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This tool was developed to provide a pragmatic framework to assist with severity classification, diagnostic workup, disposition, and treatment of patients with suspected or confirmed SARS-CoV-2 (COVID-19) in the emergency department.

- It is designed to assist with the management of adult patients (≥18 years old) with symptomatic infection.
- For information on pediatric MIS-C protocols (<u>CHOP</u>, <u>Minnesota</u>, and <u>Yale</u>)
- This tool is not intended to represent a legal standard of care for emergency physicians. Recommendations offered in this tool are not intended to represent the only diagnostic or management options available to the emergency physician. Individual physicians' judgment and consideration of patient resources/preferences is essential.
 This tool is not exhaustive in reparate to diagnostic and treatment recommendations. Patients may present with particular conditions (ML PE stroke) that could be
- This tool is not exhaustive in regards to diagnostic and treatment recommendations. Patients may present with particular conditions (MI, PE, stroke) that could be
 manifestations of severe or critical COVID-19. These conditions may require additional specific diagnostic and therapeutic interventions not discussed in this tool.
- Evidence on this topic (including differences in severity that may occur with evolving variants) is changing quickly and may alter recommendations.
- A digitized version of this tool can now be found at <u>MDCalc.</u>

Step 1 - Severity Classification - Assess the patient's severity of disease utilizing NIH criteria.

MILD	MODERATE	SEVERE	CRITICAL
Individuals who have various signs and symptoms of COVID-19 (ANY): Fever Cough Sore throat Malaise Headache Muscle pain Nausea, vomiting, diarrhea Loss of taste and smell BUT who do NOT have (ANY): Shortness of breath Dyspnea Abnormal chest imaging (if obtained)	Individuals who show evidence of lower respiratory disease during (ANY): □ Clinical assessment □ Imaging AND who have: □ Sp02 ≥94% on room air at sea level (in those with normal baseline Sp02 at rest)	Individuals who have (ANY): □ Sp02 <94% on room air at sea level (in those with normal baseline Sp02 at rest) □ Ratio of arterial partial pressure of oxygen to fraction of inspired oxygen (Pa02/Fi02) <300 mm Hg (if ABG obtained) □ RR >30 breaths/min □ Lung infiltrates >50%	Individuals with (ANY): Respiratory failure Septic shock Multiorgan dysfunction or failure
Consider Risk Prognostication and As	ssessment (see Page 3)		

Step 2 - Diagnostic Testing

The following imaging and lab tests should be considered based on your patients severity and risk for disease progression. Diagnosing acute SARS-CoV-2 infection solely on the basis of serologic test results is not recommended by NIH. Antigen Testing Algorithm for Community Settings

MILD	MODERATE	SEVERE	CRITICAL				
Based on clinician's judgement, diagnostic testing may not be necessary in patients with (ALL): □ Mild Severity □ PRIEST score ≤4 (See Page 3) □ 1 or less Risk Factors	Imaging: the optimal imaging technique people with symptomatic COVID-19. In patients may include: □ Chest X-ray □ Pulmonary Ultrason ECG: should be performed if indicated □ ECG	dditional tests to consider include: ABG Coagulation screen - (d-dimer, PT/ PTT, fibrin degradation products) Inflammatory markers - (procalcitonin / c-reactive protein)					
Exertional Sp02 may have limited ability to identify adverse outcomes in otherwise well-appearing patients:	Labs: CBC w/ differential CMP It is recommended to utilize ACEP's Laboratory Abnormalities to review	COVID-19 Field Guide section on	□ Ferritin □ LDH □ CK, CK-MB □ Troponin □ Blood and sputum cultures				

Step 3 - Disposition

The following represents a pragmatic approach for disposition of patients depending on their disease severity. Clinicians may want to consider a patient's risk for progression of disease based on PRIEST Score / Risk Assessment (see Page 3), imaging, and labs in their disposition decision. See Steps 4 and 5 on the next page for treatment guidance.

MILD	MODERATE	SEVERE	CRITICAL
 □ Discharge Home □ Supply patient with educational materials on precautions and items to be monitorinag at home In patients with PRIEST Score ≥5 and/or multiple Risk Factors □ Clinicians should consider early follow-up with primary care physician or other health system access points. □ Patient should be educated on their increased risk for severe disease and precautions to return to the ED. 	 Discharge Home, consider if ALL: PRIEST Score ≤4 1 (or less) Risk Factors No concerning Imaging or Lab results Capability and resources to care for self at home No other condition that warrants admission Admission, consider if ANY: PRIEST Score ≥5 Multiple Risk Factors Concerning Imaging or Lab results Does NOT have the capability or resources to care for self at home Admission Location: Based on clinician's judgement Observation Inpatient Floor Intermediate 	Admission Location: based on clinician's judgement Floor Bed Intermediate ICU Transfer Consider transfer if your facility does not have the resources or capacity to care for a severe COVID patient that could deteriorate.	Admission CU Transfer Consider transfer if your facility does not have the resources or capacity to care for a critically ill COVID patient. Consider transfer to an ECMO facility for patients who may benefit from this after consultation with receiving facility.
CDC Patient Educational Materials • S	AEM Patient Toolkit	\rangle	1



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Step 4 - Non-Pharmacologic Treatment

The following treatments should be considered based on your patient's severity and risk of disease progression.

MILD	MODERATE	SEVERE	CRITICAL						
 □ Consider home oxygen the may benefit) □ Monitor home 02 saturate monitor □ Progressive ambulation at contraindication) □ Resting in the prone posite □ Adequate rest/sleep □ Balanced diet □ Adequate hydration COVID-19 vaccination is refeveryone 6 months of age at of a history of symptomatic SARS-CoV-2 infection. - People who recently had 3 may consider delaying the vaccination dose by 3 mo onset or positive test (if in asymptomatic). - Additional information HE FAQs 	ion with portable as tolerated (if no tion if dyspneic commended for and older, regardless c or asymptomatic SARS-CoV-2 infection eir next COVID-19 nths from symptom fection was	 Oxygen support-nasal cannula, titrate up to 6L with an oxygenation goal of >92% High-Flow Nasal Cannula (HFNC) or high-velocity therapy (titrated up to a flow of 60L and FiO2 up to 100%) are recommended over NIPPV ^{42,43}. This intervention may reduce need for mechanical, ventilation but has not been demonstrated to reduce mortality. (AII) Non-Invasive Postive Pressure Ventilation (NIPPV) if HFNC not available Consider trial of awake prone positioning if patient can be monitored or can self rescue. Awake proning is contraindicated in patients in respiratory distress. Insufficient data recommend for or against use Nitric Oxide³⁸. Recommend against routine use of Heliox; may be considered in croup-like pediatric presentations³⁹. 	 ☐ Intubation is recommended for severe respiratory failure: ☐ Oxygenation goal for ventilated patients should be 92-96%. ☐ Consider low tidal volume (VT) ventilation (VT 4–8 mL/kg of predicted body weight) over higher VT ventilation (VT >8 mL/kg) (Al). ☐ Target plateau pressures of <30 cm H20 (All). ☐ A higher positive end-expiratory pressure (PEEP) strategy is recommended over a lower PEEP strategy (BII). ☐ For mechanically ventilated adults with refractory hypoxemia despite optimized ventilation, consider prone ventilation. ☐ Consider using a conservative fluid strategy over a liberal fluid strategy (BII). ☐ Venovenous ECMO appears to be an effective intervention in selected patients with COVID-19-related ARDS (All)⁴¹ ☐ Insufficent Data to recommend for or against ECMO in these patients. ☐ Against the routine use of inhaled nitric oxide (Al). May improve oxygenation in severe persistent hypoxia (BII)³⁸. 						

Step 5 - Pharmacologic Treatment

The following medications should be considered for treatment based on the patient's severity and risk of disease progression. Pharmacologic recommendations for patients with COVID-19 continue to evolve.

- For the latest updates and details visit the NIH or IDSA Guidelines.
- For the latest information on local availability of therapies for COVID, check your <u>State Health Department</u>.
 For tips and tricks on how to talk with patients about COVID treatment options see the <u>SAEM Provider Toolkit</u>.

DISCHARGED FROM EMERGENCY DEPARTMENT	ADMITTED TO HOSPITAL
 All patients should be offered symptom management (Alll). Based upon the emergence of the Omicron Variant of Concern (VOC), and its subvariants, the following are the current recommendations for treatment of patients with a HIGH risk of disease progression. Preferred Therapies: Use 1 of the following (listed in order of preference). 1. Nirmatrelvir 300 mg with ritonavir 100 mg (Paxlovid) orally twice daily for 5 days, initiated as soon as possible and within 5 days of symptom onset in those aged ≥12 years and weighing ≥40 kg (Alla). See Caution note, below and in the Footnote Section before prescribing. 2. Remdesivir 200 mg IV on Day 1, followed by remdesivir 100 mg IV daily on Days 2 and 3, initiated as soon as possible and within 7 days of symptom onset in those aged ≥12 years and weighing ≥40 kg (Blla). (off label use) Alternative therapy - For use when neither of the preferred therapies are available, feasible to use, or clinically appropriate: • Molnupiravir 800 mg orally twice daily for 5 days, initiated as soon as possible and within 5 days of symptom onset in those aged ≥18 years ONLY when none of the above options can be used (Clla). Molnupiravir is not recommended for pregnant or lactating females (See Footnote Section). Women who could become pregnant should use contraception during treatment and for 4 days after the last dose. Men should use contraception during treatment and for 4 days after the last dose. See molnupiravir-us.com/patients/ Bevtelovimap no longer authorized by the FDA (See Footnote Section) Providers should have CAUTION when prescribing Paxlovid due to the ritonavir component, which has significant and complex drug-drug interactions. Please see the Footnotes section for links to more information on these. See the Footnotes page for links to the EUA FDA fact sheets for these drugs 	Hospitalized for reasons other than COVID-19, but with COVID-19: See left column "Discharged from ED" for treatment recommendations. Hospitalized but does not require supplemental 02: Do not use dexamethasone (Alla) or other corticosteroids (Alll) Hospitalized and requires supplemental 02: For patients at high risk of disease progression: Remdesivir (Blla) Hospitalized and requires supplemental 02: For pts only requiring minimal supplemental 02: For pts only requiring minimal supplemental 02: For most patients: Dexamethasone plus remdesivir (Blla) If remdesivir is not available: Dexamethasone (Blla) For patients on dexamethasone with rapidly increasing oxygen needs and systemic inflammation: Add baricitinib or tocilizumab to one of the above 3 options (Blla) Hospitalized and requires 02 through hi-flow device or noninvasive ventilation: For most patients: One of the following: Dexamethasone plus baricitinib (Al) or dexamethasone plus tocilizumab (Blla) If neither baricitinib/tofacitinib nor tocilizumab/sarilumab can be procured: Dexamethasone (Al) Optional: Add remdesivir to any 1 of the above selections (Clla) Hospitalized and requires mechanical ventilation or ECMO: Upon initiation of MV or ECMO, if not already initiated: One of the following: Dexamethasone plus baricitinib (Blla) or dexamethasone plus tocilizumab (Blla) If neither baricitinib/tofacitinib nor tocilizumab/sarilum
Steroids: Dexamethasone (or other corticosteroids) should NOT be initiated in these patients in the absence of another indication. ((Allb)	Additional details on these options can be found at the <u>NIH Inpt Treatment Page</u> Anticoagulation: Unless contraindicated, anticoagulation is recommended for admitted COVID-19 patients. The recommendations and evidence for therapeutic vs. prophylactic anticoag- ulation are rapidly evolving. The latest information can be found at the <u>NIH Anticoagulation</u> <u>Page</u>
Insufficient Evidence: At this time there is insufficient data to recommend either for or against the following medications for SARS-CoV-2 (COVID-19):	- Fluvoxamine - Herbal medications - Vitamin C
DO NOT USE - The following are recommended AGAINST for the treatment of SARS-CoV-2 (COVID-19) at the tim - Anti-interluken-6 receptor monoclonal antibodies (except tocilizumab) or anti-IL-6 monoclonal antibody (siltuximab), except in a clinical trial (BIII). - Azithromycin alone (AI) - Budesonide - Chloroquine or hydroxychloroquine with or without azithromycin (AI) - Colchicine (InPt-AI) (OutPt-BIIa) - Famotidine	e of publication of this tool/itamin D - Interferons: None in hon-hospitalized patients (Alla); in hospitalized: do not use beta (Al), alpha (Alla), or lambda (Alla) - Ivermectin - Lopinavir/ritonavir (Al) or other HIV protease inhibitors (AllI) except in a clinical trial - Metformin - Nitazoxanide (Blla) - Zinc supplementation above the recommended daily dietary allowance for the prevention of COVID-19, except in a clinical trial (BII)





Supplement - Risk Prognostication and Assessment

Providers may choose to additionally utilize a risk prognostication tool and/or assess patients risk factors for complicated illness.

The COVID PRECISE Consortium living systematic review of COVID prognostic scores identifies the highest quality prognostic models as:

- The PRIEST model to predict whether patients with COVID-19 will have an adverse outcome, such as death. This model can be used to triage patients with COVID-19 that go to the ED.
- The <u>4C Mortality Score for COVID-19</u>, the Carr model³⁶ and the Xie model³⁷, to predict whether patients hospitalized with COVID-19 will have an adverse outcome, such as death, critical care or ventilatory support.
- . These models could guide physicians to make the best possible decisions for individual patients regarding, for example, intensive care support.

Optional - Risk Prognostication

Patients with MILD and MODERATE Severity should be further assessed to determine their risk of disease progression.

The PRIEST Score is a validated tool to predict a patient's risk for end organ failure and/or mortality using readily available data on initial presentation to the ED. The ACEP working group recognizes that there are other risk prognostication calculators that have been published. The PRIEST Score is included here as it offers a

pragmatic approach with variables that don't require diagnostic testing and don't overlap with medical conditions that are within the separate risk assessment section.

Variable			1 Point			2	2 Points			3 Po	ints		4	4 Points				
Respiratory rate (per minute)		12-20)			9-11		[□ 21-2	4			<9 or >2	24				
Oxygen saturation (%) See Footnote		>95				94-95		C	□ 92-9	3			<92					
Heart rate (per minute)		51-90)			41-50 or	91-110) [] 111-	130			<41 or >	130				
Systolic BP (mmHg)		111-2	219			101-110)	C	□ 91-1	00			<91 or >	219				
Temperature (°C)		36.1-	38.0			35.1-36.	0 or	[□ >39.	0			<35.1					
						38.1-39	.0											
Alertness		Alert											Confused	t				
Inspired oxygen		Room	Air					C	⊐ Supp	lementa	al Oxygei	n						
Sex		Fema	le			□ Male												
Age (years)		16-49					C	□ 50-65			1 66-80			□ >80				
Performance status		Unrestricted		Limited strenuous		us E	Limited activity,			□ Limited self-care		e C	Bed/chair bound,					
	Normal Activity			activity, can do			can self-care					L	no self-care					
Total number of boxes			light activity									\sim						
checked in each column		_ x 0 =		x 1 =			x 2 =		x 3 =			x 4 =						
Add Subtotals	C)	-	• [+			+	. [+				
Total	Scor	e	0-1	2-3	4	5	6	7	8	9	10	11	12	13	14	15	16	17+
	Risk	%	1%	2%	3%	9%	15%	18%	22%	26%	29%	34%	38%	47%	48%	50%	55%	66%

Optional - Risk Assessment

The CDC maintains a list of underlying medical conditions associated with higher risk of severe COVID-19. If your patient has one (or especially multiple) risk factors, you may want to consider in the approach taken in subsequent steps for diagnostic testing, disposition, and treatment.

The **CDC** notes that patient race/ethnicity, socioeconomic status, and healthcare resources may effect clinical outcomes and advise consideration in clinical risk assessment.

Higher Risk (conclusive)

- □ Asthma Cancer (hematologic malignancies)
- Cerebrovascular disease
- Chronic kidney disease (receiving) dialysis)
- Chronic lung diseases
 - Bronchiectasis
 - COPD
 - □ Interstitial lung disease
 - □ Pulmonary embolism
 - □ Pulmonary hypertension
- Chronic liver disease
- Cirrhosis
- □ Non-alcoholic fatty liver disease
- □ Alcoholic liver disease
- □ Autoimmune hepatitis
- Cystic fibrosis
- Diabetes mellitus (type 1 and 2)

- Disabilities (including Down syndrome)
- Heart conditions
 - Heart failure
 - Coronary artery disease
 - □ Cardiomyopathies
- - □ Mental health conditions (mood/ schizophrenia)
 - Neurologic conditions limited to dementia
 - □ Obesity (BMI ≥30 kg/m²)
 - Physical inactivity
 - □ Pregnancy and recent pregnancy
 - □ Primary immunodeficiencies
 - Smoking (current and former) □ Solid organ and blood stem cell
 - transplantation
 - □ Tuberculosis
 - Use of corticosteroids or other
- - immunosuppresive medications

Suggestive Higher Risk

- Children with certain underlying
- conditions
- \Box Overweight (BMI \geq 25 kg/m2 but <30 kg/m2)
- Sickle cell disease
- □ Substance use disorders

Mixed Evidence

- □ Alpha 1 antitrypsin deficiency
- Bronchopulmonary dysplasia
- Hepatitits B / C
- □ Hypertension
- □ Thalassemia

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FOOTNOTES

Step 1- Severity Classification

 All severity classifications are outlined by the NIH. The <u>NIH COVID-19 Treatment Guidelines</u> <u>Panel</u> is a multi-disciplinary team of experts that meets routinely to discuss the impact of new evidence on best practices in addition to providing a standardized system for classifying clinical severity.⁶

Step 2 - Diagnostic Testing

- Exertional Sp02: post-exertional Sp02 may provide modest prognostic information of adverse outcome at 30 days^{5,13,21}
- · Optimal time interval is not established.
- $\circ\,$ Some have suggested 1-2 minutes and a sit-stand option in the patient's room (due to COVID restrictions) $^{\rm 5}$
- A 3% drop has been used in several studies ^{21, 13}
- \circ Another study used a quick walk test of 6 minutes. Decrease in $\geq 3\%$ or $\geq 5\%$ (conservative cutoff or postexercise $\leq 90\%$ suggest poor outcome (need for mechanical ventilation) with LR+=3.5 and LR==0.22 21
- Diagnostic Testing: ACEP maintains a section on <u>Laboratory Abnormalities</u> in the COVID-19 Field Guide.

Step 3 - Disposition

Discharge of select COVID patients with Home Oxygen has been shown to be associated with low rates of mortality and return admission. $^{\rm 32,33,34}$

The CDC maintains Patient Educational Materials.

SAEM Patient Toolkit has materials for patients to understand more about COVID.

Helpful links from JAMA include:

- What does this mean for families?
- https://jamanetwork.com/journals/jamapediatrics/fullarticle/2763176
- Masks
 - https://jamanetwork.com/journals/jama/fullarticle/2764955
- Stopping the spread
 - https://jamanetwork.com/journals/jama/fullarticle/2763533
- What is herd immunity?
- https://jamanetwork.com/journals/jama/fullarticle/2772168

Step 4 - Non-Pharmacologic Treatment

Home Supplemental 02

Discharge of select COVID patients with Home Oxygen has been shown to be associated with low rates of mortality and return admission³²

Studies in COVID and other viral illnesses ²⁰, have shown the benefit of:

- Rest¹⁶
- Healthy diet¹⁷
- Adequate sleep 18
- Exercise ¹⁹

Issues with Sp02 measurements

- If sending patients home with instructions for pulse oximetry, be mindful that Sp02 readings should always be considered an estimate of oxygen saturation. The FDA has just issued precautions on Sp02 devices.²⁶
- Although pulse oximetry is useful for estimating blood oxygen levels, pulse oximeters may
 not accurately detect hypoxemia under certain circumstances. Pulse oximetry results can be
 affected by skin pigmentation, thickness, or temperature. In fact, an Sp02 reading of 90% may
 represent a range of Sa02 from 86% to 94%. Clinicians should keep this limitation in mind
 when making patient decisions.²⁵

Vaccination

 Additional information on current vaccinations recommendations, can be found <u>HERE</u> and <u>Vaccination FAQs</u>

• SMART Phrases from ACEP for patients can be found HERE

Treatment of Severe and Critical patients

 Recommendations for respiratory support, IV fluids, and other interventions are maintained by the NIH <u>HERE</u>.

Step 5 - Pharmacologic Treatment

Medications - recommendations are maintained by the <u>NIH</u> and <u>IDSA</u>. Recommendations for the treatment of patients discharged home, but who have a HIGH risk for disease progression is evolving quickly due to the Omicron Variant of Concern (VOC).

- Guidance can be found on the <u>NIH Outpatient Treatment Page</u>
- Paxlovid EU A Fact Sheet: <u>www.fda.gov/media/155050/download</u>
- Molnupiravir EUA Fact Sheet: www.fda.gov/media/155054/download
- The <u>SAEM Provider Toolkit</u> offers tip and tricks on how to communicate with patients about COVID treatment options.
- Bevtelovimap no longer authorized by the FDA due to high prevalence of BQ1, BQ.1.1 and XBB variants: www.fda.gov/drugs/drug-safety-and-availability/fda-announcesbebtelovimab-not-currently-authorized-any-us-region

CAUTION with prescribing Paxlovid

- Ritonavir-boosted nirmatrelvir (Paxlovid) has significant and complex drug-drug
 interactions, primarily due to the ritonavir component of the combination. Before
 prescribing, clinicians should carefully review the patient's concomitant medications,
 including over-the-counter medications and herbal supplements, to evaluate potential
 drug-drug interactions.
 - Clinicians who are not experienced in prescribing ritonavir-boosted drugs should refer to resources such as the <u>NIH Paxlovid Drug-Drug Interactions page</u>, the <u>Ontario COVID-19</u> <u>Science Advisory Table</u>, the <u>EUA fact sheet for ritonavir-boosted nirmatrelvir (Paxlovid)</u> or the <u>Liverpool COVID-19 Drug Interactions website</u> for additional guidance.
 - Consultation with an expert (e.g., clinical pharmacist, HIV specialist, and/or the patient's specialist provider[s], if applicable) should also be considered.
- Molnupiravir is not recommended for pregnant females unless there are no other options and therapy is clearly indicated (AIII). Feeding breastmilk should be avoided during molnupiravir use and for 4 days after the last dose of the drug (AIII).
- For patients with a eGFR of 30-60 ml/min, the FDA recommends nirmatrelvir 150 mg (one 150-mg tablet) with ritonavir 100 mg (one 100-mg tablet) twice daily for 5 days

Optional - Risk Prognostication

- The <u>PRIEST Score</u>: is a validated tool to predict a patient's risk for end organ failure and/or mortality.^{14,35}
- The PRIEST Score can be accessed on MDCalc.
- See notes about pulse oximetry within Section 4 footnotes.
 - 4C Mortality Score for COVID-19 is available on MDCalc.
 - The Carr model(36) and the Xie model (37) have also been validated for risk stratification of COVID-19 patients.

Optional - Risk Assessment

The CDC maintains a <u>reference</u> for medical conditions associated with high risk for severe COVID-19.

- Race/Ethnicity and access to healthcare: the <u>CDC</u> has more information on how race, ethnicity, and access to health care resources may affect outcomes ⁷
- Economic Disparity: has been shown to be an independent variable of risk¹¹
- Pregnancy: has been shown to have increased hospitalization (OR 3.5).²
- Severe cases have been shown to have pre-term labor 45.4% compared to 6.9% of mild and recovered cases.⁹

NIH

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

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CITATIONS

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