to disease severity rather than lineage



## Time is of the essence

- Early intervention is the single biggest predictor of outcome



## Treat the patient, not the variant

- Focus on age, comorbidities, and immune status when developing individualized treatment plan



## Put information into action!

Takeaways from this program can be implemented into your practice to improve patient care.

- **Utilize** guideline-concordant protocols and disease severity rather than specific variants when making inpatient SARS-CoV-2 treatment decisions in at least 80% of patients over the next 6-12 months
- **Prioritize** patient vulnerabilities over specific variants when selecting inpatient treatment plans in at least 80% of patients over the next 6-12 months
- **Recognize** the benefit of earlier versus delayed initiation of inpatient SARS-CoV-2 treatment in at least 80% of patients over the next 6-12 months

# To Receive Credit

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To receive CME/CE credit for this activity, participants must complete the post-test and evaluation online.

Participants will be able to download and print their certificate immediately upon completion.



**Other programs in this series include:**

**Part 1:**

*Guideline-Based Therapeutics  
for Hospitalized Patients with  
SARS-CoV-2 Infection*

**Part 2:**

*Early Diagnosis and Timely  
Treatment in Hospitalized  
Patients with SARS-CoV-2  
Infection*

**Part 3:**

*Risk Stratification in Hospitalized  
SARS-CoV-2 Patients*



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