

COPD

Chronic Obstructive Pulmonary Disease (COPD)

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Abbreviations

CAT = COPD Assessment Test

DPI = Dry Powder Inhaler

HF = Heart Failure

HTN = Hypertension

ICS = Inhaled Corticosteroid

IIV = Inactivated Influenza Vaccine

LABA = Long-Acting Beta₂ Agonists

LAMA = Long-Acting Muscarinic Antagonist

MDI = Metered Dose Inhaler

mMRC = Modified Medical Research Council Breathlessness Scale

NRT = Nicotine Replacement Therapy

RIV = Recombinant Influenza Vaccine

RZV = Recombinant Zoster Vaccine

SABA = Short-Acting Beta₂ Agonists

SAMA = Short-Acting Muscarinic Antagonist

 $SaO_2 = Oxygen Saturation$

SMI = Soft Mist Inhaler

ZVL = Zoster Vaccine Live

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Elements of a COPD Treatment Plan^{1,2}

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.

Brief Tobacco Cessation Counseling – the 5 As³



Ask — About Tobacco Use

 Ask about type of tobacco, how much used daily, and prior experience in quitting.



Advise — To Quit Now

 Focus on benefits of quitting for COPD and other health concerns like cardiovascular disease.





Assess — Is the Patient Ready to Quit

• Is the patient ready to quit in the next 30 days? If "Yes," then proceed. If "No," then encourage quitting.



Assist — Offer and Connect to Treatment

 Prescribe pharmacotherapy and offer behavioral support.*



Arrange — Follow Up in 1 to 2 Weeks

If patient accepts treatment, follow up in 1 to 2 weeks.
 If patient declines treatment, continue to encourage cessation at every visit.

*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 (www.pbm.va.gov).

Not Ready to Quit in the Next 30 Days? – Try Using the 5 Rs to Build Motivation³

RELEVANCE What are some things that concern you about smoking?

For example, heath concerns, affect on family, finances, etc.

RISKS What effect has tobacco had on your health?

Reviewing risks of using tobacco and then discussing the benefits of quitting helps increase motivation to quit. Focus on the health improvements from quitting.

REWARDS What will you gain by quitting tobacco?

Awareness of rewards helps maintain motivation during the quit attempt.

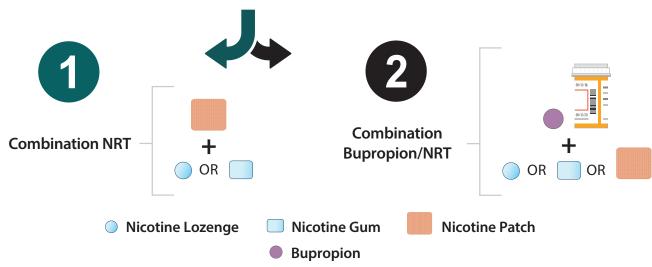
ROADBLOCKS What barriers do you see that may impact your ability to quit?

Common barriers are withdrawal symptoms, fear of failure, lack of support, depression, and being around other smokers. Identifying barriers and what has contributed to relapse in the past, can help in planning for the next guit attempt.

REPETITION Ask readiness to quit at each encounter.

Ask permission to check in at the next visit. The more healthcare providers talk about tobacco cessation, the greater the likelihood a patient may try to stop.

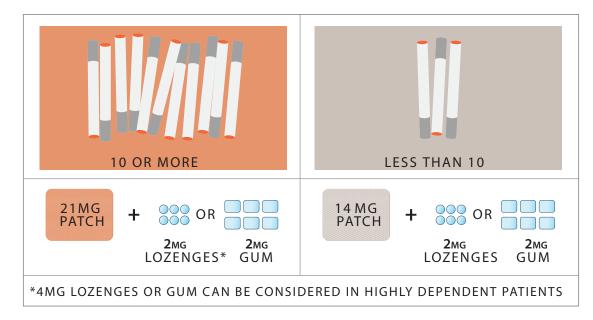
First-line Therapies for Tobacco Cessation



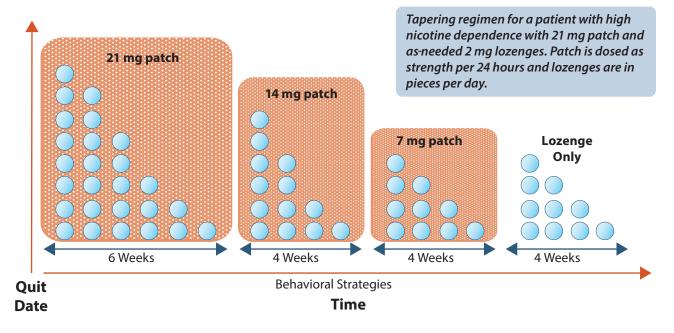
Monotherapy with Nicotine Replacement Therapy (NRT) or bupropion can be considered for patients who are unable to tolerate combination therapy or wish to use monotherapy; however, cessation rates may be lower.

Recommended Starting Dose for Combination NRT⁴

DAILY CIGARETTE CONSUMPTION



Tapering Examples for Combination NRT^{2,4}



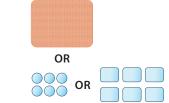
Combination Bupropion/NRT^{5,6}



SEVEN TO 14 DAYS BEFORE QUIT DAY

Start bupropion at 150mg SR once daily for three days then increase to 150mg SR twice daily.

NRT is not started until quit day.



QUIT DAY

Start NRT: Nicotine patch, lozenge, or gum.*

Continue bupropion 150mg SR twice daily.



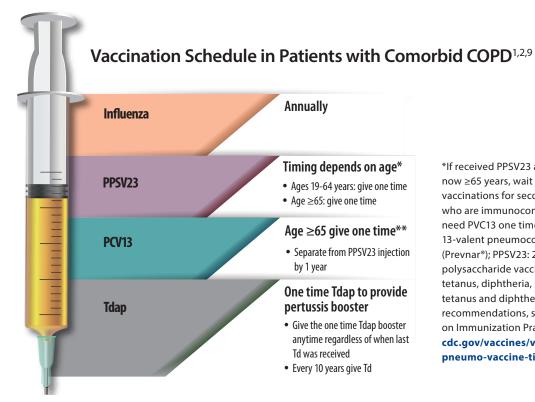
Continue bupropion for eight to 12 weeks or longer, if appropriate.**

Stop nicotine patch at the same time as bupropion. If using nicotine lozenge or gum, then taper using the same schedule as combination NRT.

*Only use one form of NRT when used in combination therapy with bupropion. Current evidence does not show that using multiple forms of NRT with bupropion is more effective. Nicotine lozenge or nicotine gum in combination with bupropion may be more effective than nicotine patch in combination with bupropion, based on current clinical evidence. **Bupropion may help with depressive symptoms, so some patients may benefit from longer term use.

Vaccines in Patients with COPD^{1,2,9}

Vaccine	Why in COPD?	Adverse Effects	Scheduling	
Influenza (IIV, RIV)	Reduces incidence of lower respiratory infections and death	Injection site reactions, myalgia, headache, diarrhea	1 dose annually	
Pneumococcal	Reduces incidence of	Fatigue, loss of appetite,	Age 19-64: 1 dose	
Polysaccharide (PPSV23)	community-acquired pneumonia	injection site reactions, fever, headache	Age 65+: 1 dose	
Pneumococcal Conjugate (PCV13)	Reduces incidence of bacteremia and invasive pneumococcal disease	Fatigue, loss of appetite, injection site reactions, fever, headache	Age 65+: 1 dose	
Zoster (RZV, ZVL)	Increased risk of shingles in patients with COPD.	Injection site reactions, myalgia, headache, nausea, shivering	Ages 50+: 2 doses given 2–6 months apart	
Tetanus, Diphtheria (Td), Pertussis (Tdap)	Increased severity of pertussis infection in patients with COPD.	Injection site reactions, GI upset, fatigue, headache	One-time booster dose with Tdap then give Td every 10 years	



*If received PPSV23 at <65 years and patient is now ≥65 years, wait 5 years between PPSV23 vaccinations for second PPSV23 dose. **Patients who are immunocompromised or asplenic also need PVC13 one time when <65 years. PCV13: 13-valent pneumococcal conjugate vaccine (Prevnar®); PPSV23: 23-valent pneumococcal polysaccharide vaccine (Pneumovax®), Tdap: tetanus, diphtheria, pertussis vaccine; Td: tetanus and diphtheria vaccine. For specific recommendations, see the Advisory Committee on Immunization Practices (ACIP): https://www.cdc.gov/vaccines/vpd/pneumo/downloads/pneumo-vaccine-timing.pdf.

Pharmacotherapy Initiation^{1,2,7,8}

Exacerbation History				
0 or 1 exacerbations not leading to hospital admission ≥2 exacerbations or ≥1 leading to hospital admission				
Assessment of Symptoms/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	

^{*}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/µL or eos \geq 100 cells/µL and \geq 2 moderate exacerbations or >1 hospitalization. ICS = inhaled corticosteroid; LABA = long-acting beta₂ agonists; LAMA = long-acting muscarinic antagonist; SABA = short-acting beta₂ agonists; SAMA = short-acting muscarinic antagonist.

Exacerbation History				
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		If persistent 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 agonists; LAMA = long-acting muscarinic antagonist; SABA = short-acting beta₂ agonists; SAMA = short-acting muscarinic antagonist.

Rescue Inhalers (Use only for Intermittent Symptoms)^{1,2,10–14}

Inhaler Formulations	Duration of Action	Dosing	Comments
		Short-Acting Beta ₂ Agonists (SABA)	
Albuterol \$	4–6 hours	MDI, DPI: 2 inhalations every 4–6 hours as needed.	Monitor for sinus tachycardia,
		Nebulizer*: 2.5 mg every 6–8 hours as needed.	tremors, nervousness,
Levalbuterol \$	6–8 hours	MDI: 2 inhalations every 4–6 hours as needed.	hypokalemia.
		Nebulizer*: 0.63 mg every 6–8 hours as needed, 3 times per day.	
		Short-Acting Muscarinic Antagonist (SAMA)	
Ipratropium \$	6–8 hours	MDI: 2 inhalations up to 4 times per day.	Monitor for dry mouth and
		Nebulizer*: 500 mcg every 6–8 hours.	urinary symptoms.
			Monitor for increased side effects in combination with LAMAs.

Cost for 30-days supply: \$ = \$0-\$49; \$\$ = \$50-\$99, \$\$\$ = \$100-\$199, \$\$\$\$ = \$200+

^{*}May be more convenient for patients who are acutely ill or patients unable to use inhaler devices; patients not responding may benefit from increased dosage. VA Formulary information at: www.pbm.va.gov/apps/VANationalFormulary.

continued from page 13 (Rescue Inhalers (Use only for Intermittent Symptoms) 1,2,10–14)

Inhaler Formulations	Duration of Action	Dosing	Comments	
	Combination SABA/SAMA			
Albuterol/ Ipratropium \$	6–8 hours	SMI: 1 inhalation up to 4 times daily. Nebulizer*: one 3 mL vial 4 times daily.	Superior to either medication alone. Monitor for side effects of individual components.	

Cost for 30-days supply: \$ = \$0-\$49; \$\$ = \$50-\$99, \$\$\$ = \$100-\$199, \$\$\$\$ = \$200+

Why Use Long-acting Bronchodilators Over Short-acting Bronchodilators for Persistent Symptoms®?

Long-acting Bronchodilators

- Improve lung function
- Improve dyspnea
- Improve health status
- Reduce exacerbations

Short-acting Bronchodilators

- Improve dyspnea
- Temporarily improve lung function

^{*}May be more convenient for patients who are acutely ill or patients unable to use inhaler devices; patients not responding may benefit from increased dosage. VA Formulary information at: www.pbm.va.gov/apps/VANationalFormulary.

Maintenance Inhalers^{1,2,10–14,15}

Inhaler Formulations	Duration of Action	Dosing	Comments
	L	ong-Acting Muscarinic Antagonist (LAMA)	
Tiotropium \$	24 hours	DPI: 2 inhalations of contents of 1 capsule daily. SMI: 2 inhalations once daily.	Monitor for increased side effects in combination with SAMAs.
Glycopyrrolate \$\$	12–24 hours	DPI: Inhale contents of 1 capsule twice daily. Nebulizer: 25 mcg every 12 hours.	May be used as initial monotherapy in all groups. Monitor for dry mouth and
Umeclidinium \$\$\$\$	24 hours	DPI: 1 inhalation once daily.	urinary symptoms.
Aclidinium \$\$\$\$	12 hours	DPI: 1 inhalation twice daily.	Use soft mist inhaler (e.g., Respimat®) as first line formulation since it is easier
Revefenacin \$\$\$\$	24 hours	Nebulizer: 175 mcg once daily.	to use.

continued from page 15 (Maintenance Inhalers1,2,10–14,15)

Inhaler Formulations	Duration of Action	Dosing	Comments
		Long-Acting Beta ₂ Agonists (LABA)	
Olodaterol \$	24 hours	SMI: 2 inhalations once daily.	May be used as initial monotherapy in groups A and
Indacaterol \$\$	24 hours	DPI: Inhale contents of 1 capsule daily.	B, however using tiotropium (LAMA) first line is a more
Salmeterol \$\$\$\$	12 hours	DPI: 1 inhalation twice daily.	cost-effective approach. Monitor for sinus tachycardia,
Arformoterol \$\$\$	12 hours	Nebulizer: 15 mcg every 12 hours.	tremors, hypokalemia. Do not use as monotherapy in patients with asthma. These
Formoterol \$\$\$\$	12 hours	Nebulizer: 20 mcg every 12 hours.	patients with astrima. These patients should also be using an inhaled corticosteroid (ICS).

Inhaler Formulations	Duration of Action	Dosing	Comments
		Inhaled Corticosteroids (ICS)*	
Mometasone \$	No bronchodilation effects – dosing	MDI: 2 inhalations twice daily. DPI: 1–2 inhalations once to twice daily.	*Not to be used as monotherapy in patients without asthma component.
Ciclesonide \$\$ Fluticasone	based on study dosing and varied drug	MDI: 1–2 inhalations by mouth twice daily. DPI: 1 inhalation once daily.	Monitor for increased risk of pneumonia, oral candidiasis
Furoate \$\$\$	half-lives.	,	(thrush), hoarse voice. Rinse mouth with water after
Fluticasone Propionate \$\$\$		MDI: 2 inhalations twice daily. DPI: 1 inhalation twice daily.	use, do not swallow water. Beclomethasone is the
Budesonide \$\$\$		DPI: 2 inhalations twice daily.	only ICS that can be safely used in combination with
Beclomethasone \$\$\$		MDI: 1 inhalation twice daily.	protease inhibitors.

continued from page 15 (Maintenance Inhalers1,2,10–14,15)

Inhaler Formulations	Duration of Action	Dosing	Comments
		Combination LABA/LAMA	
Olodaterol/ Tiotropium \$\$	24 hours	SMI: 2 inhalations once daily.	May be used as initial therapy in group D.
Indacaterol/ Glycopyrrolate \$\$	12-24 hours	DPI: Inhale contents of 1 capsule twice daily.	Do not use with other LABAs or LAMAs. Monitor for side effects of
Vilanterol/ Umeclidinium \$\$\$\$	24 hours	DPI: 1 inhalation once daily.	individual components.
Formoterol/ Glycopyrrolate \$\$\$\$	12 hours	MDI: 2 inhalations twice daily.	

Inhaler Formulations	Duration of Action	Dosing	Comments	
Combination LABA/Inhaled Corticosteroid (ICS)				
Formoterol/ Budesonide \$	12 hours	MDI: 2 inhalations twice daily.	May be used as initial therapy in group D for patients with asthma. Monitor for side effects of	
Formoterol/ Mometasone \$	12 hours	MDI: 2 inhalations twice daily.		
Salmeterol/ Fluticasone \$	12 hours	DPI: 1 inhalation twice daily. MDI: 2 inhalations twice daily.	individual components.	
Vilanterol/ Fluticasone \$\$\$\$	24 hours	DPI: 1 inhalation once daily.		
Combination LAMA/LABA/ICS				
Umeclidinium/ Vilanterol/ Fluticasone \$\$\$\$	24 hours	DPI: 1 inhalation once daily.	Monitor for side effects of individual components.	

Types of Inhalers and Use^{1,2,10-14}

Inhaler		
Metered Dose Inhalers (MDI)*		
1. Remove cap	SABA	
2. Shake well prior to use	Albuterol (ProAir HFA®, Ventolin HFA®, Proventil HFA®)	
*if using spacer, insert inhaler	Levalbuterol (Xopenex HFA®)	
into spacer	SAMA	
3. Exhale away from inhaler	Ipratropium (Atrovent HFA®)	
4. Close lips around inhaler	ICS	
5. Depress inhaler while inhaling slowly	Mometasone (Asmanex HFA®)	
6. Using with a spacer is highly	Beclomethasone (QVAR Redihaler®)	
recommended	Fluticasone (Flovent HFA®)	
	Ciclesonide (Alvesco HFA®)	

^{*}May require priming before initial use, follow device specific instructions for use. **May come as a capsule, follow device-specific instructions for use. ***Follow device-specific instructions for use. VA Formulary information at: www.pbm.va.gov/apps/VANationalFormulary.

Inhaler			
Metered Dose Inhalers (MDI)*			
7. Hold breath for at least 5 seconds	LABA/LAMA		
8. Exhale away from inhaler	Formoterol/glycopyrrolate (Bevespi Aerosphere®)		
*Remove from spacer if using	LABA/ICS		
9. Repeat if more than one dose needed	Formoterol/budesonide (Symbicort HFA®)		
10. Place cap back onto inhaler	Formoterol/mometasone (Dulera HFA®)		
	Salmeterol/fluticasone (Advair HFA®)		
Dry Powder Inhalers (DPI)**			
1. Ensure doses remaining (dose counter	SABA		
or capsules)	Albuterol (ProAir RespiClick®)		
2. Remove cap/open mouthpiece	LABA		
3. Load the medication and keep inhaler level	Indacaterol (Arcapta Neohaler®), Salmeterol (Serevent Diskus®)		

^{*}May require priming before initial use, follow device specific instructions for use. **May come as a capsule, follow device-specific instructions for use. ***Follow device-specific instructions for use. VA Formulary information at: www.pbm.va.gov/apps/VANationalFormulary.

Inhaler Dry Powder Inhalers (DPI)** 4. Exhale away from inhaler LAMA Tiotropium (Spiriva HandiHaler®), Glycopyrrolate (Seebri Neohaler®), 5. Close lips around inhaler and inhale Umeclidinium (Incruse Ellipta®), Aclidinium (Tudorza Pressair®) quickly for as long as possible **ICS** 6. Hold breath for at least 5 seconds Mometasone (Asmanex Twisthaler®), Fluticasone (Flovent Diskus®, Arnuity 7. Exhale away from inhaler Ellipta®), Budesonide (Pulmicort Flexhaler®) 8. Close inhaler and avoid moisture I ABA/I AMA 9. Repeat if more than one dose needed Indacaterol/glycopyrrolate (Utibron Neohaler®), Vilanterol/umeclidinium 10. Close cap/mouthpiece (Anoro Ellipta®) LABA/ICS Vilanterol/fluticasone (Breo Ellipta®), Salmeterol/fluticasone (Wixela Inhub™, Advair Diskus®) LAMA/LABA/ICS Umeclidinium/vilanterol/fluticasone (Trelegy Ellipta®)

^{*}May require priming before initial use, follow device specific instructions for use. **May come as a capsule, follow device-specific instructions for use. ***Follow device-specific instructions for use. VA Formulary information at: www.pbm.va.gov/apps/VANationalFormulary.

Inhaler Soft Mist Inhaler (SMI)*** 1. Hold inhaler upright, with cap closed, SABA/SAMA turn base in direction of arrows Albuterol/Ipratropium (Combivent Respimat®) on label I ARA 2. Open cap and exhale away Olodaterol (Striverdi Respimat®) from inhaler LAMA 3. Close lips around inhaler, avoid Tiotropium (Spiriva Respimat®) covering air vents with hands LABA/LAMA 4. Inhale slowly and deeply while Olodaterol/Tiotropium (Stiolto Respimat®) depressing inhaler's dose button 5 Hold breath for at least 5 seconds 6. Exhale away from inhaler 7. Close inhaler cover 8. Repeat if more than one dose needed

^{*}May require priming before initial use, follow device specific instructions for use. **May come as a capsule, follow device-specific instructions for use. ***Follow device-specific instructions for use. VA Formulary information at: www.pbm.va.gov/apps/VANationalFormulary.

Anti-Inflammatory Medications^{1,2,10,11}

Phosphodiesterase-4 (PDE4) Inhibitors			
Dosing	Comments		
Roflumilast orally once daily	Reserve for patients whose inhaler therapy has been		
Initiate at 250 mcg x 4 weeks then maintenance dose of	optimized.		
500 mcg once daily	Indicated for patients with FEV <50% with > one exacerbation requiring systemic steroids, unscheduled healthcare contact, or hospitalization in the previous year. Prescribed only by pulmonologist or designated expert.		
	Monitor for nausea, diarrhea, abdominal discomfort, unexplained weight loss, insomnia, and headaches.		
	Avoid use in patients with depression. May increase the risk of suicide. Contraindicated in moderate to severe liver impairment.		
	Extensive hepatic metabolism, so need to monitor for drug interactions.		

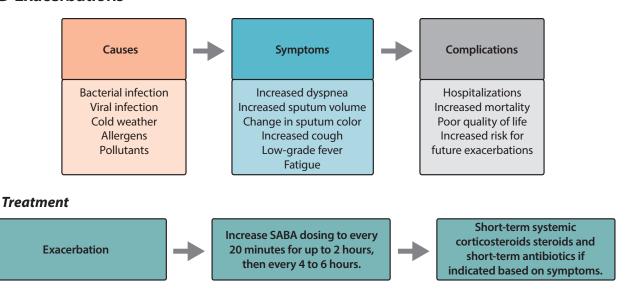
Criteria for use of roflumilast is available at https://www.pbm.va.gov. VA Formulary information at: www.pbm.va.gov. In the property in

Chronic Antibiotics^{1,2,10,11}

Chronic Azithromycin		
Dosing	Comments	
Azithromycin 250 mg PO once daily	Indicated for patients on LABA/LAMA or LABA/LAMA/ICS with recurrent exacerbations.	
500 mg PO 3x/weekly	Associated with increased bacterial resistance, hearing impairments, QTc prolongation.	
	No data showing benefit beyond one-year of treatment.	
	Only use in patients who are former smokers.	

VA Formulary information at: www.pbm.va.gov/apps/VANationalFormulary.

COPD Exacerbations^{1,2}



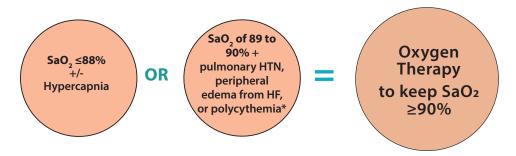
Medications for Acute Exacerbations*1,2

Drug	Dose	Comments	
Short-term Oral Antibiotics Antibiotics may be indicated in the following scenarios: increased sputum purulence, sputum volume, and dyspnea; ≥2 symptoms with at least one being increased sputum purulence; exacerbations requiring mechanical ventilation.			
Doxycycline	100 mg twice daily	Recommended length of therapy for all antibiotic use in exacerbation is 5–7 days.	
Amoxicillin/clavulanate	500 mg every 8 hours OR 875 mg every 12 hours		
Azithromycin	500 mg on day 1, then 250 mg on days 2–5		
Cefuroxime	500 mg twice daily		
Oral Corticosteroids In most cases, oral steroids are equally effective as intravenous steroids.			
Prednisone	40 mg once daily	Recommended length of therapy for steroids is 5 days.	
Methylprednisolone	40–60 mg once daily or in two divided doses		

^{*}Increased use of rescue inhaler is appropriate during exacerbations. Do not discontinue maintenance medications. **Use intravenous steroids when exacerbation is considered life-threatening. VA Formulary information at: www.pbm.va.gov/apps/VANationalFormulary.

Oxygen Therapy^{1,15–17}

Using supplemental oxygen long-term (>15 hours a day) for patients with chronic respiratory failure increases survival in patients who also have severe chronic resting hypoxemia. Oxygen is indicated when oxygen saturation (SaO_2) decreases to $\leq 88\%$. Recheck SaO_2 in 60 to 90 days after starting oxygen therapy to determine if supplemental oxygen is effective and still indicated.



*Polycythemia defined as hematocrit >55%. 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.

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This reference guide was created to be used as a tool for VA providers and is available to use from the Academic Detailing SharePoint. These are general recommendations only; specific clinical decisions should be made by the treating provider based on an individual patient's clinical condition.

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