MedStar Health

Diagnosis and Management of COPD
Clinical Practice Guideline
MedStar Health

"These guidelines are provided to assist physicians and other clinicians in making decisions regarding the care of their patients. They are not a substitute for individual judgment brought to each clinical situation by the patient's primary care provider-in collaboration with the patient. As with all clinical reference resources, they reflect the best understanding of the science of medicine at the time of publication but should be used with the clear understanding that continued research may result in new knowledge and recommendations".

The MedStar Health Ambulatory Best Practices Committee endorses and accepts the recommendations for care in *Global Strategy for the Diagnosis, Management and Prevention of COPD*, Global Initiative for Chronic Obstructive Lung Disease (GOLD) 2020. A complete copy of the document can be downloaded for personal use at:
https://goldcopd.org/wp-content/uploads/2019/12/GOLD-2020-FINAL-ver1.2-03Dec19_WMV.pdf

Below are the Key Points and key tables for each chapter of the GOLD guideline and are used with permission of the Global Initiative for Chronic Obstructive Lung Disease***. The reader is referred to the complete document for expanded information and the references behind the key points.***

KEY POINTS: Chapter 1: Definition and Overview

- *Chronic Obstructive Pulmonary Disease (COPD)is a common preventable and treatable disease that is characterized by persistent respiratory symptoms and airflow limitation that is due to airway and/or alveolar abnormalities usually caused by significant exposure to noxious particles or gases.*
- *The most common respiratory symptoms include dyspnea, cough and/or sputum production. These symptoms may be under-reported by patients*
- *The main risk factor for COPD is tobacco smoking but other environmental exposures such as biomass fuel exposure and air pollution may contribute. Besides exposures, host factors predispose individuals to develop COPD. These include genetic abnormalities, abnormal lung development and accelerated aging.*
  - *COPD may be punctuated by periods of acute worsening of respiratory symptoms, called exacerbations.*
  - *In most patients, COPD is associated with significant concomitant chronic diseases, which increase its morbidity and mortality.*

KEY POINTS: Chapter 2: Diagnosis and Assessment

- *COPD should be considered in any patient who has dyspnea, chronic cough or sputum production, a history of recurrent lower respiratory tract infections and/or a history of exposure to risk factors for the disease.*
- *Spirometry is required to make the diagnosis; the presence of a post-bronchodilator $FEV_1/FVC < 0.70$ confirms the presence of persistent airflow limitation.*
- *The goals of COPD assessment are to determine the levelof airflow limitation, the impact of disease on the patient's health status, and the risk of future events (such as exacerbations, hospital admissions, or death), in order to guide therapy.*

- *Concomitant chronic diseases occur frequently in COPD patients, including cardiovascular disease, skeletal muscle dysfunction, metabolic syndrome, osteoporosis, depression, anxiety, and lung cancer. These comorbidities should be actively sought,and treated appropriately when present as they can influence mortality and hospitalizations independently.*

DIFFERENTIAL DIAGNOSIS

A major differential diagnosis is asthma. Sometimes a clear distinction from COPD is not possible, in which case management is similar to that of asthma.

The WHO recommends that all patients with a diagnosis of COPD should be screened once for alpha-1 antitrypsin deficiency (AATD), especially in areas with high prevalence.

ASSESSMENT

COPD assessment must consider the following aspects of the disease:
- The presence and severity of the spirometric abnormality
- Current nature and magnitude of symptoms
- History of moderate and severe exacerbations and future risk
- Presence of comorbidities

Table 2.4. Classification of airflow limitation severity in COPD (Based on post-bronchodilator FEV_1)		
In patients with $FEV_1/FVC < 0.70$:		
GOLD 1:	Mild	$FEV_1 \geq 80\%$ predicted
GOLD 2:	Moderate	$50\% \leq FEV_1 < 80\%$ predicted
GOLD 3:	Severe	$30\% \leq FEV_1 < 50\%$ predicted
GOLD 4:	Very Severe	$FEV_1 < 30\%$ predicted

Spirometry should be performed post bronchodilator.

Assessment of symptoms can be measured using the MRC dyspnea scale (below)

Table 2.5. Modified MRC dyspnea scale[a] 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.	❑
mMRC Grade 1. I get short of breath when hurrying on the level or walking up a slight hill.	❑
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.	❑
mMRC Grade 3. I stop for breath after walking about 100 meters or after a few minutes on the level.	❑
mMRC Grade 4. I am too breathless to leave the house or I am breathless when dressing or undressing.	❑

[a] Fletcher CM. BMJ 1960; 2: 1662.

Because patients may experience symptoms beyond dyspnea, a comprehensive assessment of symptoms using a tool such as the CAT (below) is preferred.

Your name:

Today's date:

CAT
COPD Assessment Test

How is your COPD? Take the COPD Assessment Test™ (CAT)

This questionnaire will help you and your healthcare professional measure the impact COPD (Chronic Obstructive Pulmonary Disease) is having on your wellbeing and daily life. Your answers, and test score, can be used by you and your healthcare professional to help improve the management of your COPD and get the greatest benefit from treatment.

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.

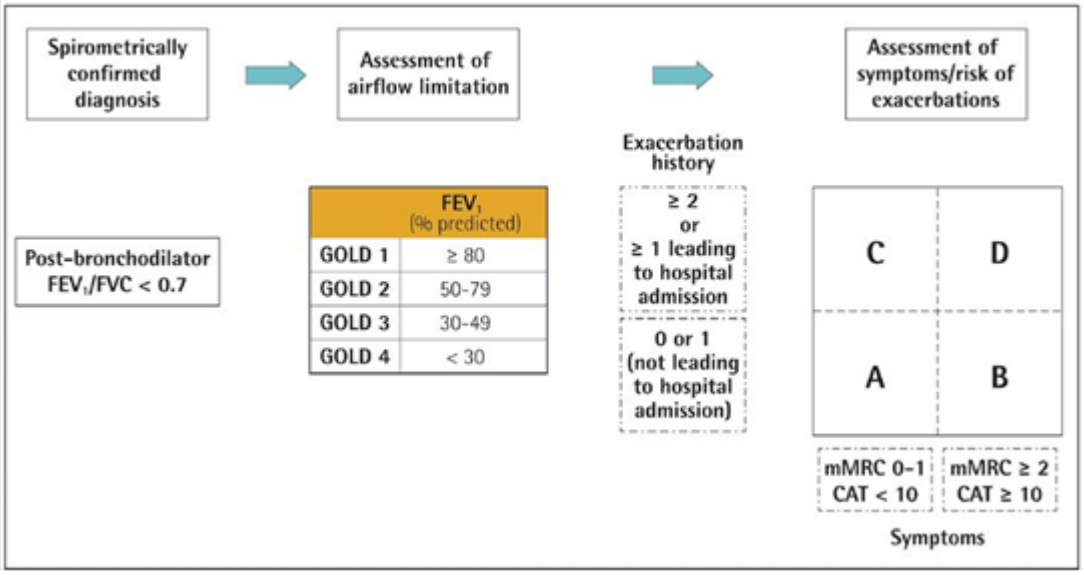
Example: I am very happy 0 1 X 2 3 4 5 I am very sad

			SCORE
I never cough	0 1 2 3 4 5	**I cough all the time**	
I have no phlegm (mucus) in my chest at all	0 1 2 3 4 5	**My chest is completely full of phlegm (mucus)**	
My chest does not feel tight at all	0 1 2 3 4 5	**My chest feels very tight**	
When I walk up a hill or one flight of stairs I am not breathless	0 1 2 3 4 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 1 2 3 4 5	**I am very limited doing activities at home**	
I am confident leaving my home despite my lung condition	0 1 2 3 4 5	**I am not at all confident leaving my home because of my lung condition**	
I sleep soundly	0 1 2 3 4 5	**I don't sleep soundly because of my lung condition**	
I have lots of energy	0 1 2 3 4 5	**I have no energy at all**	
		TOTAL SCORE	

RES/QST/09/43163/1 Date of preparation: September 2009.

Combining spirometry, symptomatic assessment and risk of exacerbation in the refined ABCD assessment tool may facilitate consideration of individual therapies for a specific patient.

Figure 2.4. The refined ABCD assessment tool



KEY POINTS: Chapter 3: Evidence Supporting Prevention and Maintenance Therapy

- *Smoking cessation is key. Pharmacotherapy and nicotine replacement reliably increase long-term smoking abstinence rates. Legislative smoking bans and counselling, delivered by healthcare professionals, improve quit rates.*
- *The effectiveness and safety of e-cigarettes as a smoking cessation aid is uncertain at present.*
- *Pharmacologic therapy can reduce COPD symptoms, reduce the frequency and severity of exacerbations, and improve health status and exercise tolerance.*
- *Each pharmacologic treatment regimen should be individualized and guided by the severity of symptoms, risk of exacerbations, side-effects, co-morbidities, drug availability and cost, and the patient's response, preference and ability to use various drug delivery devices.*
- *Inhaler technique needs to be assessed regularly.*
- *Influenza vaccination decreases the incidence of lower respiratory tract infections.*
- *Pneumococcal vaccination decreases lower respiratory tract infections.*
- *Pulmonary rehabilitation improves symptoms, quality of life, and physical and emotional participation in everyday activities.*
- *In patients with severe resting chronic hypoxemia, long-term oxygen therapy improves survival.*
- *In patients with stable COPD and resting or exercise-induced moderate desaturation, long-term oxygen treatment should not be prescribed routinely. However, individual patient factors must be considered when evaluating the patient's need for supplemental oxygen.*
- *In patients with severe chronic hypercapnia and a history of hospitalization for acute respiratory failure, long-term non-invasive ventilation may decrease mortality and prevent re-hospitalization.*
- *In select patients with advanced emphysema refractory to optimized medical care, surgical or bronchoscopic interventional treatments may be beneficial.*
- *Palliative approaches are effective in controlling symptoms in advanced COPD.*

Table 3.4. Bronchodilators in Stable COPD
• Inhaled bronchodilators in COPD are central to symptom management and commonly given on a regular basis to prevent or reduce symptoms
• Regular and as-needed use of SABA or SAMA improves FEV1 and symptoms
• Combinations of SABA and SAMA are superior compared to either medication alone in improving FEV1 and symptoms
• LABAs and LAMAs significantly improve lung function, dyspnea, health status, and reduce exacerbation rates.
• LAMAs have a greater effect on exacerbation reduction compared with LABAs and decrease hospitalizations.
• Combination treatment with a LABA and LAMA increases FEV1 and reduces symptoms compared to monotherapy
• Combination treatment with a LABA/ LAMA reduces exacerbations compared to monotherapy
• Tiotropium improves the effectiveness of pulmonary rehabilitation in increasing exercise performance
• Theophylline exerts a small bronchodilator effect in stable COPD and that is associated with modest symptomatic benefits.

Table 3.5 Anti-inflammatory therapy in stable COPD
Inhaled Corticosteroids
• An ICS combined with a LABA is more effective than the individual components in improving lung function and health status and reducing exacerbations in patients with exacerbations and moderate to very severe COPD
• Regular treatment with ICS increases the risk of pneumonia especially in those with severe disease
• Triple inhaled therapy of ICS/LAMA/LABA improves lung function, symptoms and health status and reduces exacerbations compared to ICS/LABA, LABA/LAMA or LAMA monotherapy
Oral glucocorticoids
• Long-term use of oral glucocorticoids has numerous side effects with no evidence of benefits
PDE4 inhibitors
• In patients with chronic bronchitis, severe to very severe COPD and a history of exacerbations: A PDE 4 inhibitor improves lung function and reduces moderate and severe exacerbations A PDE 4 inhibitor improves lung function and decreases exacerbations in patients who are on fixed-dose LABA/ICS combinations
Antibiotics
• Long-term azithromycin and erythromycin therapy reduce exacerbations over one year
• Treatment with azithromycin is associated with an increased incidence of bacterial resistance and hearing test impairments
Mucolytics/antioxidants
• Regular treatment with mucolytics such as erdosteine, carbocysteine and NAC reduces the risk of exacerbations in select populations
Other anti-inflammatory agents
• Simvastatin does not prevent exacerbations in COPD patients at increased risk of exacerbations and without indications for statin therapy. However, observational studies suggest that statins may have positive effects on some outcomes in patients with COPD who receive them for cardiovascular and metabolic indications
• Leukotriene modifiers have not been tested adequately in COPD patients

KEY POINTS: Chapter 4: Management of Stable COPD

- *The management strategy for stable COPD should be predominantly based on the individualized assessment of symptoms and future risk of exacerbations.*
- *All individuals who smoke should be strongly encouraged and supported to quit.*
- *The main treatment goals are reduction of symptoms and future risk of exacerbations.*
- *Management strategies include pharmacologic and non-pharmacologic interventions.*

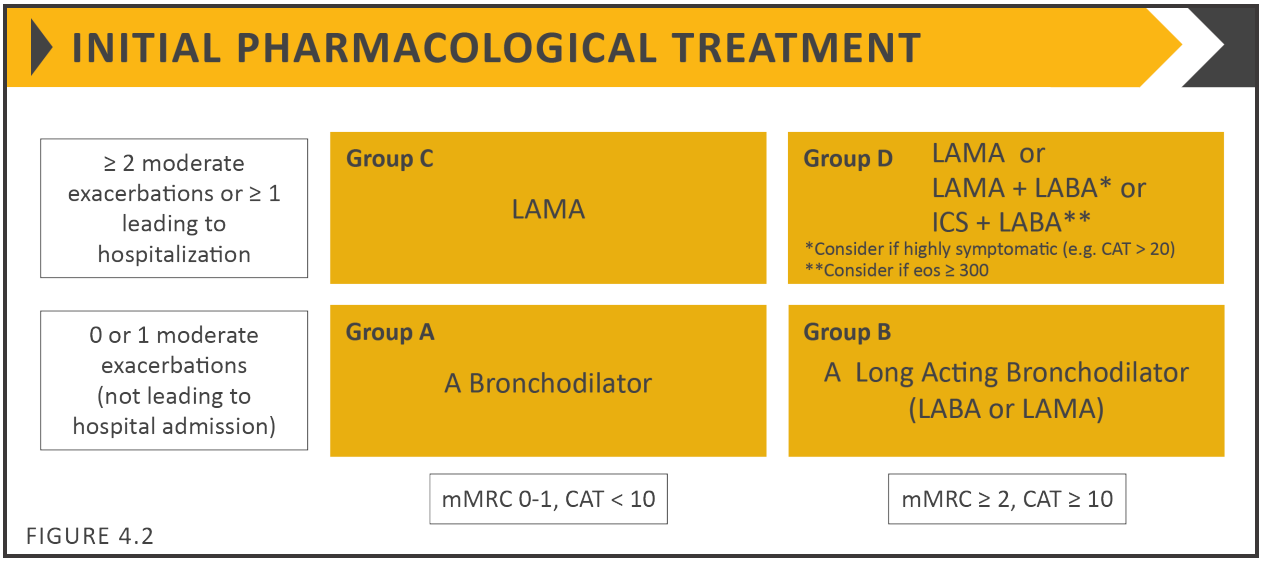
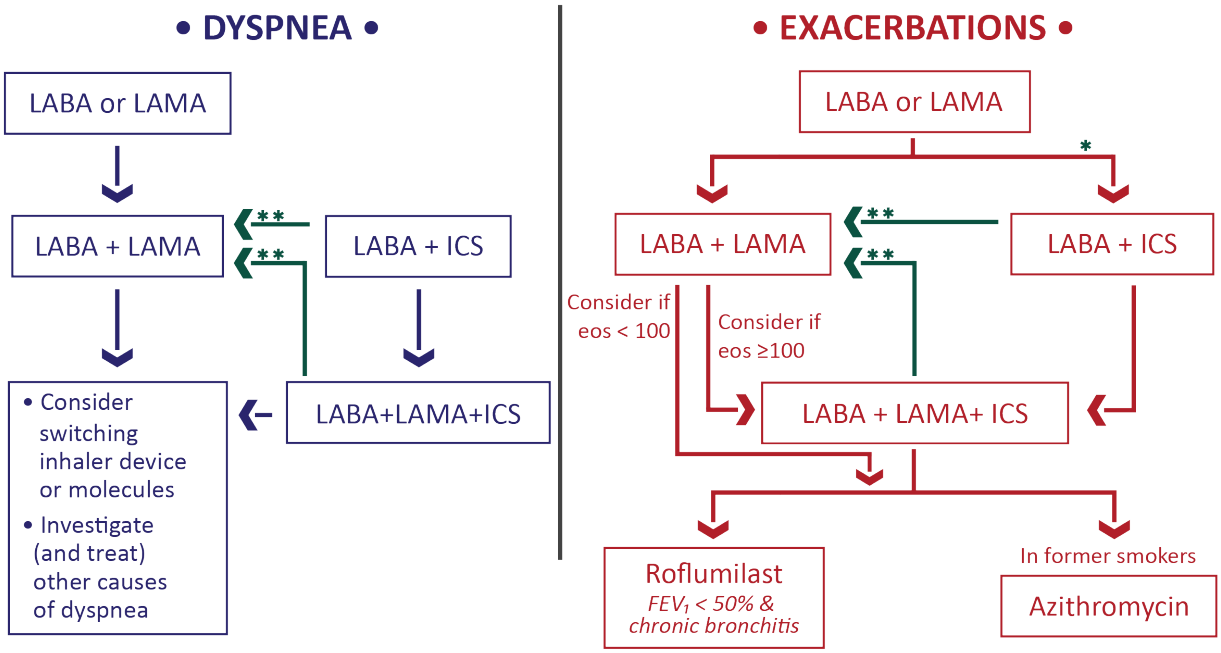


FIGURE 4.2

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



eos = blood eosinophil count (cells/µL)
**** Consider if eos ≥ 300 or eos ≥ 100 AND ≥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

NON-PHARMACOLOGIC MANAGEMENT OF COPD*

PATIENT GROUP	ESSENTIAL	RECOMMENDED	DEPENDING ON LOCAL GUIDELINES
A	Smoking Cessation (can include pharmacologic treatment)	Physical Activity	Flu Vaccination Pneumococcal Vaccination
B, C and D	Smoking Cessation (can include pharmacologic treatment) Pulmonary Rehabilitation	Physical Activity	Flu Vaccination Pneumococcal Vaccination

*Can include pharmacologic treatment.

TABLE 4.8

KEY POINTS: Chapter 5: Management of Exacerbations

- *An exacerbation of COPD is defined as an acute worsening of respiratory symptoms that results in additional therapy.*
- *As the symptoms are not specific to COPD relevant differential diagnoses should be considered.*
- *Exacerbations of COPD can be precipitated by several factors. The most common causes are respiratory tract infections.*
- *The goal for treatment of COPD exacerbations is to minimize the negative impact of the current exacerbation and to prevent subsequent events.*
- *Short-acting inhaled beta$_2$-agonists with or without short-acting anticholinergics are recommended as the initial bronchodilators to treat an acute exacerbation.*
- *Maintenance therapy with long-acting bronchodilators should be initiated as soon as possible before hospital discharge.*
- *Systemic corticosteroids can improve lung function (FEV_1), oxygenation and shorten recovery time and hospitalization duration. Duration of therapy should not be more than 5-7 days.*
- *Antibiotics, when indicated, can shorten recovery time, reduce the risk of early relapse, treatment failure, and hospitalization duration. Duration of therapy should be 5-7 days.*
- *Methylxanthines are not recommended due to increased side effect profiles.*
- *Non-invasive mechanical ventilation should be the first mode of ventilation used in COPD patients with acute respiratory failure who have no absolute contraindication because it improves gas exchange, reduces work of breathing and the need for intubation, decreases hospitalization duration and improves survival.*
- *Following an exacerbation, appropriate measures for exacerbation prevention should be initiated.*

Table 5.1 Potential Indications for hospitalization assessment*
• Severe symptoms such as sudden worsening of resting dyspnea, high respiratory rate, decreased oxygen
• 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
• Bronchodilators:
○ Increase doses and/or frequency of short-acting bronchodilators
○ Combine short-acting beta 2-agonists and anticholinergics
○ Consider use of long-active 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.

KEY POINTS: Chapter 6: COPD and Co-morbidities

- *COPD often coexists with other diseases (co-morbidities) that may have a significant impact on disease course.*
- *In general, the presence of co-morbidities should not alter COPD treatment and co-morbidities should be treated per usual standards regardless of the presence of COPD.*
- *Lung cancer is frequently seen in patients with COPD and is a main cause of death.*
- *Cardiovascular diseases are common and important co-morbidities in COPD.*
- *Osteoporosis and depression/anxiety are frequent, important co-morbidities in COPD, are often under-diagnosed, and are associated with poor health status and prognosis.*
- *Gastroesophageal reflux (GERD) is associated with an increased risk of exacerbations and poorer health status.*
- *When COPD is part of a multimorbidity care plan, attention should be directed to ensure simplicity of treatment and to minimize polypharmacy.*

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Appendix: Medications for COPD

Inhalers for COPD		
Medication[c]	**Dosing Frequency[a]**	**Cost[b]**
Short-acting Bronchodilators		
Beta-2 agonists		
Albuterol (Salbutamol) (*ProAir HFA*, *ProAir Digihaler, ProAir RespiClick, Proventil HFA*, *Ventolin HFA)*	2 inhalations every 4-6 hours as needed	*$78*
Levalbuterol (*Xopenex HFA)*	Three to four times daily as needed	$74
Anticholinergic		
Ipratropium (*Atrovent HFA*)	2 inhalations four times daily	*Atrovent HFA* $494
Combination Beta-2 agonist/ Anticholinergic		
Albuterol/ipratropium (*Combivent Respimat*)	1 inhalation four times daily	*Combivent Respimat* $512
Long-Acting Beta-2 agonists (LABAs)		
Indacaterol (*Arcapta Neohaler)*	1 capsule inhaled once daily	*Acrapta Neohaler* $310
Olodaterol (*Striverdi Respimat*)	2 inhalations once daily	*Striverdi Respimat* $269
Salmeterol (*Serevent Diskus*)	One inhalation twice daily	*Serevent Diskus* $600
Long-Acting Antimuscarinc agents (LAMAs)		
Aclidinium (*Tudorza Pressair)*	1 inhalation twice daily	*Tudorza Pressair* $343
Glycopyrrolate (*Seebri Neohaler*)	1 capsule inhaled twice daily	$473
Tiotropium (*Spiriva HandiHaler, Spiriva Respimat*)	HandiHaler: 1 capsule inhaled once daily Respimat: two inhalations once daily	*Spiriva HandiHaler* $540 *Respimat* $223
Umeclidinium (*Incruse Ellipta*)	1 inhalation once daily	$374
Combination LABA/LAMA		
Indacaterol/Glycopyrrolate (*Utibron Neohaler*)	1 capsule inhaled twice daily	$440
Olodaterol/Tiotropium (*Stiolto Respimat*)	2 inhalations once daily	$206
Vilanterol/Umeclidinium	1 inhalation once daily	$261

(*Anoro Ellipta*)		
Formoterol/Aclidinium *(Duaklir Pressair)*	1 inhalation twice daily	$1194
Formoterol/Glycopyrrolate *(Bevespi Aerosphere)*	2 inhalations twice daily	$242-438
Combination LABA/ Corticosteroid		
Formoterol/Budesonide (*Symbicort)*	2 inhalations twice daily	$352-403
Salmeterol/Fluticasone propionate (*Advair Diskus*, *Advair HFA, AirDuo RespiClick, Wixela Inhub)*	1 inhalation twice daily	$120
Vilanterol/Fluticasone furoate (*Breo Ellipta*)	1 inhalation once daily	$179
Combination LABA/Corticosteroid/Anticholinergic		
Fluticasone/Umeclidium/Vilanterol (*Trelegy Ellipta)*	1 inhalation once daily	$344
Other Agents		
Methylxanthines		
Theophylline	Variable depending on formulation Toxicity is dose related; drug levels should be monitored	$258 dollars (generic twice daily product)
Systemic Corticosteroids		
Prednisone	Once daily by mouth (usually 40mg once daily for 5 days)	($0.50 per day – therapy duration varies)
Methylprednisolone	60-125 mg 1-4 times daily IV followed by oral therapy (prednisone preferred for oral route)	($4 -$8 per day – therapy duration varies)
Phosphodiesterase-4 Inhibitor		
Roflumilast (*Daliresp*)	250 mcg once daily by mouth for 4 weeks followed by 500mcg once daily	$14

a. May differ from product labeling.
b. Wholesale cost for 30-day supply of highest strength, of generic if available unless otherwise specified. For short-acting agents, cost is for 200 inhalations.
c. Only long-acting inhalers specifically approved for COPD are included.

Inhalers for COPD adapted from Detail Document #310107. Pharmacist's Letter / Prescriber's Letter, January 2016.

Ease of use of Some Bronchodilator Inhalers				
Inhaler Type	Assembly	Indicator showing	Breath-Hand	Dependence on

		remaining doses	Coordination Needed	Strength of breath intake
Aerosphere Inhaler	Easy	Yes	Yes	No
Ellipta Inhalers	None	Yes		Yes
Respimat Inhalers	Difficult for some	Yes		No
Neohaler Inhalers	Difficult for some to remove capsules from packaging			Yes
Pressair Inhaler	None			Yes
Handihaler Inhaler	Inserting capsules into device may be difficult			Yes
Diskus Inhalers	None	Yes		Yes

(Adapted from The Medical Letter Vol 59, Issue 1518. April 10, 2017

Patient Education:

https://www.cdc.gov/copd/infographics/copd-awareness.html

https://www.cdc.gov/copd/basics-about.html

https://www.uptodate.com/contents/chronic-obstructive-pulmonary-disease-copd-the-basics?search=COPD&source=search_result&selectedTitle=1~150&usage_type=default&display_rank=1

https://www.uptodate.com/contents/medicines-for-chronic-obstructive-pulmonary-disease-copd-the-basics?search=COPD&topicRef=4649&source=see_link

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1. From the *Global Strategy for the Diagnosis, Management and Prevention of COPD*, Global Initiative for Chronic Obstructive Lung Disease (GOLD) 2020. Available from: http://www.goldcopd.org/.
2. Drugs for COPD. Med Lett Drugs Ther. 2017 Apr 10;59(1518):57-62