

# GOLD 2021

[https://goldcopd.org/wp-content/uploads/2020/11/GOLD-REPORT-2021-v1.1-25Nov20\\_WMV.pdf](https://goldcopd.org/wp-content/uploads/2020/11/GOLD-REPORT-2021-v1.1-25Nov20_WMV.pdf)

▶ KEY INDICATORS FOR CONSIDERING A DIAGNOSIS OF COPD	
<p><i>Consider COPD, and perform spirometry, if any of these indicators are present in an individual over age 40. These indicators are not diagnostic themselves, but the presence of multiple key indicators increases the probability of a diagnosis of COPD. Spirometry is required to establish a diagnosis of COPD.</i></p>	
<b>Dyspnea that is:</b>	Progressive over time. Characteristically worse with exercise. Persistent.
<b>Chronic Cough:</b>	May be intermittent and may be unproductive. Recurrent wheeze.
<b>Chronic Sputum Production:</b>	Any pattern of chronic sputum production may indicate COPD.
<b>Recurrent Lower Respiratory Tract Infections</b>	
<b>History of Risk Factors:</b>	Host factors (such as genetic factors, congenital/developmental abnormalities etc.). Tobacco smoke (including popular local preparations). Smoke from home cooking and heating fuels. Occupational dusts, vapors, fumes, gases and other chemicals.
<b>Family History of COPD and/or Childhood Factors:</b>	For example low birthweight, childhood respiratory infections etc.

TABLE 2.1

▶ CLASSIFICATION OF AIRFLOW LIMITATION SEVERITY IN COPD (BASED ON POST-BRONCHODILATOR FEV <sub>1</sub> )		
<b>In patients with FEV<sub>1</sub>/FVC &lt; 0.70:</b>		
<b>GOLD 1:</b>	Mild	FEV <sub>1</sub> ≥ 80% predicted
<b>GOLD 2:</b>	Moderate	50% ≤ FEV <sub>1</sub> < 80% predicted
<b>GOLD 3:</b>	Severe	30% ≤ FEV <sub>1</sub> < 50% predicted
<b>GOLD 4:</b>	Very Severe	FEV <sub>1</sub> < 30% predicted

TABLE 2.4

## MODIFIED MRC DYSPNEA SCALE<sup>a</sup>

PLEASE TICK IN THE BOX THAT APPLIES TO YOU | ONE BOX ONLY | Grades 0 - 4

mMRC Grade 0.	I only get breathless with strenuous exercise.	<input type="checkbox"/>
mMRC Grade 1.	I get short of breath when hurrying on the level or walking up a slight hill.	<input type="checkbox"/>
mMRC Grade 2.	I walk slower than people of the same age on the level because of breathlessness, or I have to stop for breath when walking on my own pace on the level.	<input type="checkbox"/>
mMRC Grade 3.	I stop for breath after walking about 100 meters or after a few minutes on the level.	<input type="checkbox"/>
mMRC Grade 4.	I am too breathless to leave the house or I am breathless when dressing or undressing.	<input type="checkbox"/>

<sup>a</sup> Fletcher CM. BMJ 1960; 2: 1662.

TABLE 2.5

## CAT™ ASSESSMENT

For each item below, place a mark (x) in the box that best describes you currently.  
Be sure to only select one response for each question.

EXAMPLE: I am very happy	0 <input checked="" type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	I am very sad	SCORE
I never cough	0 <input type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	I cough all the time	
I have no phlegm (mucus) in my chest at all	0 <input type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	My chest is completely full of phlegm (mucus)	
My chest does not feel tight at all	0 <input type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	My chest feels very tight	
When I walk up a hill or one flight of stairs I am not breathless	0 <input type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	When I walk up a hill or one flight of stairs I am very breathless	
I am not limited doing any activities at home	0 <input type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	I am very limited doing activities at home	
I am confident leaving my home despite my lung condition	0 <input type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	I am not at all confident leaving my home because of my lung condition	
I sleep soundly	0 <input type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	I don't sleep soundly because of my lung condition	
I have lots of energy	0 <input type="radio"/> 1 <input type="radio"/> 2 <input type="radio"/> 3 <input type="radio"/> 4 <input type="radio"/> 5	I have no energy at all	
Reference: Jones et al. ERJ 2009; 34 (3); 648-54. FIGURE 2.3			TOTAL SCORE: <input type="text"/>

## ▶ THE REFINED ABCD ASSESSMENT TOOL

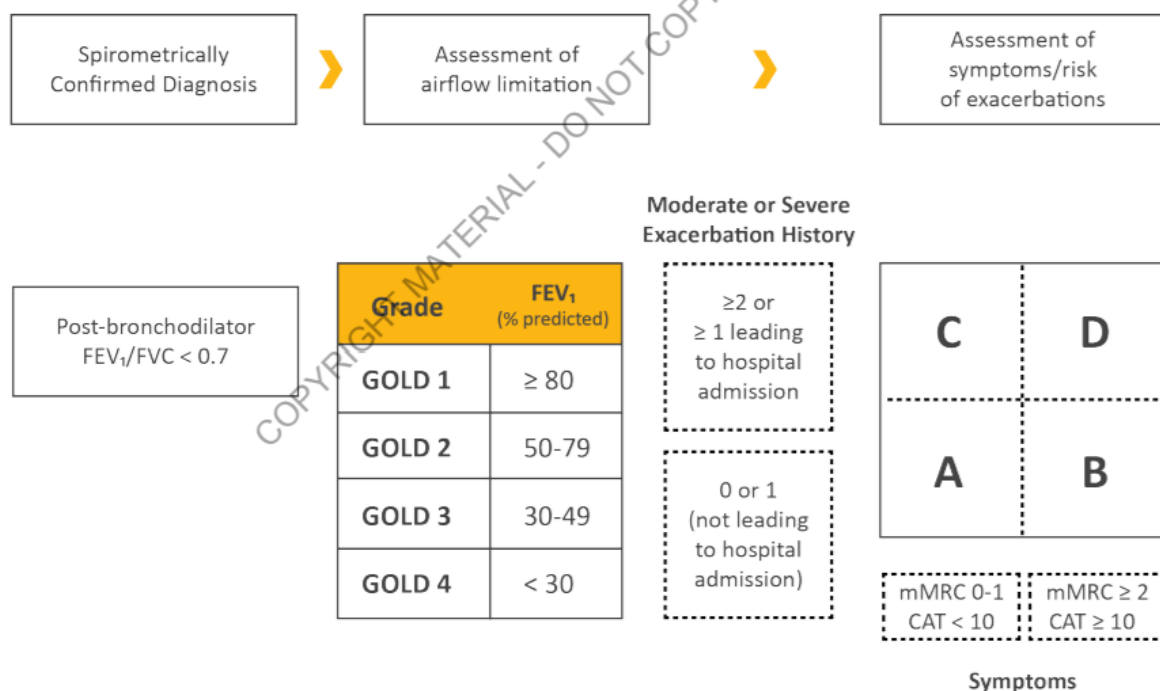


FIGURE 2.4

## ▶ VACCINATION FOR STABLE COPD

- Influenza vaccination reduces serious illness and death in COPD patients (**Evidence B**).
- The 23-valent pneumococcal polysaccharide vaccine (PPSV23) has been shown to reduce the incidence of community-acquired pneumonia in COPD patients aged ≥ 65 years with an FEV<sub>1</sub> < 40% predicted and in those with comorbidities (**Evidence B**).
- In the general population of adults ≥ 65 years the 13-valent conjugated pneumococcal vaccine (PCV13) has demonstrated significant efficacy in reducing bacteremia & serious invasive pneumococcal disease (**Evidence B**).
- The CDC recommends the Tdap (dTdap/dTpa) vaccination for adults with COPD who were not vaccinated in adolescence to protect against pertussis (whooping cough).

TABLE 3.2

## COMMONLY USED MAINTENANCE MEDICATIONS IN COPD\*

DELIVERY OPTIONS					
Generic Drug Name	Inhaler Type	Nebulizer	Oral	Injection	Duration Of Action
<b>BETA<sub>2</sub>-AGONISTS</b>					
<b>SHORT-ACTING (SABA)</b>					
Fenoterol	MDI	✓	pill, syrup		4-6 hours
Levalbuterol	MDI	✓			6-8 hours
Salbutamol (albuterol)	MDI & DPI	✓	pill, syrup, extended release tablet	✓	4-6 hours 12 hours (ext. release)
Terbutaline	DPI		pill	✓	4-6 hours
<b>LONG-ACTING (LABA)</b>					
Arformoterol		✓			12 hours
Formoterol	DPI	✓			12 hours
Indacaterol	DPI				24 hours
Olodaterol	SMI				24 hours
Salmeterol	MDI & DPI				12 hours
<b>ANTICHOLINERGICS</b>					
<b>SHORT-ACTING (SAMA)</b>					
Ipratropium bromide	MDI	✓			6-8 hours
Oxitropium bromide	MDI				7-9 hours
<b>LONG-ACTING (LAMA)</b>					
Acclidinium bromide	DPI, MDI				12 hours
Glycopyrronium bromide	DPI		solution	✓	12-24 hours
Tiotropium	DPI, SMI, MDI				24 hours
Umeclidinium	DPI				24 hours
Glycopyrrolate		✓			12 hours
Revefenacin		✓			24 hours
<b>COMBINATION SHORT-ACTING BETA<sub>2</sub>-AGONIST PLUS ANTICHOLINERGIC IN ONE DEVICE (SABA/SAMA)</b>					
Fenoterol/ipratropium	SMI	✓			6-8 hours
Salbutamol/ipratropium	SMI, MDI				6-8 hours
<b>COMBINATION LONG-ACTING BETA<sub>2</sub>-AGONIST PLUS ANTICHOLINERGIC IN ONE DEVICE (LABA/LAMA)</b>					
Formoterol/acclidinium	DPI				12 hours
Formoterol/glycopyrronium	MDI				12 hours
Indacaterol/glycopyrronium	DPI				12-24 hours
Vilanterol/umeclidinium	DPI				24 hours
Olodaterol/tiotropium	SMI				24 hours

<b>METHYLYXANTHINES</b>					
Aminophylline			solution	✓	Variable, up to 24 hours
Theophylline (SR)			pill	✓	Variable, up to 24 hours
<b>COMBINATION OF LONG-ACTING BETA<sub>2</sub>-AGONIST PLUS CORTICOSTEROID IN ONE DEVICE (LABA/ICS)</b>					
Formoterol/beclometasone	MDI, DPI				12 hours
Formoterol/budesonide	MDI, DPI				12 hours
Formoterol/mometasone	MDI				12 hours
Salmeterol/fluticasone propionate	MDI, DPI				12 hours
Vilanterol/fluticasone furoate	DPI				24 hours
<b>TRIPLE COMBINATION IN ONE DEVICE (LABA/LAMA/ICS)</b>					
Fluticasone/umeclidinium/vilanterol	DPI				24 hours
Beclometasone/formoterol/glycopyrronium	MDI				12 hours
Budesonide/formoterol/glycopyrrolate	MDI				12 hours
<b>PHOSPHODIESTERASE-4 INHIBITORS</b>					
Roflumilast			pill		24 hours
<b>MUCOLYTIC AGENTS</b>					
Erdosteine			pill		12 hours
Carbocysteine <sup>†</sup>			pill		
N-acetylcysteine <sup>†</sup>			pill		

TABLE 3.3

\*Not all formulations are available in all countries. In some countries other formulations and dosages may be available. † Dosing regimens are under discussion. MDI = metered dose inhaler; DPI = dry powder inhaler; SMI = soft mist inhaler. Note that glycopyrrolate & glycopyrronium are the same compound.

FACTORS TO CONSIDER WHEN INITIATING ICS TREATMENT		
Factors to consider when initiating ICS treatment in combination with one or two long-acting bronchodilators (note the scenario is different when considering ICS withdrawal):		
· STRONG SUPPORT ·	· CONSIDER USE ·	· AGAINST USE ·
<ul style="list-style-type: none"> <li>History of hospitalization(s) for exacerbations of COPD<sup>#</sup></li> <li>≥ 2 moderate exacerbations of COPD per year<sup>#</sup></li> <li>Blood eosinophils &gt;300 cells/μL</li> <li>History of, or concomitant, asthma</li> </ul>	<ul style="list-style-type: none"> <li>1 moderate exacerbation of COPD per year<sup>#</sup></li> <li>Blood eosinophils 100-300 cells/μL</li> </ul>	<ul style="list-style-type: none"> <li>Repeated pneumonia events</li> <li>Blood eosinophils &lt;100 cells/μL</li> <li>History of mycobacterial infection</li> </ul>
<sup>#</sup> despite appropriate long-acting bronchodilator maintenance therapy (see Table 3.4 and Figure 4.3 for recommendations); *note that blood eosinophils should be seen as a continuum; quoted values represent approximate cut-points; eosinophil counts are likely to fluctuate.		
Reproduced with permission of the © ERS 2019: <i>European Respiratory Journal</i> 52 (6) 1801219; DOI: 10.1183/13993003.01219-2018 Published 13 December 2018		

FIGURE 3.1

GOALS FOR TREATMENT OF STABLE COPD	
<ul style="list-style-type: none"> <li>Relieve Symptoms</li> <li>Improve Exercise Tolerance</li> <li>Improve Health Status</li> </ul>	REDUCE SYMPTOMS
and	
<ul style="list-style-type: none"> <li>Prevent Disease Progression</li> <li>Prevent and Treat Exacerbations</li> <li>Reduce Mortality</li> </ul>	REDUCE RISK

TABLE 4.1

INITIAL PHARMACOLOGICAL TREATMENT		
≥ 2 moderate exacerbations or ≥ 1 leading to hospitalization	Group C LAMA	Group D LAMA or LAMA + LABA* or ICS + LABA** *Consider if highly symptomatic (e.g. CAT > 20) **Consider if eos ≥ 300
0 or 1 moderate exacerbations (not leading to hospital admission)	Group A A Bronchodilator	Group B A Long Acting Bronchodilator (LABA or LAMA)
	mMRC 0-1, CAT < 10	mMRC ≥ 2, CAT ≥ 10

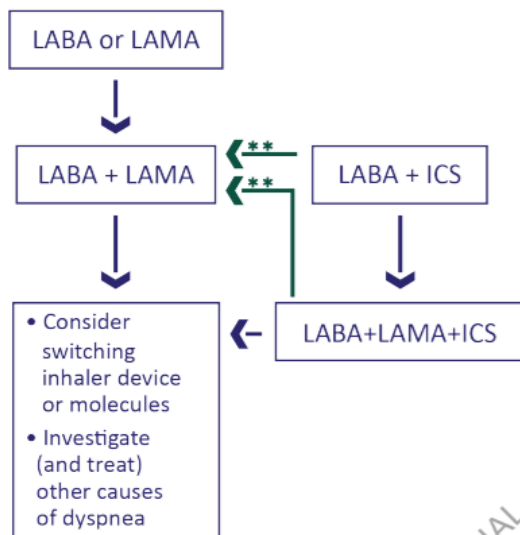
FIGURE 4.2

## FOLLOW-UP PHARMACOLOGICAL TREATMENT

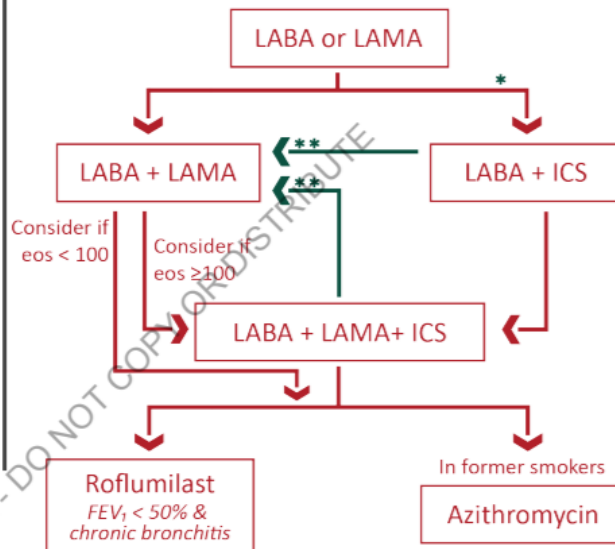
1. IF RESPONSE TO INITIAL TREATMENT IS APPROPRIATE, MAINTAIN IT.

2. IF NOT:
- ✓ Consider the predominant treatable trait to target (dyspnea or exacerbations)
  - Use exacerbation pathway if both exacerbations and dyspnea need to be targeted
  - ✓ Place patient in box corresponding to current treatment & follow indications
  - ✓ Assess response, adjust and review
  - ✓ These recommendations do not depend on the ABCD assessment at diagnosis

### • DYSPNEA •



### • EXACERBATIONS •



eos = blood eosinophil count (cells/ $\mu$ L)

\* Consider if eos  $\geq 300$  or eos  $\geq 100$  AND  $\geq 2$  moderate exacerbations / 1 hospitalization

\*\* Consider de-escalation of ICS or switch if pneumonia, inappropriate original indication or lack of response to ICS

FIGURE 4.4

▶ <b>NON-PHARMACOLOGIC MANAGEMENT OF COPD*</b>			
PATIENT GROUP	ESSENTIAL	RECOMMENDED	DEPENDING ON LOCAL GUIDELINES
<b>A</b>	Smoking Cessation (can include pharmacologic treatment)	Physical Activity	Flu Vaccination  Pneumococcal Vaccination  Pertussis Vaccination
<b>B, C and D</b>	Smoking Cessation (can include pharmacologic treatment)  Pulmonary Rehabilitation	Physical Activity	Flu Vaccination  Pneumococcal Vaccination  Pertussis Vaccination
*Can include pharmacologic treatment.			

TABLE 4.8

Exacerbations are classified as:

- ▶ Mild (treated with short acting bronchodilators only, SABDs)
- ▶ Moderate (treated with SABDs plus antibiotics and/or oral corticosteroids) or
- ▶ Severe (patient requires hospitalization or visits the emergency room). Severe exacerbations associated with acute respiratory failure.

It is now recognized that many exacerbations are not reported to healthcare professionals for therapy.

**No respiratory failure:** Respiratory rate: 20-30 breaths per minute; no use of accessory respiratory muscles; no changes in mental status; hypoxemia improved with supplemental oxygen given via Venturi mask 28-35% inspired oxygen (FiO<sub>2</sub>); no increase in PaCO<sub>2</sub>.

**Acute respiratory failure – non-life-threatening:** Respiratory rate: > 30 breaths per minute; using accessory respiratory muscles; no change in mental status; hypoxemia improved with supplemental oxygen via Venturi mask 24-35% FiO<sub>2</sub>; hypercarbia i.e., PaCO<sub>2</sub> increased compared with baseline or elevated 50-60 mmHg.

**Acute respiratory failure – life-threatening:** Respiratory rate: > 30 breaths per minute; using accessory respiratory muscles; acute changes in mental status; hypoxemia not improved with supplemental oxygen via Venturi mask or requiring FiO<sub>2</sub> > 40%; hypercarbia i.e., PaCO<sub>2</sub> increased compared with baseline or elevated > 60 mmHg or the presence of acidosis (pH ≤ 7.25).



## POTENTIAL INDICATIONS FOR HOSPITALIZATION ASSESSMENT\*

- Severe symptoms such as sudden worsening of resting dyspnea, high respiratory rate, decreased oxygen saturation, confusion, drowsiness.
- Acute respiratory failure.
- Onset of new physical signs (e.g., cyanosis, peripheral edema).
- Failure of an exacerbation to respond to initial medical management.
- Presence of serious comorbidities (e.g., heart failure, newly occurring arrhythmias, etc.).
- Insufficient home support.

\*Local resources need to be considered.

TABLE 5.2

## MANAGEMENT OF SEVERE BUT NOT LIFE-THREATENING EXACERBATIONS\*

- Assess severity of symptoms, blood gases, chest radiograph.
- Administer supplemental oxygen therapy, obtain serial arterial blood gas, venous blood gas and pulse oximetry measurements.
- Bronchodilators:
  - » Increase doses and/or frequency of short-acting bronchodilators.
  - » Combine short-acting beta 2-agonists and anticholinergics.
  - » Consider use of long-acting bronchodilators when patient becomes stable.
  - » Use spacers or air-driven nebulizers when appropriate.
- Consider oral corticosteroids.
- Consider antibiotics (oral) when signs of bacterial infection are present.
- Consider noninvasive mechanical ventilation (NIV).
- At all times:
  - » Monitor fluid balance.
  - » Consider subcutaneous heparin or low molecular weight heparin for thromboembolism prophylaxis.
  - » Identify and treat associated conditions (e.g., heart failure, arrhythmias, pulmonary embolism etc.).

\*Local resources need to be considered.

TABLE 5.3

## INDICATIONS FOR RESPIRATORY OR MEDICAL INTENSIVE CARE UNIT ADMISSION\*

- Severe dyspnea that responds inadequately to initial emergency therapy.
- Changes in mental status (confusion, lethargy, coma).
- Persistent or worsening hypoxemia ( $\text{PaO}_2 < 5.3 \text{ kPa}$  or  $40 \text{ mmHg}$ ) and/or severe/worsening respiratory acidosis ( $\text{pH} < 7.25$ ) despite supplemental oxygen and noninvasive ventilation.
- Need for invasive mechanical ventilation.
- Hemodynamic instability - need for vasopressors.

\*Local resources need to be considered.

TABLE 5.5



## ▶ INDICATIONS FOR NONINVASIVE MECHANICAL VENTILATION (NIV)

At least one of the following:

- Respiratory acidosis ( $\text{PaCO}_2 \geq 6.0 \text{ kPa}$  or  $45 \text{ mmHg}$  and arterial  $\text{pH} \leq 7.35$ ).
- Severe dyspnea with clinical signs suggestive of respiratory muscle fatigue, increased work of breathing, or both, such as use of respiratory accessory muscles, paradoxical motion of the abdomen, or retraction of the intercostal spaces.
- Persistent hypoxemia despite supplemental oxygen therapy.

TABLE 5.6

## ▶ INDICATIONS FOR INVASIVE MECHANICAL VENTILATION

- Unable to tolerate NIV or NIV failure.
- Status post - respiratory or cardiac arrest.
- Diminished consciousness, psychomotor agitation inadequately controlled by sedation.
- Massive aspiration or persistent vomiting.
- Persistent inability to remove respiratory secretions.
- Severe hemodynamic instability without response to fluids and vasoactive drugs.
- Severe ventricular or supraventricular arrhythmias.
- Life-threatening hypoxemia in patients unable to tolerate NIV.

TABLE 5.7

## ▶ INTERVENTIONS THAT REDUCE THE FREQUENCY OF COPD EXACERBATIONS

INTERVENTION CLASS	INTERVENTION
Bronchodilators	LABAs LAMAs LABA + LAMA
Corticosteroid-containing regimens	LABA + ICS LABA + LAMA + ICS
Anti-inflammatory (non-steroid)	Roflumilast
Anti-infectives	Vaccines Long Term Macrolides
Mucoregulators	N-acetylcysteine Carbocysteine Erdosteine
Various others	Smoking Cessation Rehabilitation Lung Volume Reduction Vitamin D

TABLE 5.9