

Pulmonology

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Lecture Modules

- Asthma
- PFT interpretation
- Miscellaneous
 - Sarcoidosis
 - Venous thromboembolism, DVT/PE
- Pneumonia
- CXRs
- COPD



Abbreviations

- LABA = long-acting beta agonist
- SABA = short-acting beta agonist
- LAMA = long-acting muscarinic antagonist
- SAMA = short-acting muscarinic antagonist
- ICS = inhaled corticosteroid
- LTRA = leukotriene receptor antagonist



Which of the following medication choices for steps 1-2 asthma is a **preferred** GINA treatment option?

- A. Salmeterol 50 mcg, 1 puff bid maintenance and albuterol rescue
- B. Budesonide/formoterol (80/4.5 mcg) bid maintenance and rescue
- C. Fluticasone 110 mcg, 2 puffs bid + montelukast 10 mg qHS maintenance and albuterol rescue
- D. Fluticasone + salmeterol (250/50) mcg 1 puff PRN



Correct Answer is B

- Answer A LABA/SABA is not an approved asthma treatment
- Answer C ICS/LTRA with SABA is an option but not preferred
- Answer D ICS/LABA with salmeterol is not a preferred option

A reminder – the key change in GINA 2019





Helen K. Reddel ¹, J. Mark FitzGerald², Eric D. Bateman³, Leonard B. Bacharier⁴, Allan Becker⁵, Guy Brusselle⁶, Roland Buhl⁷, Alvaro A. Cruz⁸, Louise Fleming ⁹, Hiromasa Inoue¹⁰, Fanny Wai-san Ko ¹¹, Jerry A. Krishnan¹², Mark L. Levy ¹³, Jiangtao Lin¹⁴, Søren E. Pedersen¹⁵, Aziz Sheikh¹⁶, Arzu Yorgancioglu¹⁷ and Louis-Philippe Boulet¹⁸

@ERSpublications

GINA no longer recommends treating adults/adolescents with asthma with short-acting bronchodilators alone. Instead, they should receive symptom-driven (in mild asthma) or a daily corticosteroid-containing inhaler, to reduce risk of severe exacerbations. http://bit.ly/310LLzE

Cite this article as: Reddel HK, FitzGerald JM, Bateman ED, *et al.* GINA 2019: a fundamental change in asthma management. *Eur Respir J* 2019; 53: 1901046 [https://doi.org/10.1183/13993003.01046-2019].

GINA 2019 – landmark changes in asthma management



- For safety, GINA no longer recommends SABA-only treatment for Step 1 in adults and adolescents
 - This decision was based on evidence that SABA-only treatment increases the risk of severe exacerbations, and that adding any ICS significantly reduces the risk
- GINA now recommends that all adults and adolescents with asthma should receive ICS-containing controller treatment, to reduce the risk of serious exacerbations
 - The ICS can be delivered by regular daily treatment or, in mild asthma, by as-needed low dose ICS-formoterol
- This is a population-level risk reduction strategy
 - Other examples: statins, anti-hypertensives
 - The aim is to reduce the probability of serious adverse outcomes at a population level
 - Individual patients may not necessarily experience (or be aware of) short-term clinical benefit

ICS: inhaled corticosteroids; SABA: short-acting beta₂-agonist



Global Initiative in Asthma (GINA) 2021

- International guideline for asthma
- Advocate for regular or prn ICS/LABA in step 1 patients
- Asthma classifications: mild, moderate, severe
 - They do not distinguish intermittent and persistent
- Formoterol is approved as a rescue inhaler in Europe
 - Is not approved in USA for rescue
- Excellent GINA slide set at: <u>https://ginasthma.org</u>



Abbreviations

- "SMART"
- In quality improvement, aim for goals that are:
 - Specific, Measurable, Attainable, Realistic, Timely
- In asthma care:
 - Single Maintenance And Reliever Therapy



A 23 yo female has asthma for which she uses her albuterol MDI 3-4x/week. According to GINA 2021, which medication(s) is the best to add to her current regimen?

- A. Formoterol bid and for rescue, replacing albuterol
- B. Add montelucast 10 mg daily
- C. Add ipratropium as a rescue agent alongside albuterol
- D. Budesonide/formoterol bid and for rescue, replacing albuterol



Correct answer is D

- This patient is already on a SABA and not controlled.
- Preferred medication is ICS/formoterol bid and rescue (SMART)
- If continuing SABA, an ICS should be added.
- A LTRA/montelucast is usually added when ICS/LABA is not controlling asthma
- A LABA by itself would never be indicated poor outcomes with LABA alone or LABA/SABA rescue

Adults & adolescents 12+ years

Personalized asthma management

Assess, Adjust, Review for individual patient needs

Confirmation of diagnosis if necessary Symptom control & modifiable risk factors (including lung function) Comorbidities Inhaler technique & adherence Patient preferences and goals



Treatment of modifiable risk factors and comorbidities Non-pharmacological strategies Asthma medications (adjust down/up/between tracks) Education & skills training

CONTROLLER and PREFERRED RELIEVER

(Track 1). Using ICS-formoterol as reliever reduces the risk of exacerbations compared with using a SABA reliever

STEPS 1 – 2 As-needed low dose ICS-formoterol	STEP 3 Low dose maintenance ICS-formoterol	STEP 4 Medium dose maintenance ICS-formoterol	Add-on LAMA Refer for phenotypic assessment ± anti-IgE, anti-IL5/5R, anti-IL4R Consider high dose ICS-formoterol
RELIEVER:	As-needed low-dose ICS	S-formoterol	

ASSA

ADJUST

REVIEN

Symptoms Exacerbations Side-effects

Lung function

Patient satisfaction

CONTROLLER and ALTERNATIVE RELIEVER (Track 2). Before considering a regimen with SABA reliever,	STEP 1 Take ICS whenever SABA taken	STEP 2 Low dose maintenance ICS	STEP 3 Low dose maintenance ICS-LABA	STEP 4 Medium/high dose maintenance ICS-LABA	Add-on LAMA Refer for phenotypic assessment ± anti-IgE, anti-IL5/5R, anti-IL4R Consider high dose ICS-LABA	
check if the patient is likely to be adherent with daily controller	RELIEVER: As-needed short-acting β2-agonist					
Other controller options for either track		Low dose ICS whenever SABA taken, or daily LTRA, or add HDM SLIT	Medium dose ICS, or add LTRA, or add HDM SLIT	Add LAMA or LTRA or HDM SLIT, or switch to high dose ICS	Add azithromycin (adults) or LTRA; add low dose OCS but consider side-effects	

GINA 2021, Box 3-5A

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STEP 5

STEP 5

Low, medium and high ICS doses: adults/adolescents



Adults and adolescents (12 years and older)

Inhaled corticosteroid	Total daily ICS dose (mcg) – see notes above Low Medium High		
Beclometasone dipropionate (pMDI, standard particle, HFA)	200-500	>500-1000	>1000
Beclometasone dipropionate (DPI or pMDI, extrafine particle, HFA)	100–200	>200-400	>400
Budesonide (DPI, or pMDI, standard particle, HFA)	200–400	>400-800	>800
Ciclesonide (pMDI, extrafine particle, HFA)	80–160	>160-320	>320
Fluticasone furoate (DPI)	100 20		200
Fluticasone propionate (DPI, or pMDI, standard particle, HFA)	100–250	>250-500	>500
Mometasone furoate (DPI)	Depends on DPI device – see product information		
Mometasone furoate (pMDI, standard particle, HFA)	200-400 >400		>400

This is NOT a table of equivalence. These are suggested total daily doses for the 'low', 'medium' and 'high' dose treatment options with different ICS.

DPI: dry powder inhaler; HFA: hydrofluoroalkane propellant; ICS: inhaled corticosteroid; pMDI: pressurized metered dose inhaler; * see product information

GINA 2021, Box 3-6A

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Combination ICS/LABA

- 1. Fluticasone/vilanterol DPI
 - A. 100mcg/25mcg, 200/25
 - B. 1 puff qday
- 2. Fluticasone/salmeterol DPI and MDI
 - A. 100-250-500mcg/50mcg DPI
 - B. 45-115-230mcg/21mcg MDI HFA
 - C. 2 puffs 2x/day, dose depends on severity
- 3. Budesonide/<u>formoterol</u> MDI
 - A. 80-160mcg/4.5mcg
 - B. 2 puffs 2x/day, dose depends on severity
- 4. Mometasone/<u>formoterol</u> MDI
 - A. 50-100-200mcg/5mcg
 - B. 2 puffs 2x/day, dose depends on severity



Other asthma control medications

- 1. Leukotriene Receptor Antagonists (LTRAs)
 - A. Montelukast (Singulair) 10 mg qHS (\$10-15 USD/mo)
 - B. Zafirlukast (Accolate) 40 mg daily (20mg 2x/day, \$30-50 USD/mo)
- 2. 5-Lipoxygenase Inhibitor
 - A. Zileuton 600 mg ER bid (>\$1000/mo USD... !!!)



Other asthma control medications

- 1. Immunomodulator Omalizumab (Anti-IgE)
 - A. 150-375 mg subcut. INJ q2-4 weeks, \$2000 USD/mo
 - B. Other biologic therapies
- 2. Mast cell stabilizer Cromolyn nebulizer
 - A. 1 neb 4x/day, \$160-250 USD/mo
- 3. Methylxanthine Theophylline
 - A. 300-600mg/day, divided daily to bid, \$50-60 USD/mo
 - B. Multiple toxicities



Every patient needs an asthma action plan



Peak flow meters

These two enable setting the three zones



Creative commons license at: <u>http://commons.wikimedia.org/wiki/File:Two_Peak_Flow_Meters.jpg</u>

Asthma Action Plan EMIRATES FAMILY MEDICINE SOCIETY For: Doctor: Date: لمعبة الإمارات لطب الأسرة Doctor's Phone Number Hospital/Emergency Department Phone Number Doing Well Take these long-term control medicines each day (include an anti-inflammatory). Medicine How much to take When to take It No cough, wheeze, chest tightness, or shortness of breath during the day or night Can do usual activities And, if a peak flow meter is used, Peak flow: more than _____ (80 percent or more of my best peak flow) My best peak flow is: _ Before exercise 2 or 4 puffs_ 5 minutes before exercise Asthma Is Getting Worse Add: guick-relief medicine—and keep taking your GREEN ZONE medicine. 2 or 4 puffs, every 20 minutes for up to 1 hour Cough, wheeze, chest tightness, or (short-acting beta₂-agonist) Nebulzer, once shortness of breath, or Waking at night due to asthma, or If your symptoms (and peak flow, if used) return to GREEN ZONE after 1 hour of above treatment: Can do some, but not all, usual activities Continue monitoring to be sure you stay in the green zone. -Or--Or-If your symptoms (and peak flow, if used) do not return to GREEN ZONE after 1 hour of above treatment: Peak flow: _____ to ___ 2 or a 4 puffs or a Nebulizer Take: (50 to 79 percent of my best peak flow) (short-acting bata₂-agonist) Add: ___mg per day For _____(3-10) days (oral staroid) Call the doctor in before/ in within ______ hours after taking the oral steroid. Medical Alert! Take this medicine: Very short of breath, or _____ 4 or = 6 puffs or = Nebulizer (short-acting bata--agonist) Quick-relief medicines have not helped, or Cannot do usual activities, or mg. (oral steroid) Symptoms are same or get worse after 24 hours in Yellow Zone Then call your doctor NOW. Go to the hospital or call an ambulance if: You are still in the red zone after 15 minutes AND -Or-You have not reached your doctor. Peak flow: less than _____ (50 percent of my best peak flow)

DANGER SIGNS Trouble walking and talking due to shortness of breath Lips or fingernalis are blue Take a 4 or a 6 puffs of your quick-relief medicine AND

NOW!

(phone)

Go to the hospital or call for an ambulance.

See the reverse side for things you can do to avoid your asthma triggers.

NHLBI/NIH materials in public domain: http://www.nhlbi.nih.gov/files/docs/public/lung/asthma_actplan.pdf



A 9 yo female presents to the ED with an asthma exacerbation. Her peak flow is only 50% of baseline. She does not improve after two albuterol nebulizers, steroids, supplemental oxygen, or inhaled ipratropium. Which of the following medications may have benefit in this situation?

- A. Administration of a mixture of helium and oxygen (Heliox)
- B. IV magnesium
- C. IV antibiotics
- D. Oral montelucast

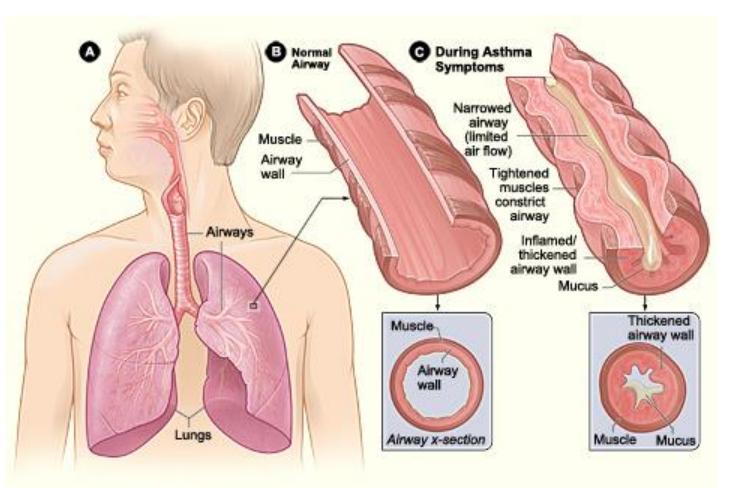


Correct answer is B

- IV magnesium has been demonstrated to help recalcitrant asthma cases when other standard therapies have not.
- This has not been shown to be beneficial in adult patients.
- Oral LTRAs have not place in the treatment of acute asthma in the ED
- IV antibiotics can be of benefit for the asthma patient with an underlying infection, but they will take hours to days to improve care
- Heliox may be beneficial in cases of upper airway obstruction (post intubation edema) but has not shown definitive benefit in patients with asthma



Importance of airway size



Bronchospasm + Inflammation

Poiseuille Equation $R^{2} \frac{1}{r^{4}}$



Asthma diagnosis

- The hallmark is reversible airway obstruction.
- Spirometry shows an increase in the FEV₁ of 12% or more after treatment with a beta agonist.
- Usually, the increase is 15-20% not subtle.
- Additionally, an absolute increase in expired volume of 200 cc or more is required.
- Diagnostic spirometry should be performed when the patient is symptomatic; o/w, the results may be normal.



Pulmonary function testing



How hard is it to do PFT's?

Hold out your hand









PFT interpretation in a Family Medicine Clinic

- **1.** Determine if the FEV₁/FVC is low: obstructive pattern
- Determine if the FVC is low: restrictive pattern (If low, consider <u>DLCO</u> – <u>pulmonology office</u>)
- 3. Grade the severity
- 4. Determine reversibility for obstructive PFT pattern: asthma vs COPD - Perform post beta agonist PFT
- 5. (Consider <u>bronchoprovocation</u> if considering <u>EIB</u> exercise induced bronchospasm – with <u>methacholine</u> in pulmonology office)

Text in black is probably best done in a pulmonology office

Johnson JD, Theurer WM. A Stepwise Approach to the Interpretation of Pulmonary Function Tests. AFP. 2014 Mar 1; 89(5): 359-366



Normal Values

- FVC is the total amount of air a person can exhale, usually measured in six seconds.
 - 80 120% of predicted is a normal value
 - 70 80% demonstrates mild reduction/restriction
 - 50 70% demonstrates moderate reduction
 - <50% demonstrates severe reduction</p>
- FEV₁ is the amount of air a person can exhale in one second.
 - 80 120% of predicted is a normal value



Normal Values

- FEV₁/FVC ratio is the percentage of FVC that can be expired in one second.
 - >80% is normal
 - 50 80% demonstrates mild obstruction
 - 30 50% demonstrates moderate obstruction
 - <30% demonstrates severe obstruction
- FEF₂₅₋₇₅ reflects small airway function
 - >80% is normal
 - 60 80% reflects mild obstruction in the small airways
 - 40 60% reflects moderate obstruction
 - <40% reflects severe obstruction



Perform test



PFT Interpretation

- Three steps in interpretation
 - Is the test valid?
 - Interpret the test
 - Classify severity of disease if present



Validity

- The test is valid is you have good patient effort and the three tests performed are internally consistent.
- You may notice a learning curve in that the latter tests are better performed than the former.
- Make sure that the tests involve maximal effort. You need to be really aggressive in coaching your patient.

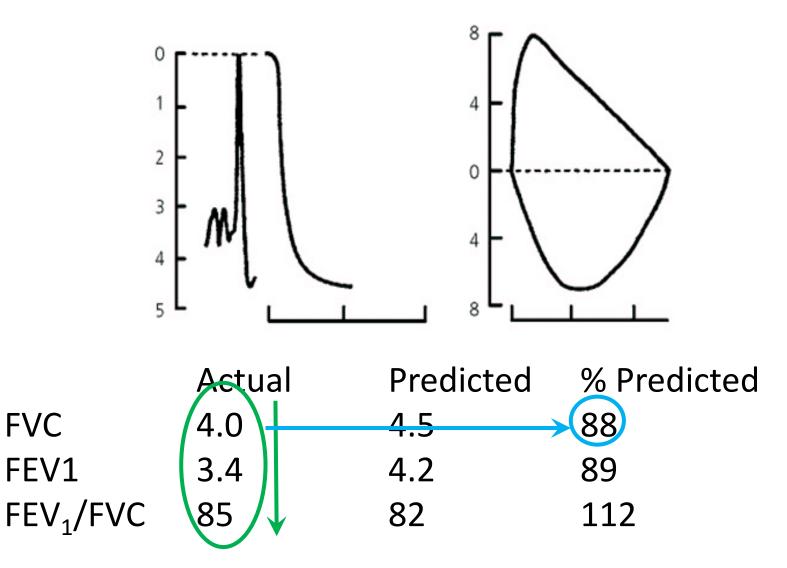




PFT Interpretation

- Assess FVC, FEV₁, and FEV₁/FVC ratio.
- FVC and FEV₁ normal, with a normal FEV₁/FVC ratio:
 - Normal Test ... yeah !!!
- FVC normal or low, FEV_1 low, and a low FEV_1/FVC ratio:
 - Obstructive lung disease
- FVC decreased, FEV₁ low or normal, and a normal to high FEV₁/FVC ratio:
 - Restrictive lung disease

PFT Interpretation





FEV1/FVC?
 Normal = 70-80%*
 (Go Top→Down)

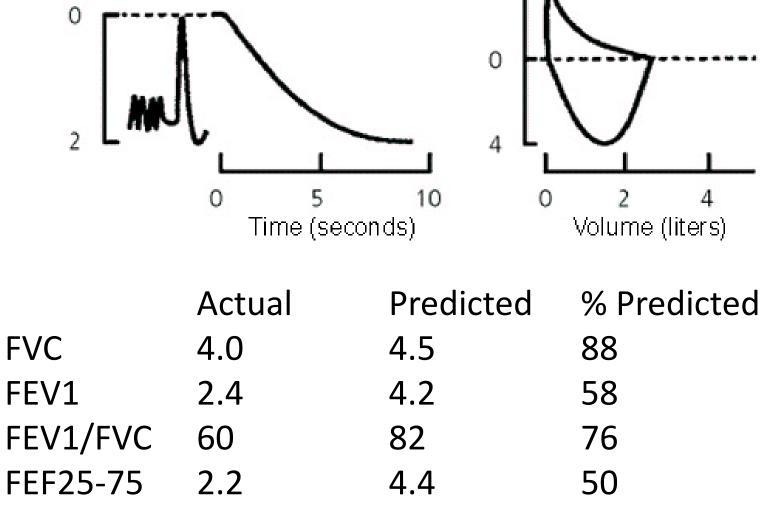
2. FVC?
Normal > 70%*
Percent predicted
(Go Left→Right)

Normal PFT

Case #2 - This PFT is best interpreted as:

emirates family medicine society تشعبة الإمارات لطب الأسرة 4

- A. Restrictive pattern
- B. Obstructive pattern
- C. Small airways disease
- D. Normal





Beta Agonist Challenge

- Perform this when there is a suspicion that the obstructive defect may be reversible –> asthma.
- Give the patient a beta agonist treatment (two puffs of an albuterol MDI or an albuterol nebulizer) and repeat the PFTs several minutes later. If you notice a 12% or more increase in FEV₁, then you have diagnosed reversible airway disease/ asthma.
- In adults > 18 yo, you must additionally have an increase in volume of 200 cc.

Case #3 - This PFT is best interpreted as:

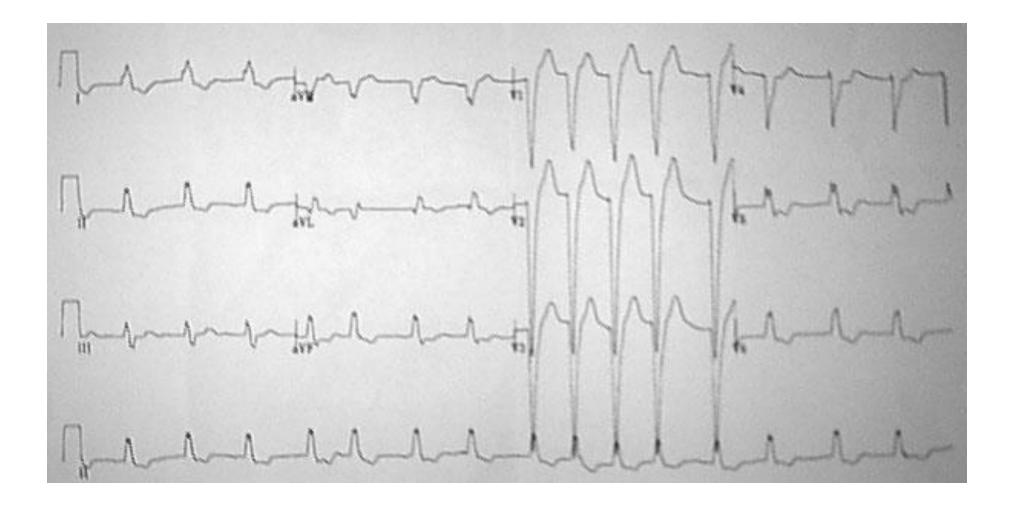


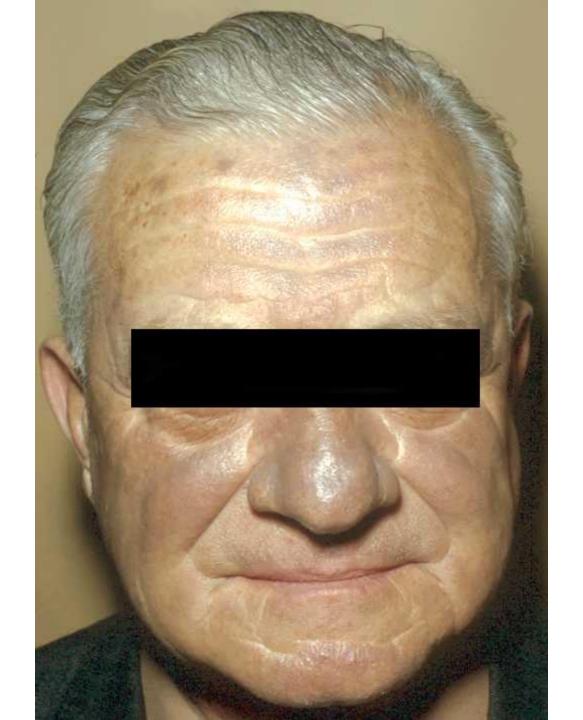
		Actual	Predicted	% Predicted
	FVC	4.0	4.5	88
	FEV1	2.6	4.2	57
	FEV1/FVC	65	82	71
A. Restrictive pattern	FEF25-75	1.7	3.6	47
B. Obstructive pattern – reversible		Beta agonist	treatme	ent
C. Obstructive pattern –		Actual	Predicted	% Predicted
not reversible	FVC	4.1	4.5	91
	FEV1	3.6	4.2	89
D. Normal	FEV1/FVC	90	82	112
	FEF25-75	3.2	3.6	91



	Actual	Predicted	% Predicted
FVC	2.9	4.5	64
FEV1	2.5	4.2	59
FEV1/FVC	89	82	113
FEF25-75	3.7	3.5	102









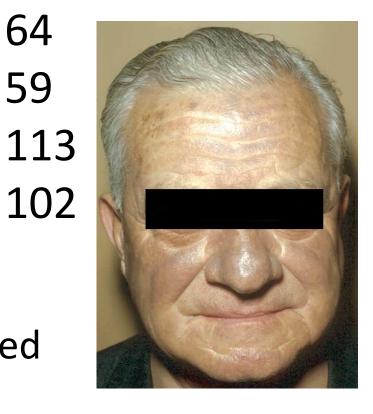


	Actual	Predicted	% F
FVC	2.9	4.5	64
FEV1	2.5	4.2	59
FEV1/FVC	89	82	113
FEF25-75	3.7	3.5	102

 DL_{CO} is decreased when measured

Restrictive lung pattern from Amiodarone

% Predicted





	Actual	Predicted	% Predicted
FVC	3.8	4.5	83
FEV1	2.2	4.2	47
FEV1/FVC	59	82	72
FEF25-75	1.6	3.7	43

Survey says ... COPD





	Actual	Predicted	% Predicted
FVC	4.0	4.5	88
FEV1	3.6	4.2	89
FEV1/FVC	90	82	112
FEF25-75	3.1	3.4	95





	Actual	Predicted	% Predicted
FVC	4.0	4.5	88
FEV1	3.3	4.2	81
FEV1/FVC	83	82	101
FEF25-75	1.7	3.5	48

Small Airways Defect



	Actual	Predicted	% Predicted
FVC	3.5	5.3	68
FEV1	3.1	4.6	68
FEV1/FVC	93	82	117
FEF25-75	3.7	3.3	120

The patient's BMI = 47

Restrictive pattern in obese patient

...



Pulmonary Miscellaneous



Pulmonary Hypertension



Pulmonary <u>arterial</u> hypertension is most likely in which disease state?

- A. Heart Failure
- B. Sickle Cell Disease
- C. COPD/Interstitial lung disease
- D. Scleroderma
- E. Chronic veno-occlusive disease (recurrent PE)

Correct answer is D Pulmonary Hypertension



- Group 1 Pulmonary <u>arterial HTN</u>
- Group 2 Left heart disease (HF and valve)
- Group 3 Lung disease/hypoxia
- Group 4 Chronic thromboembolic disease
- Group 5 Multifactorial

Dunlap B, Weyer G. Pulmonary Hypertension: Diagnosis and Treatment. Am Fam Phys. 2016 Sep 15:94 (6):463-469



Pulmonary HTN

- Pulmonary *arterial* hypertension is a primary disease of the pulmonary arteries.
- The term "Pulmonary hypertension" comprises 5 clinical categories.



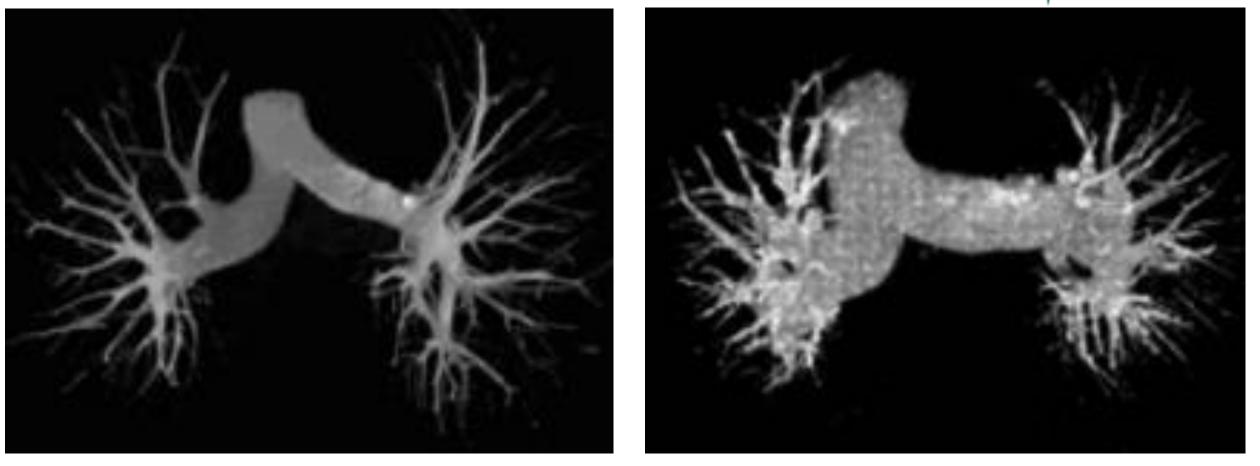
CXR findings in Pulmonary Hypertension





Normal arterial tree on Left





CC license at: https://openi.nlm.nih.gov/detailedresult.php?img=PMC3088846_kjr-12-289-g001&query=pulmonary+hypertension&it=xg&req=4&npos=9

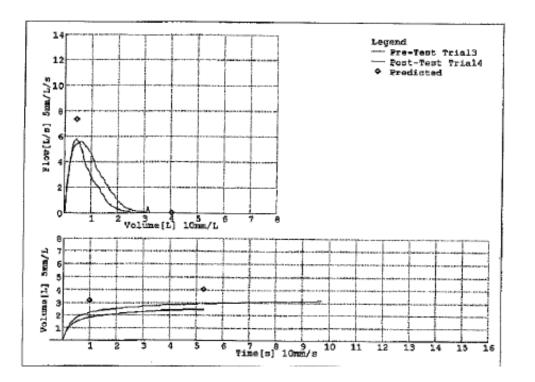


Diagnostic tests

- PFTs to confirm COPD or ILD/restrictive lung disease
- V/Q scan to detect mismatching of perfusion: normal V/Q excludes PH with sensitivity 96% and specificity 90%
 - Follow up with CT-angiogram or PA angiogram

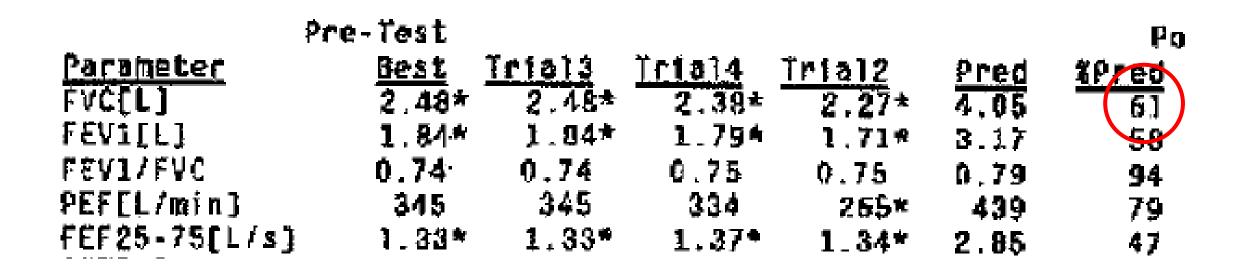


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ae 1900	Li, ba,	BMI 40.1		Predicte	d Ref	NH-	ANES II	1		
Gender	FEMALE			Yalue Se	lect	BE	ST VALU	Ē		
Ethnic	CAUCASEAN			Tech ID				-		
Snoker	#D			Automate	d QC	09	1			
Asthma	YES			BTPS (IN	/EX)		/ 1.03	2		
Test Results	Your FEV1	is 582 Pre	dicted							
Pre	- Test				Post	Test				
Parameter	Best Irial	3 Icial4	Trial2	Pred	\$Pred		Trial4	Trial2	Tria 76	Ch
FVC[L]	2.48+ 2.4		2.27*	4.05	61	3.19*	3.19*	2.94*	2.57*	29
FEVILL]	1.84*).8	4* 1.79*	1.71*	3.17	59	2.23+	2.23*	2.07*	1.81+	ži
FEV1/FVC	0.74 0.7		0.75	0.79	94	0.70	8.70	0.71	0.58*	
PEF[L/min]	345 34	5 334	255*	439	79	333	333	320*	278*	-3
FEF25-75[L/s]	1.33* 1.3	3* 1.37*	1.34*	2.85	47	1.25*	1.25*		0.89*	- ž
FET[s]	5.29 5.2		5.59			9.74	9.74	7.07	9.39	-
 Indicates Selo 	w LLN or Sig	nificant Po	st Change	e						
Pre-Test	FEV1 Var=0.	041. 2.42:	FVC	Vece0 1	0L 3.9%;		Section	Quality		
Post-Test	FEV1 Var=0.				5L 7.91;			Quality		
Interpretation	Restriction		further	examinat	ions race	and the second sec	4044100	0001103		
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PFT showing restrictive pattern





What Pulmonary Artery pressure on echocardiography is suggestive of pulmonary hypertension?

- A. > 20 mm Hg
- B. > 35 mm Hg
- C. > 45 mm Hg
- D. > 60 mm Hg



Echocardiography

- Pulmonary artery pressure > 35 mm Hg
- Frequently an incidental finding in a patient with chronic HF, or mitral/aortic valve pathology

Dunlap B, Weyer G. Pulmonary Hypertension: Diagnosis and Treatment. Am Fam Phys 2016 Sep 15;94(6):463-469



Echo vs Right Heart Cath

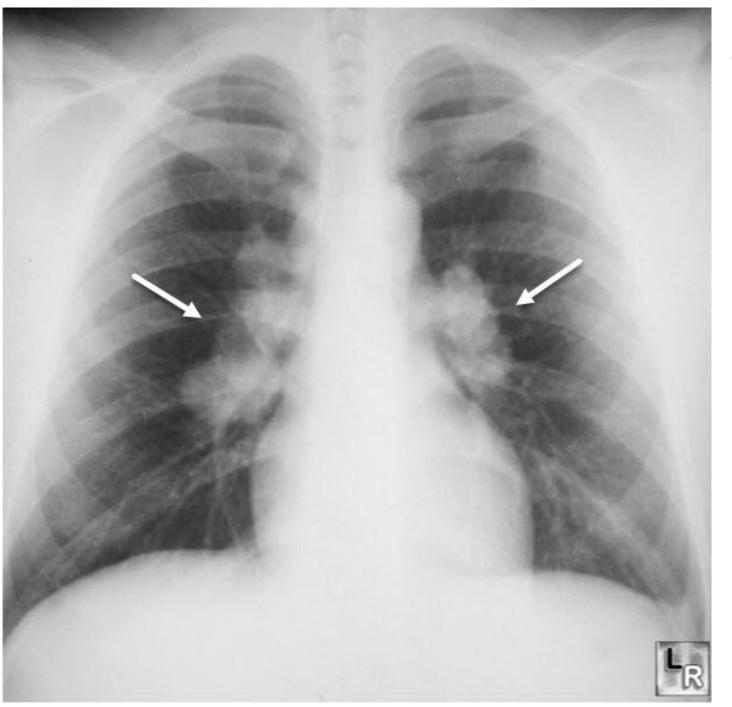
- Meta-analysis conducted for diagnosis of pulm HTN
- Sensitivity of 83%
- Specificity of 72%
- (+) likelihood ratio = 3.0
- (-) likelihood ratio = 0.24

Janda S, Shahidi N, Gin K, *et al.* Diagnostic accuracy of echocardiography for pulmonary hypertension: a systematic review and meta-analysis *Heart* 2011;**97:**612-622.



A 27 yo female presents with a painful lesion on her anterior shins for 2 weeks. Her chest xray shows mediastinal lymphadenopathy. What is the most likely diagnosis?

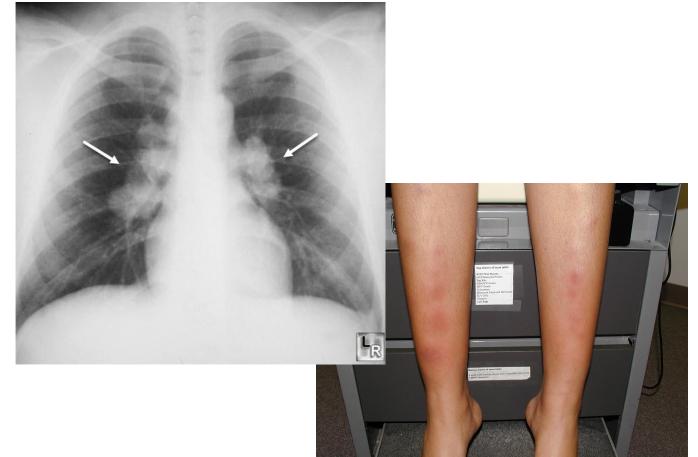






A 27 yo female presents with a painful lesion of her anterior shins for 2 weeks. Her chest xray shows mediastinal lymphadenopathy. What is the most likely diagnosis?

- A. Pulmonary TB
- B. Lymphoma
- C. Erythema multiforme
- D. Sarcoidosis





Correct answer is D

- Sarcoidosis has multiple presentations
- In Ddx of erythema nodosum a painful panniculitis commonly on the anterior shins.
- In the lungs, staging is 0-4:
- Noncaseating granulomas

STAGE	No abnormalities	5%–10%
0 STAGE 1	Lymphadenopathy (fig. A)	50%
STAGE 2	Lymphadenopathy + pulmonary infiltration (fig. B)	25%–30%
STAGE 3	Pulmonary infiltration (fig. C)	10%–12%
STAGE 4	Fibrosis	5% (up to 25% during the course of the disease)

In a 35 yo male patient who smokes regularly, 2 ppd for the past 10 years, which routine immunization is recommended by the CDC to prevent disease?

- A. Pneumococcal PPSV23
- B. Shingrix (Zoster)
- C. Meningococcal vaccine
- D. Hepatitis C vaccine



Correct answer is A

- The CDC recommends the PPSV23 vaccine for all patients who smoke.
- Shingrix to prevent zoster reactivation and post herpetic neuralgia in recommended at age 50
- Meningococcal vaccination is recommended for immunocompromised patients and those with functional asplenia, in general.
- There is not yet a vaccine for hepatitis C.



Of note, a new pneumococcal vaccine has been released – PCV15 and PCV 20.

Should not be on a board exam for at least 1-2 years



TB screening interpretation - CDC

- For low risk patients, the PPD must be > 15 mm in diameter of induration for a (+) test.
- > 10 mm is (+) for patients with high endemic states, IV drug users, high congregant settings (nursing homes), diabetes, severe CKD, children < 5 yo or children exposed to high risk adults
- > 5 mm is (+) for patients living with HIV, close contact of patient with TB, organ transplant recipients, and immunocompromised patients or patients prednisone 15 mg/day or more



A healthy 28 yo patient is screened for TB with the PPD test and it is (+). The CXR is normal and the patient is asymptomatic. What is the best therapy to decrease the reactivation of tuberculosis in this patient?

- A. Rifampin 300 mg daily for 2 months
- B. Isoniazid 300 mg daily for 9 months
- C. Streptomycin + pyrazinamide x 12 weeks
- D. INH, pyrazinamide, rifampin, and ethambutol x 12 weeks



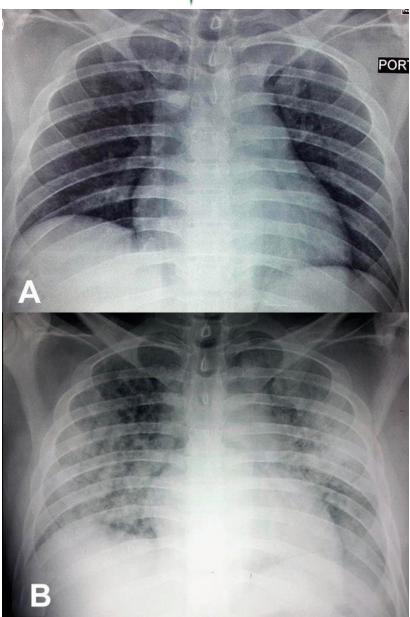
Correct answer is B

- In a healthy patient, INH is sufficient to prevent TB reactivation.
- Duration of INH therapy is 6-9 months.
- May consider using vitamin B6 to minimize risk of neuropathy from INH – 50 mg po daily
- If using rifampin alone, the dose is 600 mg and the duration is at least 4 months double the dose and duration of that in the given answer
- Four drug therapy is used to initially treat active TB until the resistance pattern is known.

A 46 yo female is in a car accident and sustains a mid femur fracture. She is stabilized, but on hospital day #2 develops progressive shortness of breath. A chest xray demonstrates diffuse fluffy infiltrates bilaterally. What is the most likely diagnosis?

- A. Pneumococcal Pneumonia
- B. Fat Embolism
- C. Acute heart failure
- D. Pulmonary embolism







Correct answer is B

- This is a classic case presentation of fat embolism syndrome.
- Fat embolism usually occurs after a long bone fracture with a significant amount of bone marrow fat entering the blood stream and embolizing to the lungs.
- This must be recognized quickly and treated with active supportive care, to include mechanical ventilation.
- The other choices are all possible but do not fit the history as well.

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A 27 yo female presents with persistent clear rhinorrhea, nasal itching, and sneezing. She does not have any pets. She had asthma as a child but has not used an inhaler in a decade. She has no dyspnea. The exam demonstrates boggy nasal turbinates but is o/w nl. The lungs are clear. What medication is best for treating this condition?

- A. Nasal fluticasone 1 puff each nostril daily to bid
- B. Oral loratadine 10 mg daily
- C. Oral pseudoephedrine 40 mg up to tid prn
- D. Oral montelucast 10 mg daily



Correct answer is A

- This patient has allergic rhinitis AR
- All the options presented are potential treatments for AR
- Nasal steroids have the best outcomes for initial treatment
- No one nasal steroid is superior to others
 - Fluticasone, mometasone, triamcinolone
- Often, patients will benefit from a combination of therapies
- Antibiotics have no place in treating AR, in general



Venous thromboembolism DVT and PE

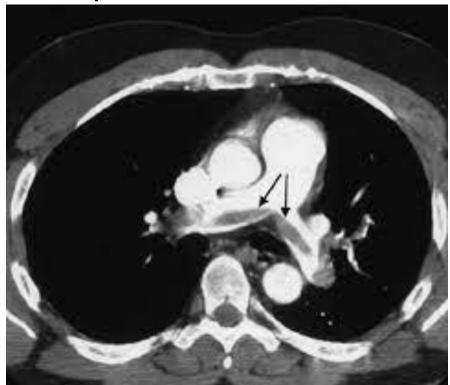
A 57 yo male underwent right inguinal herniorrhaphy 2 weeks ago. He presents with acute dyspnea, RR 32, pulse ox = 86% on room air, pulse 124, and BP 88/54. You suspect PE and this is confirmed with CT. What is the best treatment in this case?

- A. IV heparin to achieve a PTT of 60-100 (therapeutic)
- B. Low molecular weight heparin 1 mg/kg bid
- C. Apixaban 5 mg po bid
- D. Emergent thrombolysis

Correct answer is D

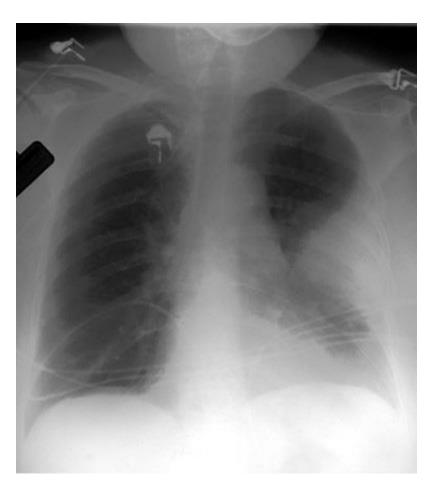


- The first three choices are all acceptable for a patient with a stable pulmonary embolus.
- However, this patient has hypotension. The echocardiogram may show elevation in the pulmonary artery/right heart pressures.
- The EKG may show signs of acute right heart strain S1Q3T3
- These patients should be considered for emergent catheter-based thrombolysis.
- (Picture saddle embolus)





Pulmonary Embolus Hampton's Hump







DVT

- Use the Well's DVT criteria to assess patient risk
- \leq 0, low risk, 5% prevalence of DVT
- 1-2, moderate risk, 17% prevalence
- ≥ 3, high risk, 17-53% prevalence of DVT
- If pt is low-moderate risk, check d-dimer \rightarrow negative, d/c
 - Positive d-dimer \rightarrow U/S
- If pt is high risk, check U/S as first step.



VTE treatment

- Stable patients can be assessed and discharged in the outpatient realm/ED
- Start Factor X inhibitor or thrombin inhibitor
 - Rivaroxaban, apixaban, edoxaban, and others
 - Dabigatran
- Can be considered in stable PE patients as well, although many are still admitted
- Warfarin still an option
 - Requires initiation and transient hypercoagulable state
 - LMWH until INR therapeutic at 2.0 3.0



Pulmonary infections



A 67 yo female with steroid dependent COPD on prednisone 20 mg daily for the past two months develops pneumonia. Which pathogen below should be especially targeted in the choice of initial antibiotics?

- A. Mycoplasma pneumonia
- B. Pseudomonas aeruginosa
- C. Staphylocococcus aureas
- D. Streptococcus pneumonia



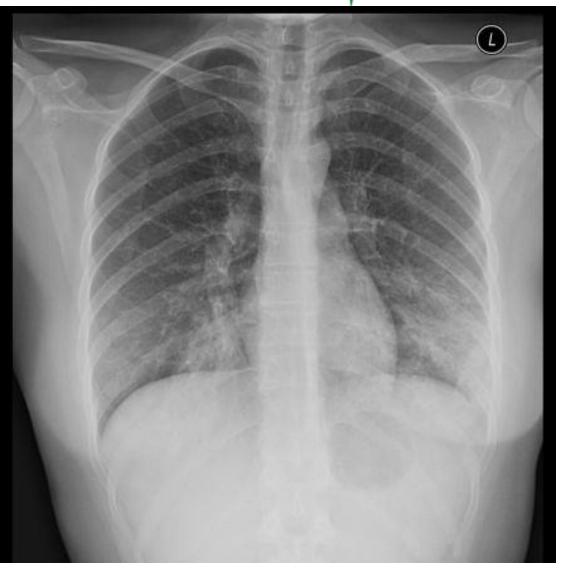
Correct answer is B

- This patient is high risk and should be treated beyond the usual standards for CAP – community acquired pneumonia.
- The selection of initial antibiotics should resemble that for health-care associated pneumonia.
- The most significant/pathologic organism to target is pseudomonas.
- Mycoplasma is commonly associated with "walking pneumonia" CAP
- Streptococcus may have acute onset of rigors/chills and have a lobar pattern.
- Staph aureus may be associated with IV drug use and be associated with lung abscess.

A 39 yo female has had a cough for 2 weeks – o/w feels well. She has no other PMHx. Works in banking. No COVID exposures and is Trivaxxed. She has a temp of 37.7 and has bilateral inspiratory rales in the mid lung fields. CXR as seen. Which of the following bacteria is least likely to cause this pneumonia?

- A. Mycoplasma pneumonia
- B. Legionella pneumonia
- C. Chlamydia pneumonia
- D. Streptococcus pneumonia







Correct answer is D

- This patient has mild pneumonia, or "walking pneumonia."
- Her CURB-65 score is 0, meaning low risk \rightarrow outpatient treatment
- Her PORT/PSI score is 29 (age -10) \rightarrow risk class I, 0.1% mortality
- The first three bacteria may have an insidious onset and can potentially resolve on their own.
- Treatments for CAP specifically target these bacteria and include macrolides (azithromycin, clarithromycin) and tetracycline class meds (doxycycline).

Community-acquired pneumonia Treatment

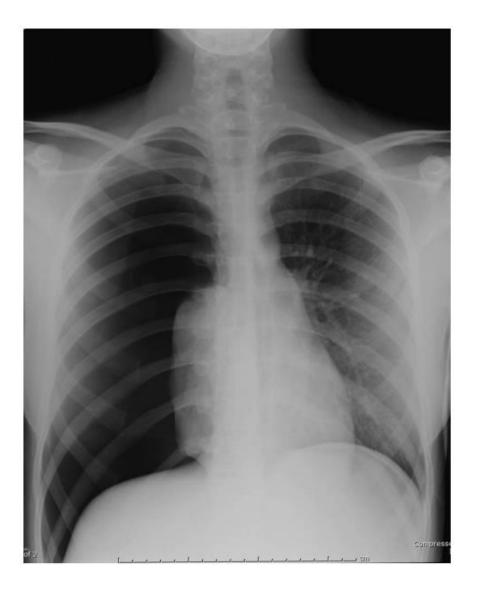


- Oral penicillins are still excellent first line therapies
 - High dose amoxicillin (amox+clavulanate)
 - Some authorities add a macrolide to this (preferred over doxycycline)
- If suspicion for non-pneumococcal infection, consider macrolide/ doxycycline first
- If allergic to PCN and can tolerate cephalosporin:
 - Cefpodoxime preferred (oral third generation)
- If above agents cannot be used, consider respiratory quinolone:
 - Levofloxacin, moxifloxacin, gemifloxacin



Chest radiography

Acute shortness of breath and chest pain





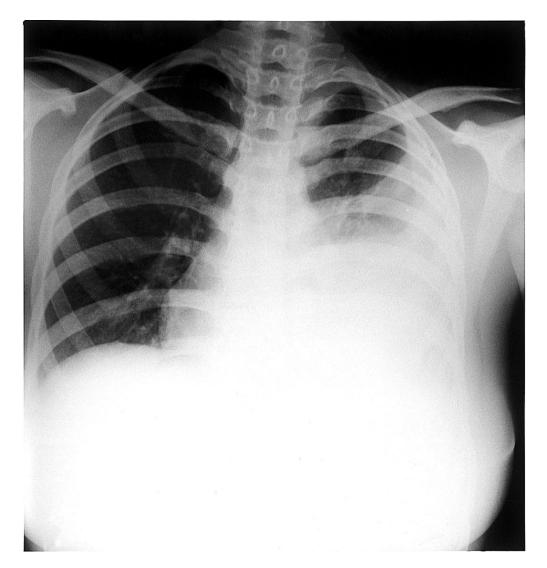
A. Tension pneumothorax

B. Pulmonary embolus

- C. Emphysema
- D. Mass c/w cancer



With no layering of the fluid, the most likely diagnosis is:



A. Pleural effusion

B. Tumor

C. Empyema

D. Cardiomegaly



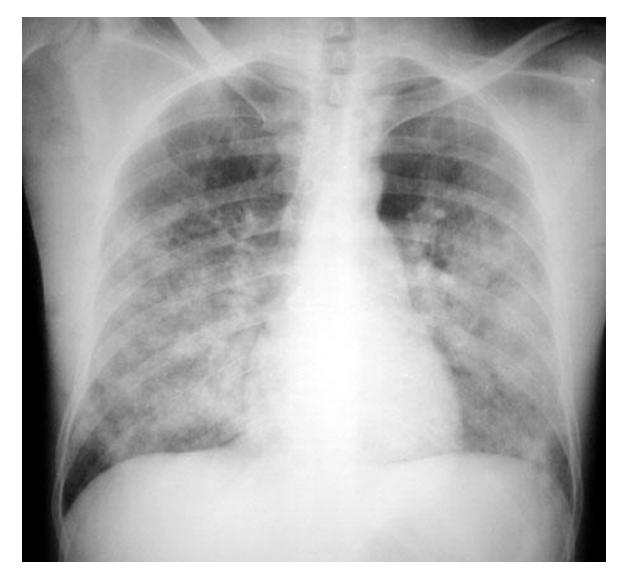
Pulmonary edema



- Enlarged cardiac silhouette
- Cephalization
- No effusion
- Sharp heart borders

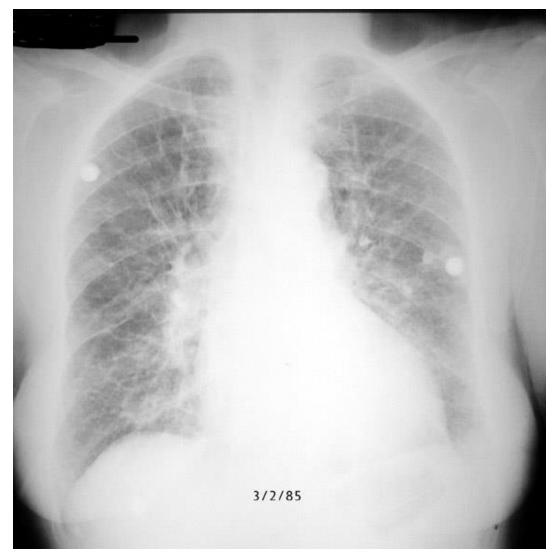


Acute pulmonary edema





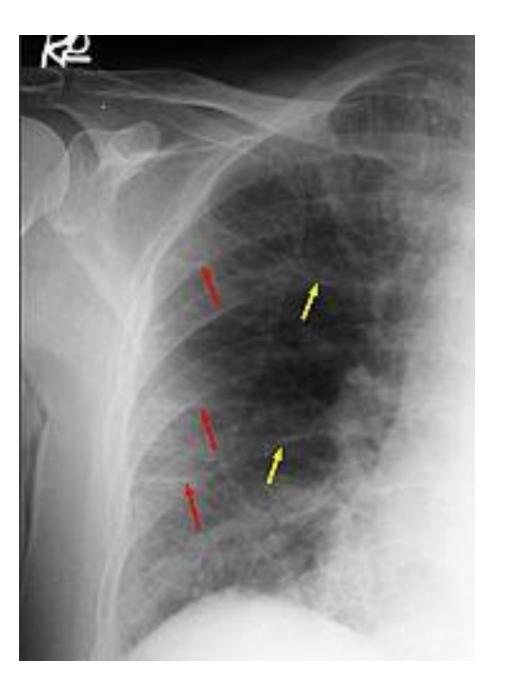
Cephalization





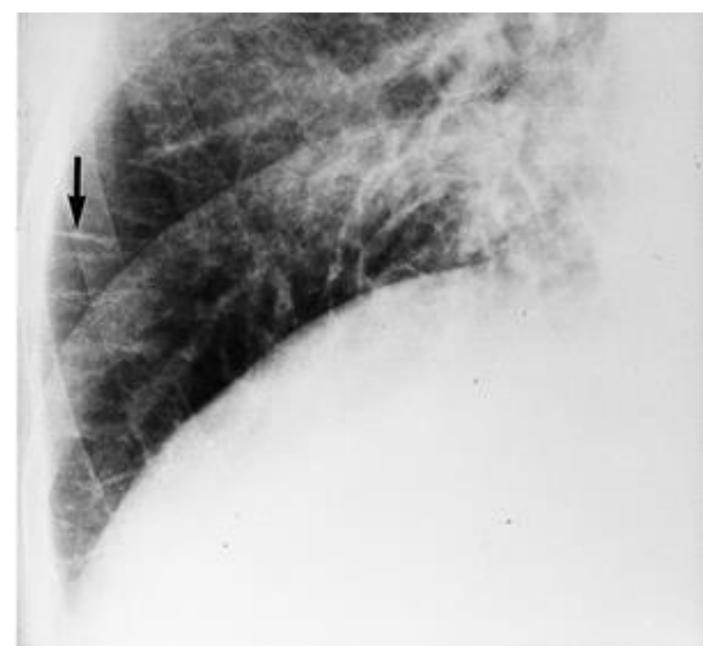
Vascular/hilar fullness



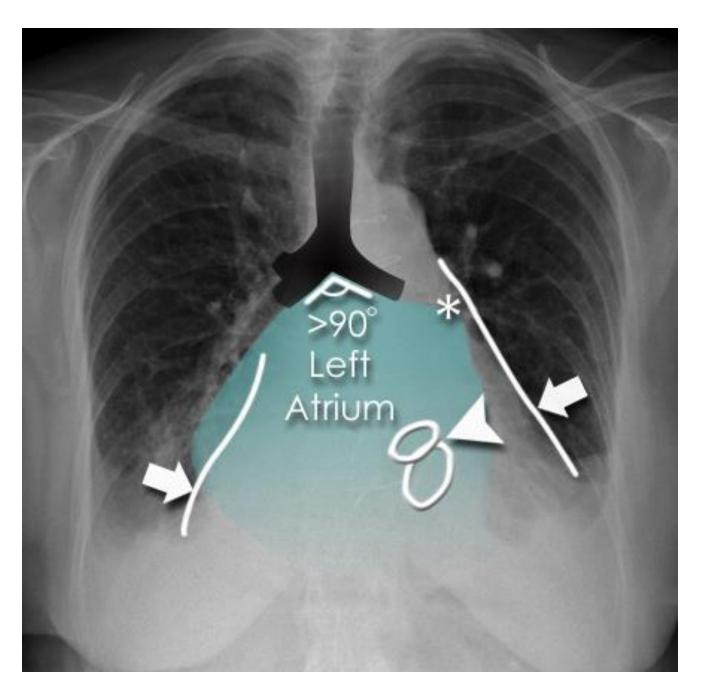




Kerley B lines Interstitial Fluid











Elderly man with hypotension



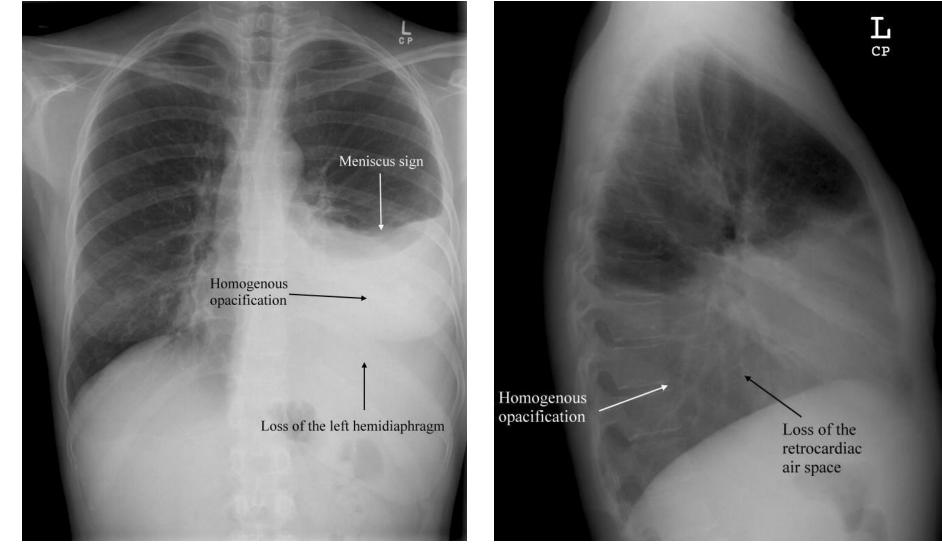
Free air under the diaphragm

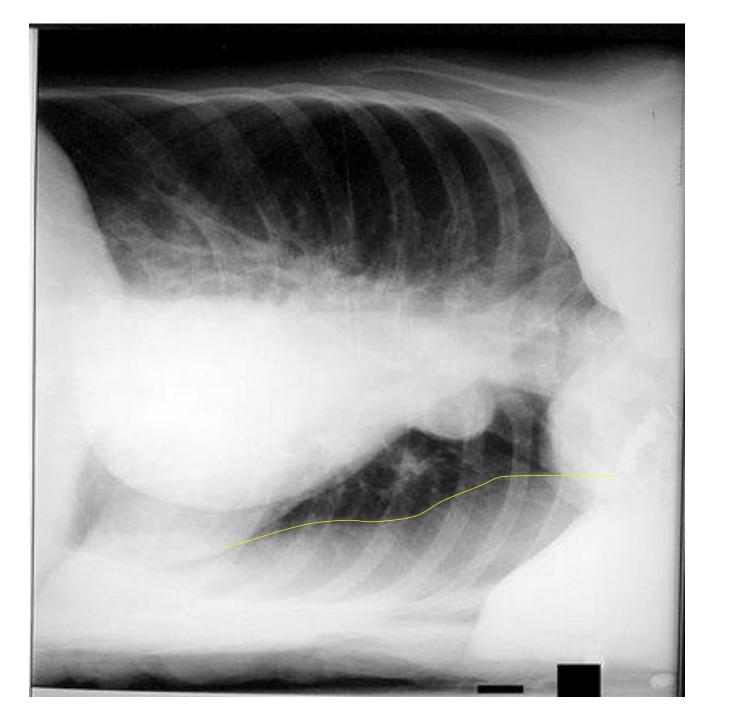






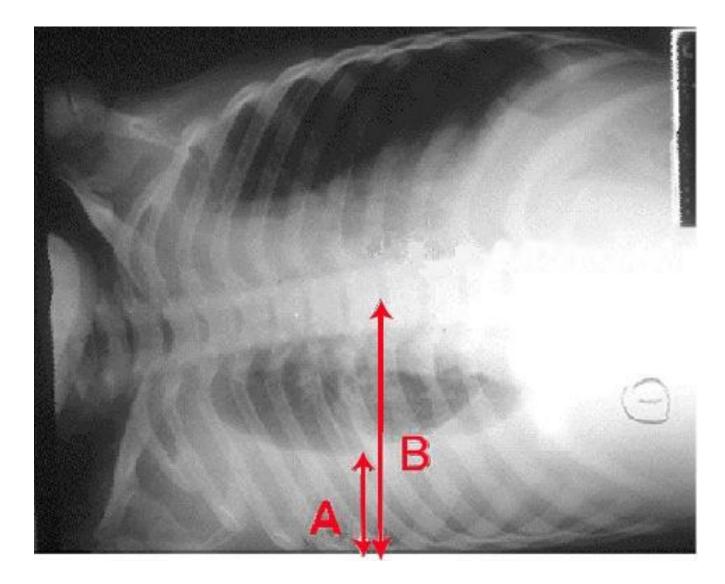
Pleural Effusion











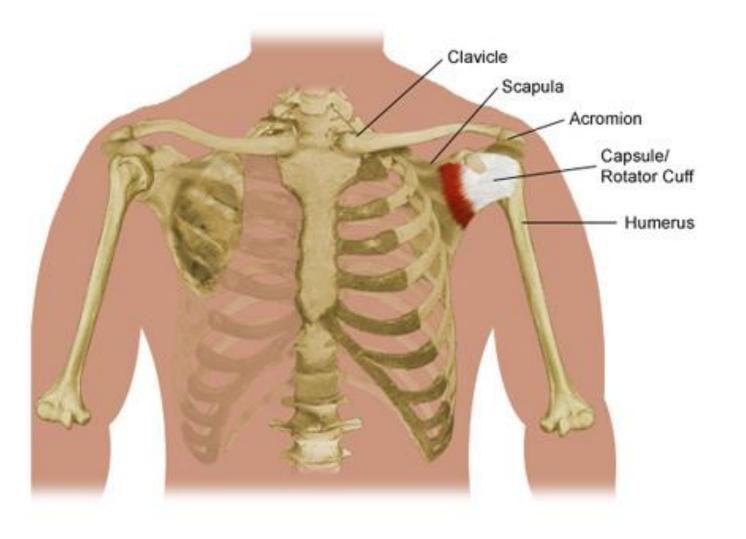


Technique for percussion

- Standard is to percuss with stethoscope or reflex hammer
 - Difficult in patients with increased BMI
- Percuss the ipsilateral clavicle
 - The tap transmits quite well

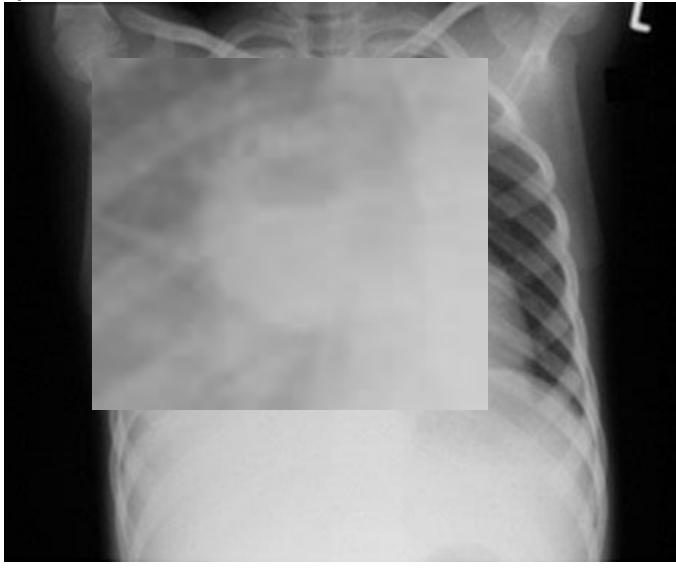


Clavicular Percussion





Pulmonary Abscess



Interstitial Lung Disease



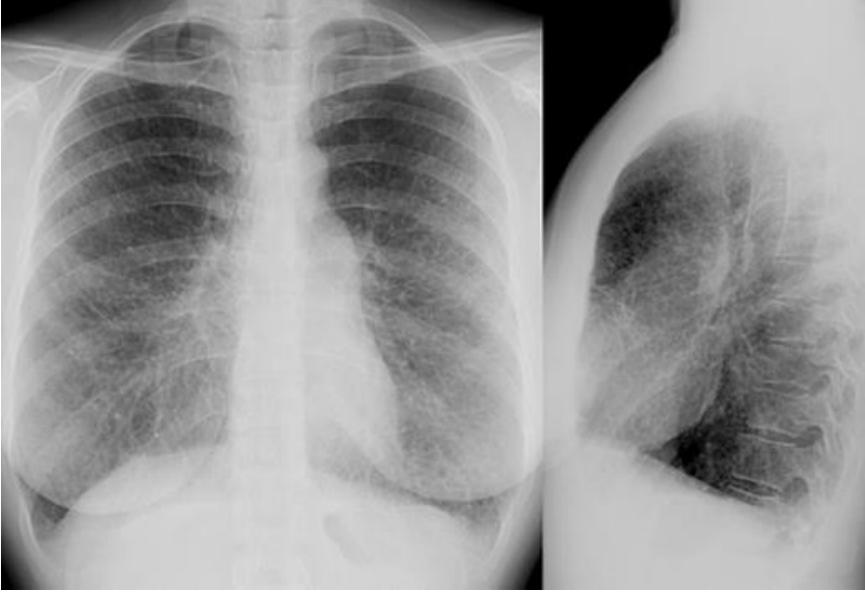


Honeycomb lung













COPD



The gold standard for diagnosing COPD is:

- A. History and physical exam
- B. Chest x-ray
- C. Spirometry
- D. Assay for alpha-1-antitrypsin levels
- E. CT scan of the chest



Steps in evaluation of COPD

- 1. Spirometry to diagnosis/confirm
- 2. Grade airflow limitation
- 3. Assess disease severity
- 4. Assess exacerbation risk

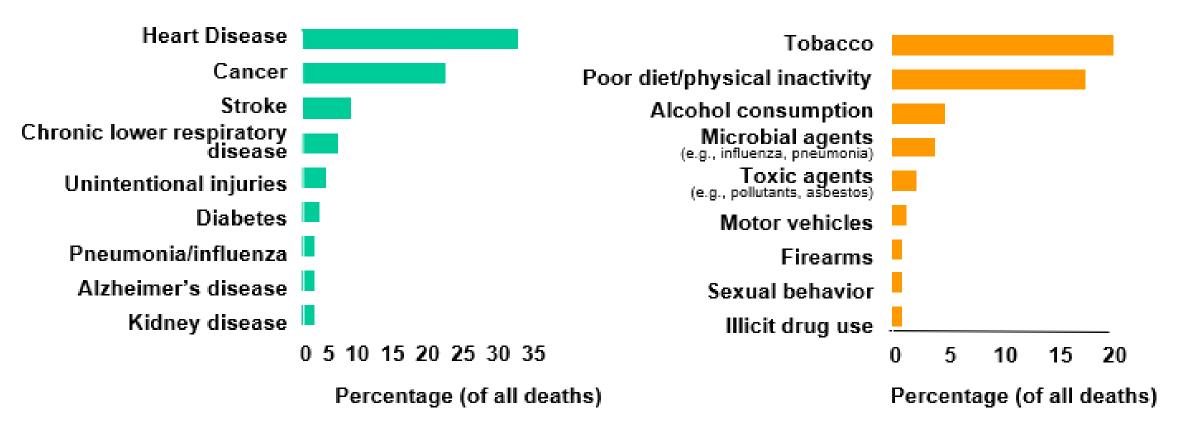


Step 1 - Spirometry

- In patients with dyspnea, chronic cough, or sputum production and a hx of exposure to risk factors, **spirometry is a must to make the diagnosis**.
- Spirometry provides vital information in assessing the severity of disease and to assist with disease management and should be **repeated annually**.
- Spirometry is positive if the post-bronchodilator FEV1/FVC ratio is < 0.70 indicating persistent airflow limitation.
- Spirometry is underutilized potentially leading to misdiagnosis

Leading Causes of Death* United States, 2000

Actual Causes of Death[†] United States, 2000



*Source: Minino AM, Arias E, Kochanek KD, Murphy SL, Smith BL. Deaths: final data for 2000. National Vital Statistics Reports 2002; 50(15):1-20. [†]Source: Mokdad AH, Marks JS, Stroup DF, Gerberding JL. Actual causes of death in the United States, 2000. JAMA. 2004;291 (10): 1238-1246.



DDx of a <u>fixed</u> obstructive pattern

- COPD chronic obstructive pulmonary disease
- Asthma (Can become fixed with a reversible component)
- Alpha-1 antitrypsin deficiency
- Bronchiectasis
- Bronchiolitis obliterans
- Cystic fibrosis

Johnson JD, Theurer WM. A Stepwise Approach to the Interpretation of Pulmonary Function Tests. Am Fam Physician. 2014 Mar 1;89(5):359-366



Step 2 – Grade Airflow Limitation

- COPD severity determined by degree of airflow limitation on spirometry:
 - GOLD 1: mild FEV1 >
 - GOLD 2: moderate
 - GOLD 3: severe
 - GOLD 4: very severe

- FEV1 > 80% predicted
- FEV1 51-80% predicted
- FEV1 31-50% predicted
- FEV1 < 30% predicted

Executive Summary: Global Initiative for Chronic Obstructive Lung Disease, Updated 2021

COPD Assessment Test* (or similar assessment)



- Each statement is rated from 0 to 5 max of 40 points; ≥ 10 is high risk
 - I never cough
 - I have no phlegm in my chest at all
 - My chest does not feel tight at all
 - When I walk up a hill or flight of stairs I am not breathless
 - I am not limited doing any activities at home
 - I am confident leaving my home despite my lung condition
 - I sleep soundly
 - I have lots of energy

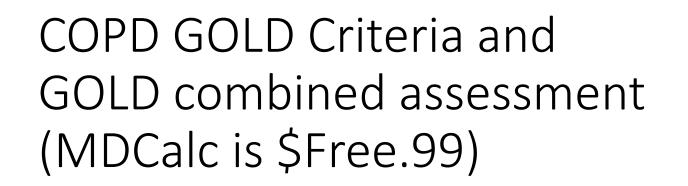
*COPD Assessment Test is a trademark of GlaxoSmithKline

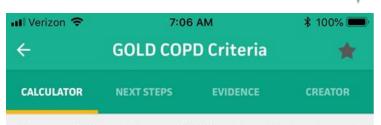




Modified Medical Research Council Dyspnea Scale (mMRC)

- **0 points -** dyspnea only with strenuous exercise
- **1 point** dyspnea when hurrying or walking up a slight hill
- 2 points walks slower than people of same age because of dyspnea or has to stop for breath when walking at own pace
- **3 points** stops for breath after walking 100 yards or after a few minutes
- 4 points too dyspneic to leave house or dyspneic when dressing





Assesses different stages of COPD and provides treatment recommendations.

INSTRUCTIONS

Use in patients >18 years of age with alreadydiagnosed COPD by spirometry (FEV $_1$ /FVC <0.7) with baseline symptoms and lung function. Do not use to diagnose COPD and do not use in patients with acute exacerbation.

When to Use V Pearls/Pitfalls V

lls ✔ Why Use ✔

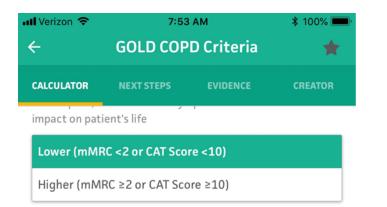
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Symptom burden

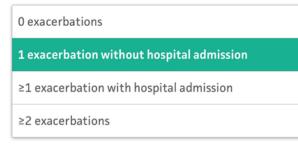
<u>mMRC</u> 2 = Walks slower than people of the same age because of dyspnea or has to stop for breath when walking at own pace; CAT 10 = COPD symptoms have low-medium impact on patient's life

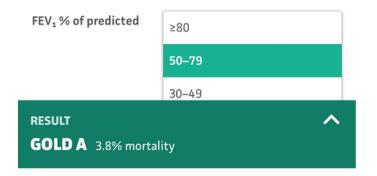
Lower (mMRC <2 or CAT Score <10)

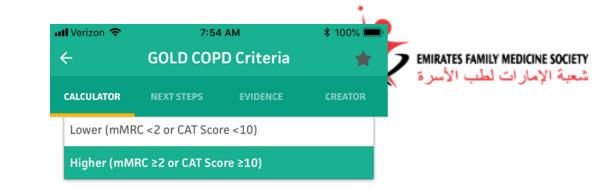
Higher (mMRC ≥ 2 or CAT Score ≥ 10)











Exacerbation history

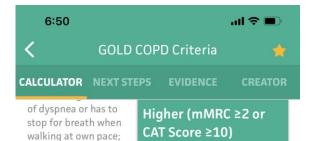
0 exacerbations

1 exacerbation without hospital admission

 \geq 1 exacerbation with hospital admission

≥2 exacerbations





RESULT

GOLD D

20.1%

3-year mortality

Grade 3 airflow obstruction

Exacerbation history

CAT 10 = COPDsymptoms have lowmedium impact on patient's life

1 exacerbation without

0 exacerbations

≥1 exacerbation with hospital admission

≥2 exacerbations

hospital admission

RESULT

>>> **>>**

GOLD D

Grade 3 airflow obstruction

20.1%

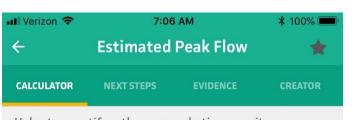
3-year mortality

LAMA + LABA recommended, plus ICS if frequent exacerbations

LAMA + LABA recommended, plus ICS if frequent exacerbations



Helpful apps Peak Flow estimates





Helps to quantify asthma exacerbation severity.

When to Use 🗸	Pearls/Pitfalls 🗸	Why Use 🛩
Sex	Female	Male
Age		51 years
Height		58 in 4

RESULT	~
454 L/min Estimated Peak Flow	



GOLD Groups

- A. Low risk/less symptoms GOLD 1-2 FEV1 > 50%, ≤ 1 exacerbation/yr, CAT < 10 or mMRC 0-1
- B. Low risk/more symptoms GOLD 1-2 FEV1 > 50%, \leq 1 exacerbation/yr, CAT \geq 10 or mMRC \geq 2
- C. High risk/less symptoms GOLD 3-4 FEV1 < 50%, ≥ 2 exacerbations/yr, CAT < 10 or mMRC 0-1
- D. High risk/more symptoms GOLD 3-4 FEV1 < 50%, \ge 2 exacerbations/yr, CAT \ge 10 or mMRC \ge 2



A 57 yo male with COPD currently uses his rescue albuterol/ipratropium inhalers 3-4x/day most days. No recent exacerbations. The addition of which medication would be the next best step?

- A. Prednisone 20 mg po daily
- B. Fluticasone inhaled 220 mcg 2 puffs bid
- C. Tiotropium 2 puffs daily
- D. Montelukast 10 mg daily



Correct answer is C

- This patient is currently on SABA/SAMA
- Adding either a LABA or a LAMA to this patient's care would be the next best step.
- He has not had any recent exacerbations. Thus, no indication for ICS
- Montelucast/LTRA can be used in asthma but are not routinely used in COPD w/o other medical conditions (SAR)



Initial Pharmacologic Management of COPD

- Group A
 - FEV1 > 50%, ≤1 exacerbation/yr, CAT <10 or mMRC 0-1
 - First choice: SABA or SAMA as needed
 - Alternatives:
 - LAMA
 - LABA
 - SABA/SAMA combination
 - Other possibility: theophylline



Management (continued)

- Group B
 - FEV1 > 50%, \leq 1 exacerbation/yr, CAT \geq 10 or mMRC \geq 2
 - First choice: LABA or LAMA
 - Alternative: LABA and LAMA
 - Other possibilities: SABA and/or SAMA, theophylline



Management (continued)

- Group C
 - FEV1 < 50%, \geq 2 exacerbations/yr, CAT < 10 or mMRC 0-1
 - First choice: LAMA, or LABA/ICS
 - Alternatives:
 - LAMA and LABA
 - LAMA and PDE-4 I
 - LABA and PDE-4 I
 - Other possibilities: SABA and/or SAMA, theophylline



Management (continued)

- Group D
 - FEV1 < 50%, \geq 2 exacerbations/yr, CAT \geq 10 or mMRC \geq 2
 - First choice: LABA/ICS and/or LAMA
 - Alternatives:
 - LABA/ICS and LAMA
 - LABA/ICS and PDE-4 I
 - LAMA/LABA
 - LAMA and PDE-4 I
 - Other possibilities: SABA and/or SAMA, theophylline







A 73 yo female has a 60 pk yr history of tobacco abuse. According to the USPSTF (United States Preventive Services Task Force), this patient should be screened with PFTs to assess for COPD.

- A. True
- B. False



Correct answer is B

- Even with a high pk yr history, there is no patient benefit to screen for COPD in an asymptomatic patient.
- Conversely, this patient should be offered lung cancer screening with low dose chest CT.
- USPSTF Recommendations for screening
- http://epss.ahrq.gov/ePSS/GetResults.do?new=true
- Screening for COPD in asymptomatic patients is a Grade D recommendation
 - Not recommended
 - Last updated 2016
- ACP and GOLD do not recommend screening in asymptomatic patients → lack of clinical benefit



Short-Acting Beta Agonists

- Albuterol, metaproterenol, levalbuterol
 - Albuterol/ipratropium combination
- Mechanism of action: relax smooth muscles
- Duration of action: approx. 4-6 hours
- Benefits: decrease dyspnea, improve ability to function
- Side effects: tachycardia, jitteriness, cardiac dysrhythmias in some, increase in tremor, hypokalemia in patient's on kaliuretic drugs
- Misc: Teach proper technique; recommend use with **spacer**
- MDI preferred: avoid oral SABAs if possible
- No decrease in mortality



Long-Acting Beta Agonists

- Salmeterol, formoterol inhalers used twice daily
- Indacaterol, olodaterol inhalers used once daily
- Arformoterol nebulizer solution used twice daily
- Mechanism of action: relax smooth muscles
- Duration of action: approx. 12-24 hours
- Benefits: decrease dyspnea, improve ability to function, decrease hospitalization rates (salmeterol, formoterol), decrease exacerbation rates (salmeterol, formoterol, indacaterol)
- Side effects: similar to SABAs
- No decrease in mortality; increase in mortality in COPD/asthma overlap and in asthma



Short-Acting Muscarinic Antagonists

• Ipratropium

- Mechanism of action: bronchodilation by blocking M2 and M3 receptors
- Duration of action: approx. 6-8 hours
- Benefits: decrease dyspnea, improve ability to function
- Side effects: dry mouth, bitter taste
- Misc: teach proper technique; recommend use with spacer
- No decrease in mortality

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Long-Acting Muscarinic Antagonists

- Tiotropium, umeclidinium, glycopyrronium bromide inhalers used once daily
- Aclidinium inhaler used twice daily
- Mechanism of action: bronchodilation by blocking M2 and M3 receptors
- Duration of action: approx. 12-24 hours
- Benefits: decrease dyspnea, improve ability to function, decrease in exacerbations and hospitalizations
- Side effects: dry mouth, bitter taste
- No decrease in mortality

Inhaled Corticosteroids



- Fluticasone furoate duration of action is 24 hours
- Mechanism of action: uncertain
- Duration of action: 12 hours
- Benefits: improves symptoms and quality of life, reduces exacerbations in patients with an FEV1 < 60% predicted
- Does not modify long-term decline of FEV1 or decrease mortality



Other Agents

- Antibiotics Use for acute exacerbations. Avoid chronic use
- Mucolytics may reduce exacerbations per Cochrane review. No improvement in QOL
- Antitussives avoid use in stable COPD as cough has a protective role
- Opioids effective for dyspnea in patients with severe disease



A 63 yo male has COPD and currently uses tiotropium bid, and albuterol/ipratropium combination for dyspnea. Which blood test would suggest benefit from the addition of an ICS?

- A. Blood eosinophilia > 300
- B. WBC > 15,000 cells
- C. Platelet count > 450K
- D. Creatinine > 1.5 mg/dL



Correct answer is A

- Recent evidence suggests potential benefit from starting ICS when blood eosinophil count is > 300.
- Additionally, for patients having ≥ 1 exacerbations in a year, ICS has benefit
- The other lab values, while abnormal, are not indications to add ICS

COPD Care in 2020 recommendations



- Blood eosinophil count: if > 300, consider ICS for therapy
- For escalating therapy beyond SABA/SAMA:
 - Start with LAMA
 - Then add LABA
 - Add ICS if > 1 exacerbation/year or > 300 Eosinophil count
- For recalcitrant cases (consult pulmonology) :
 - Consider adding roflumilast/ Daliresp, a PDE4I
 - Consider lung reduction surgeries or lung transplant

Labaki WW and Rosenberg SR. Chronic obstructive pulmonary disease. *Ann Intern Med* 2020 Aug 4; 173:ITC17. (<u>https://doi.org/10.7326/AITC202008040</u>)



American Thoracic Society Guidelines April 2020

- For COPD patients with dyspnea or exercise intolerance, use LABA/LAMA rather than monotherapy with either medication.
- For patients on LABA/LAMA with persistent sx and 1 or more COPD exacerbations in the prior year, add ICS.
- For stable patients on triple therapy with LABA/LAMA/ICS who have not had an exacerbation in the prior year, consider deprescribing ICS.
- For patients on optimal therapy with frequent exacerbations, the group advises against maintenance oral steroid therapy

COPD Triple therapy vs Dual therapy



- 8500 patients with at least 1 COPD exacerbation in prior year.
- Four groups
 - Budesonide 160 ug + formoterol + glycopyrrolate (low dose triple therapy)
 - Budesonide 320 ug + formoterol + glycopyrrolate (high dose triple therapy)
 - Budesonide 320 ug + formoterol (ICS/LABA)
 - Formoterol + glycopyrrolate (LABA/LAMA)
- At one year follow, the triple therapy groups had less exacerbations:
 - 1.08 and 1.07 versus 1.42 and 1.24
- All cause mortality was lower with the high dose triple therapy vs LABA/LAMA at 1.3% vs 2.3% (NNT = 100)
- Incidence of PNA higher in triple therapy groups at 3.5 and 4.5% vs 2.3% in the LABA/LAMA group (NNH = 83 and 45 respectively)

ICS and fracture risk



- Meta-analysis of 16 trials with 9143 ICS users and 8370 controls
- Most patients had severe COPD and were men
- Mean duration of studies = 90 weeks (~ 2 years)
- Fracture risk rose significantly with ICS use
 - 180 fractures in ICS users; 141 in controls (OR 1.27, p = 0.04)
- Authors also reviewed 7 controlled observational studies with 69,000 patients similar fx risk (OR 1.21, p<0.0001)
- Use ICS as appropriate be aware of risk of fracture

Loke YK et al. Risk of fractures with inhaled corticosteroids in COPD: Systematic review and meta-analysis of randomised controlled trials and observational studies. *Thorax* 2011 Aug; 66:699



Recent evidence suggests that the use of beta blockers in patients with COPD is not harmful, and perhaps protective.

A. True

B. False



Correct answer is A

- BB use in patients with COPD is neutral to positive on outcomes.
- BB use in asthma can still be harmful in a minority of patients and should only be prescribed when the perceived benefits outweigh harms.

β-blockers in COPD patients #1



- Traditional concern is worsening of COPD control in patients placed on β -blocker.
- Retrospective cohort study with 6000 Scottish patients with COPD.
- 819 patients used β -blockers over a 4.4 year period.
 - 88% of β -blockers were cardioselective
- 1/3 of patients died
- β-blocker use was associated with a 22% lower all cause mortality.
- β-blockers had no adverse effect on lung function

Short PM et al. Effect of β blockers in treatment of chronic obstructive pulmonary disease: A retrospective cohort study. BMJ 2011 May 10; 342:d2549

β-blockers in COPD patients #2



- 2230 COPD patients in the Netherlands on β-blockers followed for 7 years.
- The mean (SD) age of the patients with COPD was 64.8 (11.2) years at the start of the study, and 53% of the patients were male
- The adjusted hazard ratios of β-blocker use for mortality was 0.68 (95% CI, 0.56-0.83).
- The adjusted hazard ratios for exacerbation of COPD was 0.71 (95% CI, 0.60-0.83).
- Patients on β-blockers were 30% less likely to die from any cause or experience a COPD exacerbation

Rutten FH et al. β-blockers may reduce mortality and risk of exacerbations in patients with chronic obstructive pulmonary disease. *Arch Intern Med* 2010 May 24; 170:880.



COPD highlights

- No USPSTF indication to screen asymptomatic patients.
- May start with either LABA or LAMA when patients have moderate-tosevere COPD
- Use ICS to reduce frequency of exacerbations
- Consider pulmonary rehab in any patient with mild COPD or worse
- B-blockers not harmful in COPD patients.

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