

Chronic Obstructive Pulmonary Disease (COPD) A VA Clinician's Guide



Chronic Obstructive Pulmonary Disease (COPD)

A VA Clinician's Guide



VA Pharmacy Benefits Management Academic Detailing Service Real Provider Resources Real Patient Results

Your Partner in Enhancing Veteran Health Outcomes

VA PBM Academic Detailing Service Email Group PharmacyAcademicDetailingProgram@va.gov

VA PBM Academic Detailing Service SharePoint Site https://vaww.portal2.va.gov/sites/ad

VA PBM Academic Detailing Service Public Website http://www.pbm.va.gov/PBM/academicdetailingservicehome.asp

Table of Contents

Key Messages	2
Key Facts	3
Initial Assessment and Diagnosis	4
Developing a Treatment Plan for COPD	5
Acute Exacerbations	18
Summary of Treatment Approaches	23
Appendix A	24
Appendix B	25
References	26

Key Messages

Evaluate all patients with symptoms and/or a history of exposure to risk factors for COPD and use spirometry to make the diagnosis.	4
Encourage all patients to quit smoking regardless of COPD severity.	6
Give all Veterans with COPD the influenza vaccine annually and pneumococcal vaccines as indicated by age.	7
Use a LAMA or LABA as initial therapy for most patients in Group A and Group B. Short-acting agents, SAMA or SABA, should only be used for immediate relief of symptoms.	10
Use LAMA as first line therapy for patients in Group C and D. Progress to combination therapy, usually LAMA + LABA, if monotherapy is not effective.	11
Treat an exacerbation with short-acting beta-agonist(s), continue long-acting bronchodilator(s), and give short courses of oral corticosteroids (5-days) and antibiotics (5-7 days) when indicated.	21

Key Facts

Almost 16 million Americans have been diagnosed with chronic obstructive pulmonary disease (COPD). The actual prevalence is likely much higher since many people with low pulmonary function are not aware of their condition and remain undiagnosed.^{1,2,3}

COPD was the 4th leading cause of death in the United States in 2017 and the 2nd most common cause of admission for an ambulatory care sensitive condition (ACSC) from 2018–2019.^{4,5}

Figure 1. One Person Dies Every Four Minutes from Complications Related to COPD⁶

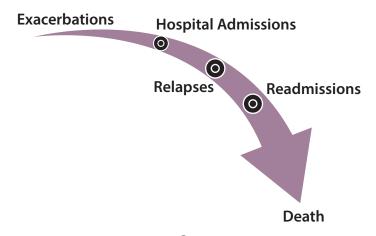


The Importance of COPD Exacerbations

An acute COPD exacerbation is any worsening of a patient's dyspnea, cough, or sputum from their baseline symptom variability.⁷ Acute exacerbations are significant since the risk of death increases as the number and severity of exacerbations increase.^{7,8}

Exacerbations are acute events that can change the trajectory of the disease in a person with COPD^{7,8}

Figure 2. Trajectory of COPD After Experiencing Exacerbations^{7,8}



Initial Assessment and Diagnosis

COPD should be considered in any patient with symptoms (e.g., chronic cough, dyspnea) or a history of exposure to risk factors. Airway obstruction is the hallmark of COPD, which is typically confirmed using spirometry in patients with risk factors and symptoms consistent with COPD.

Figure 3. Risk Factors for and Symptoms of COPD⁹⁻¹²

• Smoking tobacco
• Occupational dust or fumes
• Environmental (e.g., smoke from fires)
• Genetic: alpha-1 antitrypsin deficiency (AATD)*

OR

• Chronic cough
• Shortness of breath/dyspnea
• Frequent respiratory infections
• Fatigue
• Sputum production
• Wheezing/chest tightness

Spirometry Testing**

COPD is confirmed by a $\frac{\text{FEV}}{\text{FVC}}$ < 0.70

OR

If the Veteran's results are below the Lower Limit of Normal (LLN) they have COPD.

*Testing (serum and/or genetic) is required to diagnose AATD. Consider testing in patients < age 45 at onset of symptoms who have minimal exposure to tobacco or other environmental risk factors. **Tested after administration of bronchodilator. If FEV_1/FVC is between 0.6 to 0.8 range, repeat spirometry to confirm diagnosis. FEV_1 = forced expiratory volume in one second; FVC = forced vital capacity.



Chronic inhalation of anything combustible can cause COPD. This includes cannabis and tobacco in any inhaled forms or respiratory routes of administration (e.g., pipe, cigarette, water pipe/hookah, and vaping). Eliminating exposure to tobacco smoke and reducing exposure to air pollution can prevent COPD in most individuals. 79

Evaluate all patients with symptoms and/or a history of exposure to risk factors for COPD and use spirometry to make the diagnosis.

Developing a Treatment Plan for COPD

Simply prescribing an inhaler for a patient with COPD is not a treatment plan. Treatment decisions for COPD patients should be individualized, multi-modal, and guided by symptoms, symptom severity, comorbidities, and the frequency of acute exacerbations. (Note: a patient's ability to use an inhaler device and their preference should be incorporated into the treatment plan.)

Figure 4. Elements of a COPD Treatment Plan⁹

Smoking Cessation	Quitting has the greatest impact on slowing COPD progression.
Vaccines	Influenza vaccine helps reduce COPD exacerbations and hospitalizations. Pneumococcal vaccine PPSV23 reduces the rates of community-acquired pneumonia in patients with COPD.
Pharmacotherapy for COPD	Pharmacotherapy reduces symptoms, frequency and severity of exacerbations, and improves exercise tolerance and health status.
Non-pharmacologic Therapies	Proper nutrition, exercise, and use of pulmonary rehabilitation helps improve quality of life and reduce exacerbations.
Treating Other Comorbidities	The most common cause of death in Veterans with COPD is cardiovascular disease. Addressing this, along with other common comorbidities, like depression, lung cancer, obesity, and osteoporosis, is vital to the overall health of patients with COPD.

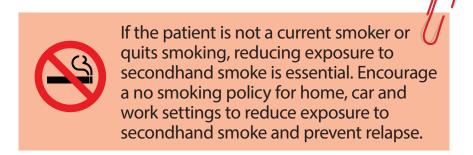
Smoking Cessation

Ask every patient with COPD if they smoke tobacco. If they do, smoking cessation should be addressed. Smoking cessation improves survival in patients with COPD, including older patients. Veterans who say they are ready to quit in the next 30 days should receive pharmacotherapy support (e.g., nicotine replacement, bupropion, or varenicline). 16

Figure 5. Pathway for Addressing Tobacco Use – The 5 A's^{16–18}



*Behavioral supports with evidence for benefit include individual sessions, group sessions, or provider support via telephone or Quitline (the VA National Quitline is: 1-855-QUIT-VET (1-855-784-8838). See the Academic Detailing Service (ADS) **Tobacco Use Disorder Provider Guide** and ADS **Tobacco Use Disorder Quick Reference Guide** for more detailed information on pharmacotherapy for tobacco cessation (www.pbm.va.gov).

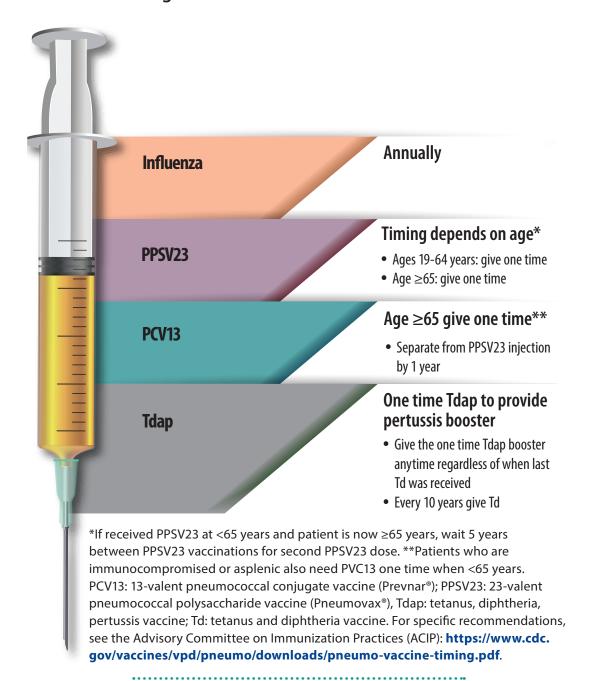


Encourage all patients to quit smoking regardless of COPD severity.

Vaccines

Annual influenza vaccination can significantly reduce the incidence of lower respiratory infections and death in people with COPD.^{9,19,20} The pneumococcal vaccine PPSV23 (23-valent) reduces the incidence of community-acquired pneumonia in COPD patients under 65 with an FEV <40% and in those with comorbidities.^{21,22} Pneumococcal vaccines (13- or 23-valent), are recommended for all patients 65 years and older.

Figure 6. Vaccine Timing for Adults with COPD^{9,21,22}



Give all Veterans with COPD the influenza vaccine annually and pneumococcal vaccines as indicated by age.

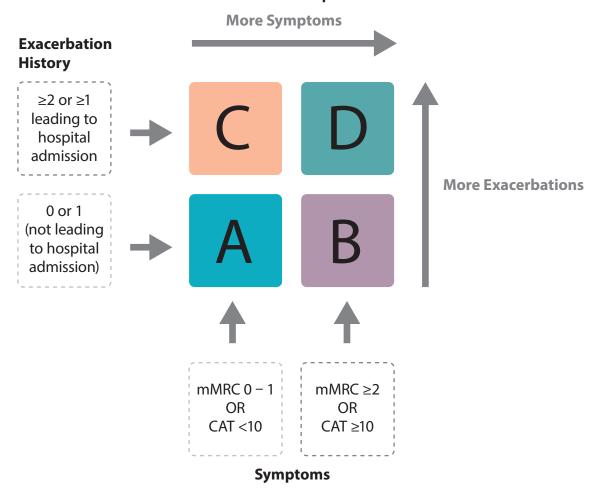
Pharmacotherapy for Stable (Non-exacerbating) COPD

Pharmacotherapy can reduce symptoms, decrease the risk and severity of exacerbations, and improve health status and exercise tolerance. Review the Veteran's exacerbation history and assess his or her symptoms to guide pharmacotherapy initiation and the need for medication changes.

Determining Treatment Group (GOLD A, B, C or D)^{9,23–26}

The most widely used therapeutic strategy for treating COPD is from the Global Initiative for Chronic Obstructive Lung Disease (GOLD).9

Figure 7. ABCD Assessment Tool – GOLD Groups^{25,26}



For this diagram, think of the Exacerbation History as the Y-axis and Symptoms as the X-axis.

- Groups A and C have a lower symptom burden (lower mMRC and CAT scores).
- Groups B and D have a greater symptom burden.
- Groups A and B have infrequent exacerbations.
- Groups C and D have frequent exacerbations.

To categorize a patient into a group (A–D), include both exacerbation history and symptoms. For example, if a patient has had three exacerbations in the past year and a CAT score of nine, they would be in group C.

CAT = COPD Assessment Test; mMRC = Modified Medical Research Council Breathlessness Scale; See **Appendix** for the full questionnaires. Exacerbation history is the number of exacerbations in the past year.

Table 1. Initial Treatment by GOLD Group⁹

Exacerbation History			
0 or 1 exacerbations not leading to hospital admission ≥2 exacerbations or ≥1 leading to hospital admission			eading to hospital
	Assessment of Sympton	ns/Risk of Exacerbations	
Milder Symptoms	Worsening Symptoms	Milder Symptoms	Worsening Symptoms
mMRC 0–1 or CAT <10	mMRC ≥2 or CAT ≥10	mMRC 0-1 or CAT <10	mMRC ≥2 or CAT ≥10
Group A	Group B Group C		Group D
Bronchodilator (long-acting or short-acting)	LAMA or LABA LAMA		LAMA or LAMA + LABA*
Persistent symptoms – use a LAMA or LABA Occasional dyspnea – use a SAMA or SABA	If persistent symptoms on long-acting monotherapy then use LAMA + LABA symptoms on maximal inhaler therapy, consultation with a pulmonologist is recommended.		
Short-acting agents (SAMA or SABA) should be considered for patients on long-acting			

bronchodilators who need immediate relief.

*Consider starting with LAMA + LABA if patient is highly symptomatic (e.g., CAT >20). Consider starting with LABA + ICS if patient has a history of asthma or CAT score >20 and eosinophil count (eos) \geq 300 cells/ μ L or eos \geq 100 cells/ μ L and \geq 2 moderate exacerbations or >1 hospitalization. ICS = inhaled corticosteroid; LABA = long-acting beta₂ agonist; LAMA = long-acting muscarinic antagonist; SABA = short-acting beta₂ agonist; SAMA = short-acting muscarinic antagonist.



Long-acting bronchodilators are preferred over short-acting agents for initial treatment in Group A patients with persistent symptoms because they improve lung function, dyspnea, and health status as well as reduce exacerbations while short-acting agents only improve dyspnea and temporarily improve lung function.^{9,27–30}

- First line: LAMA or LABA for persistent symptoms.
- SAMA or SABA can be considered in patients with only occasional dyspnea.
- SAMA or SABA can be added to long-acting bronchodilators in patients who need immediate relief, prescribed on an as-needed schedule.9



■ First line: LAMA or LABA

 In Group B patients, there is no evidence to recommend one class of long-acting bronchodilator over another for initial treatment.⁹ Use a LAMA or LABA as initial therapy for most patients in Group A and Group B. Short-acting agents, SAMA or SABA, should only be used for immediate relief of symptoms.

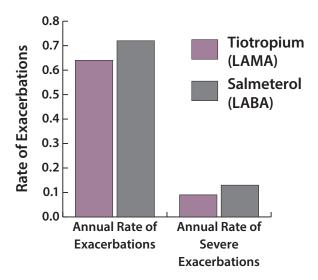


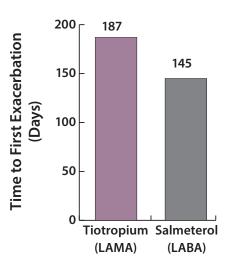
■ First line: LAMA^{30,31,32}

■ LAMA + LABA can be used in patients whose symptoms are not controlled on monotherapy. Inhaled corticosteroids (ICS) should be reserved for patients with asthma or if symptoms are not controlled on LAMA + LABA.

Figure 8. Reduced Annual Rate of Exacerbations and Increased Time to First Exacerbation with Tiotropium (LAMA) Monotherapy Compared to Salmeterol (LABA)³¹

Tiotropium Reduced Annual Rate of Exacerbations by 11% and Severe Exacerbations by 27% Tiotropium Increased the Time to First Exacerbation by 42 Days





POET-COPD trial comparing tiotropium to salmeterol in patients with moderate-to-very-severe COPD and a history of exacerbations in the previous year (p < 0.05 for all comparisons in figures).

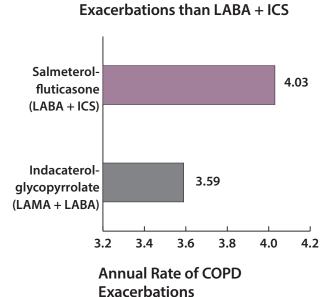


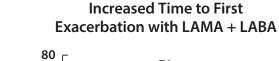
- First line: LAMA, consider LAMA + LABA if patient has more severe symptoms (e.g., CAT score >20)^{9,33}
 - Combination of LAMA + LABA reduced COPD exacerbations and increases the time to first exacerbation compared to LABA + ICS.³³
- Initial therapy with LABA + ICS can be considered in patients with:9
 - A history of asthma

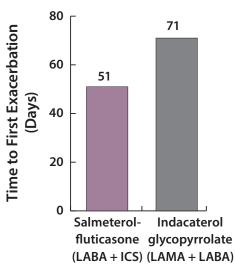
LAMA + LABA had a Lower Rate of COPD

 Severe symptoms and eos ≥300 cells/µL Blood eos ≥100 cells/µL and ≥2 moderate exacerbations or >1 hospitalization

Figure 9. Combination of LAMA + LABA Reduced COPD Exacerbations and Increased the Time to First Exacerbation Compared to LABA + ICS³³







FLAME trial evaluating indacaterol-glycopyrrolate (LABA+LAMA) compared to salmeterol-fluticasone (LABA+ICS) in patients with COPD and a history of at least one exacerbation in the previous year. The trial was designed to evaluate the annual rate of all COPD exacerbations (p <0.001 for all comparisons in figure).

Use LAMA as first line therapy for patients in Group C and D. Progress to combination therapy, usually LAMA + LABA, if monotherapy is not effective.

•••••

Follow Up Assessment and Treatment of COPD

Follow up treatment is based on how the patient responded:9

- ✓ If the patient had an adequate response to the initial treatment, then maintain it.
- ★ If not, modify treatment based on whether patient is having continued dyspnea or if they are having exacerbations. If they are having both, then follow the recommendations for exacerbations.⁹

Figure 10. Assessing Treatment Response for All GOLD Groups⁹

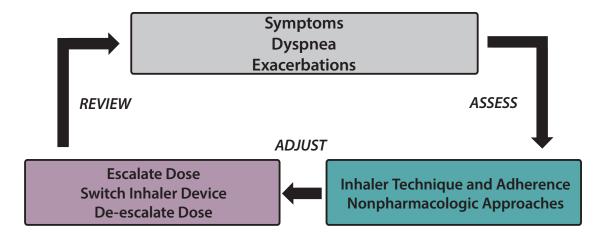


Figure 11. Proper Inhaler Technique is Critical for Effective COPD Pharmacotherapy

- ✓ Provide instructions and demonstrate proper technique when prescribing an inhaler device.
- ✓ Spacers improve drug delivery when used with metered dose inhalers (MDI).
- ✓ Inhaler technique and adherence to therapy should be assessed before considering dose adjustments and/or changing therapy. (Links to VA Instructional Videos in the Veteran Health Library: Combivent Respimat, Pressurized Metered Dose Inhaler, HandiHaler, Mometasone Twisthaler, How to Use a Nebulizer)

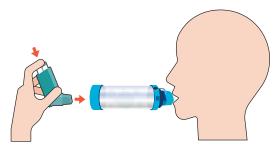


Table 2. At Follow-up Visit, Modify Treatment Based on Dyspnea and/or Exacerbations^{9,10,34,35}

Dyspnea Pathway			
Initial Treatment	New Treatment	Continued Dyspnea?	
LAMA or LABA LABA + ICS	Determine if ICS is needed. For most patients, LAMA+ LABA should be used. If	Consider switching inhaler device(s) and/or using a different drug in the same class.	
	ICS is needed, then use LAMA+LABA+ICS Evaluate other causes of dyspnea. Consider reto pulmonologist.		
Exacerbation Pathway			
Initial Treatment	New Treatment	Continued Exacerbations?	
LAMA or LABA	LAMA + LABA	LAMA + LABA + ICS	
	LABA + ICSIn patients with a history of asthma*	LAMA + LABA + ICS	
LABA + ICS	Determine if ICS is needed. For most patients, LAMA+ LABA should be used. If ICS is needed, then use LAMA+LABA+ICS	Refer to pulmonologist for consideration of roflumilast or azithromycin.	

^{*}Consider if eos \geq 300 cells/ μ L or if eos \geq 100 cells/ μ L and \geq 2 moderate exacerbations or >1 hospitalization. eos = blood eosinophil count; ICS = inhaled corticosteroid; LABA = long-acting beta₂ agonist; LAMA = long-acting muscarinic antagonist.

Table 3. Considerations for Starting Treatment with Inhaled Corticosteroids (ICS)^{9,36}

 History of asthma History of hospitalization(s) for exacerbations of COPD ≥2 moderate exacerbations of COPD/year Blood eosinophils 100–300 cells/μL Repeated pneumonia events Blood eosinophils 100–300 cells/μL History of mycobacterial infection 	Encourage Use	Consider Use	Advise Against Use
· blood edsirioprilis /300 celis/ µL	 History of hospitalization(s) for exacerbations of COPD ≥2 moderate exacerbations of 	of COPD/year • Blood eosinophils	 Blood eosinophils <100 cells/μL History of mycobacterial

Patient should be appropriately using one or two long-acting bronchodilators as maintenance therapy prior to considering ICS.

Use of Inhaled Corticosteroids

Monotherapy with ICS is not recommended in patients with COPD unless the patient also has asthma, since it does not improve FEV₁ or decrease exacerbations. Evidence shows that inhaled corticosteroids are most effective when used with long-acting bronchodilators in patients with severe COPD (Grades C and D). Blood eosinophil counts (eos) may help predict response to ICS (eos >300 cells/ μ L = greater likelihood to respond, eos <100 cells/ μ L = lower likelihood to respond).

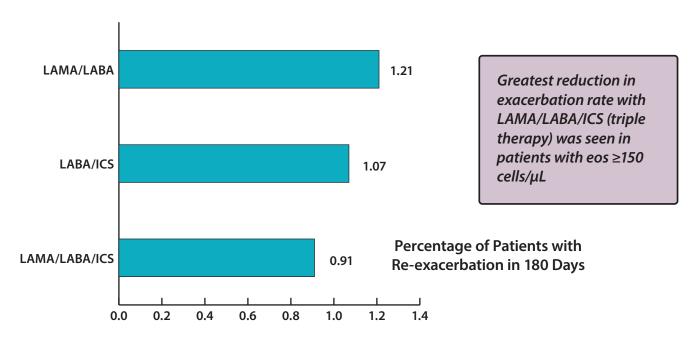
ICS inhalers have adverse effects which need to be considered and discussed before prescribing. ICS adverse effects include:9,40-42

- Oral candidiasis
- Hoarse voice
- Skin bruising
- Pneumonia

Patients at highest risk of pneumonia related to ICS include patients:

- Who smoke
- Are >55 years of age
- Have a history of pneumonia
- Have a body mass index <25 kg/m²
- Have severe airflow limitation

Figure 12. Triple Therapy Versus Dual Therapy in Patients with Grade C or D COPD³⁷
Lower Exacerbation Rate Using LAMA/LABA/ICS



Rate of Moderate or Severe COPD Exacerbation Per Year

IMPACT trial evaluated the use of LAMA/LABA, LABA/ICS, and LAMA/LABA/ICS (single inhaler triple combination) in patients with COPD. LAMA/LABA/ICS had the lowest rate of exacerbations at 0.91/year (p< 0.001). In a sub-group analysis, patients with an eos \geq 150 cells/µL showed the greatest reduction in exacerbation rates using LAMA/LABA/ICS. LAMA/LABA = umeclidinium (LAMA)/ vilanterol (LABA); LABA/ICS = vilanterol (LABA)/fluticasone furoate (ICS); LAMA/LABA/ICS = umeclidinium (LAMA)/vilanterol (LABA)/fluticasone furoate (ICS).

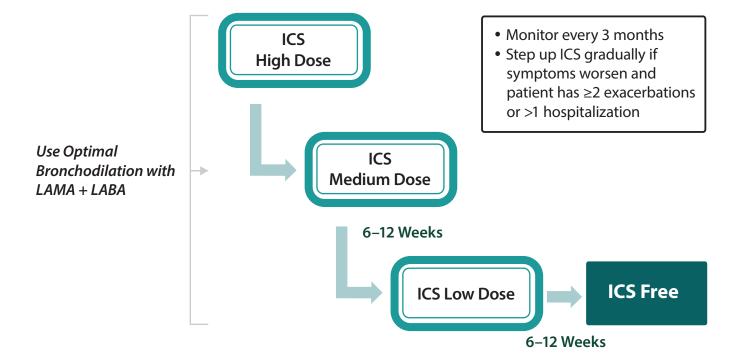
Withdrawal of Inhaled Corticosteroids

Corticosteroid inhalers may need to be discontinued. The dose should be slowly tapered to avoid worsening of symptoms. Caution should be used when withdrawing ICS in patients with blood eos ≥ 300 cells/µL due to a higher risk of experiencing more exacerbations after stopping ICS. $^{9,40-42}$

Possible candidates for ICS discontinuation include patients with:9

- Pneumonia
- Inappropriate original indication for ICS
- Lack of response to ICS
- Stable COPD without exacerbation in past year

Figure 13. Tapering Inhaled Corticosteroids^{43–44}



When Inhalers are not Enough

In patients with continued exacerbations despite maximizing inhaler therapy, a referral to a pulmonologist is recommended. Roflumilast or azithromycin may be indicated.⁹

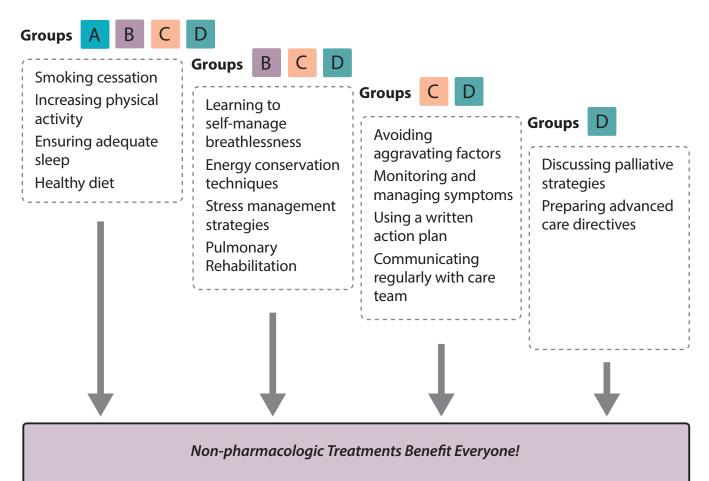
Table 4. Options for Group D Patients on LABA + LAMA +/- ICS who Still Have Exacerbations^{9,45-47}

Roflumilast*	Macrolide Antibiotic
Indicated for patients with FEV <50% with >1 recorded exacerbation requiring systemic steroids, unscheduled healthcare contact, or hospitalization in the previous year.	Azithromycin is the macrolide of choice for this indication. Best evidence is for use in patients who are former smokers. May reduce exacerbation rates in some patients.
Patients should be on optimal bronchodilatory therapy (LAMA + LABA) +/- ICS. Prescribing should be done by a pulmonologist or designated expert.	Patients should be on optimal bronchodilatory therapy (LAMA + LABA) +/- ICS. Prescribing should be done by a pulmonologist or designated expert.
Avoid use in patients with depression; may increase risk of suicide. Contraindicated in patients with moderate to severe liver impairment.	Associated with an increased incidence of bacterial resistance, prolongation of QTc interval, and impaired hearing tests.
Roflumilast 500 mg tablet orally once daily; may initiate at 250 mg daily for 4 weeks to reduce adverse effects.	Azithromycin 250 mg orally once daily or 500mg three times weekly.

^{*}Criteria for use of roflumilast is available at **https://www.pbm.va.gov/**; see ADS COPD Quick Reference Guide for more detailed medication information.

Non-pharmacologic Therapies

Figure 14. Non-pharmacologic Treatments Recommended by Group Classification 9,48,49



Non-pharmacologic treatments are a vital part of the treatment plan for all patients with COPD.

Pulmonary Rehabilitation

Pulmonary rehabilitation is an evidence-based, multidisciplinary intervention for patients with chronic respiratory disease who are symptomatic and have a reduction in daily life activities.^{9,12,50,51}

Pulmonary Rehabilitation:

 Reduces symptoms, optimizes functional status, increases participation, and reduces healthcare costs through stabilizing or reversing systemic manifestations of the disease.

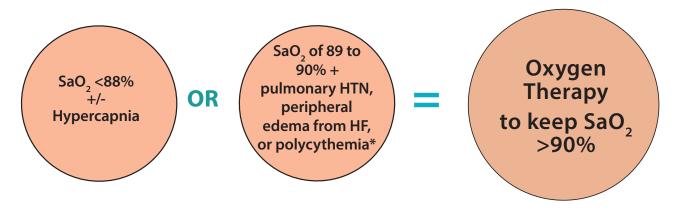


- May benefit patients with COPD groups B, C, and D⁹
- Should be provided at diagnosis, at discharge after hospitalization for an exacerbation, or when symptoms are progressively deteriorating⁹. Check with your local facility about the referral process for Pulmonary Rehabilitation.

Oxygen Therapy

Using supplemental oxygen long-term (>15 hours a day) increases survival in patients with chronic respiratory failure and severe chronic resting hypoxemia.⁵² In patients with stable COPD and moderate resting or exercise-induced arterial desaturation, however, long-term oxygen does not prolong survival or time to first hospitalization or provide sustained benefit in health status, lung function, or 6-minute walk distance.⁵³

Figure 15. Oxygen is Indicated When Oxygen Saturation (SaO₂) Falls Below 88%^{9,12}



CHF = congestive heart failure; HTN = hypertension; $SaO_2 = Oxygen$ saturation. *Polycythemia defined as hematocrit >55%.

Recheck SaO₂ 60 to 90 days after starting oxygen therapy to decide if:

- Supplemental oxygen is still indicated
- Prescribed supplemental oxygen is effective

Acute Exacerbations

A COPD exacerbation is defined as an acute worsening of respiratory symptoms that requires additional therapy. This section will focus on medication management and will not include non-invasive or invasive ventilation techniques for severe exacerbations of COPD.

Exacerbations can be classified as:6,9,12

- Mild treated with short-acting bronchodilators only
- Moderate treated with short-acting bronchodilators plus antibiotics and/or oral corticosteroids
- Severe requires hospitalization or visit to the emergency department

The most common causes of exacerbations are respiratory tract infections. Long-term prognosis following hospitalization for COPD exacerbation is poor with a 5-year mortality rate of about 50%.^{54–57}

Figure 16. Factors Associated with Poor Outcome After an Exacerbation^{54–57}

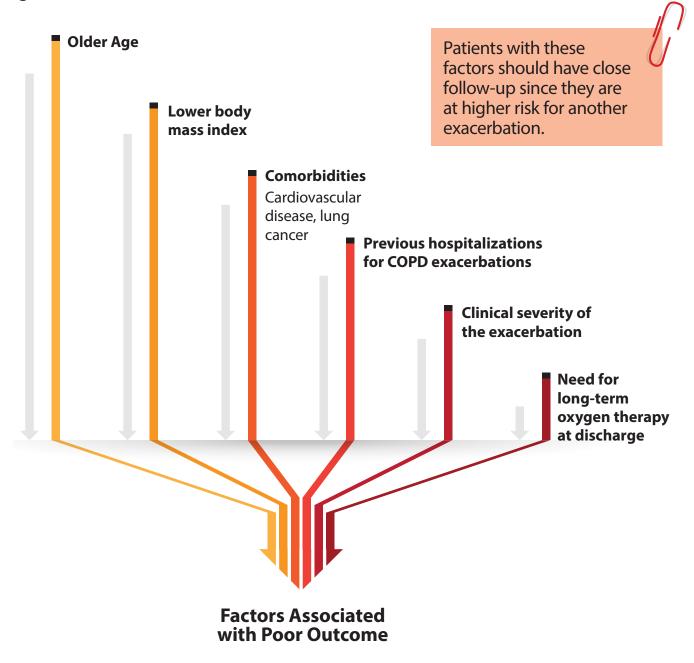
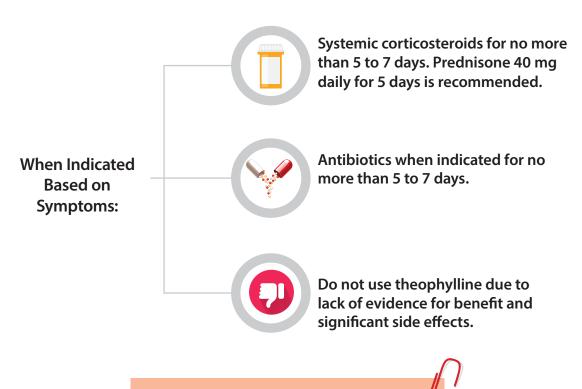


Figure 17. Recommended Approach to Managing COPD Exacerbations^{6,9,54–57}



All Patients:

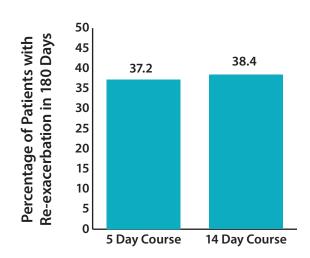
Short-acting inhaled beta2-agonist (SABA) with or without short-acting muscarinic antagonist (SAMA).

Continue long-acting bronchodilators.

Oral Steroid Use^{56,57}

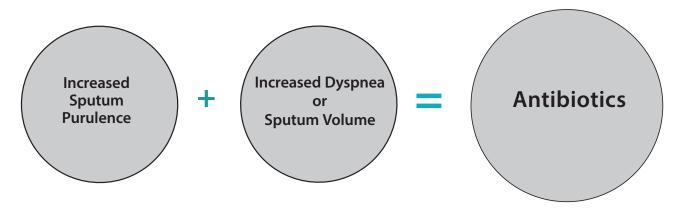
- Typical oral steroid regimen is prednisone 40 mg orally daily for 5 days.
- Steroid use >5 days was not shown to improve outcomes and will increase side effects.
- Use for patients with a poor response to bronchodilators.
- Prednisone does not need to be tapered when using a 5-day course.

Figure 18. Re-exacerbation Rates are Similar Using Prednisone for 5 Days Compared to 14 Days⁵⁶



Antibiotics are indicated when there is an increase in sputum purulence along with increased dyspnea or an increase in sputum volume.

Figure 19. When to Use Antibiotics^{7,9,12}



Choice of antibiotic should be based on local resistance patterns, but empirical therapy is usually amoxicillin + clavulanic acid, macrolide (e.g., azithromycin), or second generation cephalosporin (e.g., cefuroxime).

Treat an exacerbation with short-acting beta-agonist(s), continue long-acting bronchodilator(s), and give short courses of oral corticosteroids (5-days) and antibiotics (5–7 days) when indicated.

Figure 20. Interventions that May Reduce the Frequency of COPD Exacerbations⁹

Long-acting Bronchodilators	Corticosteroid- containing Regimens	Anti- inflammatory (non-steroid)	Anti-infectives	Non- pharmacologic
LAMA	LABA + ICS	Roflumilast	Vaccines	Smoking Cessation
LABA LAMA + LABA	LAMA + LABA + ICS		Long-term Macrolides	Pulmonary Rehabilitation

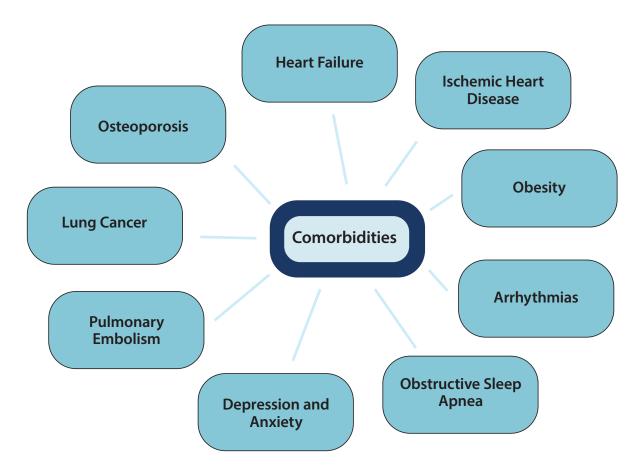


In the VA COPD CARE Service Project, having a clinical pharmacy specialist and a registered nurse involved in seeing patients after hospital discharge for a COPD exacerbation resulted in 0% of patients being readmitted to the hospital or emergency department with a COPD exacerbation at 30-days post discharge.⁵⁸

Treating Other Comorbidities

Treating comorbidities is very important for optimal management of COPD. Conditions that are of greatest concern:⁹

Figure 21. Important Comorbidities to Treat for Optimal COPD Management



Summary of Treatment Approaches

Prevention and treatment of exacerbations involves many different types of pharmacologic and non-pharmacologic treatments. Ensuring that all patients diagnosed with COPD receive the recommended treatments will help reduce exacerbations and hospitalizations. When a Veteran does have a COPD exacerbation, providing evidence based treatments can reduce the rates of future exacerbations and readmissions. A summary of the recommendations are listed below.

Figure 22. Review of the Recommendations for Patients with Stable COPD and Those with COPD Exacerbations

All Patients with Stable COPD

- Smoking Cessation
- Spirometry for diagnosis
- Long-acting bronchodilators
 - LAMA
 - LABA
- Influenza vaccine
- Short-acting beta-agonist for acute symptoms
- Pulmonary Rehabilitation
- Oxygen therapy
 - For severe resting hypoxemia

All Patients Having a COPD Exacerbation

- Short-acting and long-acting bronchodilators
- Corticosteroids
- Antibiotics if indicated
- → After the exacerbation
 - Pulmonary rehabilitation
 - Escalation of therapy
 - Adding second long-acting bronchodilator
 - If on LAMA and LABA, then add ICS



Therapies that May be Overused:

- Oxygen in non-severe resting hypoxemia, especially after an exacerbation
- Inhaled corticosteroids (ICS)
- Antibiotics in treatment of exacerbations

Appendix A

When I walk up a hill or a

breath

condition

I sleep soundly

I have lots of energy

flight of stairs I am not out of

I am not limited to doing any

I am confident leaving my

home despite my lung

activities at home

Your name:	CAT
Today's date:	
How is your COPD? Take the COPD Assessment Test™ (CAT) This questionnaire will help you and your healthcare professional to measure the impact the COPD (Chronic Obstructive Pulmonary Disease) is having on your wellbeing and daily life answers and test score can be used by you and your healthcare professional to help improte the management of your COPD and gain the greatest benefit from the treatment.	. Your
For each item below, place a mark (X) in the box that best describes your current situation Please ensure that you only select one response for each question.	
Example: I am very happy	
	SCORE
I never cough I never cough I cough all the time	
I have no phlegm (mucus) on my chest at all O 1 2 3 4 5 My chest is full of phlegm (mucus)	
My chest does not feel tight at all	

When I walk up a hill or a

completely out of breath

I am completely limited to

doing all activities at home

I am not confident leaving

my home at all because of

because of my lung condition

TOTAL SCORE

my lung condition

I do not sleep soundly

I have no energy at all

flight of stairs I am

A COPD assessment test was developed by an interdisciplinary group of international COPD experts with support from GSK. GSK's activities in connection with the COPD assessment test are monitored by a supervisory council that includes external, independent experts, one of which is chair of the council.

CAT, the COPD assessment test and the CAT logo are trademarks that belong to the GSK group of companies. ©2009 GSK. All rights reserved.

Appendix B

Modified Medical Research Council Breathlessness Scale

Grade	Description of Breathlessness
0	I only get breathless with strenuous exercise
1	I get short of breath when hurrying on level ground or walking up a slight hill
2	On level ground, I walk slower than people of the same age because of breathlessness, or have to stop for breath when walking at my own pace
3	I stop for breath after walking about 100 yards or after a few minutes on level ground
4	I am too breathless to leave the house or I am breathless when dressing

This summary was written by:

Julianne Himstreet, Pharm D., BCPS **Daina L Wells**, Pharm D., MBA, BCPS, BCPP **Sarah J. Popish**, Pharm D., BCPP

We thank our expert reviewers:

David H. Au, MD, MS
Allen Blaivas, MD
Donald Curran, MD
Debbie Khachikian, Pharm.D.
Rahul Khosla, MD, MBA
Cain Eric Kirk, Pharm.D. BCACP
Michael McFarland, Pharm.D., FCCP, BCACP
Mary Parker, Pharm.D., FCCP, BCPS, BCCP
Edward Portillo, Pharm.D.
Faryal Qureshi, Pharm.D.
Robert M. Reed, MD
Govindan Sushant, MD
J. Michael Wells, MD, MSPH

REFERENCES

- Croft JB, Wheaton AG, Liu Y, et al. Urban-Rural County and State Differences in Chronic Obstructive Pulmonary Disease — United States, 2015. MMWR Morb Mortal Wkly Rep. 2018; 67:205–211. DOI: http://dx.doi.org/10.15585/mmwr.mm6707a1
- 2. Mannino, DM, Homa, DM, Akinbami, LJ, Ford, ES, and Redd, SC. Centers for Disease Control and Prevention. Chronic obstructive pulmonary disease surveillance—United States, 1971–2000. MMWR Surveill Summ. 2002; 51: 1–16
- 3. Hill K, Golstein RS, Guyatt GH, et al. Prevalence and underdiagnosis of chronic obstructive pulmonary disease among patients at risk in primary care. *CMAJ*. 2010; 182(7):673–8.
- 4. Murphy SL, Xu JQ, Kochanek KD, Arias E. Mortality in the United States, 2017. NCHS Data Brief, no 328. Hyattsville, MD: National Center for Health Statistics. 2018.
- 5. US Department of Veterans Affairs. 172VA10P2: VHA Corporate Data Warehouse VA. 79 FR4377. Accessed May 16, 2019.
- 6. U.S. Department of Health and Human Services. National Institutes of Health. COPD: Are you at risk? NIH Publication No. 07-5840. September 2006. https://www.cdc.gov/copd/pdfs/fact_sheet-COPD-Are_You_at_Risk.pdf. Accessed 3/11/2019.
- 7. Criner GJ, Bourbeau M, Diekemper RL, et al. Prevention of Acute Exacerbations of COPD. American College of Chest Physicians and Canadian Thoracic Society Guidelines. *Chest*. 2015;147(4):894–942.
- 8. Connors, AF Jr, Dawson, NV, Thomas, C et al. Outcomes following acute exacerbation of severe chronic obstructive lung disease. The SUPPORT investigators (Study to Understand Prognoses and Preferences for Outcomes and Risks of Treatments). *Am J Respir Crit Care Med.* 1996; 154: 959–967b.
- Global Initiative for Chronic Obstructive Lung Disease. Global strategy for the diagnosis, management, and prevention of chronic obstructive pulmonary disease. Updated 2020. Global initiative for Chronic Obstructive Lung Disease website. Accessed 11/12/2019. https://goldcopd.org/wp-content/uploads/2019/11/GOLD-2020-REPORT-ver1.0wms.pdf.

- 10. Siu AL, Bibbins-Domingo K, Grossman DC, et al. Screening for chronic obstructive pulmonary disease: US Preventive Services Task Force Recommendation Statement. *JAMA*. 2016; 315(13):1372–7.
- 11. Qaseem A, Snow V, Shekelle P, et al. Diagnosis and management of stable chronic obstructive pulmonary disease: a clinical practice guideline from the American College of Physicians. *Ann Intern Med.* 2007; 147(9):633–8.
- 12. U.S. Department of Veterans Affairs, Department of Defense. VA/DoD Clinical Practice Guidelines for the Management of Chronic Obstructive Pulmonary Disease. Veterans' Health Administration, Office of Quality, Safety and Value and Office of Evidence Based Practice. The Management of Chronic Obstructive Pulmonary Disease Working Group; December 2014.
- 13. Raad D, Gaddam S, Schunemann HJ, et al. Effects of water-pipe smoking on lung function: a systematic review and meta-analysis. *Chest*. 2011; 139(4):764–74.
- 14. Tan WC, Lo C, Jong A, et al. Marijuana and chronic obstructive lung disease: a population-based study. *CMAJ*. 2009; 180(8):814–20.
- 15. Jha P, Ramasundarahettige C, Landsman V, et. al. 21st-Century hazards of smoking and benefits of cessation in the United States. *NEJM*. 2013; 368:341–350.
- 16. Fiore MC, Jaén CR, Baker TB, et al. Treating Tobacco Use and Dependence: 2008 Update. Clinical Practice Guideline. Rockville, MD: U.S. Department of Health and Human Services. Public Health Service. May 2008.
- 17. Barua RS, Rigotti NA, Benowitz NL, et al. 2018 ACC expert consensus decision pathway on tobacco cessation treatment. *J Am Coll Cardio.l* 2018; 72:3332–65.
- 18. U.S. Department of Health and Human Services. The Health Consequences of Smoking—50 Years of Progress. A Report of the Surgeon General. Atlanta: U.S. Department of Health and Human Services, Centers for Disease Control and Prevention, National Center for Chronic Disease Prevention and Health Promotion, Office on Smoking and Health, 2014.
- 19. Poole PJ, Chacko E, Wood-Baker RW, Cates CJ. Influenza vaccine for patients with chronic obstructive pulmonary dis-ease. *Cochrane database of systematic reviews (Online)*. 2006(1):CD002733.
- 20. Wongsurakiat P, Maranetra KN, Wasi C, et al. Acute respiratory illness in patients with COPD and the effectiveness of influenza vaccination: a randomized controlled study. *Chest.* 2004; 125(6):2011–20.
- 21. Nichol KL, Baken L, Wuorenma J, Nelson A. The health and economic benefits associated with pneumococcal vaccina-tion of elderly persons with chronic lung disease. *Archives of internal medicine*. 1999; 159(20):2437–2442.
- 22. Walters JA, Tang JN, Poole P, et al. Pneumococcal vaccines for preventing pneumonia in chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2017;1: Cd001390.
- 23. Miller MR, Hankinson J, Brusasco V, et al. Standardisation of spirometry. *Eur Respir J*. 2005; 26(2):319–38.
- 24. Pellegrino R, Viegi G, Brusasco V, et al. Interpretive strategies for lung function tests. *Eur Respir J*. 2005; 26(5):948–68.
- 25. Jones PW, Harding G, Berry P, et al. Development and first validation of the COPD Assessment Test. *Eur Respir J*. 2009; 34(3):648–54.

- 26. Fletcher CM. Standardised questionnaire on respiratory symptoms: a statement prepared and approved by the MRC Committee on the Aetiology of Chronic Bronchitis (MRC breathlessness score). BMJ. 1960; 2:1662.
- 27. Sestini P, Renzoni E, Robinson S, et al. Short-acting beta 2 agonists for stable chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2002;(4):CD001495.
- 28. Kew KM, Mavergames C, Walters JA. Long-acting beta2 agonists for chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2014;10(10):CD010177.
- 29. Appleton S, Poole P, Smith B, et al. Long-acting beta 2-agonists for poorly reversible chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2006; 3(3):CD001104.
- 30. Barr RG, Bourbeau J, Camargo CA, et al. Inhaled tiotropium for stable chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2005;(2):CD002876.
- 31. Vogelmeir C, Hederer B, Glaab T, et al. Tiotropium versus salmeterol for the prevention of exacerbations of COPD. *NEJM*. 2011; 364(12):1093–1103.
- 32. Decramer ML, Chapman KR, Dahl R, et al. Once-daily indacaterol versus tiotropium for patients with severe chronic obstructive pulmonary disease (INVIGORATE): a randomized, blinded, parallel-group study. *The Lancet Respiratory Medicine*. 2013; 1(7):524–33.
- 33. Wedzicha J.A., Banerji D., Chapman K.R., et al: Indacaterol-glycopyrronium versus salmeterol-fluticasone for COPD. *NEJM*. 2016; 374: pp. 2222–2234.
- 34. Magnussen H, Disse B, Rodriguez-Roisin R, et al. Withdrawal of inhaled glucocorticoids and exacerbations of COPD. *NEJM*. 2014; 371:1285–94.
- 35. Chapman KR, Hurst JR, Frent SM, et al. Long-term triple therapy de-escalation to indacaterol/glycopyrronium in COPD patients (SUNSET): a randomized, double-blind, triple-dummy clinical trial. *Am J Respir Crit Care Med*. 2018; 198(3):329–339.
- 36. Jørgen Vestbo, Leonardo Fabbri, Alberto Papi, Stefano Petruzzelli, Mario Scuri, Alessandro Guasconi, Stefano Vezzoli, Dave Singh European Respiratory Journal 2018 52: 1801230; DOI: 10.1183/13993003.01230-201.
- 37. Lipson DA, Barnhart F, Brealey N, et al. Once daily single inhaler triple versus dual therapy in patients with COPD. *NEJM*. 2018; 378(18):1671–80.
- 38. Siddiqui SH, Guasconi A, Vestbo J, et al. Blood eosinophils: A biomarker of response to extra fine beclomethasone/formoterol in chronic obstructive pulmonary disease. *Am J Respir Crit Care Med*. 2015; 192(4):523–5.
- 39. Bafadhel M, Peterson S, De Blas MA, et al. Predictors of exacerbation risk and response to budesonide in patients with chronic obstructive pulmonary disease: a post-hoc analysis of three randomized trials. *The Lancet Respiratory Medicine*. 2018; 6(2):117–26.
- 40. Crim C, Dransfield MT, Bourbeau J, et al. Pneumonia risk with inhaled fluticasone furoate and vilanterol compared with vilanterol alone in patients with COPD. *Annals of the American Thoracic Society.* 2015; 12(1):27–34.
- 41. Crim C, Calverley PMA, Anderson JA, et al. Pneumonia risk with inhaled fluticasone furoate and vilanterol in COPD patients with moderate airflow limitation: The SUMMIT trial. *Respir Med*. 2017; 131:27–34.
- 42. Yang IA, Clarke MS, Sim EH, et al. Inhaled corticosteroids for stable chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2012; 7(7):CD002991.

- 43. Cataldo D, Derom E, Liistro G, et al. Overuse of inhaled corticosteroids in COPD: five questions for withdrawal in daily practice. *Int J Chron Obstruct Pulmon Dis*. 2018; 13:2089–2099. Published 2018 Jul 5. doi:10.2147/COPD.S164259
- 44. Magnussen H, Disse B, Rodriguez-Roisin R, et al. Withdrawal of inhaled glucocorticoids and exacerbations of COPD. *NEJM*. 2014; 371(14):1285–1294.
- 45. Martinez FJ, Rabe KF, Sethi S, et al. Effect of roflumilast and inhaled corticosteroid/long-acting beta-2-agonist on chronic obstructive pulmonary disease exacerbations (RE2SPOND) A randomized clinical trial. *Am J Respir Crit Car Med*. 2016; 194(5):599–67.
- 46. Rabe KF, Calverley PMA, Martinz FJ, et al. Effect of roflumilast in patients with severe COPD and a history of hospitalization. *Eur Respir J.* 2017; 50(1) 1700158; DOI: 10.1183/13993003.
- 47. Albert RK, Connett J, Bailey WC, et al. Azithromycin for prevention of exacerbations of COPD. *NEJM*. 2011; 365(8):689–98.
- 48. Lahham A, McDonald CF, Holland AE. Exercise training alone or with the addition of activity counseling improves physical activity level in COPD: a systematic review and meta-analysis of randomized controlled trials. *Int J Chron Obstruct Pulmon Dis*. 2016; 11:3121–36.
- 49. Ortega F, Toral J, Cejudo P, et al. Comparison of effects of strength and endurance training in patients with chronic obstructive pulmonary disease. *Am Je Respir Crit Care Med*. 2002; 166(5):669–74.
- 50. Nici L, Donner C, Wouters E, et al. ATS/ERS Pulmonary Rehabilitation Writing Committee. American Thoracic Society/European Respiratory Society Statement on Pulmonary Rehabilitation. *Am J Respir Crit Care Med*. 2006; 173:1390–1413.
- 51. McCarthy B, Casey D, Devane D, Murphy K, Murphy E, Lacasse Y. Pulmonary rehabilitation for chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2015; 2(2): CD003793.
- 52. Cranston JM, Crockett AJ, Moss JR, et.al. Domiciliary oxygen for chronic obstructive pulmonary disease. *Cochrane Database Syst Rev.* 2005; (4): CD001744.
- 53. Long-term Oxygen Treatment Trial Research Group. A randomized trial of long-term oxygen for COPD with moderate desaturation. *NEJM*. 2016; 375(17):1617.
- 54. Miravitlles M, Anzueto A. Chronic Respiratory Infection in Patients with Chronic Obstructive Pulmonary Disease: What is the Role of Antibiotics. *Intl J Mol Sci.* 2017, 18, 1344.
- 55. Hurst JR, Wedzicha JA. Chronic obstructive pulmonary disease: the clinical management of an acute exacerbation. *Postgradu-ate medical journal*. 2004; 80(947):497–505. http://pmj.bmj.com/content/80/947/497.full.pdf
- 56. Leuppi JD, Schuetz P, Bingisser R, et al. Short-term vs conventional glucocorticoid therapy in acute exacerbations of chronic obstructive pulmonary disease: the REDUCE randomized clinical trial. *JAMA*. 2013; 309(21):2223 2231. http://www.ncbi.nlm.nih.gov/pubmed/23695200
- 57. Walters JA, Tan DJ, White CJ, Gibson PG, Wood-Baker R, Walters EH. Systemic corticosteroids for acute exacerbations of chronic obstructive pulmonary disease. *Cochrane database of systematic reviews (Online)*. 2014(9):CD001288. https://www.ncbi.nlm.nih.gov/pubmed/25178099.
- 58. VA COPD CARE Service Project. Diffusion of Excellence Gold Status Project. VA Pulse: https://www.vapulse.va.gov/groups/copd-care.

U.S. Department of Veterans Affairs

This reference guide was created as a tool for VA providers and is available from the Academic Detailing Service SharePoint.

These are general recommendations only. The treating provider should make clinical decisions based on an individual patient's clinical condition.

VA PBM Academic Detailing Service Email Group PharmacyAcademicDetailingProgram@va.gov

VA PBM Academic Detailing Service SharePoint Site https://vaww.portal2.va.gov/sites/ad/SitePages/Home.aspx

VA PBM Academic Detailing Public WebSite http://www.pbm.va.gov/PBM/academicdetailingservicehome.asp

Revised November 2019 IB 10-1155, P96917 **www.va.gov**