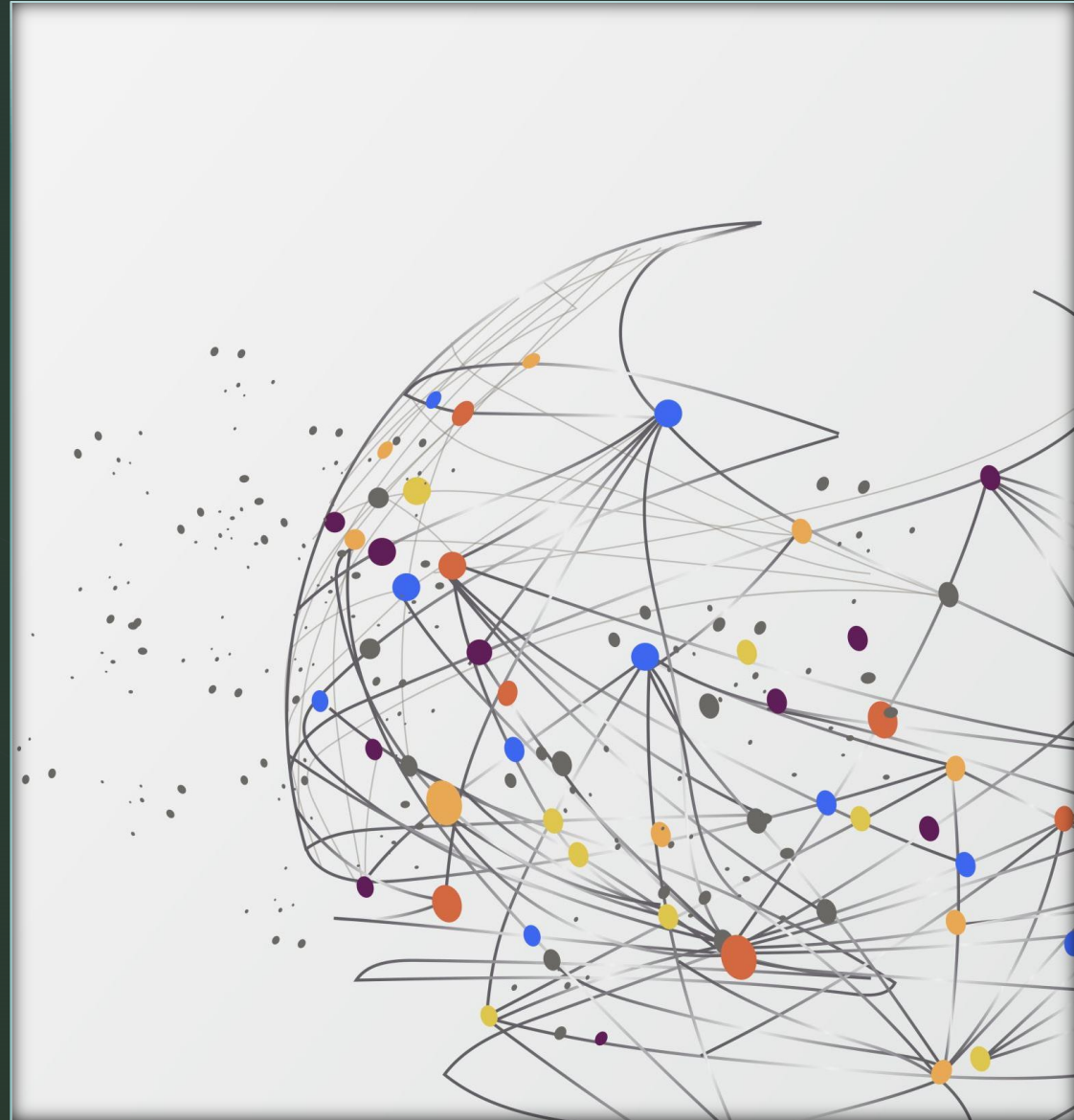


Pulmonary Medicine

Mallory Hatmaker, APRN.CNP

Cleveland Clinic Respiratory Institute



OBJECTIVES

Participants should gain basic knowledge of interpreting spirometry tests to diagnose asthma, COPD and other chronic lung diseases

COPD- participants should be able to diagnose, treat, and prescribe proper pharmacological medications for managing COPD patients both maintenance and exacerbations

Asthma- participants should be able to diagnose, treat and prescribe proper pharmacological medications for managing Asthma patients both maintenance and exacerbations

Participants should gain knowledge on prescribing proper pharmacological treatment for tobacco cessation

Patients should gain knowledge on interstitial lung diseases and updates related to respiratory conditions post COVID19



PULMONARY FUNCTION TESTS

PFTS

SPIROMETERY

LUNG FUNCTION TESTING



INDICATIONS FOR PULMONARY FUNCTION TESTING

- Preoperative evaluation
- Evaluating patients with pulmonary complaints
- Assessing treatment effectiveness
- Monitoring disease progress
- Screening certain patient populations
- Research

Ferguson GT,
Enright PL, Buist
S, Higgins MW.;
Office spirometry
for lung health
assessment in
adults. Chest
2000 1146-1161.

Consensus statement of the
National Lung Health
Education Program (NLHEP)

- -input from ACCP and NHLBI

Recommended spirometry for:

- -smokers >45 years old
- -patients with pulmonary complaints
- -global health assessment

In 2016, USPSTF/AAFP
recommended against
screening with

spirometry for asymptomatic
adults even if they smoke

SPIROMETRY

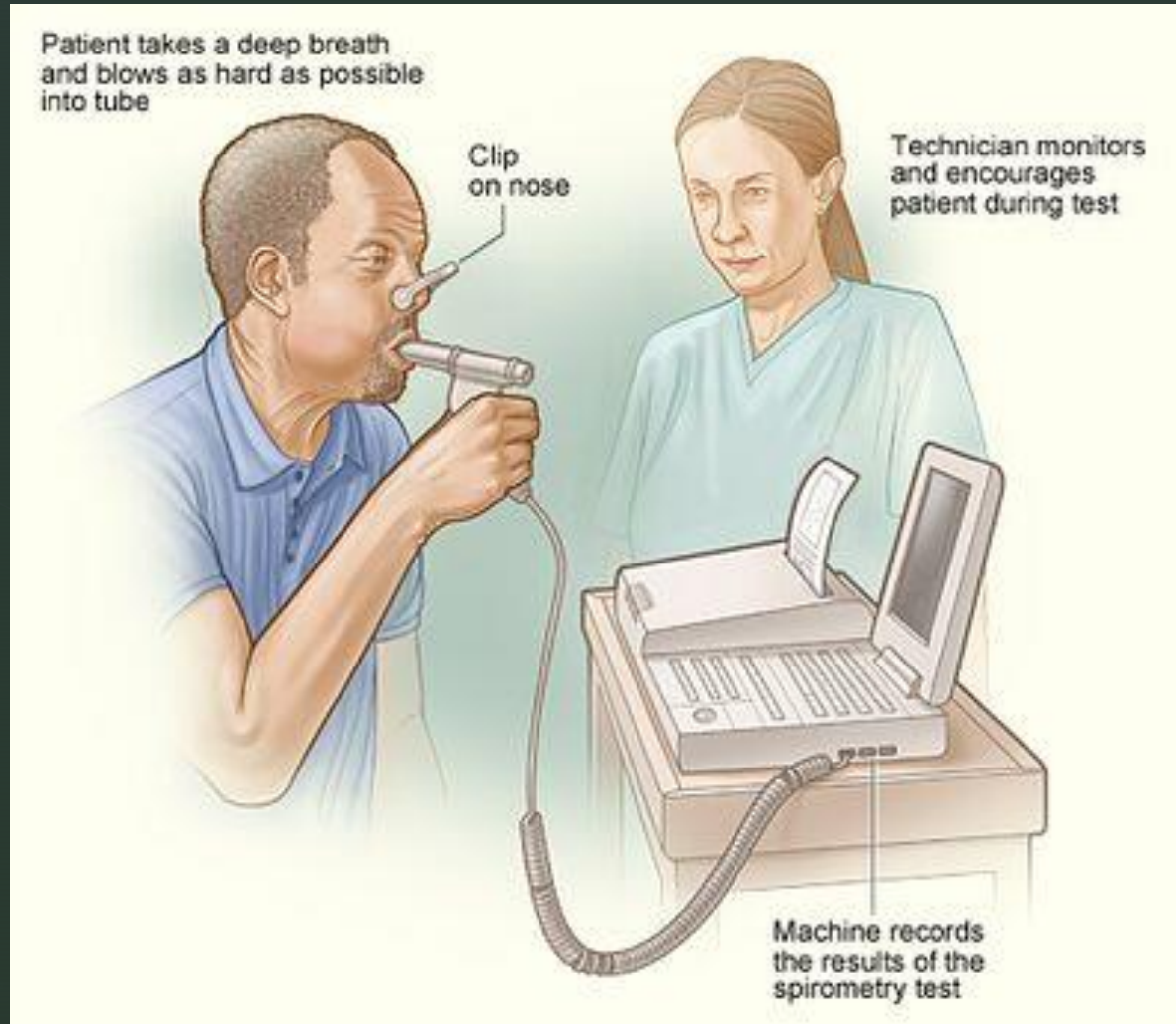
- Most commonly ordered test
- Appropriate for most situations
- Procedure:
 - -a seated patient, usually wearing noseclips, inhales maximally to Total Lung Capacity (TLC) and then exhales maximally
 - -3 attempts completed to meet standards for reliability and reproducibility
 - -inhaled and exhaled flow/volume measured via pneumotach and numerical and graphical printouts generated
 - -bronchodilator may be given by aerosol or MDI if warranted and the testing is repeated to assess the effectiveness of the treatment

Patient takes a deep breath
and blows as hard as possible
into tube

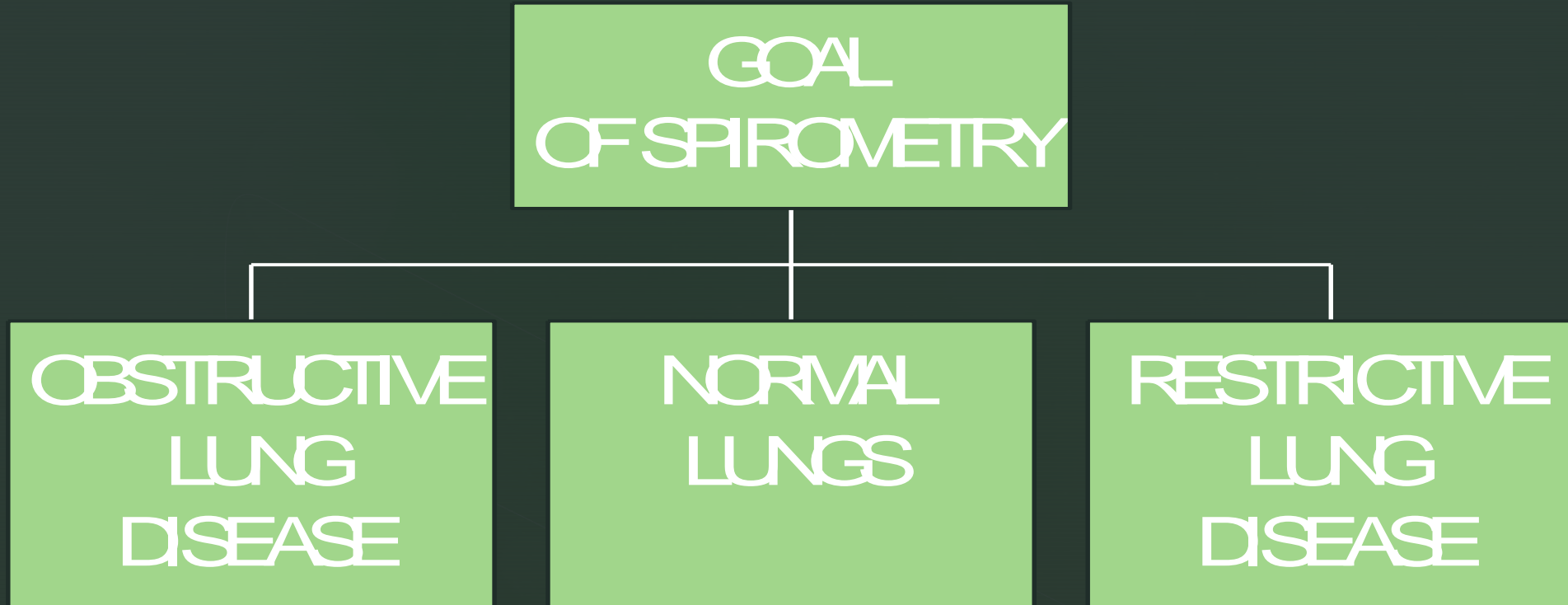
Clip
on nose

Technician monitors
and encourages
patient during test

Machine records
the results of the
spirometry test







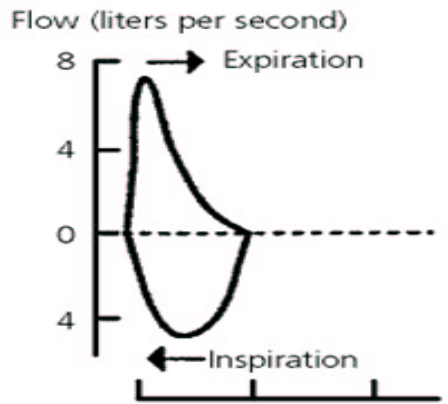
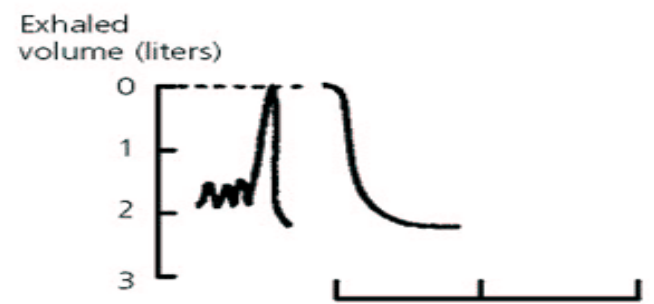
... on the day of testing



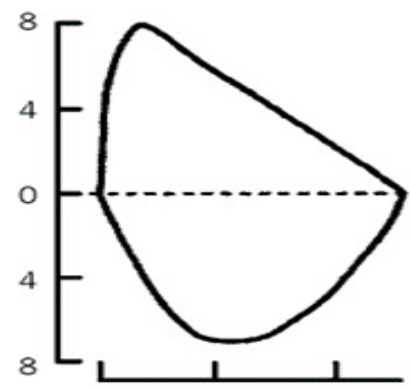
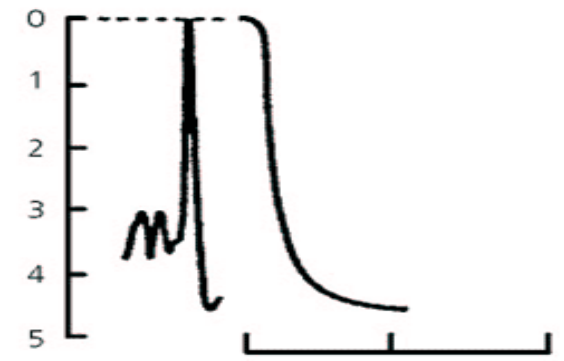
OBSTRUCTIVE LUNG DISEASE

- Decreased airflow
- Normal or increased air volume

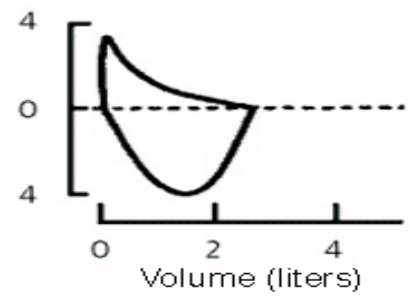
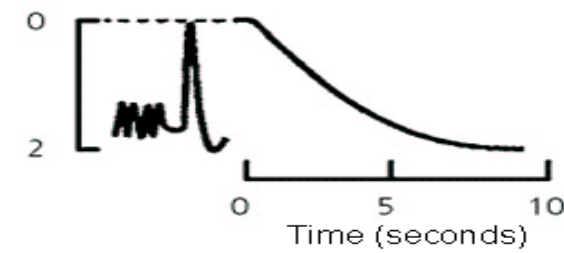
- Chronic Obstructive Pulmonary Disease
 - Emphysema
 - Chronic Bronchitis
 - Asthma
 - Bronchiectasis
 - Cystic fibrosis



A



B

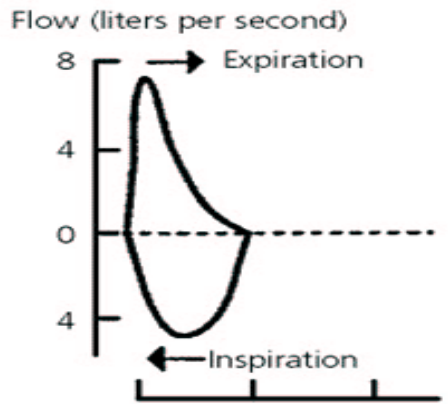
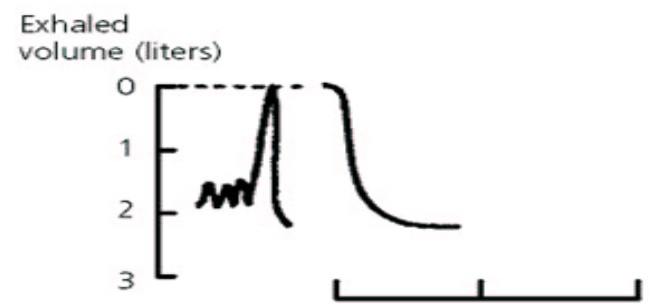


C

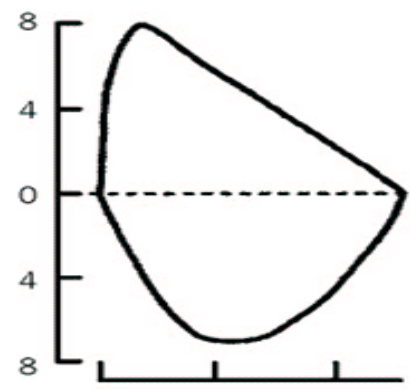
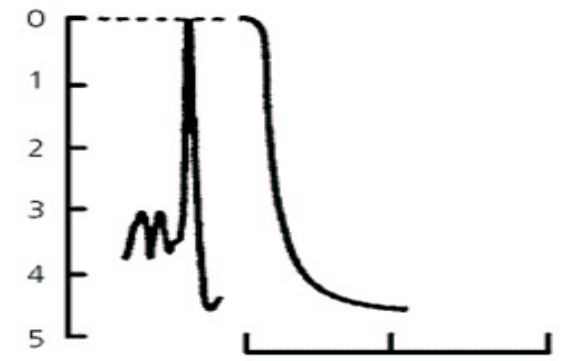


RESTRICTIVE LUNG DISEASE

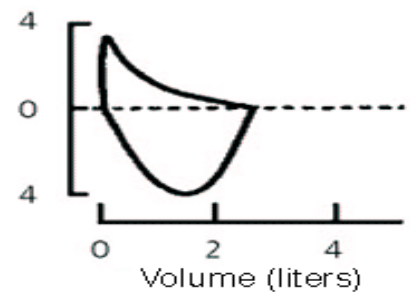
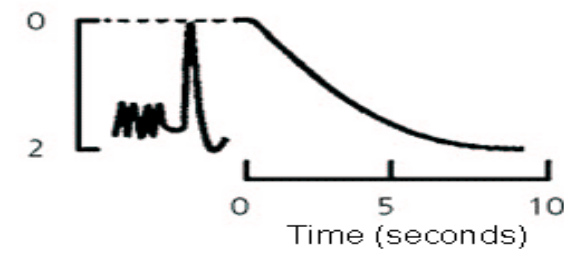
- Normal airflow
- Decreased air volume
- “PAINT” mnemonic



A

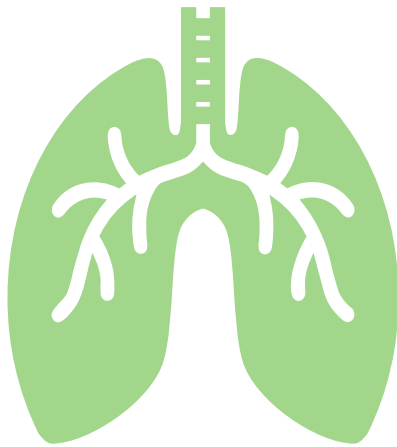


B



C

RESTRICTIVE LUNG DISEASE “PAINT”



- Pleural-pleural effusion/mass/thickening
- Alveolar-pulmonary edema, pneumonia, cancer
- Interstitial-pulmonary fibrosis, sarcoidosis, silicosis pneumoconiosis etc.
- Neuromuscular-myasthenia gravis, ALS, Guillain-Barre, diaphragmatic paralysis
- Thoracic Cage-kyphoscoliosis, obesity

FULL SPIROMETRY YIKES !!

	Pred	LLN	Pre-	%	Post-	%	% chg
FVC	3.34	2.59	1.86	55	1.83	55	-1
FEV 1	2.44	1.81	0.75	31	0.73	30	-3
FEV1%F	73.81	64.13	40.36	55	39.72	54	-2
FEV 3	3.18	2.16	1.14	36	1.13	36	0
FEV3%E	92.05	87.41	61.22	67	61.71	67	1
FEV6			1.47		1.45		-1
PEF	6.63	4.27	2.62	40	2.70	41	3
MEF 50	2.88	1.25	0.19	6	0.22	8	19
FIF 50			3.14		3.17		1
F25/75	1.90	0.61	0.19	10	0.18	10	-3
FE%FIF	?????						
FET			13.75		14.64		6

FULL SPIROMETRY

FOCUS ON THE MAIN 3

	Pred	LLN	Pre-	%	Post-	%	% chg
FVC	3.34	2.59	1.86	55	1.83	55	-1
FEV 1	2.44	1.81	0.75	31	0.73	30	-3
FEV1%F	73.81	64.13	40.36	55	39.72	54	-2
FEV 3	3.18	2.16	1.14	36	1.13	36	0
FEV3%E	92.05	87.41	61.22	67	61.71	67	1
FEV6			1.47		1.45		-1
PEF	6.63	4.27	2.62	40	2.70	41	3
MEF 50	2.88	1.25	0.19	6	0.22	8	19
FIF 50			3.14		3.17		1
F25/75	1.90	0.61	0.19	10	0.18	10	-3
FE%FIF							
FET			13.75		14.64		6

THE MAIN 3

Fev1/FVC Ratio

-The amount of air exhaled in the first second compared to the total amount of air exhaled



Forced Vital Capacity (FVC)

-The maximum volume of air that can be exhaled following a maximal inhalation



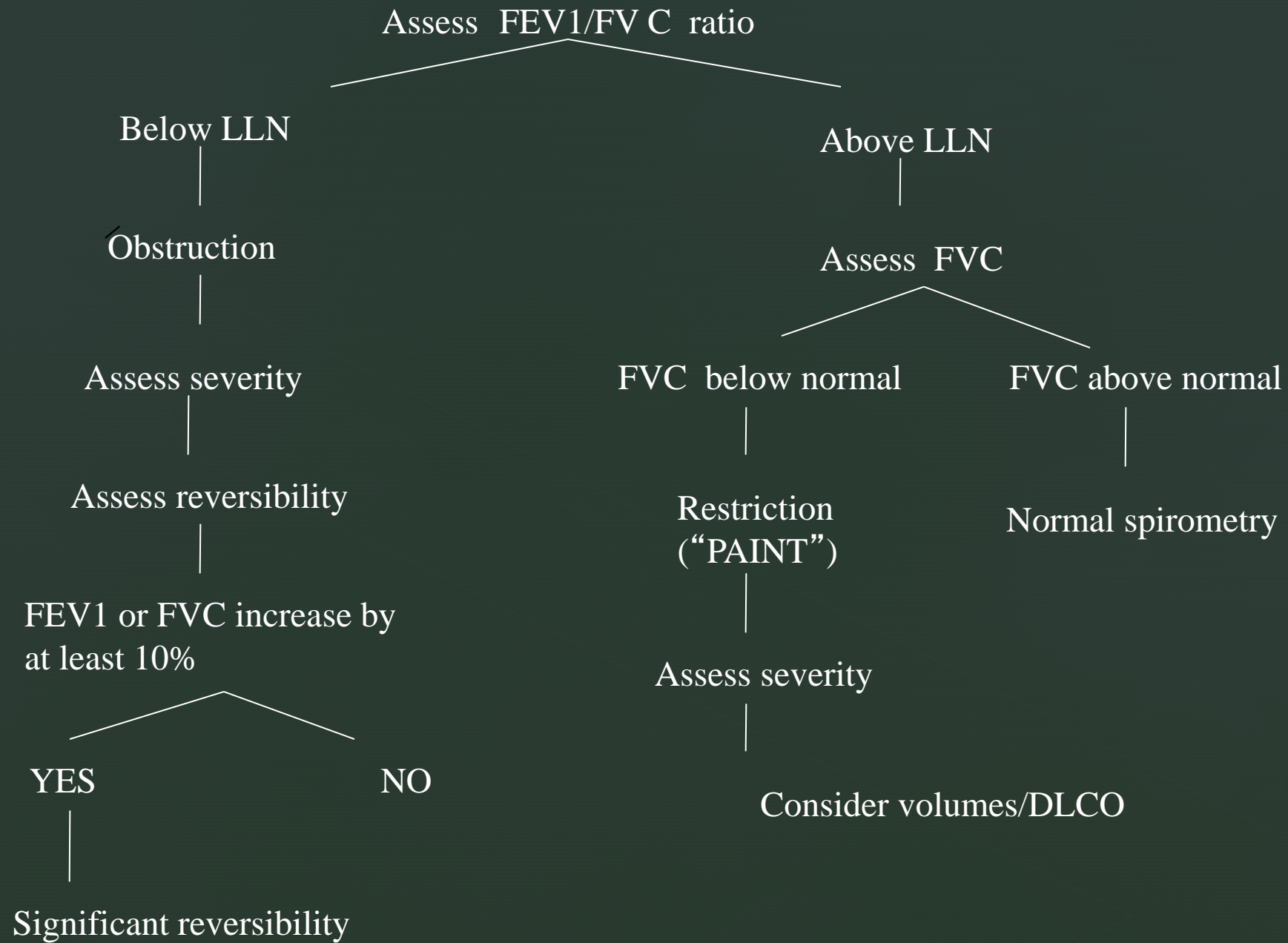
Forced Expiratory Volume in 1 Second (Fev1)

-The volume of air exhaled in the first second of the FVC

AAH, THAT'S BETTER

	Pred	LLN	Pre-	%	Post-	%	% chg
FVC	3.34	2.59	1.86	55	1.83	55	-1
FEV 1	2.44	1.81	0.75	31	0.73	30	-3
FEV1%F	73.81	64.13	40.36		39.72		

↓
<.70 AND LLN

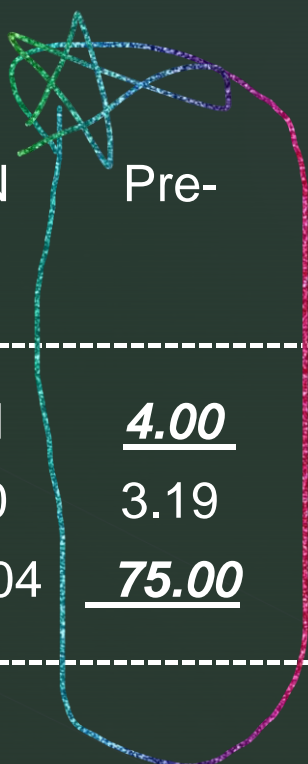


SPIROMETRY

	Pred	LLN	Pre-	%	Post-	%
▪ FVC	4.52	3.41	4.00	88	4.20	92
▪ FEV 1	3.54	2.70	3.19	90	3.24	91
▪ FEV1%F	78.32	70.04	75.00			

- Assess FEV1/FVC ratio
- Below LLN----this reveals obstruction(or below 70%-GOLD)
- Above LLN----this can be restrictive or normal (or above 70%-GOLD)
 - Assess FVC—if normal, spirometry is normal
---if below normal, spirometry is restrictive

SPIROMETRY



	Pred	LLN	Pre-	%	Post-	%

▪ FVC	4.52	3.41	<u>4.00</u>	88		
▪ FEV 1	3.54	2.70	3.19	90		
▪ FEV1%F	78.32	70.04	<u>75.00</u>			

- FEV1/FVC ratio is 75% which is above the LLN 70.04
- Spirometry is either normal or restrictive
- FVC is 4.00 liters, which is above LLN 3.41 liters
- This is a normal spirometry

Spirometric Severity Classification

(based on post-BD FEV₁)

Stage	For patients with FEV ₁ /FVC < 0.70:
1: Mild	FEV ₁ >80% predicted
2: Moderate	50% ≤ FEV ₁ < 80% predicted
3: Severe	30% ≤ FEV ₁ < 50% predicted
4: Very Severe	FEV ₁ < 30% predicted

SPIROMETRY

	Pred	LLN	Pre-	%	Post-	%	%chg

FVC	4.52	3.41	3.59	79	3.76	83	4
FEV 1	3.54	2.70	<u>1.18</u>	<u>33</u>	1.18	33	0
FEV1%F	78.32	70.04	<u>32.84</u>		31.51		

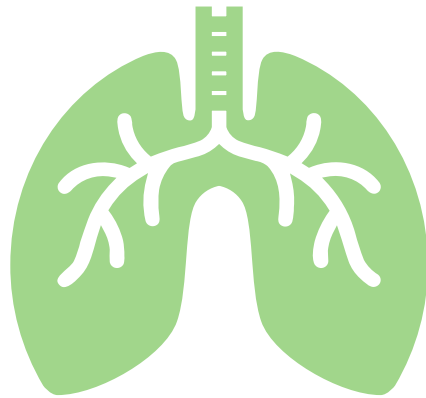
- FEV1/FVC ratio is 32.84% which is below LLN 70.04%
- Spirometry is obstructive
- FEV1 is 1.18 liters which is 33% of predicted
- Spirometry is severe obstruction
- FEV1 showed no change and FVC improved 4%
- There is no significant reversibility

SPIROMETRY

	Pred	LLN	Pre-	%

FVC	4.52	3.41	<u>2.60</u>	<u>60</u>
FEV 1	3.54	2.70	2.10	60
FEV1%F	78.32	70.04	<u>80.00</u>	

- FEV1/FVC ratio 80% is above LLN 70.04%
- Spirometry is either normal or restrictive
- FVC is 2.60 liters, below LLN 3.41 liters
- Spirometry is restrictive



SEVERITY of RESTRICTION

- **Total Lung Capacity (TLC) (Normal range: 80 - 120% of predicted)**
 - TLC > 120 = Hyperinflation
 - TLC < 80% = Restrictive disease (ATS criteria for severity):
 - 70-80 % predicted: mild
 - 60-70% predicted: moderate
 - 50-60% predicted: moderately severe
 - **< 50% predicted: severe**
- **• Step 5: RV/TLC ratio (Normal range: < 35% or < predicted)**
 - RV/TLC > 35% or > predicted indicates Air trapping

OBSTRUCTION vs. RESTRICTION

Obstruction

- decreased airflow
(reduced FEV1/FVC)
- normal/increased lung volume
- normal/decreased DLCO

Restriction

- normal airflow
(normal FEV1/FVC)
- decreased lung volume
- normal or decreased DLCO

DIFFUSION CAPACITY OF CARBON MONOXIDE

Spirometry
tells us how
well we move
air in and out
of the lungs

Lung
volumes tell
us how much
air we have
in the lungs

How do we
know how
well we
exchange
oxygen?

-how well
oxygen
diffuses or
transfers

DECREASED DLCO

Emphysema- destroyed alveoli

Fibrosis-scarring of alveoli and interstitium

Pulmonary vascular disease

Anemia

PULMONARY FUNCTION

	Pred	LLN	Pre-	%	Post-	%	%chg

FVC	4.52	3.41	4.00	88	4.20	92	5
FEV 1	3.54	2.70	1.60	45	1.70	48	6
FEV1%F	78.32	70.04	40.00		41.00		

1. Severe obstructive disease with no significant BD response

2. emphysema/COPD

chronic bronchitis

chronic asthma

bronchiectasis

sarcoidosis possible

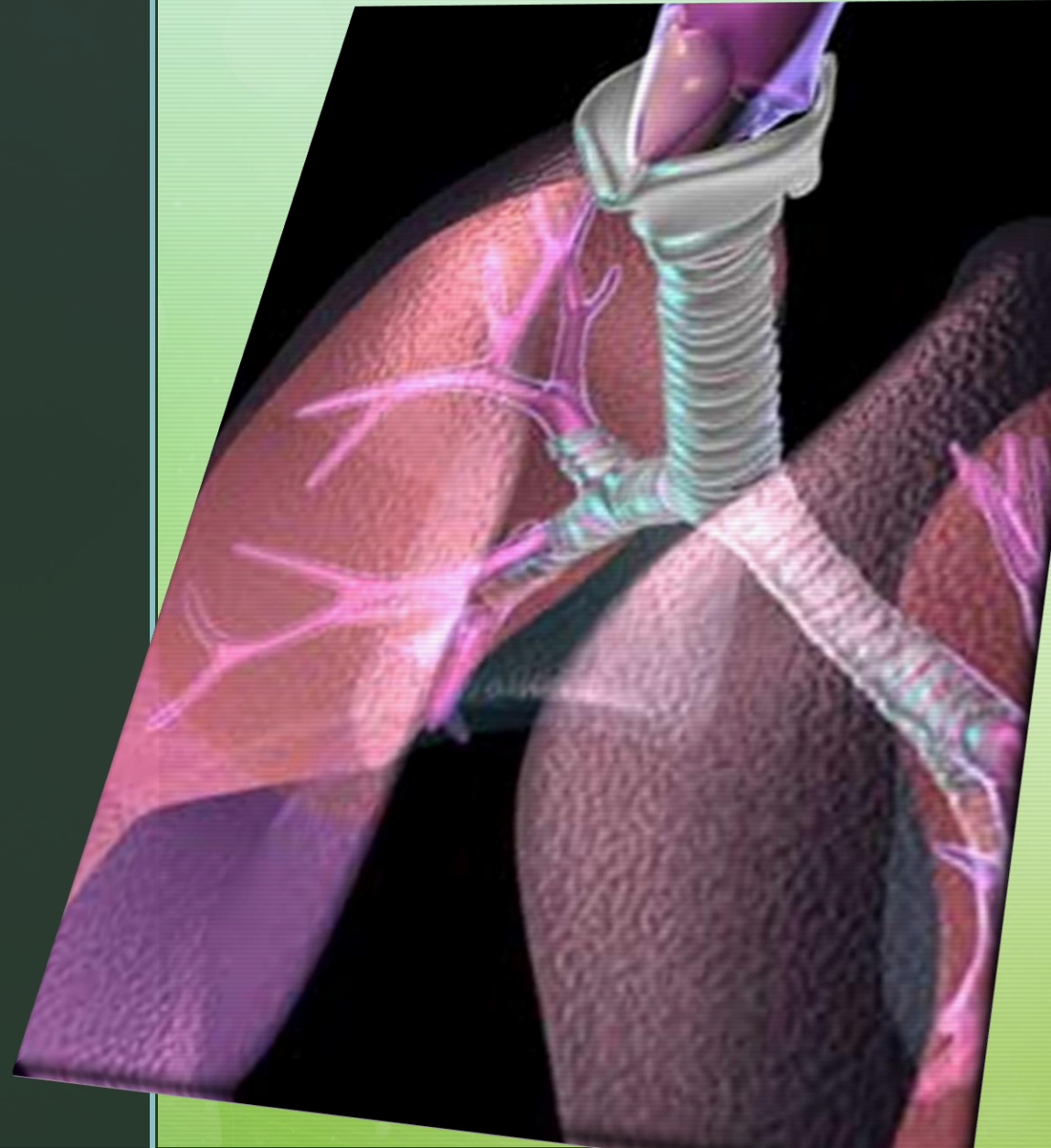


THE BREATH SOUND THEY DIDN'T
TEACH YOU IN SCHOOL.

Asthma & COPD

Asthma and COPD are important and common chronic conditions, seen by primary care

> 50 million affected/underdiagnosed



Asthma and COPD

Asthma
Sensitizing agent

COPD
Noxious agent

Asthmatic airway inflammation
CD4+ T lymphocytes
Eosinophils

COPD airway inflammation
CD8+ T lymphocytes
Macrophages Neutrophils



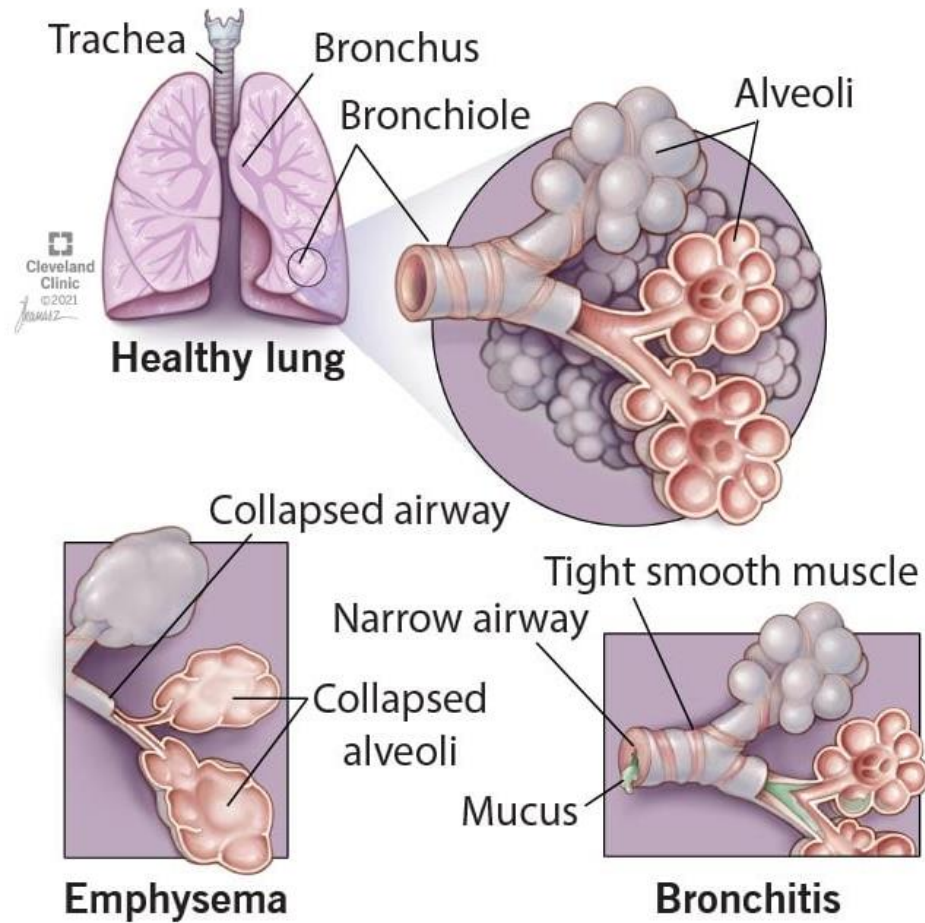
Reversible

Airflow Limitation

Irreversible

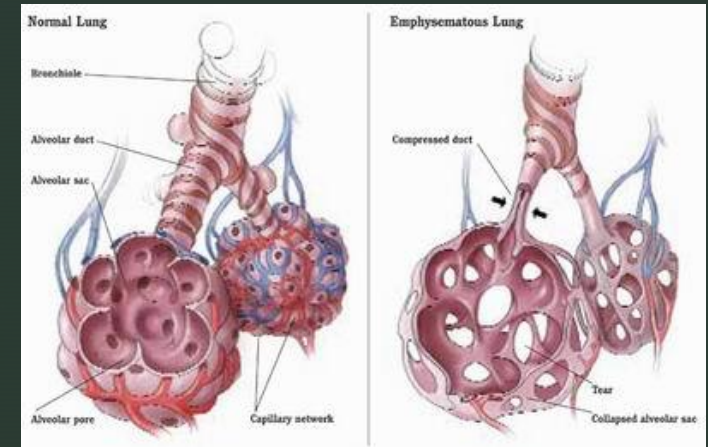
COPD

Chronic Obstructive Pulmonary Disease (COPD)



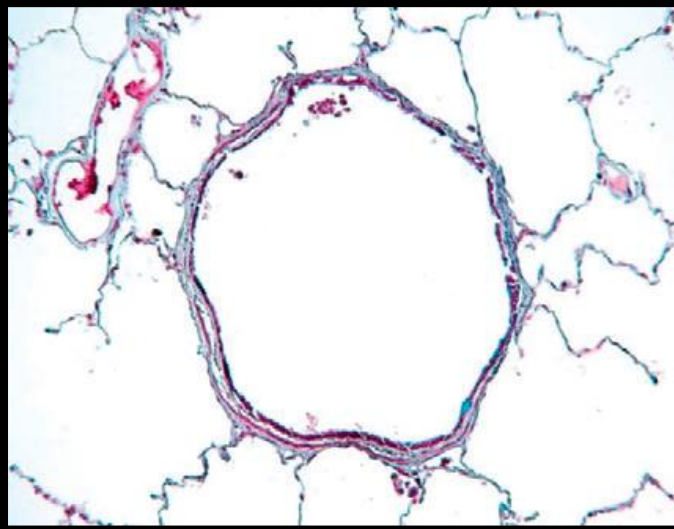
COPD

A common preventable and treatable disease, characterized by *persistent airflow limitation* that is usually progressive and associated with an *enhanced chronic inflammatory response* in the airways and the lung to noxious particles or gases.



Exacerbations and **comorbidities** contribute to the overall severity in individual patients.

Airway features in a healthy individual and in patient with COPD



Normal Airway



COPD: Narrowed airways
infiltration of inflammatory cells,
mucosal hyperplasia and
deposition of connective tissue
in peribronchiolar space

Symptoms related to airway disease and loss of alveolar spaces

- Shortness of breath with activity
- Frequent cough
- Cough with mucus (phlegm)
- Limiting activity due to shortness of breath
- Frequent colds/nose/throat infections
- Limiting activities due to shortness of breath or tiredness.

Diagnosis of COPD

Dyspnea, chronic cough or sputum, risk factors or family history of COPD

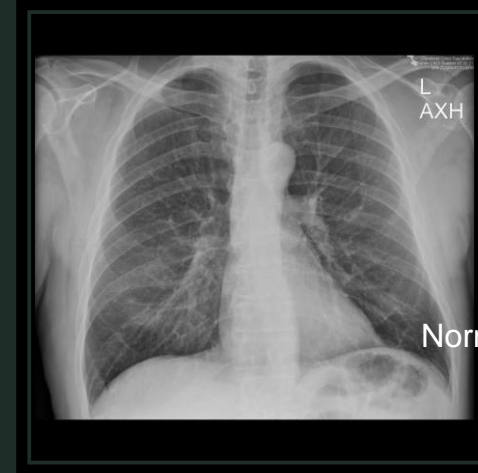
SPIROMETRY is required to make the diagnosis

Post bronchodilator FEV_1/FVC ratio < 0.70 or $< LLN$

Irreversible or partially reversible airflow obstruction

CXR not useful to diagnose COPD, useful to rule out

Exclude alternative explanation for obstructive lung disease



COPD

GOLD Guidelines: Composite Assessment



1) Determine SYMPTOM BURDEN



2) Severity of obstruction on
SPIROMETRY



3) RISK of EXACERBATIONS



COMBINE to guide the
management of COPD

Gold Diagnostics

GOLD ABE Assessment Tool

Figure 2.3

Spirometrically confirmed diagnosis

Assessment of airflow obstruction

Assessment of symptoms/risk of exacerbations

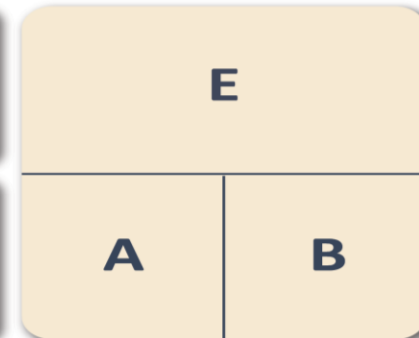
Post-bronchodilator
FEV1/FVC < 0.7

GRADE	FEV1 (% predicted)
GOLD 1	≥ 80
GOLD 2	50-79
GOLD 3	30-49
GOLD 4	< 30

EXACERBATION HISTORY

≥ 2 moderate exacerbations or
≥ 1 leading to hospitalization

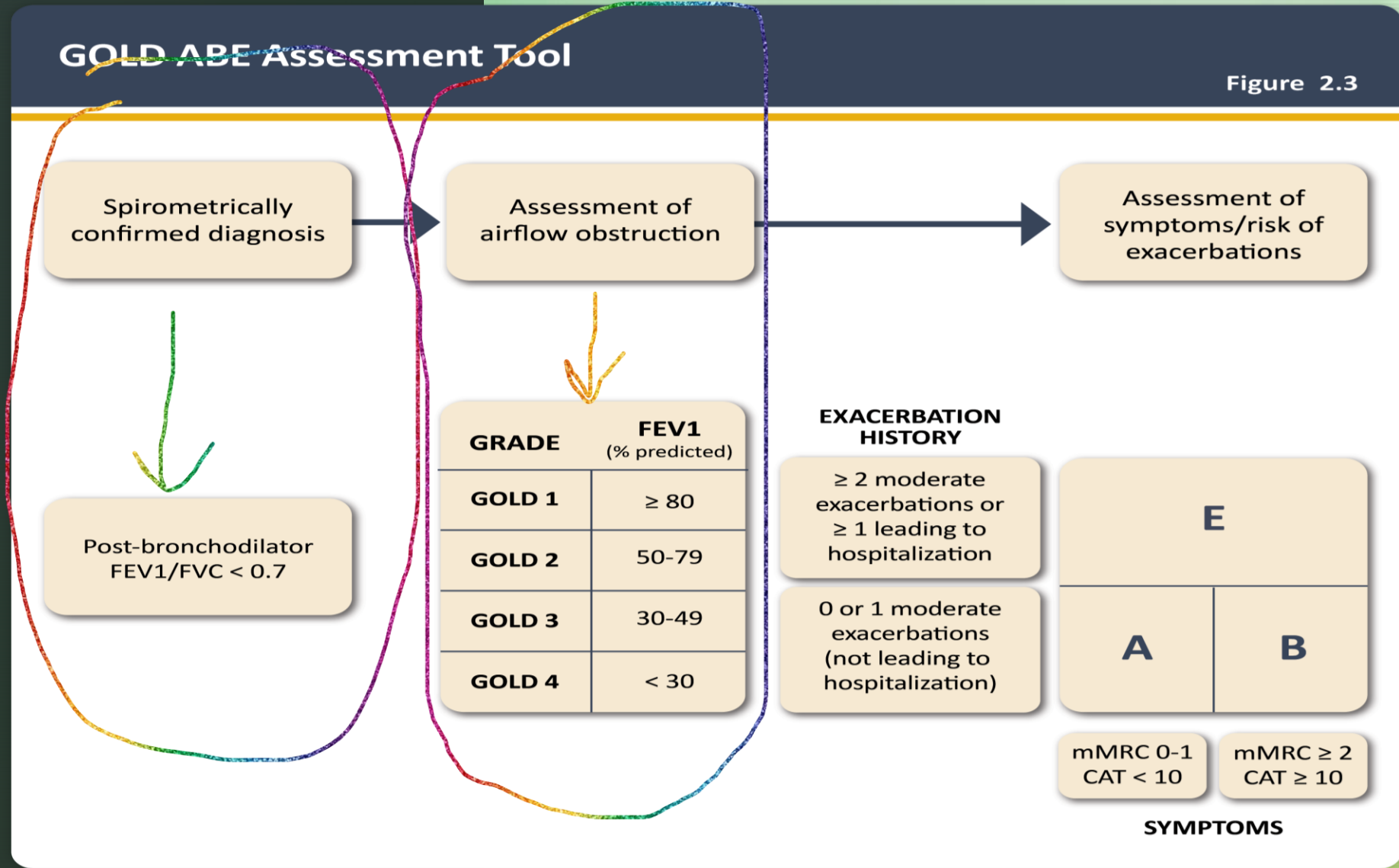
0 or 1 moderate exacerbations
(not leading to hospitalization)



mMRC 0-1
CAT < 10

mMRC ≥ 2
CAT ≥ 10

SYMPTOMS





SCORE

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

mMRC 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

COPD - Symptoms

Spirometric Severity Classification

(based on post-BD FEV₁)

Stage	For patients with FEV ₁ /FVC < 0.70:
1: Mild	FEV ₁ >80% predicted
2: Moderate	50% ≤ FEV ₁ < 80% predicted
3: Severe	30% ≤ FEV ₁ < 50% predicted
4: Very Severe	FEV ₁ < 30% predicted

COPD Assessment Test

I never cough

0 1 2 3 4 5

I cough all the time

I have no phlegm (mucus)
in my chest at all

0 1 2 3 4 5

My chest is completely
full of phlegm (mucus)

My chest does not
feel tight at all

0 1 2 3 4 5

My chest feels
very tight

When I walk up a hill or
one flight of stairs I am
not breathless

0 1 2 3 4 5

When I walk up a hill or
one flight of stairs I am
very breathless

I am not limited doing
any activities at home

0 1 2 3 4 5

I am very limited doing
activities at home

I am confident leaving
my home despite my
lung condition

0 1 2 3 4 5

I am not at all confident
leaving my home because
of my lung condition

I sleep soundly

0 1 2 3 4 5

I don't sleep soundly
because of my lung
condition

I have lots of energy

0 1 2 3 4 5

I have no energy at all

Symptoms- Modified Medical Research Council (mMRC)

PLEASE TICK IN THE BOX THAT APPLIES TO YOU
(ONE BOX ONLY)

mMRC Grade 0. I only get breathless with strenuous exercise.

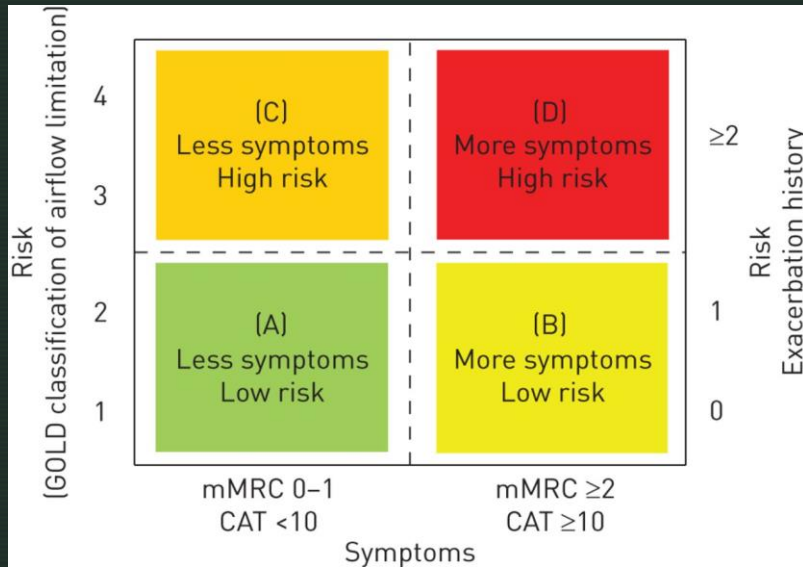
mMRC Grade 1. I get short of breath when hurrying on the level
or walking up a slight hill.

mMRC Grade 2. I walk slower than people of the same age on the
level because of breathlessness, or I have to stop for breath when
walking on my own pace on the level.

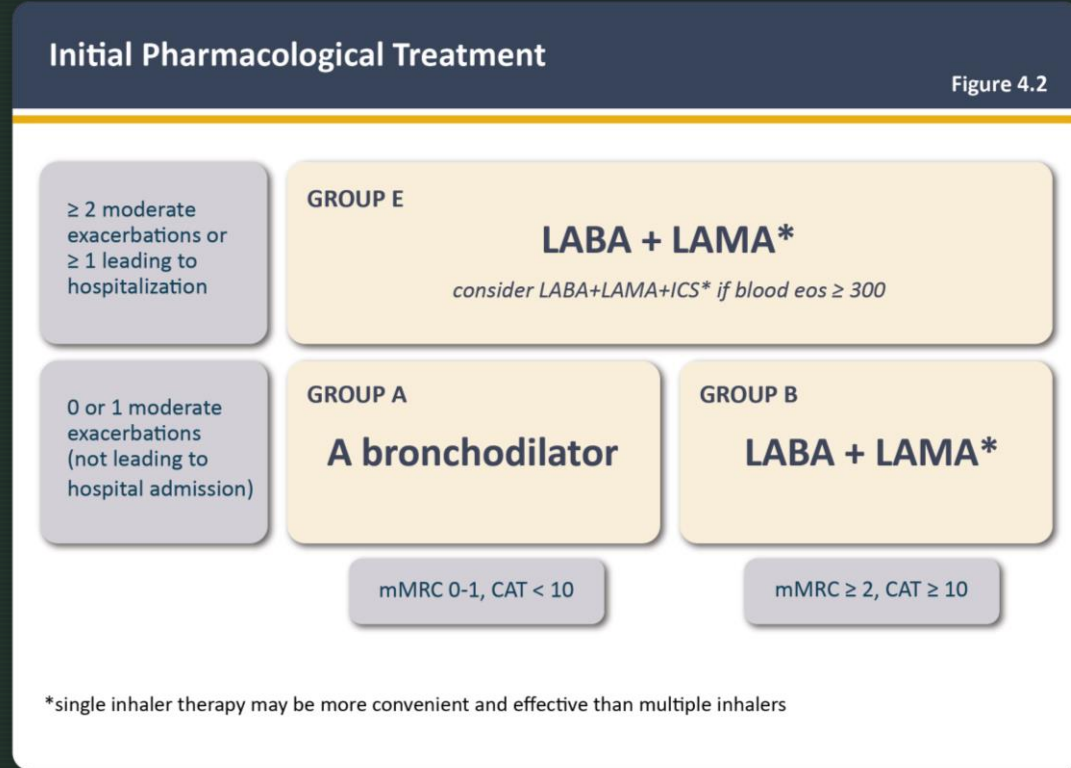
mMRC Grade 3. I stop for breath after walking about 100 meters or
after a few minutes on the level.

mMRC Grade 4. I am too breathless to leave the house or I am
breathless when dressing or undressing.

Old



2023 – Highlight importance of exacerbations



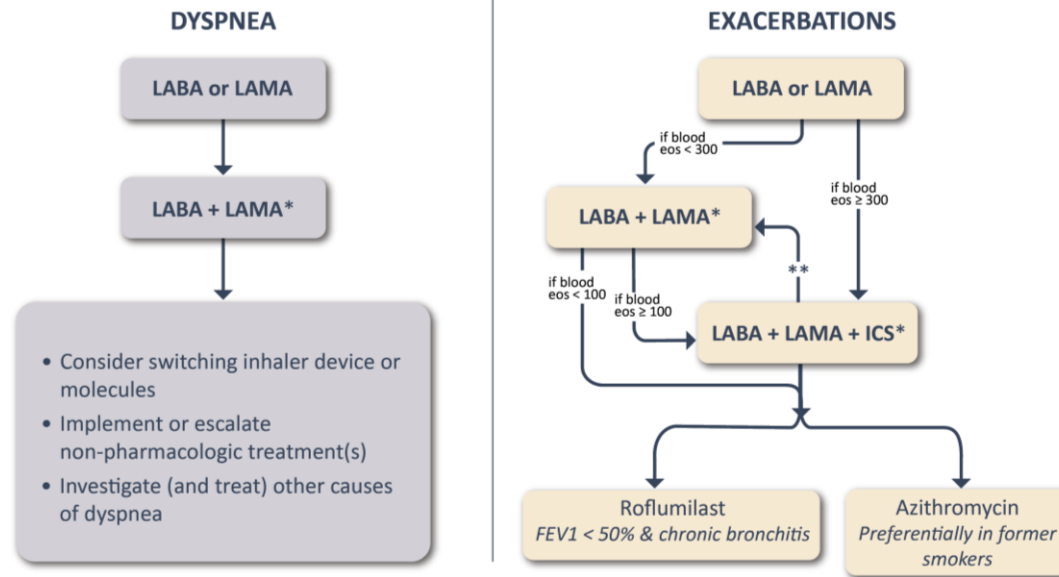
Treatment COPD

Treatment COPD

Follow-up Pharmacological Treatment

Figure 4.4

- 1 IF RESPONSE TO INITIAL TREATMENT IS APPROPRIATE, MAINTAIN IT.
- 2 IF NOT:
 - Check adherence, inhaler technique and possible interfering comorbidities
 - Consider the predominant treatable trait to target (dyspnea or exacerbations)
 - Use exacerbation pathway if both exacerbations and dyspnea need to be targeted
 - Place patient in box corresponding to current treatment & follow indications
 - Assess response, adjust and review
 - These recommendations do not depend on the ABE assessment at diagnosis



*Single inhaler therapy may be more convenient and effective than multiple inhalers

**Consider de-escalation of ICS if pneumonia or other considerable side-effects. In case of blood eos ≥ 300 cells/μl de-escalation is more likely to be associated with the development of exacerbations

- Many patients will be on LAMA/LABA
- The minority of patients will be on LAMA/LABA/ICS
- Few patients will be on LABA/ICS
- Blood eosinophils are an essential biomarker for COPD

Treatment COPD Exacerbations

- Moderate exacerbation = need of oral steroids
- Severe exacerbation = hospitalized (due to respiratory failure i.e. need oxygen vs. BiPAP)

Factors to Consider when Initiating ICS Treatment

Figure 3.1

Factors to consider when adding ICS to long-acting bronchodilators:

(note the scenario is different when considering ICS withdrawal)

STRONGLY FAVORS USE

History of hospitalization(s) for exacerbations of COPD[#]
≥ 2 moderate exacerbations of COPD per year[#]
Blood eosinophils ≥ 300 cells/μL
History of, or concomitant asthma

FAVORS USE

1 moderate exacerbation of COPD per year[#]
Blood eosinophils 100 to < 300 cells/μL

AGAINST USE

Repeated pneumonia events
Blood eosinophils < 100 cells/μL
History of mycobacterial infection

[#]despite appropriate long-acting bronchodilator maintenance therapy (see Table 3.4 and Figure 4.3 for recommendations);

*note that blood eosinophils should be seen as a continuum; quoted values represent approximate cut-points; eosinophil counts are likely to fluctuate.

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EXACERBATIONS

Smooth muscle dysfunction

- Bronchoconstriction
- Bronchial hyperactivity
- Hyperplasia/hypertrophy
- Inflammatory mediator release

Airway inflammation

- Mucosal edema
- Inflammatory cell activation
- Cellular proliferation
- Epithelial damage
- Basement membrane thickening

Managing Exacerbations

CAUSES/Risk Factors

Prior exacerbations

Infections

- 50-80% viral
- 45% bacterial H flu, M catarrhalis, S pneumonia

Lower BMI, Continued smoking

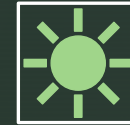
Air pollution, extremes in weather

Certain cancers

Heart failure, PE, arrhythmia, PTX

(Unknown in 1/3 of cases)

SIGNS



Change in mucus



Difficulty coughing up mucus



Cough



Shortness of breath



Wheezing

Treatment of COPD Exacerbation

SABA or in combo
with Atrovent
(DUONEB)
PRN

Consider adding
LABA+LAMA

Systemic
Corticosteroids

Tobacco
Cessation

Antimicrobial
Therapy



SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir® Digihaler™
90 mcg albuterol sulfate inhalation powder
DBI A



ProAir® RespiClick®
90 mcg albuterol sulfate inhalation powder
DBI A



Proventil® HFA
90 mcg albuterol sulfate
DBI A G



Ventolin® HFA
90 mcg albuterol sulfate
DBI A G



Xopenex® HFA™
45 mcg levalbuterol tartrate
A G



LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent® Diskus®
50 mcg salmeterol xinafoate inhalation powder
DBI A C



Striverdi® Respimat®
2.5 mcg olodaterol hydrochloride
DBI C



INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they do not relieve sudden symptoms of coughing, wheezing or shortness of breath

Alvesco® HFA
80, 160 mcg ciclesonide
DBI A



ArmonAir® Digihaler™
55, 113, 232 mcg fluticasone propionate inhalation powder
DBI A



Arnuity® Ellipta®
50, 100, 200 mcg fluticasone furoate inhalation powder
DBI A



Asmanex® HFA
50, 100, 200 mcg mometasone furoate
DBI A



Asmanex® Twisthaler®
110, 220 mcg mometasone furoate inhalation powder
DBI A



Flovent® Diskus®
50, 100, 250 mcg fluticasone propionate inhalation powder
DBI A



Flovent® HFA
44, 110, 220 mcg fluticasone propionate
DBI A



Pulmicort Flexhaler®
90, 180 mcg budesonide inhalation powder
DBI A



QVAR® Redihaler™
40, 80 mcg beclomethasone dipropionate
DBI A



MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheeze and chest tightness associated with chronic lung diseases

Short-acting
Aтровент® HFA
17 mcg ipratropium bromide
DBI C



Long-acting
Increase® Ellipta®
82.5 mcg umecidinium inhalation powder
DBI C



Spiriva® HandiHaler®
18 mcg tiotropium bromide inhalation powder
G



Spiriva® Respimat®
1.25, 2.5 mcg tiotropium bromide
DBI A G



Tudorza® Pressair™
600 mcg acclidium bromide inhalation powder
DBI G



COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent® Respimat®
20/100 mcg ipratropium bromide and albuterol
DBI C



COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair Diskus®
100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder
DBI A C G



Advair® HFA
45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate
DBI A G



AirDuo® RespiClick®
55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder
DBI A G



Breo® Ellipta®
100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder
DBI A G



Dulera®
50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate
DBI A



Symbicort®
80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate
DBI A C G



Wixela™ Inhub™
100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder
DBI A C



Anoro® Ellipta®
62.5/25 mcg umecidinium and vilanterol inhalation powder
DBI C



Bevespi Aerosphere®
9/4.8 mcg glycopyrrolate and formoterol fumarate
DBI C



Duaklir® Pressair™
400, 12 mcg acclidium bromide and formoterol fumarate
DBI C



Stiolto™ Respimat®
2.5/2.5 mcg tiotropium bromide and olodaterol
DBI C

Trelegy® Ellipta®
200/10/2.5/25 mcg, 100/50/2.5/25 mcg fluticasone furoate, umecidinium and vilanterol inhalation powder
DBI A C

Breztri Aerosphere™
160/9/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate
DBI C

BIOLOGICS

target cells and pathways that cause airway inflammation; delivered by injection or IV

Cinqair®
reslizumab
A



Dupixent®
dupilumab
A




Fasenra™
bambucumab
A



Nucala®
mepolizumab
A



Tezspire™
tezepelumab-ekko
A



Xolair®
omalizumab
A



BRONCHIAL THERMOPLASTY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities.
www.bronchoasthma.com



PDE4 INHIBITORS

ease lung inflammation and reduce exacerbations

Daliresp®
250, 500 mcg roflumilast
C



INHALERS

Short Acting Bronchodilators (SABA)

“Relievers” treat bronchospasm,
fast acting

- Albuterol (ProAir®)
- Levalbuterol (Xopenex®)

Long-acting bronchodilators (LABA)

Prevent development of
bronchospasm

- Salmeterol (Serevent
Diskus®)
- Formoterol (Foradil®)





SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir® Digihaler™ 90 mcg albuterol sulfate inhalation powder DIN A	ProAir® RespiClick® 90 mcg albuterol sulfate inhalation powder DIN A	Proventil® HFA 90 mcg albuterol sulfate DIN A, B	Ventolin® HFA 90 mcg albuterol sulfate DIN A, B	Xopenex® HFA 45 mcg levalbuterol tartrate DIN A, C
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent® Diskus™ 50 mcg salmeterol xinafoate inhalation powder DIN A, C	Striverdi® RespiMat™ 2.5 mcg olodaterol hydrochloride DIN C
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INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they do not relieve sudden symptoms of coughing, wheezing or shortness of breath

Alvesco® HFA 80, 180 mcg ciclesonide DIN A	ArmonAir® Digihaler™ 55, 113, 232 mcg fluticasone propionate inhalation powder DIN A	Arnuity® Ellipta™ 50, 100, 200 mcg fluticasone furoate inhalation powder DIN A	Asmanex® HFA 50, 100, 200 mcg mometasone furoate DIN A	Asmanex® Twisthaler™ 110, 220 mcg mometasone furoate inhalation powder DIN A	Flovent® Diskus™ 50, 100, 250 mcg fluticasone propionate inhalation powder DIN A	Flovent® HFA 44, 110, 220 mcg fluticasone propionate DIN A	Pulmicort Flexhaler™ 90, 180 mcg budesonide inhalation powder DIN A	QVAR® Redihaler™ 40, 80 mcg beclomethasone dipropionate DIN A
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MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheeze and chest tightness associated with chronic lung diseases

Atrovent® HFA 17 mcg ipratropium bromide DIN C	Increase® Ellipta™ 12.5 mcg umecidinium inhalation powder DIN C	Spiriva® HandiHaler™ 18 mcg tiotropium bromide inhalation powder DIN C	Spiriva® RespiMat™ 1.25, 2.5 mcg tiotropium bromide DIN A, C	Tudorza™ Pressair™ 600 mcg aclidinium bromide inhalation powder DIN C
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COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent® RespiMat™ 20/100 mcg ipratropium bromide and albuterol DIN C
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COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair Diskus™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DIN A, C, G	Advair® HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate DIN A, G	AirDuo® Digihaler™ 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DIN A	AirDuo® RespiClick® 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DIN A, G	Breo® Ellipta™ 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder DIN A, E	Dulera® 50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate DIN A	Symbicort® 80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate DIN A, C, G	Wixela™ Inhub™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DIN A, C
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COMBINATION MEDICATIONS

contain both long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Anoro® Ellipta™ 62.5/25 mcg umecidinium and vilanterol inhalation powder DIN C	Bevespi Aerosphere® 9/4.8 mcg glycopyrrolate and formoterol fumarate DIN C	Duaklir® Pressair™ 400, 12 mcg aclidinium bromide and formoterol fumarate DIN C
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COMBINATION MEDICATIONS

contain inhaled corticosteroid, long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Stiolto® RespiMat™ 2.5/2.5 mcg tiotropium bromide and olodaterol DIN C	Trelegy® Ellipta™ 200/12.5/25 mcg, 100/62.5/25 mcg fluticasone furoate, umecidinium and vilanterol inhalation powder DIN A, C	Bretri Aerosphere™ 160/9/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate DIN C
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BIOLOGICS

target cells and pathways that cause airway inflammation; delivered by injection or IV

Cinqair® reslizumab A	Dupixent® dupilumab A	Fasenra™ bivalizumab A	Nucala® mepolizumab A	Tezspire™ tezepelumab-ekko A	Xolair® omalizumab A
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BRONCHIAL THERMOPLASTY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities. www.bronchoasthma.com

PDE4 INHIBITORS

ease lung inflammation and reduce exacerbations

Daliresp® 250, 500 mcg roflumilast C

INHALERS

Inhaled Corticosteroids

Inhibit production of inflammatory agents

- Budesonide (Pulmicort®)
- Fluticasone (Flovent®)

Long-acting muscarinic antagonists (LAMAS)

Long-acting bronchodilation, potentially anti-eosinophilic/anti-inflammatory effect

- Tiotropium Bromide (Spiriva®)

SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir® Digihaler® 90 mcg albuterol sulfate DBI A	ProAir® RespiClick® 90 mcg albuterol sulfate inhalation propellant DBI A G	Proventil® HFA 90 mcg albuterol sulfate DBI A G	Ventolin® HFA 90 mcg albuterol sulfate DBI A G	Xopenex® HFA 45 mcg levalbuterol tartrate A G
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent® Diskus® 50 mcg salmeterol xinafoate inhalation powder DBI A C	Striverdi® RespiMat® 2.5 mcg olodaterol hydrochloride DBI C
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INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they do not relieve sudden symptoms on their own

Alvesco® HFA 80, 180 mcg ciclesonide DBI A	ArmonAir® Digihaler® 55, 113, 232 mcg fluticasone propionate inhalation powder DBI A	Arnuity® Ellipta® 50, 100, 200 mcg fluticasone furoate inhalation powder A	Asmanex® HFA 50, 100, 200 mcg mometasone furoate DBI A	Asmanex® Twister® 110, 220 mcg mometasone furoate inhalation powder DBI A	Flovent® Diskus® 50, 100, 250 mcg fluticasone propionate inhalation powder DBI A	Flovent® HFA 44, 110, 220 mcg fluticasone propionate DBI A	Pulmicort Flexhaler® 90, 180 mcg budesonide inhalation powder DBI A	QVAR® Redihaler™ 40, 80 mcg beclomethasone dipropionate DBI A
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MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheeze and chest tightness associated with chronic lung diseases

Atrovent® HFA 17 mcg ipratropium bromide DBI C	Increase® Ellipta® 12.5 mcg umedclidium inhalation powder DBI C	Spiriva® HandiHaler® 18 mcg tiotropium bromide inhalation powder C	Spiriva® RespiMat® 1.25, 2.5 mcg tiotropium bromide DBI A C	Tudorza® Pressair™ 600 mcg aclidinium bromide inhalation powder DBI C
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COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and long-acting beta₂-agonist and anticholinergic

Ambivent® RespiMat® 200 mcg tiotropium bromide and albuterol DBI C

COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair Diskus® 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DBI A C G	Advair® HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate DBI A G	AirDuo® Digihaler™ 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DBI A	AirDuo® RespiClick® 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DBI A G	Breo® Ellipta® 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder DBI A G	Dulera® 50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate DBI A	Symbicort® 80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate DBI A C G	Wixela™ Inhub™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DBI A C	Anoro® Ellipta® 62.5/25 mcg umedclidium and formoterol inhalation powder DBI C	Bevespi Aerosphere® 94.8 mcg glycopyrrolate and formoterol fumarate DBI C	Duaklir® Pressair™ 400, 12 mcg aclidinium bromide and formoterol fumarate DBI C	Stiolto® RespiMat® 2.5/2.5 mcg tiotropium bromide and olodaterol DBI C	Trelegy® Ellipta® 200/12.5/25 mcg, 100/62.5/25 mcg fluticasone furoate, umedclidium and vilanterol inhalation powder DBI A C	Breztri Aerosphere™ 160/94.8 mcg budesonide, glycopyrrolate and formoterol fumarate DBI C
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COMBINATION MEDICATIONS

contain inhaled corticosteroid, long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Ambivent® RespiMat® 200 mcg tiotropium bromide and albuterol DBI C

BIOLOGICS

target cells and pathways that cause airway inflammation; delivered by injection or IV

Cinqair® reslizumab A	Dupixent® dupilumab A	Fasenra™ bivalizumab A	Nucala® mepolizumab A	Tezspire™ tezepelumab-ekko A	Xolair® omalizumab A
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BRONCHIAL THERMOPLASTY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities.

www.bcforgasthma.com



PDE4 INHIBITORS

ease lung inflammation and reduce exacerbations

Daliresp® 250, 500 mcg roflumilast C

COMBO INHALERS

ICS + LABA

- Advair Diskus® (Flovent +Salmeterol)
- Symbicort® (Pulmicort + Formoterol)
- Dulera: mometasone-formoterol

SAMA +SABA

- Combivent
- Duoneb

SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

<p>ProAir® Digihaler® 90 mcg albuterol sulfate inhalation powder DBI A</p> 	<p>ProAir® RespiClick® 90 mcg albuterol sulfate inhalation powder DBI A</p> 	<p>Proventil® HFA 90 mcg albuterol sulfate DBI A G</p> 	<p>Ventolin® HFA 90 mcg albuterol sulfate DBI A G</p> 	<p>Xopenex® HFA 45 mcg levalbuterol tartrate A G</p> 
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

<p>Serevent® Diskus® 50 mcg salmeterol xinafoate inhalation powder DBI A C</p> 	<p>Striverdi® RespiMat® 2.5 mcg olodaterol hydrochloride DBI C</p> 
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INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they do not relieve sudden symptoms of coughing, wheezing or shortness of breath

<p>Alvesco® HFA 80, 160 mcg ciclesonide DBI A</p> 	<p>ArmonAir® Digihaler® 55, 113, 232 mcg fluticasone propionate inhalation powder DBI A</p> 	<p>Arnuity® Ellipta® 50, 100, 200 mcg fluticasone furoate inhalation powder DBI A</p> 	<p>Asmanex® HFA 50, 100, 200 mcg mometasone furoate DBI A</p> 	<p>Asmanex® Twisthaler® 110, 220 mcg mometasone furoate inhalation powder DBI A</p> 	<p>Flovent® Diskus® 50, 100, 250 mcg fluticasone propionate inhalation powder DBI A</p> 	<p>Flovent® HFA 44, 110, 220 mcg fluticasone propionate DBI A</p> 	<p>Pulmicort Flexhaler® 90, 180 mcg budesonide inhalation powder DBI A</p> 	<p>QVAR® Redihaler™ 40, 80 mcg beclomethasone dipropionate DBI A</p> 
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MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheeze and chest tightness associated with chronic lung diseases

<p>Atrovent® HFA 17 mcg ipratropium bromide DBI C</p> 	<p>Increase® Ellipta® 12.5 mcg umedidinium inhalation powder DBI C</p> 	<p>Spiriva® HandiHaler® 18 mcg tiotropium bromide inhalation powder C</p> 	<p>Spiriva® RespiMat® 1.25, 2.5 mcg tiotropium bromide DBI A C</p> 	<p>Tudorza® Pressair™ 600 mcg aclidinium bromide inhalation powder DBI C</p> 
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COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

<p>Combivent® RespiMat® 20/100 mcg ipratropium bromide and albuterol DBI C</p> 

COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist

<p>Advair Diskus® 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DBI A C G</p> 	<p>Advair® HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate DBI A G</p> 	<p>AirDuo® Digihaler™ 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DBI A</p> 	<p>AirDuo® RespiClick® 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DBI A G</p> 	<p>Breo® Ellipta® 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder DBI A G</p> 	<p>Dulera® 50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate DBI A</p> 	<p>Symbicort® 80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate DBI A C G</p> 	<p>Wixela™ Inhub™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DBI A C</p> 
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COMBINATION MEDICATIONS

contain both long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

<p>Lenoro® Ellipta® 2.5/2.5 mcg umedidinium and glycopyrrolate inhalation powder DBI C</p> 	<p>Bevespi Aerosphere® 9/4.8 mcg glycopyrrolate and formoterol fumarate DBI C</p> 	<p>Duaklir® Pressair™ 400, 12 mcg aclidinium bromide and formoterol fumarate DBI C</p> 	<p>Stiolto® RespiMat® 2.5/2.5 mcg tiotropium bromide and olodaterol DBI C</p> 	<p>Trelegy® Ellipta® 200/12.5/25 mcg, 100/62.5/25 mcg fluticasone furoate, umedidinium and vilanterol inhalation powder DBI A C</p> 	<p>Breztri Aerosphere™ 160/9/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate DBI C</p> 
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BIOLOGICS

<p>Cinqair® reslizumab A</p> 	<p>Dupixent® dupilumab A</p> 	<p>Fasenra™ bambucumab A</p> 	<p>Nucala® mepolizumab A</p> 	<p>Tezspire™ tezepelumab-ekko A</p> 	<p>Xolair® omalizumab A</p> 
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BRONCHIAL THERMOPLASTY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities.
www.bronchoasthma.com



PDE4 INHIBITORS

ease lung inflammation and reduce exacerbations

<p>Daliresp® 250, 500 mcg roflumilast C</p> 
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COMBO INHALERS

LAMA + LABA

- Anoro Ellipta
- Stiolto Respimat
- Bevespi

TRIPLE THERAPY

ICS/LAMA/LABA

- Trelegy
- Breztri



SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir® Digihaler™
90 mcg albuterol sulfate inhalation powder
DBI A



ProAir® RespiClick®
90 mcg albuterol sulfate inhalation powder
DBI A



Proventil® HFA
90 mcg albuterol sulfate
DBI A G



Ventolin® HFA
90 mcg albuterol sulfate
DBI A G



Xopenex® HFA
45 mcg levalbuterol tartrate
A G



LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent® Diskus®
50 mcg salmeterol xinafoate inhalation powder
DBI A C



Striverdi® Respimat®
2.5 mcg olodaterol hydrochloride
DBI C



INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they do not relieve sudden symptoms of coughing, wheezing or shortness of breath

Alvesco® HFA
80, 160 mcg ciclesonide
DBI A



ArmonAir® Digihaler™
55, 113, 232 mcg fluticasone propionate inhalation powder
DBI A



Arnuity® Ellipta®
50, 100, 200 mcg fluticasone furoate inhalation powder
DBI A



Asmanex® HFA
50, 100, 200 mcg mometasone furoate
DBI A



Asmanex® Twister™
110, 220 mcg mometasone furoate inhalation powder
DBI A



Flovent® Diskus®
50, 100, 250 mcg fluticasone propionate inhalation powder
DBI A



Flovent® HFA
44, 110, 220 mcg fluticasone propionate
DBI A



Pulmicort® Flexhaler™
90, 180 mcg budesonide inhalation powder
DBI A



QVAR® Redihaler™
40, 80 mcg beclomethasone dipropionate
DBI A



MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheeze and chest tightness associated with chronic lung diseases

Short-acting
Aтровент® HFA
17 mcg ipratropium bromide
DBI C



Long-acting
Increase® Ellipta®
12.5 mcg umecidinium inhalation powder
DBI C



Spiriva® HandiHaler™
18 mcg tiotropium bromide inhalation powder
G



Spiriva® Respimat®
1.25, 2.5 mcg tiotropium bromide
DBI A G



Tudorza™ Pressair™
600 mcg acclidium bromide inhalation powder
DBI C



COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent® Respimat®
20/100 mcg oxvatropium bromide and albuterol
DBI C



COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair Diskus®
100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder
DBI A C G



Advair® HFA
45/21, 115/21, 232/21 mcg fluticasone propionate and salmeterol xinafoate
DBI A G



AirDuo® RespiClick®
55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder
DBI A G




Breo® Ellipta®
100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder
DBI A G



Dulera®
50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate
DBI A



Symbicort®
80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate
DBI A C G



Wixela™ Inhaler
100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder
DBI A C



COMBINATION MEDICATIONS

contain both long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Anoro® Ellipta®
62.5/25 mcg umecidinium and vilanterol inhalation powder
DBI C



Bevespi Aerosphere®
9/4.8 mcg glycopyrrolate and formoterol fumarate
DBI C



Duaklir® Pressair™
400, 12 mcg acclidium bromide and formoterol fumarate
DBI C



Stiolto® Respimat®
2.5/2.5 mcg tiotropium bromide and olodaterol
DBI C



Trelegy® Ellipta®
200/12.5/25 mcg, 100/62.5/25 mcg budesonide, fluticasone furoate, and vilanterol inhalation powder
DBI A C



Breztri Aerosphere™
160/9/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate
DBI C



BIOLOGICS

target cells and pathways that cause airway inflammation; delivered by injection or IV

Cinqair®
reslizumab
A



Dupixent®
dupilumab
A



Fasenra™
bambucicab
A



Nucala®
mepolizumab
A



Tezspire™
tezepelumab-ekko
A



Xolair®
omalizumab
A



BRONCHIAL THERMOPLASTY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities.
www.bronchothermy.com



PROLONGED ACTION INHALERS

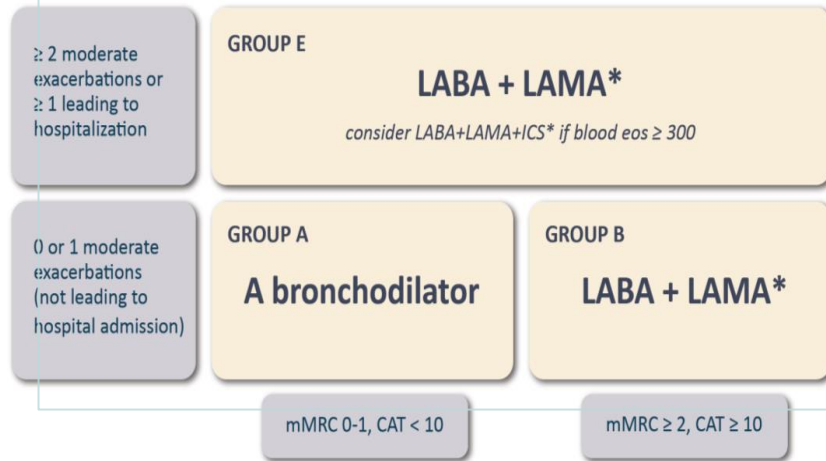
ease lung inflammation and reduce exacerbations

Daliresp®
250, 500 mcg roflumilast
C



Initial Pharmacological Treatment

Figure 4.2

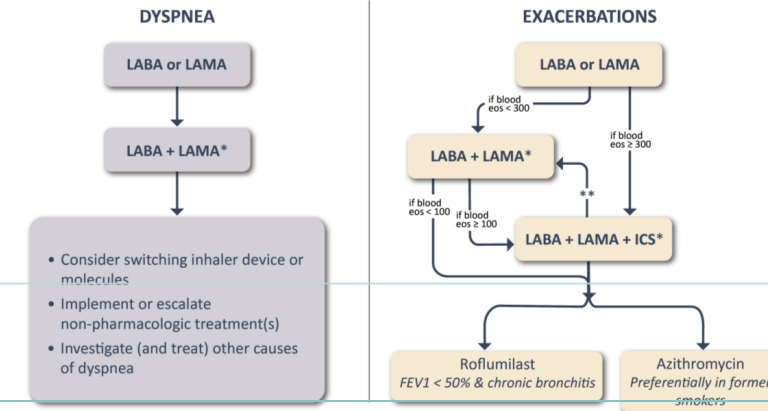


*single inhaler therapy may be more convenient and effective than multiple inhalers

Follow-up Pharmacological Treatment

Figure 4.4

- IF RESPONSE TO INITIAL TREATMENT IS APPROPRIATE, MAINTAIN IT.
- IF NOT:
 - Check adherence, inhaler technique and possible interfering comorbidities
 - Consider the predominant treatable trait to target (dyspnea or exacerbations)
 - Use exacerbation pathway if both exacerbations and dyspnea need to be targeted
 - Place patient in box corresponding to current treatment & follow indications
 - Assess response, adjust and review
 - These recommendations do not depend on the ABE assessment at diagnosis



*Single inhaler therapy may be more convenient and effective than multiple inhalers

**Consider de-escalation of ICS if pneumonia or other considerable side-effects. In case of blood eos ≥ 300 cells/ μ l de-escalation is more likely to be associated with the development of exacerbations

COPD TREATMENT



allergyandasthma.org
800.878.4403

DBP = DOSE INDICATOR G = GENERIC AVAILABLE DISEASE STATES: A = ASTHMA C = COPD

SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir® Digihaler™ 90 mcg albuterol sulfate inhalation powder DBP A	ProAir® RespiClick® 90 mcg albuterol sulfate inhalation powder DBP A	Proventil® HFA 90 mcg albuterol sulfate DBP A G	Ventolin® HFA 90 mcg albuterol sulfate DBP A G	Xopenex® HFA® 45 mcg levalbuterol tartrate A G
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent® Diskus® 50 mcg salmeterol xinafoate inhalation powder DBP A G	Striverdi® RespiMat® 2.5 mcg olodaterol hydrochloride DBP C
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Alvesco® HFA 80, 160 mcg ciclesonide DBP A	ArmonAir® Digihaler™ 55, 113, 232 mcg fluticasone propionate inhalation powder DBP A	Arnuity® Ellipta® 50, 100, 200 mcg fluticasone furoate inhalation powder DBP A	Asmanex® HFA 50, 100, 200 mcg mometasone furoate DBP A	Asmanex® Twisthaler® 110, 220 mcg mometasone furoate inhalation powder DBP A	Flovent® Diskus® 50, 100, 250 mcg fluticasone propionate inhalation powder DBP A	Flovent® HFA 44, 110, 220 mcg fluticasone propionate DBP A	Pulmicort Flexhaler® 90, 180 mcg budesonide inhalation powder DBP A	QVAR® Redihaler™ 40, 80 mcg beclomethasone dipropionate DBP A
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MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheeze and chest tightness associated with chronic lung diseases

Atrovent® HFA 17 mcg ipratropium bromide DBP C	Increase® Ellipta® 82.5 mcg umeclidinium inhalation powder DBP C	Spiriva® HandiHaler® 18 mcg tiotropium bromide inhalation powder DBP C	Spiriva® Respimat® 1.25, 2.5 mcg tiotropium bromide DBP A G	Tudorza® Pressair™ 603 mcg aclidinium bromide inhalation powder DBP C
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COMBINATION MEDICATIONS

contains both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent® RespiMat® 10/100 mcg oxtripropium bromide and albuterol DBP C

COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair® Diskus® 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol inhalation powder DBP A C G	Advair® HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol inhalation powder A G	AirDuo® Digihaler™ 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder DBP A G	AirDuo® RespiClick® 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder DBP A G	Breo® Ellipta® 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder DBP A G	Dulera® 200/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate DBP A	Symbicort® 80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate DBP A C G	Wixela™ Inhub™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate DBP A C
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COMBINATION MEDICATIONS

contain both long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Anoro® Ellipta® 82.5/25 mcg umeclidinium and vilanterol inhalation powder DBP C	Bevespi Aerosphere® 3/4.8 mcg glycopyrrolate and formoterol fumarate DBP C	Duaklir® Pressair™ 600, 12 mcg aclidinium bromide and formoterol fumarate DBP C
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COMBINATION MEDICATIONS

contain inhaled corticosteroid, long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Trelegy® Ellipta® 200/82.5/25 mcg fluticasone furoate, umeclidinium and vilanterol inhalation powder DBP A C	Breztri Aerosphere™ 160/3/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate DBP C
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BIOLOGICS

target cells and pathways that cause airway inflammation, delivered by injection or IV

Cinqair® reslizumab A	Dupixent® dupilumab A	Fasenra® bevalizumab A	Nucala® mepolizumab A	Tezspire™ tezepelumab-ekko A	Xolair® omalizumab A
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BRONCHIAL THERMOPLASTY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities.
www.bcfors asthma.com

PDE4 INHIBITORS

ease lung inflammation and reduce exacerbations

Daliresp® 250, 500 mcg roflumilast C

SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir® Digihaler® 90 mcg albuterol sulfate inhalation powder DRB A	ProAir® RespiClick® 90 mcg albuterol sulfate inhalation powder DRB A	Proventil® HFA 90 mcg albuterol sulfate DRB A G	Ventolin® HFA 90 mcg albuterol sulfate DRB A G	Xopenex® HFA 45 mcg levalbuterol tartrate A G
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent® Diskus® 50 mcg salmeterol xinafoate inhalation powder DRB A C	Striverdi® RespiMat® 2.5 mcg olodaterol hydrochloride DRB C
--	--

INHALED CORTICOSTEROIDS

relieve inflammation in the airways, but they do not relieve symptoms of coughing, wheezing or shortness of breath

Alvesco® HFA 80, 180 mcg ciclesonide DRB A	ArmonAir® Digihaler® 55, 113, 232 mcg fluticasone propionate inhalation powder DRB A	Arnuity® Ellipta® 50, 100, 200 mcg fluticasone furoate inhalation powder DRB A	Asmanex® HFA 50, 100, 200 mcg mometasone furoate DRB A	Asmanex® Twisthaler® 110, 220 mcg mometasone furoate inhalation powder DRB A	Flovent® Diskus® 50, 100, 250 mcg fluticasone propionate inhalation powder DRB A	Flovent® HFA 44, 110, 220 mcg fluticasone propionate DRB A	Pulmicort Flexhaler® 90, 180 mcg budesonide inhalation powder DRB A	QVAR® Redihaler™ 40, 80 mcg beclomethasone dipropionate DRB A
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MUSCARINIC ANTAGONISTS

relieve cough, sputum production, wheezing and shortness of breath associated with chronic lung diseases

Atrovent® HFA 17 mcg ipratropium bromide DRB C	Increase® Ellipta® 12.5 mcg umedclidium inhalation powder DRB C	Spiriva® HandiHaler® 18 mcg tiotropium bromide inhalation powder C	Spiriva® Respimat® 1.25, 2.5 mcg tiotropium bromide DRB A G	Tudorza® Pressair™ 600 mcg aclidinium bromide inhalation powder DRB C
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COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent® RespiMat® 20/100 mcg tiotropium bromide and albuterol DRB C

COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair Diskus® 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DRB A C G	Advair® HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate DRB A G	AirDuo® RespiClick® 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder DRB A	AirDuo® RespiClick® 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder DRB A G	Breo® Ellipta® 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder DRB A G	Dulera® 50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate DRB A	Symbicort® 80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate DRB A C G	Wixela® Inhub™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder DRB A C	Anoro® Ellipta® 62.5/25 mcg umedclidium and vilanterol inhalation powder DRB C	Besepi Aerosphere® 9/4.8 mcg glycopyrrolate and formoterol fumarate DRB C	Duaklir® Pressair™ 400, 12 mcg aclidinium bromide and formoterol fumarate DRB C	Stiolto® RespiMat® 2.5/2.5 mcg tiotropium bromide and olodaterol DRB C	Trelegy® Ellipta® 200/12.5/25 mcg, 100/62.5/25 mcg fluticasone furoate, umedclidium and vilanterol inhalation powder DRB A C	Breztri Aerosphere™ 160/9/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate DRB C
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COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA) and short-acting muscarinic antagonist (LAMA)

Combivent® RespiMat® 20/100 mcg tiotropium bromide and albuterol DRB C

BIOLOGICS

target cells and pathways that cause airway inflammation; delivered by injection or IV

Cinqair® reslizumab A	Dupixent® dupilumab A	Fasenra™ bambucumab A	Nucala® mepolizumab A	Tezspire™ tezepelumab-ekko A	Xolair® omalizumab A
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BRONCHIAL THERMOPLASTY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities.
www.bcforgasthma.com
A



PDE4 INHIBITORS

ease lung inflammation and reduce exacerbations

Daliresp® 250, 500 mcg roflumilast C
--

SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir[®] Digihaler[®] 90 mcg albuterol sulfate inhalation powder 	ProAir[®] RespiClick[®] 90 mcg albuterol sulfate inhalation powder 	Proventil[®] HFA 90 mcg albuterol sulfate 	Ventolin[®] HFA 90 mcg albuterol sulfate 	Xopenex[®] 45 mcg levalbuterol tartrate 
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent[®] Diskus[®] 50 mcg salmeterol xinafoate inhalation powder 	Striverdi[®] Respimat[®] 2.5 mcg olodaterol hydrochloride 
--	--

INHALED CORTICOSTEROIDS

relieve coughing, wheezing or shortness of breath

Alvesco[®] HFA 80, 160 mcg ciclesonide 	ArmonAir[®] Digihaler[®] 55, 113, 232 mcg fluticasone propionate inhalation powder 	Arnuity[®] Ellipta[®] 50, 100, 200 mcg fluticasone furoate inhalation powder 	Asmanex[®] HFA 50, 100, 200 mcg mometasone furoate 	Asmanex[®] Twisthaler[®] 110, 220 mcg mometasone furoate inhalation powder 	Flovent[®] Diskus[®] 50, 100, 250 mcg fluticasone propionate inhalation powder 	Flovent[®] HFA 44, 110, 220 mcg fluticasone propionate 	Pulmicort Flexhaler[®] 90, 180 mcg budesonide inhalation powder 	QVAR[®] Redihaler[™] 40, 80 mcg beclomethasone dipropionate 
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MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheeze and chest tightness associated with chronic lung diseases

Atrovent[®] HFA 17 mcg ipratropium bromide 	Increase[®] Ellipta[®] 82.5 mcg umedclidium inhalation powder 	Spiriva[®] HandiHaler[®] 18 mcg tiotropium bromide inhalation powder 	Spiriva[®] Respimat[®] 1.25, 2.5 mcg tiotropium bromide 	Tudorza[®] Pressair[™] 600 mcg acclidium bromide inhalation powder 
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COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent[®] Respimat[®] 20/100 mcg ipratropium bromide and albuterol 
--

COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair Diskus[®] 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol inhalation powder 	Advair[®] HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate 	AirDuo[®] Digihaler[™] 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder 	AirDuo[®] RespiClick[™] 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder 	Breo[®] Ellipta[®] 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder 	Dulera[®] 50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate 	Symbicort[®] 80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate 	Wixela[™] Inheor[®] 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate 
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COMBINATION MEDICATIONS

contain both long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Anoro[®] Ellipta[®] 82.5/25 mcg umedclidium and vilanterol inhalation powder 	Bevespi Aerosphere[®] 9/4.8 mcg glycopyrrolate and formoterol fumarate 	Duaklir[®] Pressair[™] 400, 12 mcg acclidium bromide and formoterol fumarate 	Stiolto[™] Respimat[®] 2.5/2.5 mcg tiotropium bromide and olodaterol 
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COMBINATION MEDICATIONS

contain inhaled corticosteroid, long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Alegry[®] Ellipta[®] 80/2.5/25 mcg, 160/3/4.8 mcg budesonide, glycopyrrolate and vilanterol inhalation powder 	Breztri Aerosphere[™] 160/3/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate 
--	--

BIOLOGICS

target cells and pathways that cause airway inflammation; delivered by injection or IV

Cinqair[®] reslizumab 	Dupixent[®] dupilumab 	Fasenra[™] bernalzumab 	Nucala[®] mepolizumab 	Tezspire[™] tezepelumab-ekko 	Xolair[®] omalizumab 
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BRONCHIAL THERMY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities.



www.bronchothermy.com

PHOSPHODIESTERASE INHIBITORS

ease lung inflammation and reduce exacerbations

Daliresp[®] 250, 500 mcg roflumilast 
--



SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir® Digihaler® 90 mcg albuterol sulfate inhalation powder Ⓜ ⓐ	ProAir® RespiClick® 90 mcg albuterol sulfate inhalation powder Ⓜ ⓐ	Proventil® HFA 90 mcg albuterol sulfate Ⓜ ⓐ ⓐ	Ventolin® HFA 90 mcg albuterol sulfate Ⓜ ⓐ ⓐ	Xopenex® HFA 45 mcg levalbuterol tartrate Ⓜ ⓐ ⓐ
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent® Diskus® 50 mcg salmeterol xinafoate inhalation powder Ⓜ ⓐ ⓐ	Striverdi® RespiMat® 2.5 mcg olodaterol hydrochloride Ⓜ ⓐ ⓐ
--	--

INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they decrease the frequency and severity of symptoms of coughing, wheezing or shortness of breath

Alvesco® HFA 80, 180 mcg ciclesonide Ⓜ ⓐ	Asmanex® Twister® 110, 220 mcg mometasone furoate inhalation powder Ⓜ ⓐ	Flovent® Diskus® 50, 100, 250 mcg fluticasone propionate inhalation powder Ⓜ ⓐ	Flovent® HFA 44, 110, 220 mcg fluticasone propionate Ⓜ ⓐ	Pulmicort Flexhaler® 90, 180 mcg budesonide inhalation powder Ⓜ ⓐ	QVAR® Redihaler™ 40, 80 mcg beclomethasone dipropionate Ⓜ ⓐ
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MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheeze and chest tightness associated with chronic lung diseases

Atrovent® HFA 17 mcg ipratropium bromide Ⓜ ⓐ ⓐ	Increase® Ellipta® 12.5 mcg umecidinium inhalation powder Ⓜ ⓐ ⓐ	Spiriva® HandiHaler® 18 mcg tiotropium bromide inhalation powder ⓐ	Spiriva® RespiMat® 1.25, 2.5 mcg tiotropium bromide Ⓜ ⓐ ⓐ	Tudorza® Pressair™ 600 mcg acclidium bromide inhalation powder Ⓜ ⓐ
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COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent® RespiMat® 20/100 mcg ipratropium bromide and albuterol Ⓜ ⓐ ⓐ
--

COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair Diskus® 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder Ⓜ ⓐ ⓐ ⓐ	Advair® HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate Ⓜ ⓐ ⓐ	AirDuo® Digihaler™ 55/14, 112/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder Ⓜ ⓐ	AirDuo® RespiClick® 55/14, 112/14, 232/14 mcg fluticasone propionate and salmeterol xinafoate inhalation powder Ⓜ ⓐ	Breo® Ellipta® 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder Ⓜ ⓐ ⓐ	Dulera® 50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate Ⓜ ⓐ	Symbicort® 80/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate Ⓜ ⓐ ⓐ ⓐ	Wixela™ Inhub™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhalation powder Ⓜ ⓐ ⓐ
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COMBINATION MEDICATIONS

contain both long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Anoro® Ellipta® 62.5/25 mcg umecidinium and vilanterol inhalation powder Ⓜ ⓐ ⓐ	Bevespi Aerosphere® 9/4.8 mcg glycopyrrolate and formoterol fumarate Ⓜ ⓐ ⓐ	Duaklir® Pressair™ 400, 12 mcg acclidium bromide and formoterol fumarate Ⓜ ⓐ ⓐ	Stiolto® RespiMat® 2.5/2.5 mcg tiotropium bromide and olodaterol Ⓜ ⓐ ⓐ	Trelegy® Ellipta® 200/12.5/25 mcg, 100/6.25/25 mcg fluticasone furoate, umecidinium and vilanterol inhalation powder Ⓜ ⓐ ⓐ	Breztri Aerosphere™ 160/9/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate Ⓜ ⓐ ⓐ
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BIOLOGICS

target cells and pathways that cause airway inflammation; delivered by injection or IV

Cinqair® reslizumab ⓐ	Dupixent® dupilumab ⓐ	Fasenra™ bambucumab ⓐ	Nucala® mepolizumab ⓐ	Tezspire™ tezepelumab-ekko ⓐ	Xolair® omalizumab ⓐ
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BRONCHIAL THERMOPLASTY

A minimally invasive procedure that uses mild heat to reduce airway smooth muscle, leading to fewer severe asthma flares, ER visits, and days lost from activities.
www.bronchoasthma.com



PDE4 INHIBITORS

ease lung inflammation and reduce exacerbations

Daliresp® 250, 500 mcg roflumilast ⓐ

COPD- Triple Therapy - Dual Therapy = ICS

Do we Need ICS?

Pros

- Potential impact on exacerbations
- Biological plausibility for certain phenotypes
 - Reduce airway inflammation
 - Upregulate beta receptors
 - Mild improvement in FEV₁ (initial sustained)

Cons

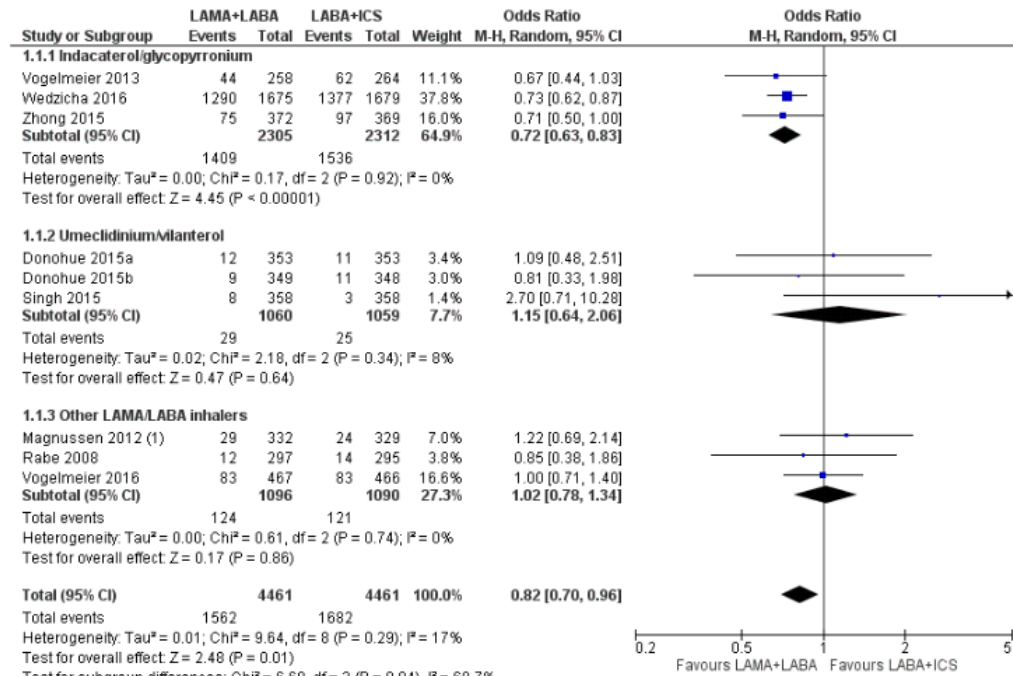
- Skin thinning and easy bruising
- Oral thrush, and hoarseness
- Increased risk of pneumonia
- Osteoporosis
- Early onset diabetes
- Cataracts
- Cost
- No effect on FEV₁ decline

LAMA-LABA vs. LABA - ICS

More Exacerbations

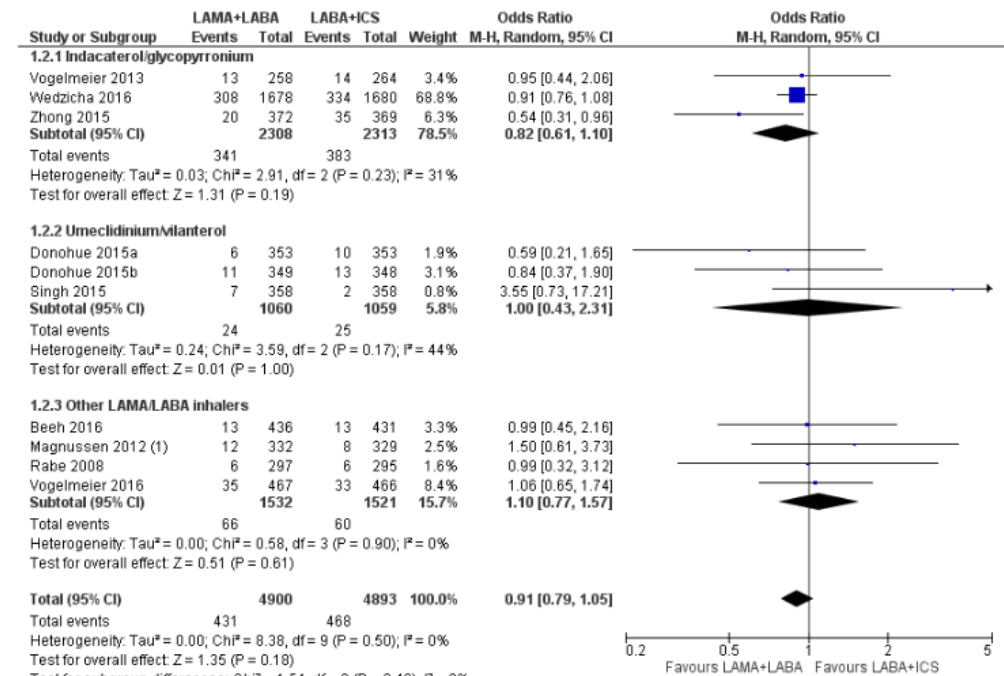
Likely More Adverse Effects

Figure 3. Forest plot of comparison: 1 Long-acting muscarinic antagonist (LAMA) plus long-acting beta-agonist (LABA) versus LABA plus ICS (inhaled corticosteroid), outcome: 1.1 Exacerbation.



Footnotes
(1) Magnussen 2012: Due to a unit-of-analysis error introduced by crossover design, confidence intervals might be wider in our analysis.

Figure 4. Forest plot of comparison: 1 Long-acting muscarinic antagonist (LAMA) plus long-acting beta-agonist (LABA) versus LABA plus ICS (inhaled corticosteroid), outcome: 1.2 Serious adverse events.



Footnotes
(1) Magnussen 2012: Due to a unit-of-analysis error introduced by crossover design, confidence intervals might be wider in our analysis.

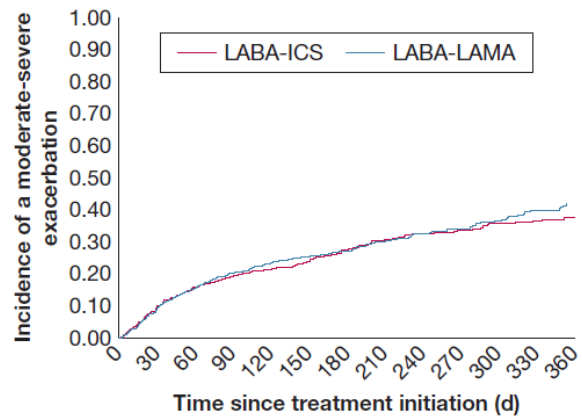


Figure 2 – One-year cumulative incidence of the first moderate or severe COPD exacerbation, comparing current treatment with LABA-LAMA and LABA-ICS, estimated using the Kaplan-Meier method. See Figure 1 legend for expansion of abbreviations.

TABLE 3] Crude and Adjusted Hazard Ratios of Severe Pneumonia Comparing LABA-LAMA Initiation With LABA-ICS Initiation in Patients With COPD in the First Year After Treatment Initiation, From the As-Treated Analysis

Treatment Exposure	No. of Patients	No. of Events	Person-Years	Rate per 100 per Year	Crude ^a HR	Adjusted ^b HR (95% CI)
As-treated exposure						
LABA-LAMA	1,977	32	629	5.1	0.66	0.66 (0.41-1.05)
LABA-ICS	1,977	41	535	7.7	1.00	1.00 (Reference)
On-treatment exposure^c						
LABA-LAMA	1,977	49	907	5.4	0.65	0.66 (0.48-0.92)
LABA-ICS	1,977	143	1,778	8.0	1.00	1.00 (Reference)

See Table 1 and 2 legends for expansion of abbreviations.

^aCrude, after matching on high-dimensional propensity scores and sex.

^bAfter matching on high-dimensional propensity scores and sex, adjusted further for the decile of propensity score.

^cOn-treatment exposure based on analysis of current use during the entire 1-year follow-up, allowing patients to switch or add treatments.

DOI: 10.1016/j.chest.2019.03.005

LAMA-LABA vs. LABA – ICS – Real world

SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir[®] Digihaler[™]
90 mcg
albuterol sulfate
inhalation powder
DRG A



ProAir[®] RespiClick[™]
90 mcg
albuterol sulfate
inhalation powder
DRG A



Proventil[®] HFA
90 mcg
albuterol sulfate
DRG A G



Ventolin[®] HFA
90 mcg
albuterol sulfate
DRG A G



Xopenex[®] HFA[®]
45 mcg
levalbuterol tartrate
A G



LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent[®] Diskus[®]
50 mcg
salmeterol xinafoate
inhalation powder
DRG A C



Striverdi[®] Respimat[®]
2.5 mcg
olodaterol hydrochloride
DRG C



INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they do not relieve sudden symptoms of coughing, wheezing or shortness of breath

Alvesco[®] HFA
80, 160 mcg
ciclesonide
DRG A



ArmonAir[®] Digihaler[™]
55, 113, 232 mcg
fluticasone propionate
inhalation powder
DRG A



Arnuity[®] Ellipta[®]
50, 100, 200 mcg
fluticasone furoate
inhalation powder
DRG A



Asmanex[®] HFA
50, 100, 200 mcg
mometasone furoate
DRG A



Asmanex[®] Twisthaler[®]
110, 220 mcg
mometasone furoate
inhalation powder
DRG A



Flovent[®] Diskus[®]
50, 100, 250 mcg
fluticasone propionate
inhalation powder
DRG A



Flovent[®] HFA
44, 110, 220 mcg
fluticasone propionate
DRG A



Pulmicort Flexhaler[®]
90, 180 mcg
budesonide
inhalation powder
DRG A



QVAR[®] Redihaler[™]
40, 80 mcg
beclomethasone dipropionate
DRG A



MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheezes and chest tightness associated with chronic lung diseases

Short-acting
Atrovent[®] HFA
17 mcg
ipratropium bromide
DRG C



Long-acting
Incruse[®] Ellipta[®]
82.5 mcg
umeclidinium
inhalation powder
DRG C



Spiriva[®] HandiHaler[®]
18 mcg
tiotropium bromide
inhalation powder
G



Spiriva[®] Respimat[®]
1.25, 2.5 mcg
tiotropium bromide
DRG A C



Tudorza[™] Pressair[™]
600 mcg
acetylcholinesterase
inhalation powder
DRG C



COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent[®] Respimat[®]
10/100 mcg
vixelanguin bromide and albuterol
DRG C



COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair[®] Diskus[®]
100/50, 250/50, 500/50 mcg
fluticasone propionate and salmeterol xinafoate
DRG A C G



Advair[®] HFA
45/21, 115/21, 230/21 mcg
fluticasone propionate and salmeterol xinafoate
DRG A G



AirDuo[®] Digihaler[™]
55/14, 113/14, 232/14 mcg
fluticasone propionate and salmeterol xinafoate
DRG A



AirDuo[®] RespiClick[™]
55/14, 113/14, 232/14 mcg
fluticasone propionate and salmeterol xinafoate
DRG A G



Breo[®] Ellipta[®]
100/25, 200/25 mcg
fluticasone furoate and vilanterol
inhalation powder
DRG A G



Dulera[®]
50/5, 100/5, 200/5 mcg
mometasone furoate and formoterol fumarate dihydrate
DRG A



Symbicort[®]
60/4.5, 160/4.5 mcg
budesonide and formoterol fumarate dihydrate
DRG A C G



Wixela[™] Inhub[™]
100/50, 250/50, 500/50 mcg
fluticasone propionate and salmeterol xinafoate
DRG A C



Anoro[®] Ellipta[®]
62.5/25 mcg
umeclidinium and vilanterol
inhalation powder
DRG C



Bevespi Aerosphere[®]
9/4.8 mcg
glycopyrrate and formoterol fumarate
DRG C



Duakir[®] Pressair[™]
600, 12 mcg
acetylcholinesterase and formoterol fumarate
DRG C



Stiolto[®] Respimat[®]
2.1/2.5 mcg
tiotropium bromide and olodaterol
DRG C



Trelegy[®] Ellipta[®]
200/62.5/25 mcg
fluticasone furoate, umeclidinium and vilanterol
inhalation powder
DRG A C



Bretri Aerosphere[™]
160/9/4.8 mcg
budesonide, glycopyrrate and formoterol fumarate
DRG C



contain inhaled corticosteroid, long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)





Controllers:

- Inhaled steroids: Qvar, Arnuity, Asmanex, Flovent
- ICS/LABAs: Dulera, Breo, Advair, AirDuo, Symbicort, Wixela
- LAMA: Spiriva
- Triple therapy approved for asthma: Trelegy (200 mg fluticasone dose)



Relievers:

- Short acting beta2-agonist: Albuterol (brands are Proventil, ProAir, Ventolin); Levalbuterol (brand is Xopenex)
- Short acting anti-muscarinic: Ipratropium bromide- Atrovent
- Combination SABA/SAMA – Duoneb solution for nebulizer, Combivent inhaler

Non – Pharmacologic treatment

**Smoking
Cessation**

Inhaler training

**Pulmonary
rehab – for pts
in group B & E**

**Oxygen therapy
for resting
hypoxia**

**Vaccination: flu,
pneumococcal,
COVID, RSV**

**Spirometry
yearly**

Exercise Training has benefits: increase efficiency, less sedentary

Pulmonary Rehabilitation aims to

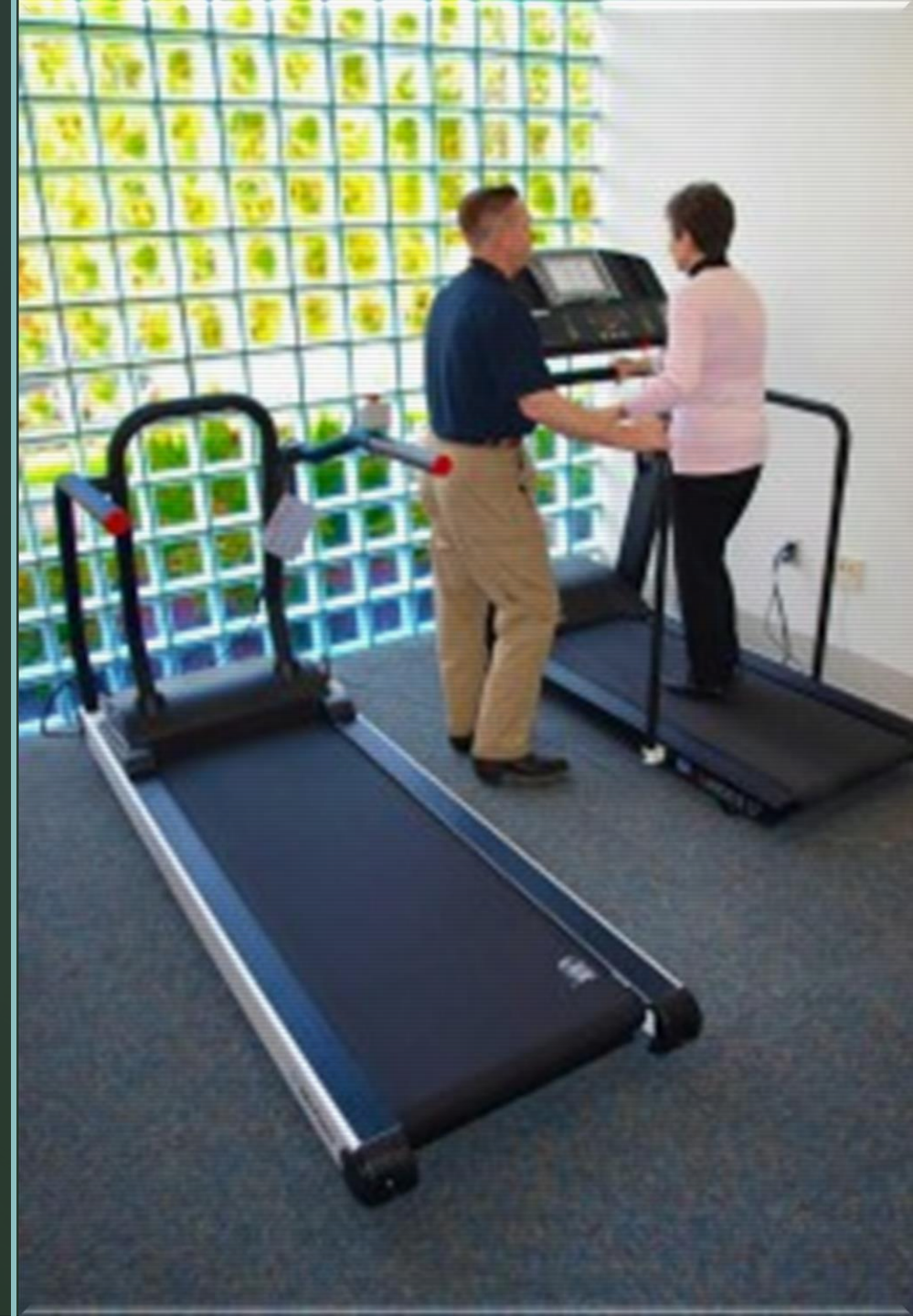
- 1) Target extra-pulmonary effects of lung disease
- 2) Ensure patients learn optimally manage disease
- 3) Provide patient with skills to lead healthy life

Physical Activity

Nutrition

Emotions/psychological well-being

***effective after exacerbations



Lung Cancer screening

50-80 year old

>20 pack year history

quit <15 years ago



100

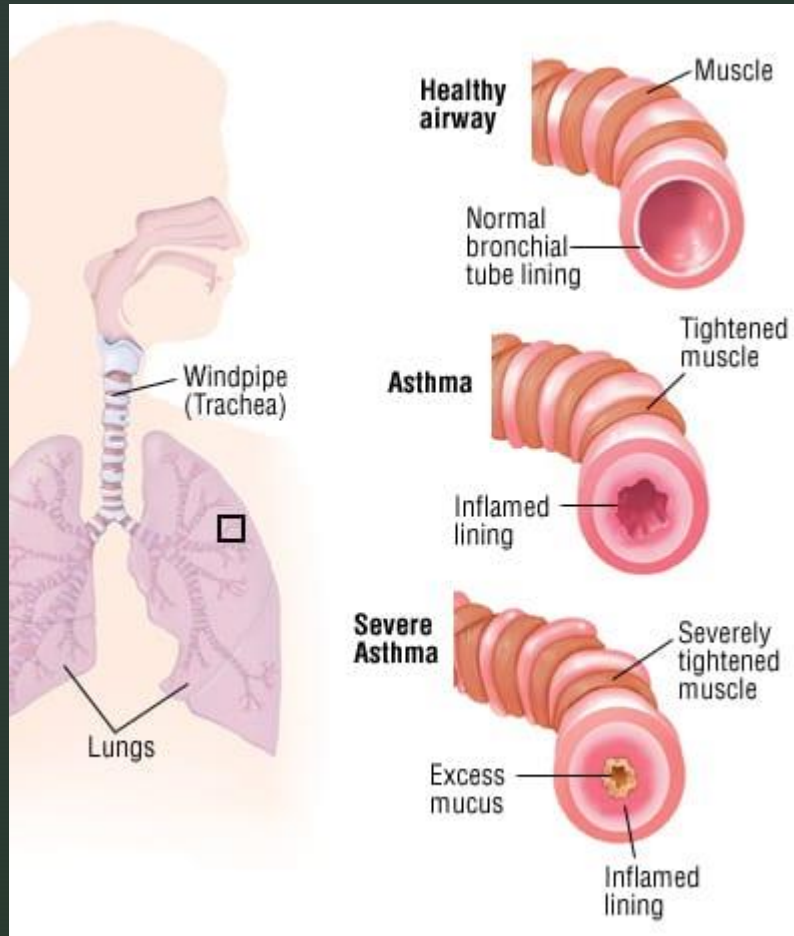


DARTH VADER

Making asthma scary since '77

ASTHMA

Chronic airway inflammation Asthma



What happens to your lungs when you have asthma

The diagram shows a human torso with the lungs and airways. The left lung is shown in a normal state, and the right lung is shown in a state with asthma. The asthma-affected lung shows narrowed airways, tightened muscles, and mucus clogging the airways.

LUNG WITHOUT ASTHMA

- Muscles relaxed
- Normal airways
- Normal amount of mucus

LUNG WITH ASTHMA

- Muscles tighten
- Airways swell
- Mucus clogs the airways
- Lungs have difficulty moving air in and out

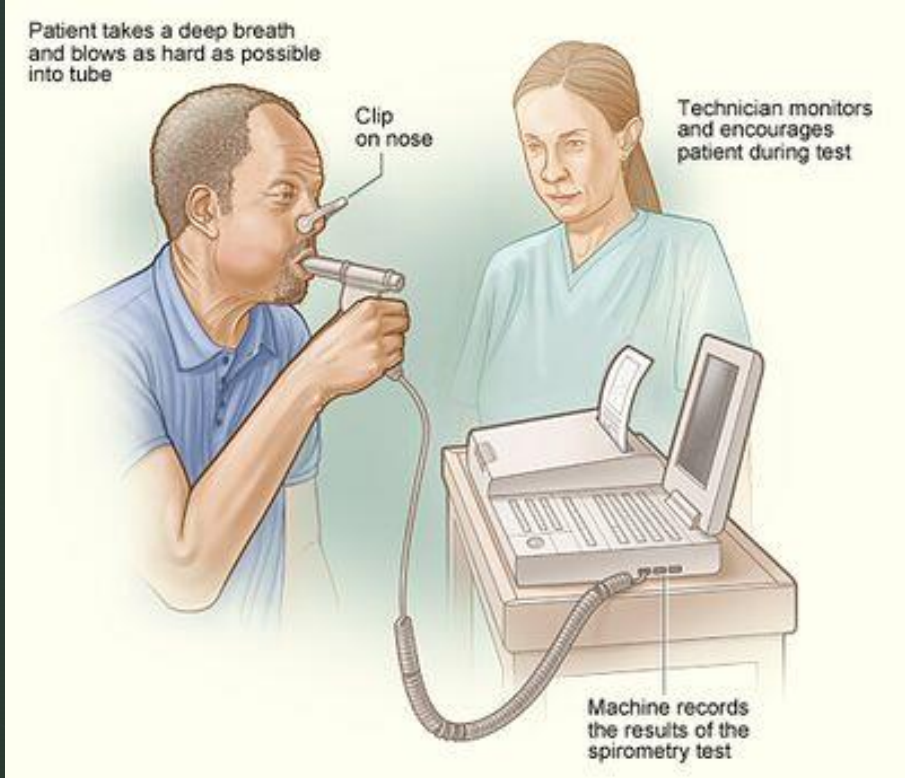
aaafa Asthma and Allergy Foundation of America
aafa.org



Asthma Triggers



You better be ready to blow the best breaths you have ever breathed in your entire life.



Patient takes a deep breath and blows as hard as possible into tube

Clip on nose

Technician monitors and encourages patient during test

Machine records the results of the spirometry test

Variable flow obstruction seen in asthma

This can be measured by spirometry with a baseline measurement and testing after bronchodilator is administered

Significant if-

- FEV₁ increases >10% and at least 200 ml

		Meas	LLN	Pred	ULN	%	Post	%	%Chg
FVC	L	3.79	2.92	3.65	4.41	103.6	3.78	103.4	-0.2
FEV1	L	2.32	2.47	3.08	3.67	75.3	2.87	93.1	23.7
FEV1/FVC	%	61	73	85	94	72.3	76	89.6	23.9

Guidelines



National Asthma
Education and
Prevention Program
Coordinating
Committee
(NAEPPCC) (the
NAEPP Guidelines)

- Began in 1989
- Big update in 2007, then nothing until 2021!

Global Initiative for
Asthma (GINA)

- Began in 1993
- Most recent update in 2022

Diagnosing Asthma



Pulmonary function testing



CLINICAL SYNDROME!

(can be confirmed/supported
with testing)

Spirometry - positive for
asthma if airway obstruction
is found with reversibility, and
significant bronchodilator
response




Bronchial Provocation
Testing – Methacholine
Challenge or Mannitol
Challenge

FeNO – Fractionated exhaled
nitric oxide - >50 ppb often
seen in poorly controlled
asthma.



Labs

Elevated IgE
peripheral eosinophilia

Components of Severity		Classification of Asthma Severity ≥12 years of age			
		Intermittent	Persistent		
			Mild	Moderate	Severe
Impairment Normal FEV ₁ /FVC: 8–19 yr 85% 20–39 yr 80% 40–59 yr 75% 60–80 yr 70%	Symptoms	≤2 days/week	>2 days/week but not daily	Daily	Throughout the day
	Nighttime awakenings	≤2x/month	3–4x/month	>1x/week but not nightly	Often 7x/week
	Short-acting beta ₂ -agonist use for symptom control (not prevention of EIB)	≤2 days/week	>2 days/week but not daily, and not more than 1x on any day	Daily	Several times per day
	Interference with normal activity	None	Minor limitation	Some limitation	Extremely limited
	Lung function	<ul style="list-style-type: none"> • Normal FEV₁ between exacerbations • FEV₁ >80% predicted • FEV₁/FVC normal 	<ul style="list-style-type: none"> • FEV₁ >80% predicted • FEV₁/FVC normal 	<ul style="list-style-type: none"> • FEV₁ >60% but <80% predicted • FEV₁/FVC reduced 5% 	<ul style="list-style-type: none"> • FEV₁ <60% predicted • FEV₁/FVC reduced >5%
Risk	Exacerbations requiring oral systemic corticosteroids	0–1/year (see note)	≥2/year (see note) 		
		Consider severity and interval since last exacerbation. Frequency and severity may fluctuate over time for patients in any severity category.  			
Recommended Step for Initiating Treatment (See figure 4–5 for treatment steps.)		Step 1	Step 2	Step 3	Step 4 or 5
		and consider short course of oral systemic corticosteroids In 2–6 weeks, evaluate level of asthma control that is achieved and adjust therapy accordingly.			

Source: NAEPP Asthma Guidelines, 2018

Intermittent Asthma

Management of Persistent Asthma in Individuals Ages 12+ Years

Treatment	STEP 1	STEP 2	STEP 3	STEP 4	STEP 5	STEP 6 [■]
Preferred	PRN SABA	Daily low-dose ICS and PRN SABA or PRN concomitant ICS and SABA [▲]	Daily and PRN combination low-dose ICS-formoterol [▲]	Daily and PRN combination medium-dose ICS-formoterol [▲]	Daily medium-high dose ICS-LABA + LAMA and PRN SABA [▲]	Daily high-dose ICS-LABA + oral systemic corticosteroids + PRN SABA
Alternative		Daily LTRA* and PRN SABA or Cromolyn,* or Nedocromil,* or Zileuton,* or Theophylline,* and PRN SABA	Daily medium-dose ICS and PRN SABA or Daily low-dose ICS-LABA, or daily low-dose ICS + LAMA, [▲] or daily low-dose ICS + LTRA,* and PRN SABA or Daily low-dose ICS + Theophylline* or Zileuton,* and PRN SABA	Daily medium-dose ICS-LABA or daily medium-dose ICS + LAMA, and PRN SABA [▲] or Daily medium-dose ICS + LTRA,* or daily medium-dose ICS + Theophylline,* or daily medium-dose ICS + Zileuton,* and PRN SABA	Daily medium-high dose ICS-LABA or daily high-dose ICS + LTRA,* and PRN SABA	
		Steps 2-4: Conditionally recommend the use of subcutaneous immunotherapy as an adjunct treatment to standard pharmacotherapy in individuals ≥ 5 years of age whose asthma is controlled at the initiation, build up, and maintenance phases of immunotherapy [▲]			Consider adding Asthma Biologics (e.g., anti-IgE, anti-IL5, anti-IL5R, anti-IL4/IL13)**	

Adults & adolescents 12+ years

Personalized asthma management:
Assess, Adjust, Review response



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

Symptoms
Exacerbations
Side-effects
Lung function
Patient satisfaction

Treatment of modifiable risk factors & comorbidities
Non-pharmacological strategies
Education & skills training
Asthma medications

A holistic approach – not just symptom control

ICS-containing controller is recommended across all severities to reduce exacerbation risk

“Preferred” and “other” options are provided at each step, based on evidence

Asthma medication options:
Adjust treatment up and down for individual patient needs

PREFERRED CONTROLLER

to prevent exacerbations and control symptoms

Other controller options

PREFERRED RELIEVER

Other reliever option

STEP 1

As-needed low dose ICS-formoterol*
Low dose ICS taken whenever SABA is taken†

STEP 2

Daily low dose inhaled corticosteroid (ICS), or as-needed low dose ICS-formoterol*
Leukotriene receptor antagonist (LTRA), or low dose ICS taken whenever SABA taken †

STEP 3

Low dose ICS-LABA

Medium dose ICS, or low dose ICS+LTRA #

STEP 4

Medium dose ICS-LABA

High dose ICS, add-on tiotropium, or add-on LTRA #

STEP 5

High dose ICS-LABA
Refer for phenotypic assessment ± add-on therapy, e.g. tiotropium, anti-IgE, anti-IL5/5R, anti-IL4R

Add low dose OCS, but consider side-effects

As-needed low dose ICS-formoterol*

As-needed low dose ICS-formoterol for patients prescribed maintenance and reliever therapy ‡

As-needed short-acting β₂-agonist (SABA)

* Off-label; data only with budesonide-formoterol (bud-form)
† Off-label; separate or combination ICS and SABA inhalers

‡ Low-dose ICS-formoterol is the reliever for patients prescribed bud-form or BDP-form maintenance and reliever therapy
Consider adding HDM SLIT for sensitized patients with allergic rhinitis and FEV₁ >70% predicted

See 2019 GINA Severe Asthma Pocket Guide for more details about Steps 4–5

Maintenance OCS is not a preferred option at Step 5 because of serious side-effects

SABA is not a preferred reliever because of the risks of SABA-only treatment, including if adherence is poor



SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

<p>ProAir® Digihaler™ 90 mcg albuterol sulfate inhalation powder DR A</p> 	<p>ProAir® RespiClick® 90 mcg albuterol sulfate inhalation powder DR A</p> 	<p>Proventil® HFA 90 mcg albuterol sulfate DR A G</p> 	<p>Ventolin® HFA 90 mcg albuterol sulfate DR A G</p> 	<p>Xopenex® HFA 45 mcg levalbuterol tartrate A G</p> 
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

<p>Serevent® Diskus® 50 mcg salmeterol xinafoate inhalation powder DR A C</p> 	<p>Striverdi® RespiMat® 2.5 mcg olodaterol hydrochloride DR C</p> 
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INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they do not relieve sudden symptoms of coughing, wheezing or shortness of breath

<p>Alvesco® HFA 80, 180 mcg ciclesonide DR A</p> 	<p>ArmonAir® Digihaler™ 55, 113, 232 mcg fluticasone propionate inhalation powder DR A</p> 	<p>Arnuity® Ellipta® 50, 100, 200 mcg fluticasone furoate inhalation powder DR A</p> 	<p>Asmanex® HFA 50, 100, 200 mcg mometasone furoate DR A</p> 	<p>Asmanex® Twisthaler® 110, 220 mcg mometasone furoate inhalation powder DR A</p> 	<p>Flovent® Diskus® 50, 100, 250 mcg fluticasone propionate inhalation powder DR A</p> 	<p>Flovent® HFA 44, 110, 220 mcg fluticasone propionate DR A</p> 	<p>Pulmicort Flexhaler® 90, 180 mcg budesonide inhalation powder DR A</p> 	<p>QVAR® Redihaler™ 40, 80 mcg beclomethasone dipropionate DR A</p> 
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MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheezes and chest tightness associated with chronic lung diseases

<p>Short-acting</p> <p>Atrovent® HFA 17 mcg ipratropium bromide DR C</p> 	<p>Long-acting</p> <p>Incruse® Ellipta® 82.5 mcg umeclidinium inhalation powder DR C</p> 	<p>Spiriva® HandiHaler® 18 mcg tiotropium bromide inhalation powder G</p> 	<p>Spiriva® RespiMat® 1.25, 2.5 mcg tiotropium bromide DR A C</p> 	<p>Tudorza™ Pressair™ 600 mcg aclidinium bromide inhalation powder DR C</p> 
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COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent® RespiMat®
10/100 mcg ipratropium bromide and albuterol
DR C



COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

<p>Advair® Diskus® 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol inhalation powder DR A C G</p> 	<p>Advair® HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate DR A G</p> 	<p>AirDuo® Digihaler™ 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder DR A</p> 	<p>AirDuo® RespiClick® 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder DR A G</p> 	<p>Breo® Ellipta® 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder DR A G</p> 	<p>Dulera® 50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate DR A</p> 	<p>Symbicort® 60/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate DR A C G</p> 	<p>Wixela™ Inhub™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhaled pressurized canister DR A C</p> 
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COMBINATION MEDICATIONS

contain both long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

<p>Anoro® Ellipta® 32.5/25 mcg umeclidinium and vilanterol inhalation powder DR C</p> 	<p>Bevespi Aerosphere® 9/4.8 mcg glycopyrrolate and budesonide fumarate DR C</p> 	<p>Duakir® Pressair® 600, 12 mcg aclidinium bromide and formoterol fumarate DR C</p> 	<p>Stiolto® RespiMat® 2.1/2.5 mcg tiotropium bromide and olodaterol DR C</p> 	<p>Trelegy® Ellipta® 200/62.5/25 mcg, 100/62.5/25 mcg fluticasone furoate, umeclidinium and vilanterol inhalation powder DR A C</p> 	<p>Bretri Aerosphere™ 160/9/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate DR C</p> 
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Medications- Step 1

- NAEPP Guidelines: PRN SABA

Brands

Proventil, Ventolin, ProAir (all albuterol)

Xopenex (levalbuterol)

- GINA Guidelines: PRN low-dose ICS/LABA (but only formoterol!) – budesonide/formoterol most studied, but other combination with formoterol is fine

Brands

Symbicort (budesonide/formoterol)

Dulera (mometasone/formoterol)

Perforomist (nebulized formoterol – can be combined with ICS)

Medications – Step 2

NAEPP

- Daily low dose ICS, and as needed SABA for quick relief therapy

OR

- Intermittent as-needed SABA + an ICS use concomitantly (one after the other)

GINA

- Daily low- dose ICS

OR

- As needed low-dose ICS/formoterol



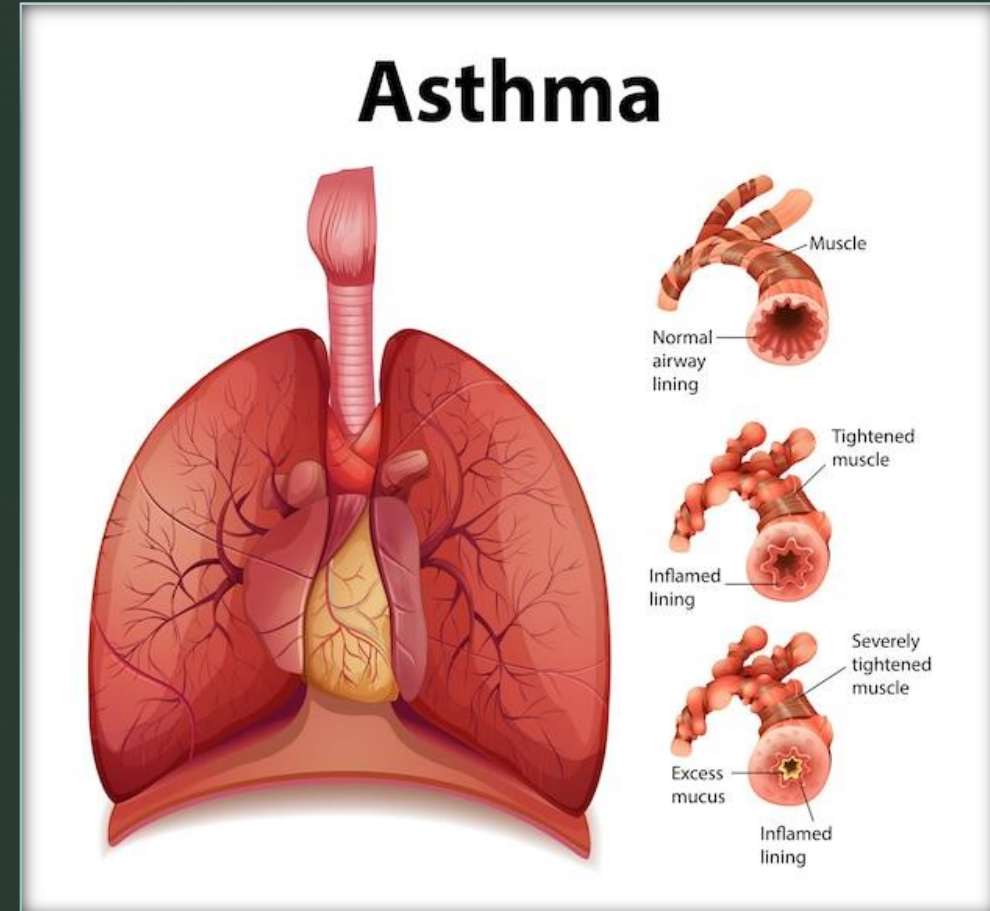
Medication Step 3

NAEPP

- Daily and PRN low-dose ICS/LABA (Also called SMART (Single Maintenance and Reliever Therapy) – suitable for Steps 3 and 4.)

GINA

- Low-dose ICS/LABA, with PRN ICS/formoterol as rescue. (Essentially this is SMART therapy too! But leaves the daily inhaler up to you if you want to use another LABA type)
- Consider adding on leukotriene modifier (LTRA) – Singulair (montelukast), Accolate (zarfirlukast), Zflo (Zileuton)



Leukotriene Modifiers

- Montelukast (Singulair) 10 mg once a day (PO)
- Zafirlukast (Accolate) 20 mg BID (PO)

Bronchoconstriction, vascular permeability, eosinophil recruitment, and chronic inflammation are mediated through the G protein-coupled activation of cysteinyl leukotriene receptors. SO – Leukotriene stimulate airway contraction, and inflammatory cascade ft. mucosal edema, mucous secretion

Montelukast and zafirlukast block cysteinyl leukotriene CysLT1 receptors, resulting in reduced eosinophilic recruitment to airway tissue.

Great for: Exercise-induced bronchospasm, asthma with allergic rhinitis, cold-exacerbated asthma, aspirin-exacerbated respiratory disease (AERD).

- Zileuton (Zyflo) 600 mg QID, or ER dosing 1200 mg BID

Works differently to other LTRAs, Specific 5-lipoxygenase inhibitor which inhibits leukotriene formation.



SHORT-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer quick relief of symptoms such as coughing, wheezing and shortness of breath for 3-6 hours

ProAir® Digihaler™ 90 mcg albuterol sulfate inhalation powder DR A	ProAir® RespiClick® 90 mcg albuterol sulfate inhalation powder DR A	Proventil® HFA 90 mcg albuterol sulfate DR A G	Ventolin® HFA 90 mcg albuterol sulfate DR A G	Xopenex® HFA 45 mcg levalbuterol tartrate A G
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LONG-ACTING BETA₂-AGONIST BRONCHODILATORS

relax tight muscles in airways and offer lasting relief of symptoms such as coughing, wheezing and shortness of breath for at least 12 hours

Serevent® Diskus® 50 mcg salmeterol xinafoate inhalation powder DR A C	Striverdi® RespiMat® 2.5 mcg olodaterol hydrochloride DR C
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INHALED CORTICOSTEROIDS

reduce and prevent swelling of airway tissue; they do not relieve sudden symptoms of coughing, wheezing or shortness of breath

Alvesco® HFA 80, 180 mcg ciclesonide DR A	ArmonAir® Digihaler™ 55, 113, 232 mcg fluticasone propionate inhalation powder DR A	Arnuity® Ellipta® 50, 100, 200 mcg fluticasone furoate inhalation powder DR A	Asmanex® HFA 50, 100, 200 mcg mometasone furoate DR A	Asmanex® Twisthaler® 110, 220 mcg mometasone furoate inhalation powder DR A	Flovent® Diskus® 50, 100, 250 mcg fluticasone propionate inhalation powder DR A	Flovent® HFA 44, 110, 220 mcg fluticasone propionate DR A	Pulmicort Flexhaler® 90, 180 mcg budesonide inhalation powder DR A	QVAR® Redihaler™ 40, 80 mcg beclomethasone dipropionate DR A
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MUSCARINIC ANTAGONISTS (ANTICHOLINERGIC)

relieve cough, sputum production, wheezes and chest tightness associated with chronic lung diseases

Short-acting Atrovent® HFA 17 mcg ipratropium bromide DR C	Long-acting Incruse® Ellipta® 82.5 mcg umeclidinium inhalation powder DR C	Spiriva® HandiHaler® 18 mcg tiotropium bromide inhalation powder G	Spiriva® RespiMat® 1.25, 2.5 mcg tiotropium bromide DR A C	Tudorza™ Pressair™ 600 mcg aclidinium bromide inhalation powder DR C
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COMBINATION MEDICATIONS

contain both short-acting beta₂-agonist and short-acting muscarinic antagonist

Combivent® RespiMat® 10/100 mcg ipratropium bromide and albuterol DR C

COMBINATION MEDICATIONS

contain both inhaled corticosteroid and long-acting beta₂-agonist (LABA)

Advair® Diskus® 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol inhalation powder DR A C G	Advair® HFA 45/21, 115/21, 230/21 mcg fluticasone propionate and salmeterol xinafoate DR A G	AirDuo® Digihaler™ 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder DR A	AirDuo® RespiClick® 55/14, 113/14, 232/14 mcg fluticasone propionate and salmeterol inhalation powder DR A G	Breo® Ellipta® 100/25, 200/25 mcg fluticasone furoate and vilanterol inhalation powder DR A G	Dulera® 50/5, 100/5, 200/5 mcg mometasone furoate and formoterol fumarate dihydrate DR A	Symbicort® 60/4.5, 160/4.5 mcg budesonide and formoterol fumarate dihydrate DR A C G	Wixela™ Inhub™ 100/50, 250/50, 500/50 mcg fluticasone propionate and salmeterol xinafoate inhaled powder at Actavis Inhaler DR A C
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COMBINATION MEDICATIONS

contain both long-acting beta₂-agonist (LABA) and long-acting muscarinic antagonist (LAMA)

Anoro® Ellipta® 32.5/25 mcg umeclidinium and vilanterol inhalation powder DR C	Bevesi Aerosphere® 9/4.8 mcg glycopyrrolate and formoterol fumarate DR C	Duakir® Pressair® 600, 12 mcg aclidinium bromide and formoterol fumarate DR C	Stiolto® RespiMat® 2.1/2.5 mcg tiotropium bromide and olodaterol DR C	Trelegy® Ellipta® 200/62.5/25 mcg, 100/62.5/25 mcg fluticasone furoate, umeclidinium and vilanterol inhalation powder DR A C	Bretri Aerosphere™ 160/9/4.8 mcg budesonide, glycopyrrolate and formoterol fumarate DR C
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Medications - Step 4

NAEPP

- Daily and PRN medium-dose ICS/LABA (Also called SMART (Single Maintenance and Reliever Therapy) – suitable for Steps 3 and 4.)
- Add LTRA

GINA

- Medium-dose ICS/LABA, with PRN ICS/formoterol as rescue
- LTRA
- Consider use/addition of LAMA – tiotropium (currently the only FDA approved LAMA for asthma in the US)



Medications – Step 5

NAEPP

- Daily medium to high-dose ICS/LABA
- LTRA
- Albuterol PRN as rescue
- Add LAMA
- Consider biologics

GINA

- High-dose ICS/LABA, with PRN ICS/formoterol as rescue
- LTRA
- LAMA
- Consider biologics
- Systemic glucocorticoids



Medications - Step 6

NAEPP

- Daily medium to high-dose ICS/LABA
- LTRA
- Albuterol PRN as rescue
- Add LAMA
- Consider biologics
- Systemic glucocorticoids

Inhaler Types

Controllers/Relievers?!

- Low dose inhaled steroid + LAMA
- Low dose ICS + SABA, Approved Jan 2023- FDA approves drug combination treatment for adults with asthma
 - Airsupra (albuterol and budesonide) inhalation aerosol
 - For the as-needed ***treatment or prevention*** of bronchoconstriction and to reduce the risk of asthma attacks in patients with asthma 18 years of age and older.
 - Combination of albuterol (a beta-2 adrenergic agonist) and budesonide (a corticosteroid)
 - Dose: 2 inhalations of a combination of albuterol 90 mcg and budesonide 80 mcg per inhalation (total dose albuterol 180 mcg and budesonide 160 mcg) as needed for asthma symptoms.

It is the first combination of an inhaled corticosteroid (ICS) and a short-acting beta-agonist to be approved in the U.S. Additionally, *Airsupra is the first product containing an ICS to be approved in the U.S. as a **reliever treatment** (rather than as a controller) for asthma.*

The treatment works to relax the muscles and reduce inflammation in the lung airways to reduce the risk of severe asthma attacks.



GLUCOCORTICOIDS

“Systemic corticosteroids use is associated with increased risk of infections and cardiovascular events; of chronic conditions such as type 2 diabetes mellitus, osteoporosis, and cataracts; of metabolic effects such as weight gain; and of neuropsychiatric effects such as insomnia, depression, and behavioral disturbances.

A dose-response relationship with systemic corticosteroid exposure has been documented for many of these events.-

Even short-term use of oral corticosteroids (OCS) in a large population-based study has been associated with increased rates of sepsis, thromboembolism, and fracture within 30 days of OCS initiation. Of note, decreases in serum cortisol and markers of bone formation, and changes in white blood cell counts, are evident within hours of oral prednisone administration to healthy subjects.”

(Price, et al., 2018)

GLUCOCORTICOIDS

If you have patients needing even 1-2 bursts of steroids/year for their asthma,

**PLEASE CONSIDER SENDING TO ASTHMA
SPECIALIST!!!**

We can help with the following

- Disease education
- Inhaler training
- Exploring and mitigating comorbidities
- Advancing to higher steps of care

So, what is severe asthma?

- Asthma that requires treatment with high dose ICS + second controller and/or systemic corticosteroids to prevent it from becoming 'uncontrolled' or that remains 'uncontrolled' despite this therapy.

Defining "Uncontrolled"

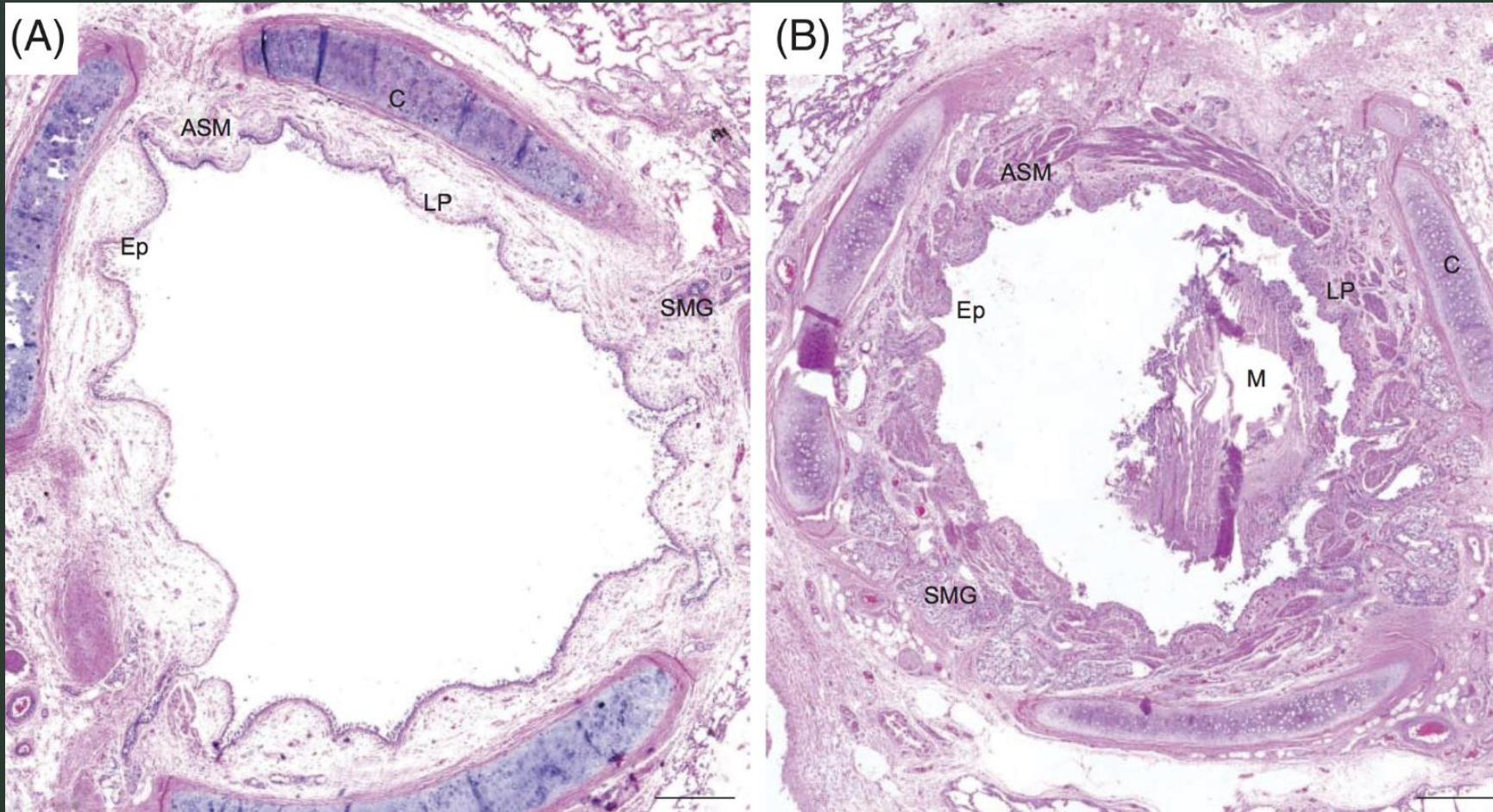
- Poor symptom control (frequent use of rescue/reliever, night time waking, missing work or school)
- Frequent severe exacerbations (≥ 2 bursts of systemic steroid in the past year, 1 ED visit or hospitalization for asthma in the past year, history of ICU stay or mechanical ventilation for asthma, or FEV1 that is $<80\%$.) Relatively large proportion of resource expenditure, **only 10% prevalence**

(Chung et al., 2013)

Chronic airway inflammation seen in Asthma

Healthy patient without asthma

Case of fatal asthma



(King et al., 2018)

Treating Severe Asthma

- First, make sure it's asthma – Not every wheeze is asthma!
 - Differentials in adults:
 - Chronic obstructive pulmonary disease
 - Bronchiectasis (Asthma treatment could make this worse!)
 - Congestive heart failure
 - Gastroesophageal reflux disease,
 - Mechanical obstruction of the airways (e.g., tumors, foreign bodies, subglottic stenosis)
 - Vocal cord dysfunction or other laryngeal dysfunction
 - Pulmonary embolism
 - Pulmonary infiltrates with eosinophilia or other parenchymal disease
 - Upper airway disease
 - Differentials in children:
 - Foreign body aspiration causing airway obstruction
 - Pneumonia/bronchiolitis
 - Cystic fibrosis
 - Bronchopulmonary dysplasia (in premature infants)
 - Primary ciliary dyskinesia syndrome
 - Immune deficiency

NOT Asthma

Severe thrush Fungal laryngitis



Severe polypoid corditis from smoking and reflux

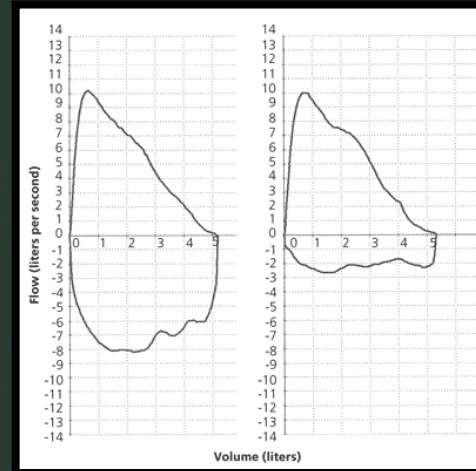




NOT Asthma

Sinusitis

Paradoxical Vocal Fold Motion



Courtesy: C. Milstein, PhD



Treating Severe Asthma

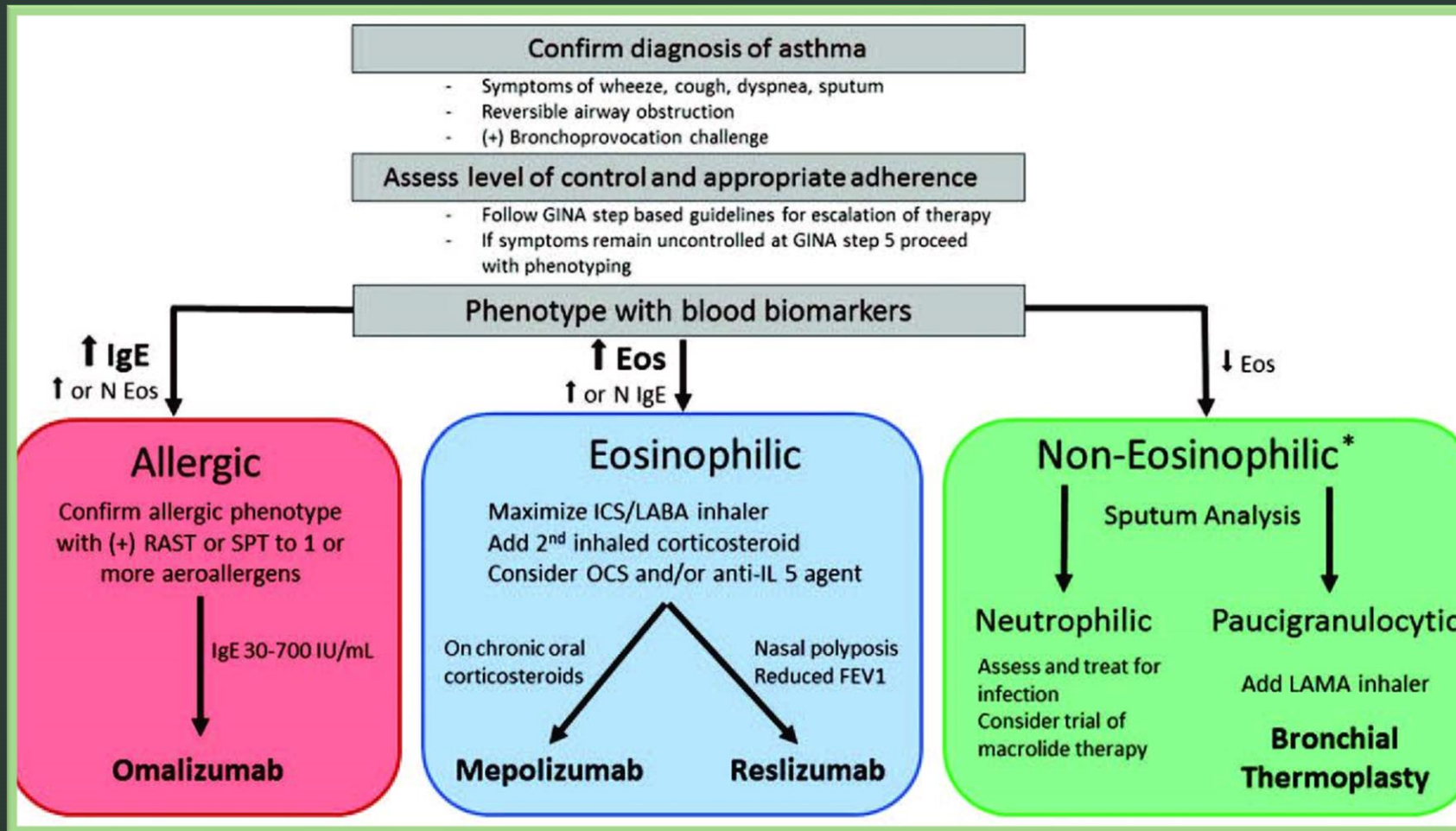
- Airway regimen optimized?
- Patients using inhalers correctly?
- Patients even HAVE their inhalers?!
- Comorbid conditions managed? (OSA, sinusitis, GERD, etc etc)

Treating Severe Asthma

- PHENOTYPE IT! Phenotype is the variable clinical presentation of a disease state related to both genetic and environmental influences.
 - Clinical characteristics (gender, age of onset, severity, response to steroids)
 - Physiology (lung function, airway hyperresponsiveness)
 - Triggers (respiratory infections, allergens, pollution, tobacco, aspirin sensitivity)

We eventually hope to be able to “endotype” using validated biomarkers too – endotype describing distinct pathophysiologic mechanisms at a cellular and molecular level

(Kuruvilla et al., 2019)



Algorithm to guide selection advanced therapies for severe asthma

(Oberle and Mathur, 2017)

Pregnancy and Asthma – Are Meds Safe?

- Primary goals of asthma treatment during pregnancy are preventing exacerbations and minimizing maternal impairment.

“The benefits of good adherence to asthma regimens during pregnancy outweigh the risks associated with the medications used.”

(Kher & Mota, 2017)

Uncontrolled asthma in pregnancy: Effects on mother and fetus

Maternal effects

Preeclampsia
Pregnancy-induced hypertension
Gestational diabetes
Premature rupture of membranes
Cesarean birth
Chorioamnionitis
Hyperemesis
Postpartum hemorrhage

Fetal effects

Perinatal death
Preterm birth
Low birth weight
Intrauterine growth restriction
Congenital malformations
Admission to neonatal intensive care unit
Hyperbilirubinemia
Respiratory distress syndrome
Transient tachypnea of the newborn
Asphyxia
Increased risk of intracerebral hemorrhage, anemia

Drug therapy of maternal asthma: Former pregnancy risk categories

Drug	Category ^a
Short-acting beta-agonist	
Albuterol	C
Long-acting beta-agonists	
Formoterol	C
Salmeterol	C
Inhaled corticosteroids	
Budesonide (inhalation)	B
Fluticasone (inhalation)	C
Leukotriene modifiers	
Montelukast, zafirlukast	B
Zileuton	C
Monoclonal antibody	
Omalizumab	B
Xanthine derivative	
Theophylline	C
Intranasal corticosteroids	
Intranasal budesonide	B
Intranasal fluticasone	C
Intranasal mometasone	C
Intranasal triamcinolone	C

^a Former US Food and Drug Administration risk category (see **Drugs**, this page).

Management of asthma exacerbations in pregnancy

Monitor to ensure the maternal oxygen saturation remains above 95%

Consider systemic (oral) corticosteroids

Administer rescue therapy with a short-acting beta-agonist

Consider hospital admission if symptoms do not improve with emergency room care

Consider admission to intensive care unit and mechanical ventilation if symptoms are severe or if patient shows signs of impending respiratory failure

Carefully monitor fetal health

(Kher & Mota, 2017)

TOBACCO CESSATION

Smoke a **FRESH** cigarette"

IF the cigarette you have been smoking stings or burns your throat, switch to Camels and see the difference.

It's the peppery dust left in tobacco by inefficient cleaning methods that makes you cough.

It's the unkindly hot smoke of harsh, dried-out tobacco that burns and irritates your throat.

There is no peppery dust in Camels—that's whisked away by a special vacuum-cleaning process.

There are no stale, crumbly, parched tobaccos—the fine Turkish and mild Domestic tobaccos of which Camels are blended come to you in prime, factory-fresh condition, thanks to the Humidor Pack.

This scientific germ-safe wrapping—not plain ordinary Cellophane, but *moisture-*

proof Cellophane which costs nearly twice as much—seals in all the natural aroma and freshness, seals it so tightly that wet weather cannot make Camels damp, nor drought weather make them dry.

Camels are milder and more throat-friendly because they are dust-free and fresh.

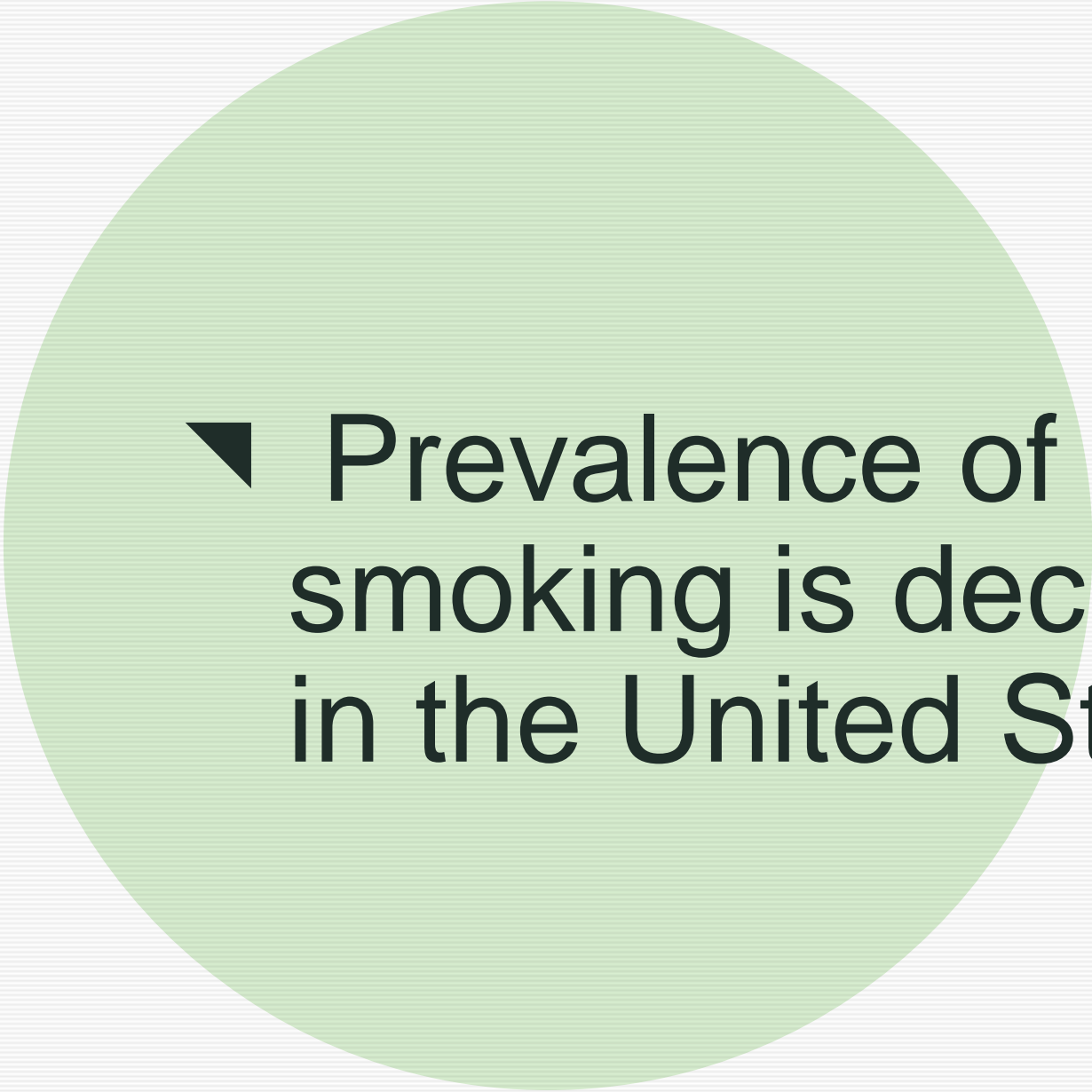
Give your throat a vacation, switch to Camels for just one day. Then leave them—if you can.

Tune in CAMEL QUARTER HOUR featuring Morton Downey and Tony Wons — Camel Orchestra, direction Jacques Renard — Columbia System—every night except Sunday



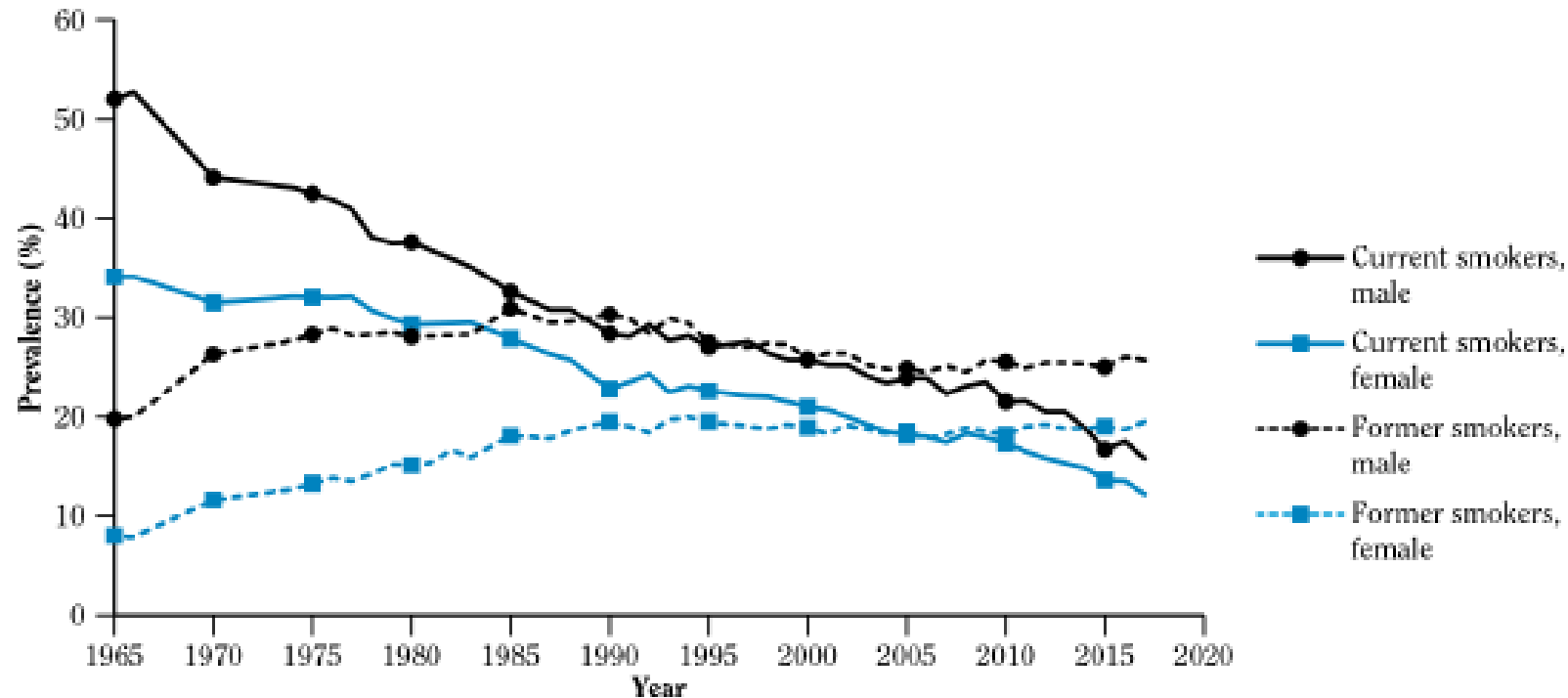
Don't remove the moisture-proof wrapping from your package of Camels after you open it. The Humidor Pack is protection against dust and germs. In offices and homes, even in the dry atmosphere of artificial heat, the Humidor Pack delivers fresh Camels and keeps them right until the last one has been smoked

CAMELS

- 
- ▼ Prevalence of cigarette smoking is decreasing in the United States

Trends in cigarette smoking among adults in the United States 1965 to 2017

Figure ES.1 Trends in prevalence (%) of current and former cigarette smoking among adults 18 years of age and older, by sex; National Health Interview Survey (NHIS) 1965–2017; United States



Source: NHIS, National Center for Health Statistics, public use data, 1965–2017.

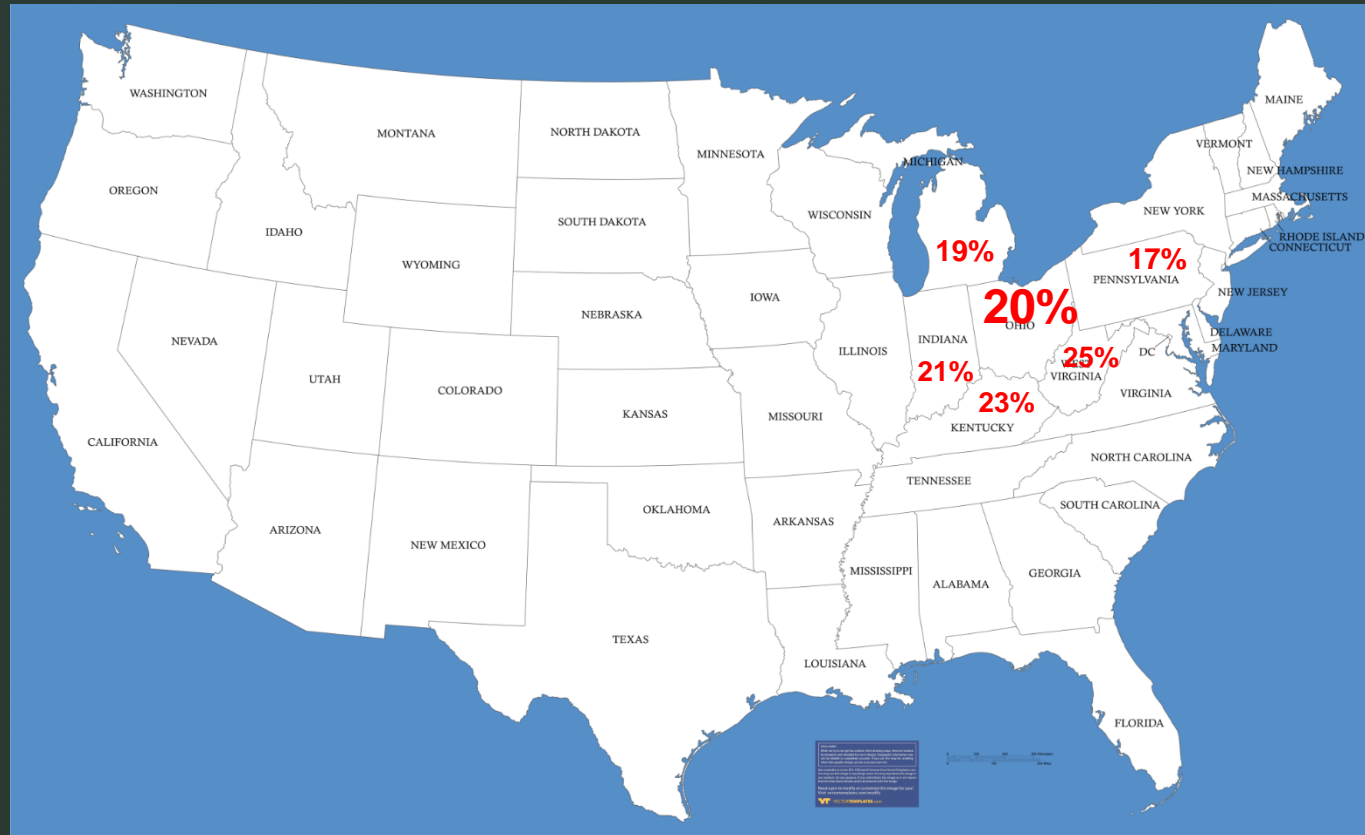
Note: From 1965 to 2017, data were reported for the following years: 1965, 1966, 1970, 1974, 1976–1980, 1983, 1985, 1987, 1988, 1990–1995, and 1997–2017.

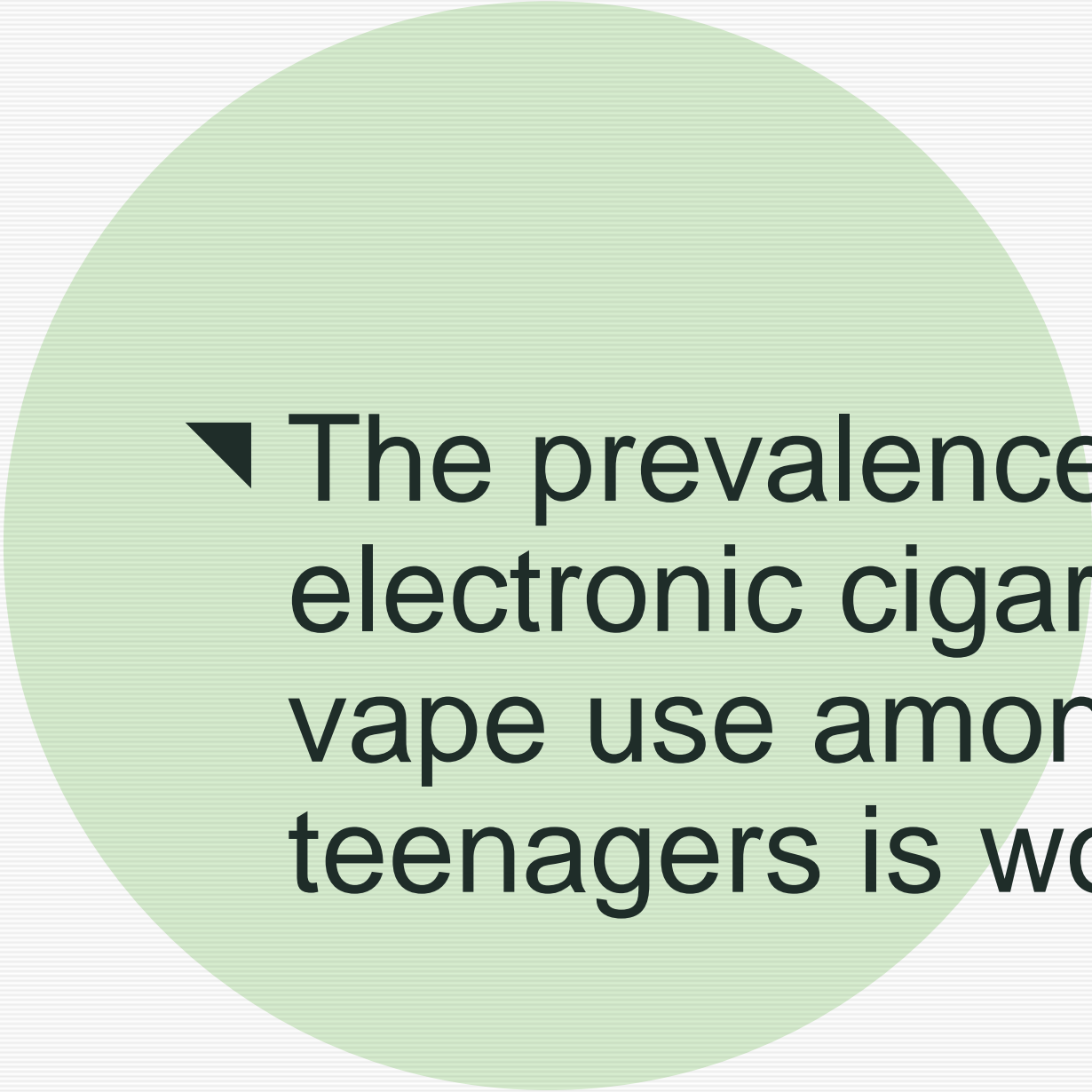
Current cigarette smoking among adults in the United States

US: 12.5% of adults currently smoke cigarettes ~30 million

OH: 20%

Cleveland: 35 % (2015)



- 
- ▼ The prevalence of electronic cigarette or vape use among teenagers is worrisome

14.1% of high school students and 3.3% of middle school students reported current use of e-cigarettes

Characteristic	Overall		High school		Middle school	
	Estimated weighted no. [†]	% (95% CI)	Estimated weighted no. [†]	% (95% CI)	Estimated weighted no. [†]	% (95% CI)
Among all students (N = 28,291)						
Current use of e-cigarettes	2,550,000	9.4 (8.0–11.1)	2,140,000	<u>14.1 (12.4–16.0)</u>	380,000	<u>3.3 (2.6–4.2)</u>
Among current e-cigarette users						
Frequency of use during past 30 days						
1–5 days	1,030,000	40.6 (37.2–44.1)	790,000	37.2 (33.4–41.1)	230,000	<u>60.0 (53.3–66.3)</u>
6–19 days	430,000	17.1 (14.2–20.4)	360,000	16.8 (13.9–20.2)	70,000	<u>19.3 (12.7–28.3)</u>
20–30 days	1,080,000	42.3 (38.5–46.3)	980,000	46.0 (41.6–50.4)	80,000	20.8 (15.8–26.8)
Daily e-cigarette use [§]	700,000	27.6 (24.5–31.0)	640,000	<u>30.1 (26.6–33.9)</u>	40,000	<u>11.7 (8.0–16.7)</u>

Disposable and “unknown” brands are the most common

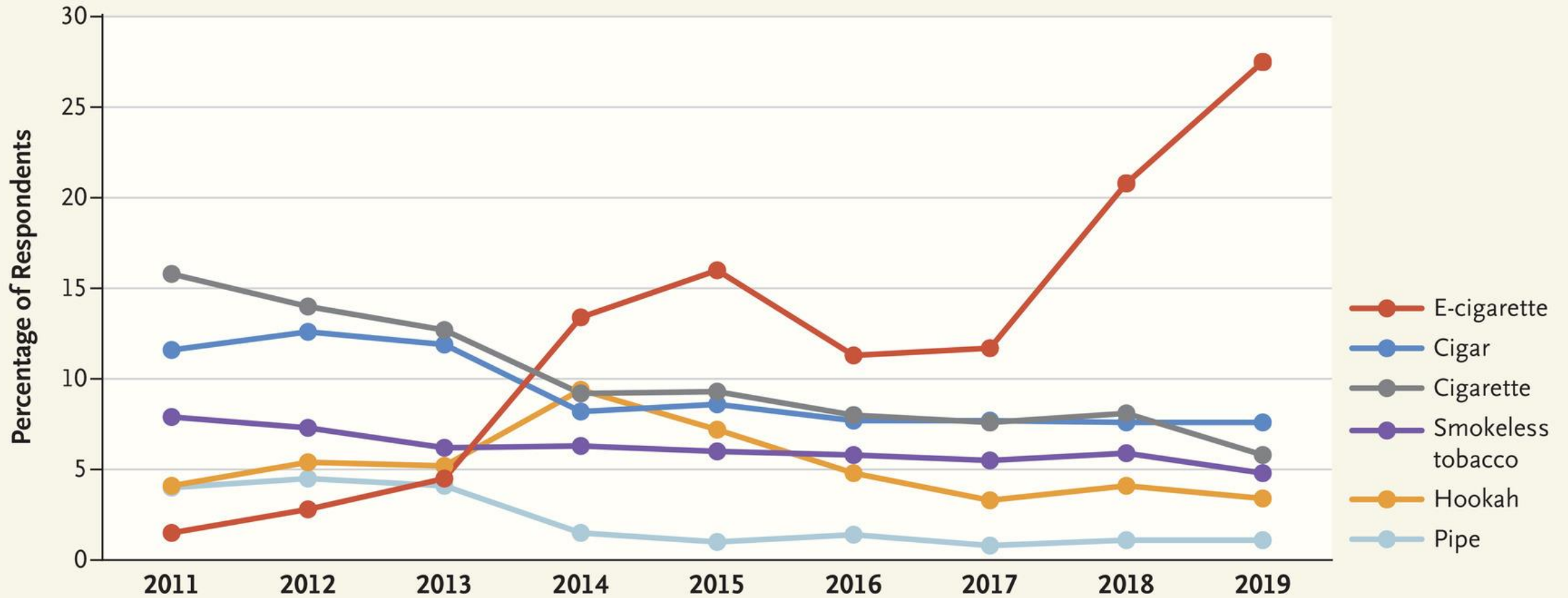
Characteristic	Overall		High school		Middle school	
	Estimated weighted no.†	% (95% CI)	Estimated weighted no.†	% (95% CI)	Estimated weighted no.†	% (95% CI)
Device type most often used¶						
Disposables	1,390,000	55.3 (49.5–61.0)	1,210,000	57.2 (51.7–62.6)	170,000	45.8 (34.5–57.6)
Prefilled or refillable pods or cartridges	630,000	25.2 (19.7–31.5)	540,000	25.7 (20.2–32.0)	80,000	21.6 (12.8–33.9)
Tanks or mod system	160,000	6.7 (5.3–8.4)	120,000	5.9 (4.5–7.8)	30,000	9.8 (7.1–13.5)
Don't know the type	320,000	12.8 (10.2–16.1)	230,000	11.2 (8.6–14.4)	80,000	22.8 (17.0–29.9)
Any brand**						
Puff Bar	730,000	29.7 (25.5–34.4)	610,000	29.3 (25.0–34.0)	110,000	30.9 (21.3–42.4)
Vuse	580,000	23.6 (17.9–30.3)	490,000	23.8 (17.9–30.9)	70,000	20.9 (13.2–31.3)
JUUL	540,000	22.0 (17.8–26.9)	440,000	21.2 (16.3–27.1)	80,000	23.8 (17.8–30.9)
SMOK (including NOVO)	330,000	13.5 (10.8–16.6)	290,000	14.3 (11.4–17.9)	20,000	7.8 (4.4–13.5)
NJOY	200,000	8.3 (6.0–11.4)	170,000	8.2 (5.6–11.7)	20,000	7.3 (4.3–12.1)
Hyde††	180,000	7.3 (4.4–12.0)	160,000	7.9 (4.6–13.3)	—§§	—§§
blu	160,000	6.5 (4.9–8.6)	110,000	5.6 (3.9–7.8)	30,000	10.2 (5.7–17.6)
STIG	120,000	5.0 (3.6–6.8)	90,000	4.7 (3.2–6.7)	—§§	—§§
Suorin	110,000	4.8 (3.6–6.5)	90,000	4.8 (3.5–6.5)	—§§	—§§
Logic	100,000	4.3 (3.0–6.1)	70,000	3.8 (2.5–5.6)	—§§	—§§
Mojo	90,000	4.0 (2.8–5.5)	70,000	3.7 (2.6–5.3)	—§§	—§§
Leap	90,000	3.7 (2.6–5.2)	60,000	3.0 (2.0–4.4)	—§§	—§§
For smoke	80,000	3.2 (2.4–5.2)	60,000	2.9 (1.8–4.7)	—§§	—§§
Some other brand not listed	790,000	32.2 (27.8–37.0)	670,000	32.2 (27.4–37.4)	120,000	32.8 (25.5–41.0)
Not sure/Don't know the brand	700,000	28.3 (24.8–32.0)	550,000	26.7 (22.7–31.1)	140,000	37.4 (29.7–45.8)

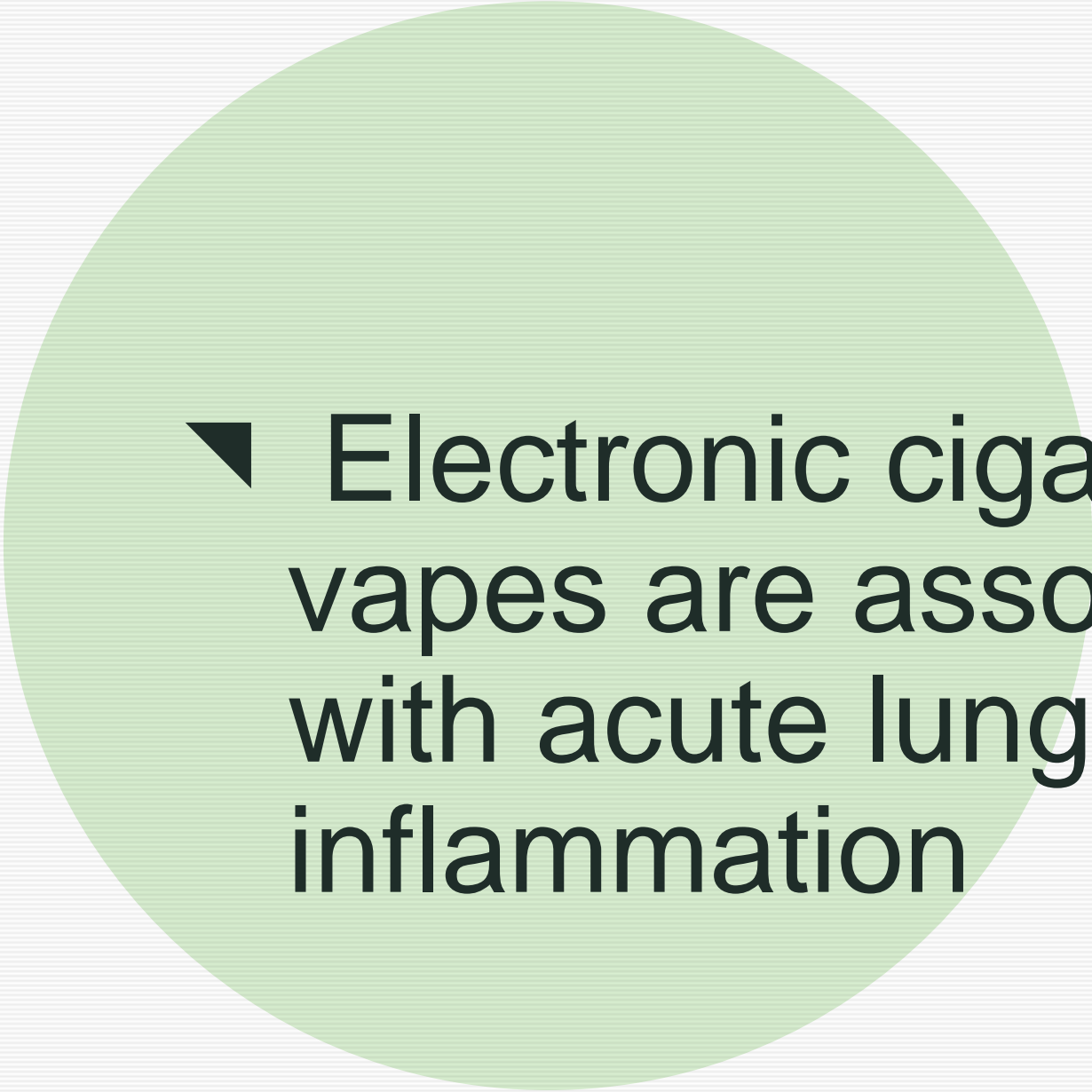
Strong preference for fruit and candy flavors

85% used flavored e-cigarettes

Characteristic	Overall		High school		Middle school	
	Estimated weighted no. [†]	% (95% CI)	Estimated weighted no. [†]	% (95% CI)	Estimated weighted no. [†]	% (95% CI)
Flavor type used^{†††}						
Fruit	1,450,000	<u>69.1 (65.4–72.6)</u>	1,220,000	<u>68.5 (64.4–72.3)</u>	210,000	<u>71.1 (63.9–77.3)</u>
Candy, desserts, or other sweets	800,000	38.3 (33.8–42.9)	660,000	37.3 (32.6–42.2)	130,000	43.6 (36.3–51.3)
Mint	610,000	29.4 (25.6–33.5)	540,000	30.3 (25.9–35.1)	70,000	23.7 (18.9–29.3)
Menthol	550,000	26.6 (21.0–33.1)	500,000	28.2 (22.2–35.2)	40,000	16.2 (10.3–24.6)
Alcoholic drinks	150,000	7.6 (5.6–10.2)	120,000	6.8 (4.7–9.8)	30,000	10.8 (7.0–16.1)
Chocolate	80,000	4.3 (3.1–5.9)	60,000	3.8 (2.7–5.3)	— ^{§§}	— ^{§§}
Clove or spice	60,000	2.9 (1.9–4.6)	40,000	2.6 (1.6–4.2)	— ^{§§}	— ^{§§}
Some other flavor not listed	240,000	11.7 (10.1–13.6)	200,000	11.7 (9.9–13.7)	30,000	11.9 (8.0–17.5)

Current tobacco product use among U.S. High School students 2011 to 2019



- 
- ▼ Electronic cigarettes or vapes are associated with acute lung inflammation

CDC - EVALI hospitalizations and deaths

Number of hospitalized EVALI cases and deaths in the United States

As of February 18, 2020

Hospitalizations

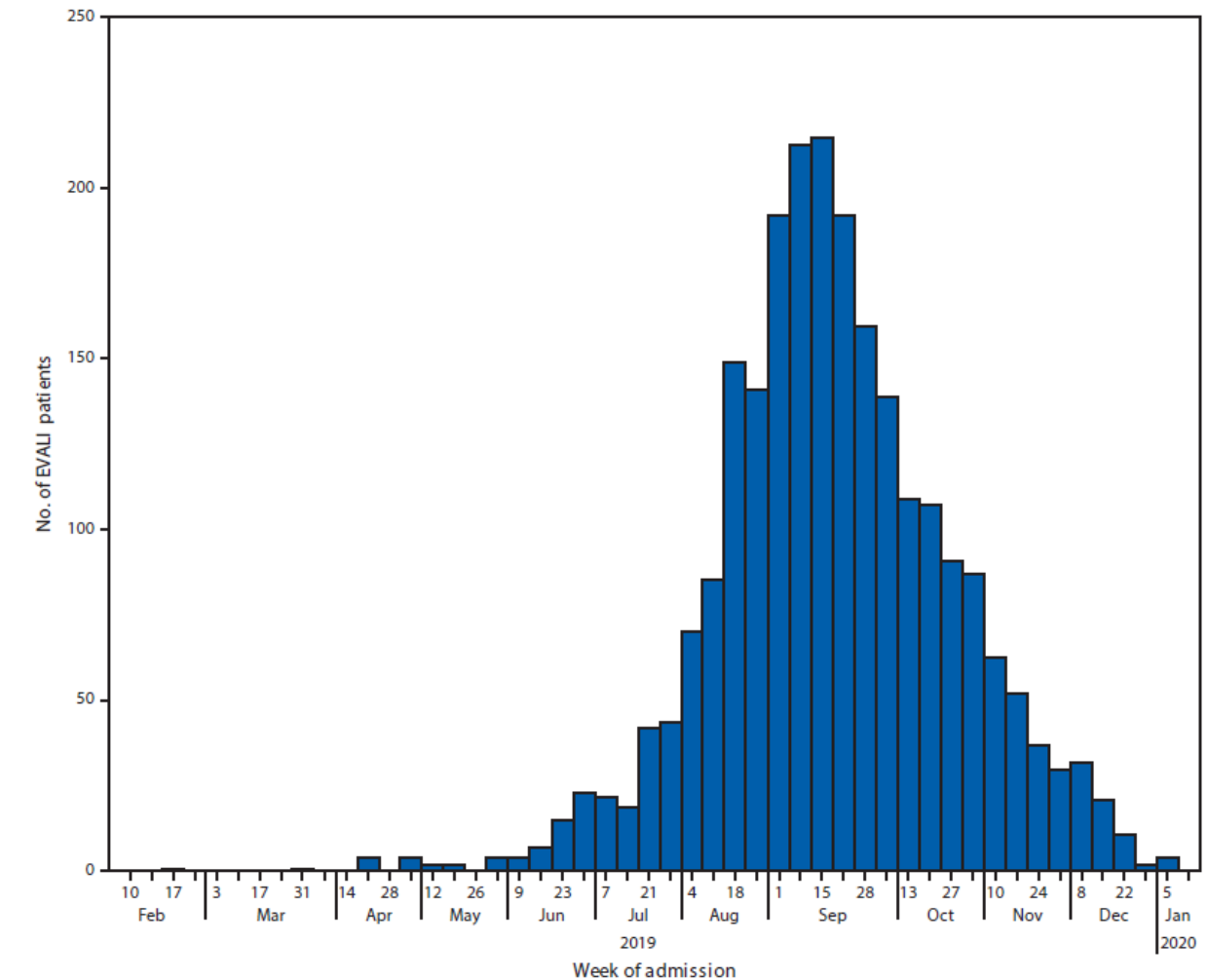
- 2,807 cases

Deaths

- 68 deaths have been confirmed in 29 states

EVALI hospitalizations declined after sharp increase in August-September 2019

FIGURE 1. Number of patients (N = 2,398) with e-cigarette, or vaping, product use–associated lung injury (EVALI) by week of hospital admission — United States, February 10, 2019–January 14, 2020



EVALI hospitalizations and deaths

- 82% reported using THC-containing products

└─→ 50% provided information about source -> 78% informal sources

- 57% reported using nicotine-containing products

└─→ 54% provided information about source -> 17% informal sources

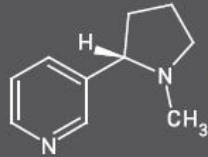
Which substance was associated with EVALI?

Vitamin E acetate is an additive was linked to the EVALI outbreak.

- A study external icon analyzed samples from 51 EVALI cases and a comparison group of samples from 99 comparison individuals without EVALI
- Vitamin E acetate was identified in BAL fluid samples from 48 of the 51 EVALI patients

That`s only part of the story

**VOLATILE
ORGANIC
COMPOUNDS**



NICOTINE

**ULTRAFINE
PARTICLES**



**HEAVY METALS SUCH AS
NICKEL, TIN, AND LEAD**

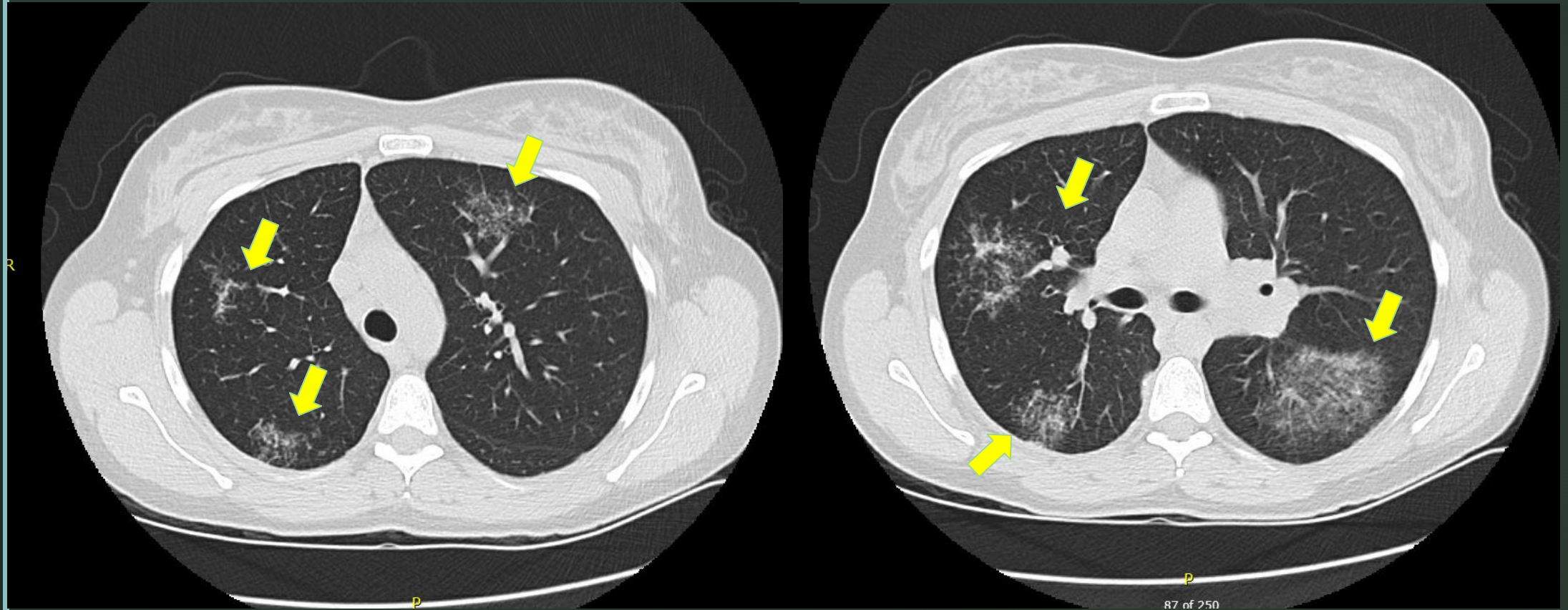


**FLAVORING SUCH AS
DIACETYL, A CHEMICAL
LINKED TO A SERIOUS
LUNG DISEASE**

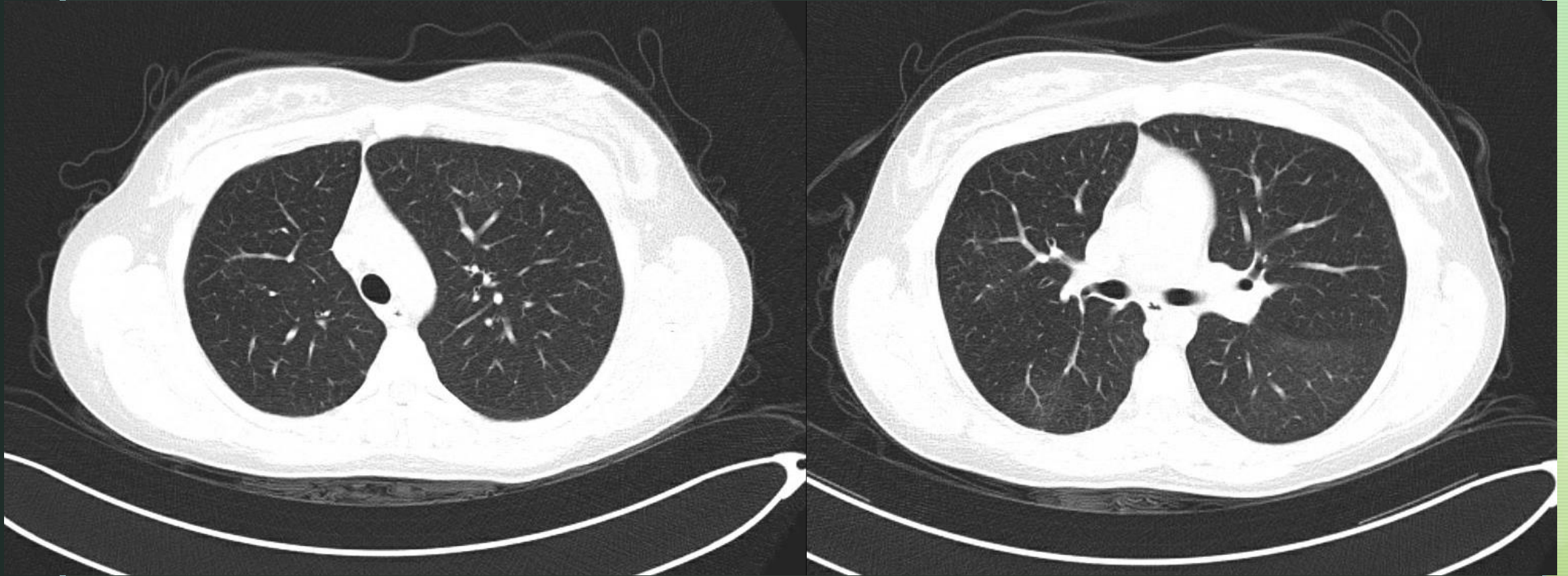


**CANCER-CAUSING
CHEMICALS**

Acute lung inflammation after vaping



3 months after stopping vaping



EVALI requiring ICU admission

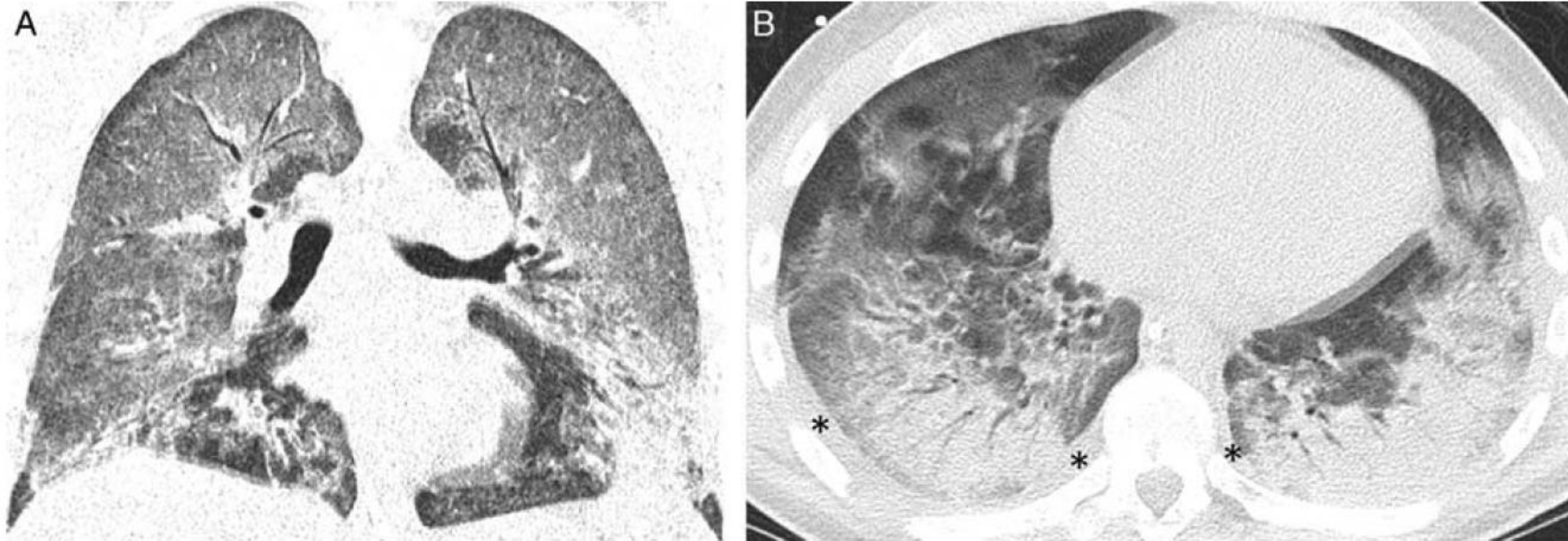
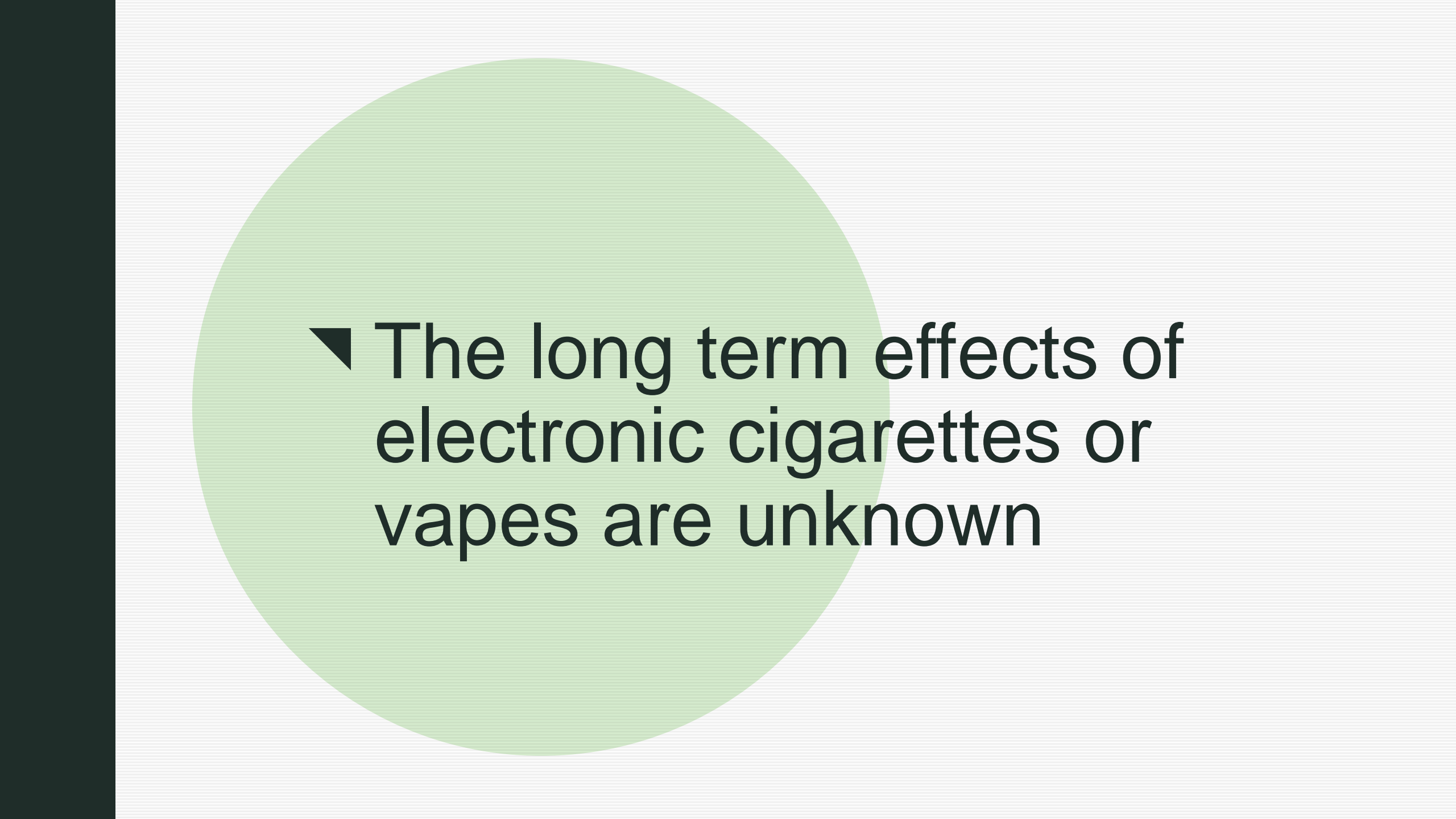


FIGURE 1. Coronal (A) and axial (B) CT images in a 21-year-old man show diffuse ground-glass opacities with dense consolidations in the posterior lower lobes, suggestive of DAD. Trace pleural effusions (asterisks) are noted.

- 
- ▼ The long term effects of electronic cigarettes or vapes are unknown



▪ It is unknown whether vaping can cause

- COPD
- Lung fibrosis
- Heart disease
- Cancer

E-cigarettes contain harmful chemicals

Table 1. Levels of Toxicants in E-Cigarette Aerosol Compared With Nicotine Inhaler and Cigarette Smoke

Toxicant	Range in Content in Aerosol From 12 E-Cigarette Samples per 15 Puffs*	Range in Content in Conventional Cigarette Micrograms in Mainstream Smoke From 1 Cigarette	Content in Nicotine Inhaler Mist per 15 Puffs*
Formaldehyde, μg	0.2–5.61	1.6–52	0.2
Acetaldehyde, μg	0.11–1.36	52–140	0.11
Acrolein, μg	0.07–4.19	2.4–62	ND
o-Methylbenzaldehyde, μg	0.13–0.71	...	0.07
Toluene, μg	ND–0.63	8.3–70	ND
p,m-xylene, μg	ND–0.2	...	ND
NNN, ng	ND–0.00043	0.0005–0.19	ND
NNK, ng	ND–0.00283	0.012–0.11	ND
Cadmium, ng	ND–0.022	...	0.003
Nickel, ng	0.011–0.029	...	0.019
Lead, ng	0.003–0.057	...	0.004



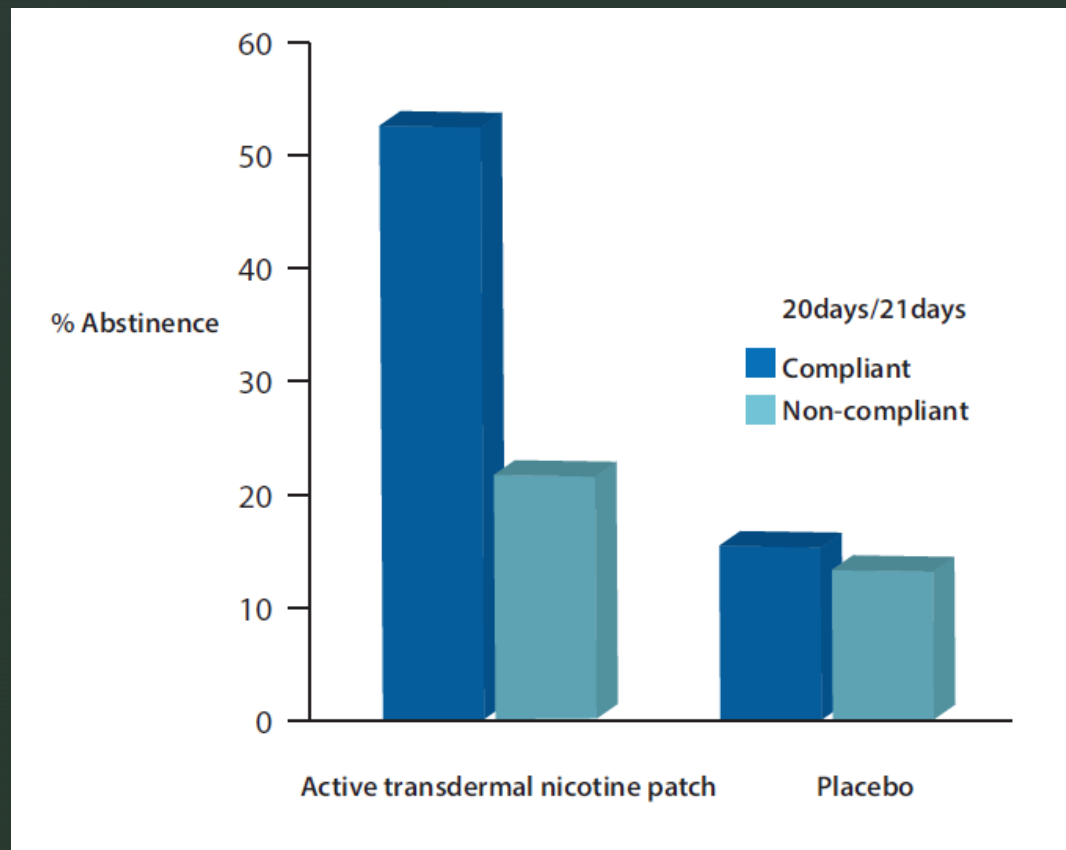
- Pharmacotherapy and behavioral support are each effective when used alone, but combining them is more effective.

Effectiveness of methods used to treat tobacco dependence

Method	Versus placebo Odds ratio (95% CI)	Estimated abstinence rate
Patch	1.64 (1.52-1.78)	23.4%
Gum	1.49 (1.40-1.60)	19%
Lozenge	1.95 (1.61-2.36)	24.2%
Inhaler	1.90 (1.36-2.67)	24.8%
Nasal spray	2.02 (1.49-2.73)	26.7%
Bupropion	2.0 (1.8-2.2)	24.2%
Varenicline	3.1 (2.5-3.8)	33.2%
Long+short NRT	1.25 (1.15-1.36) vs single NRT	
Varenicline+NRT	1.62 (1.18-2.23) vs varenicline alone	

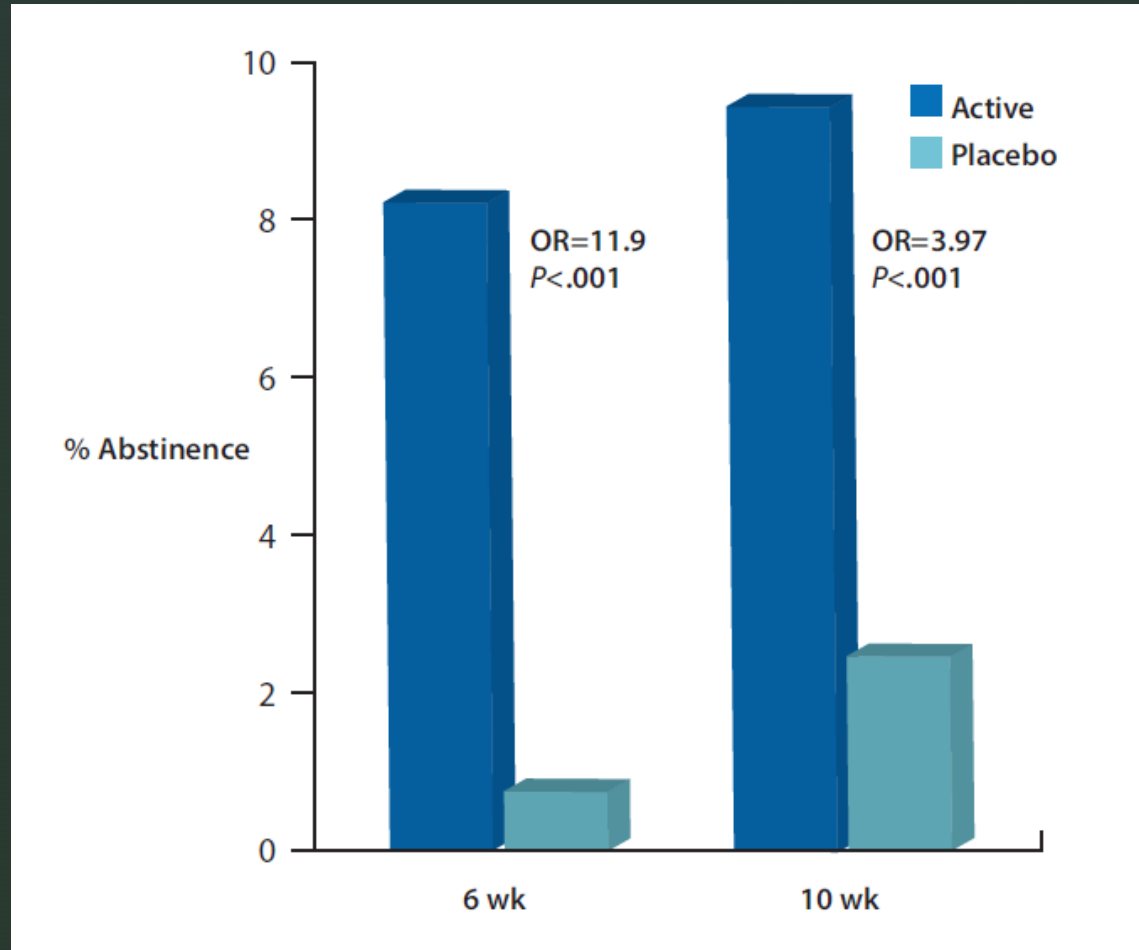
Wearing NRT patches consistently increases quit rates

Advise not to skip dates -> they won't work if they are not used.

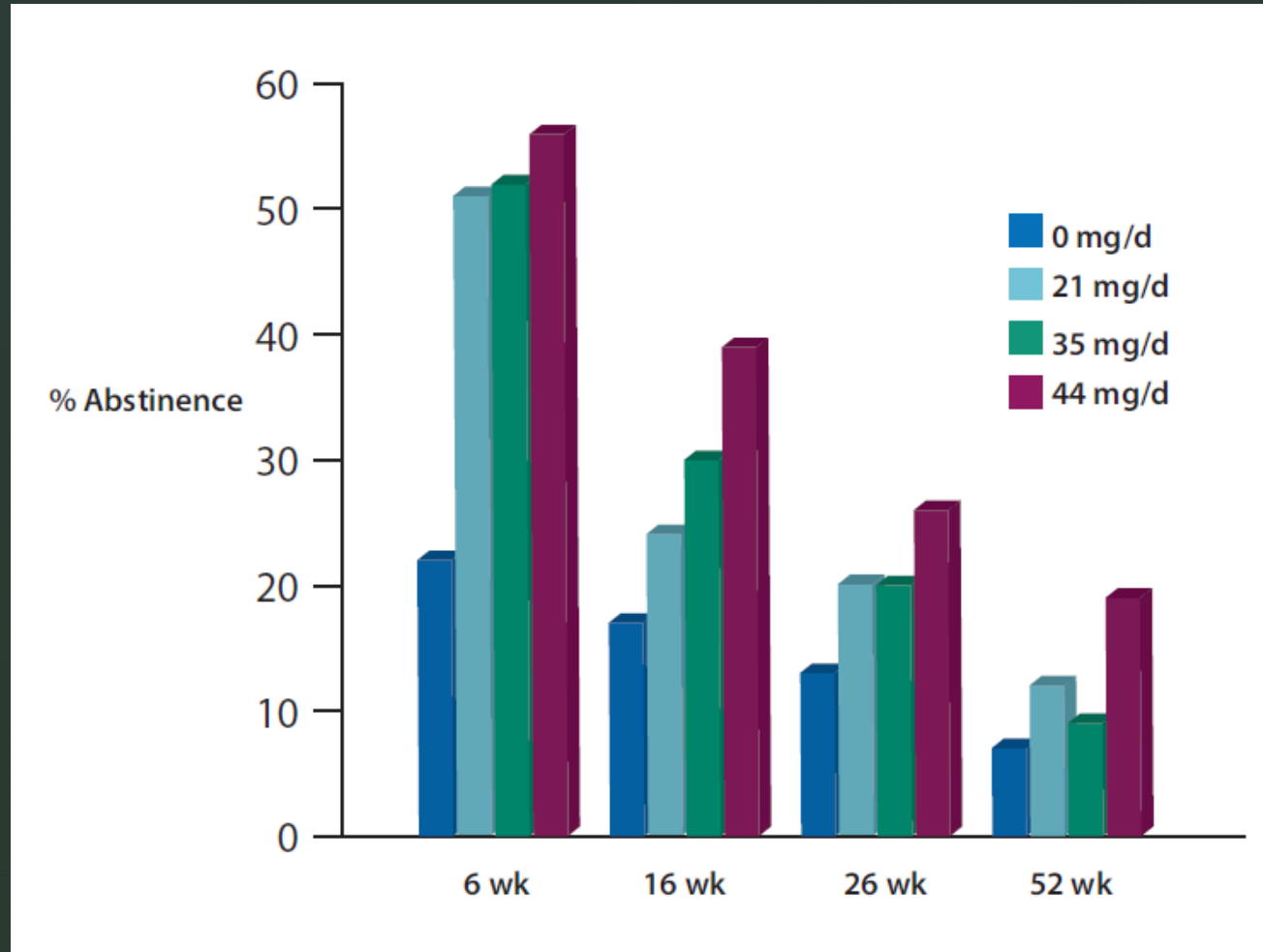


Continually wearing patches after lapse promotes recovery

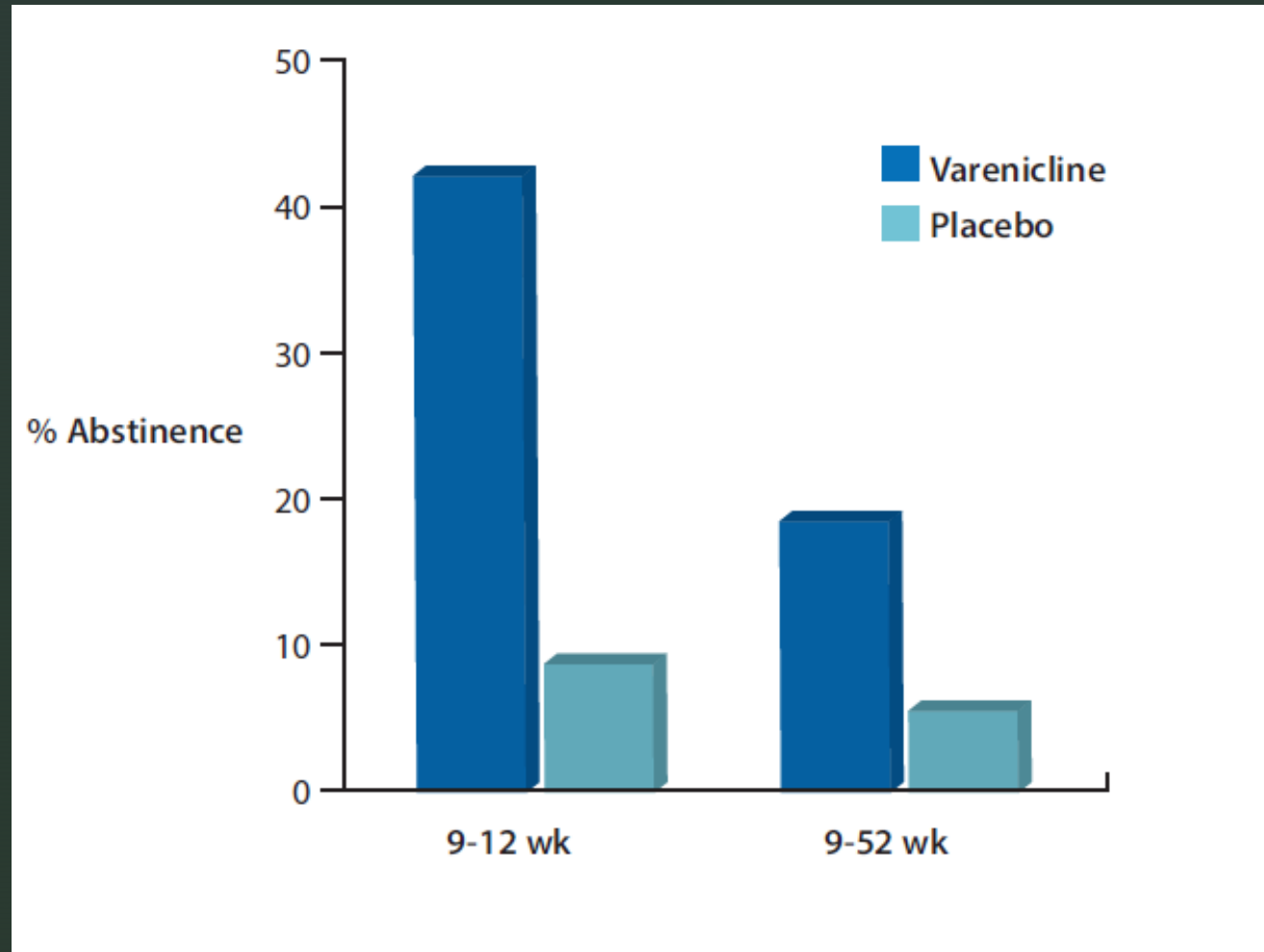
It won't work if they are not used.



Higher-dose nicotine patches are safe and may be more effective



Extending varenicline for 6 months to prevent relapse is effective



2018 ACC Expert Consensus Decision Pathway on Tobacco Cessation Treatment

TABLE 4 Recommended Pharmacotherapy for Smoking Cessation in Patients with CVD

	Outpatient With Stable CVD	Inpatient With ACS
1st line	Varenicline OR combination NRT*	<i>In-hospital to relieve nicotine withdrawal:</i> Nicotine patch OR combination NRT* <i>At discharge:</i> Combination NRT or varenicline†
2nd line	Bupropion OR single NRT product	<i>At discharge:</i> Single NRT product
3rd line	Nortriptyline‡	Bupropion§
If single agent is insufficient to achieve abstinence	Combine categories of FDA-approved drugs: Varenicline + NRT (single agent) Varenicline + bupropion Bupropion + NRT (single agent)	n/a



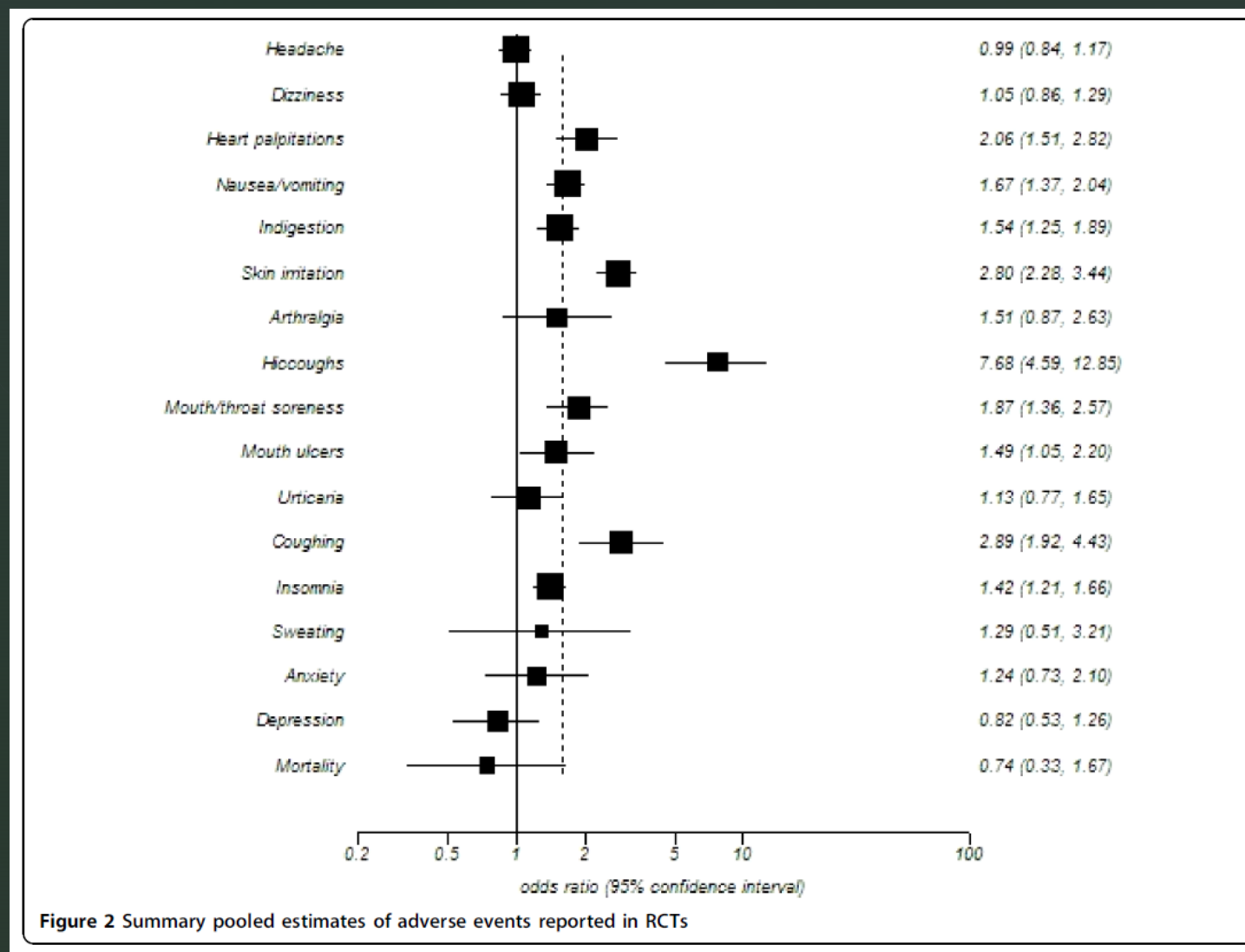
▼ Tobacco
treatment is safe

There is no significant increase in the risk of serious adverse events with the use of nicotine-replacement therapy

Meta- regression analysis

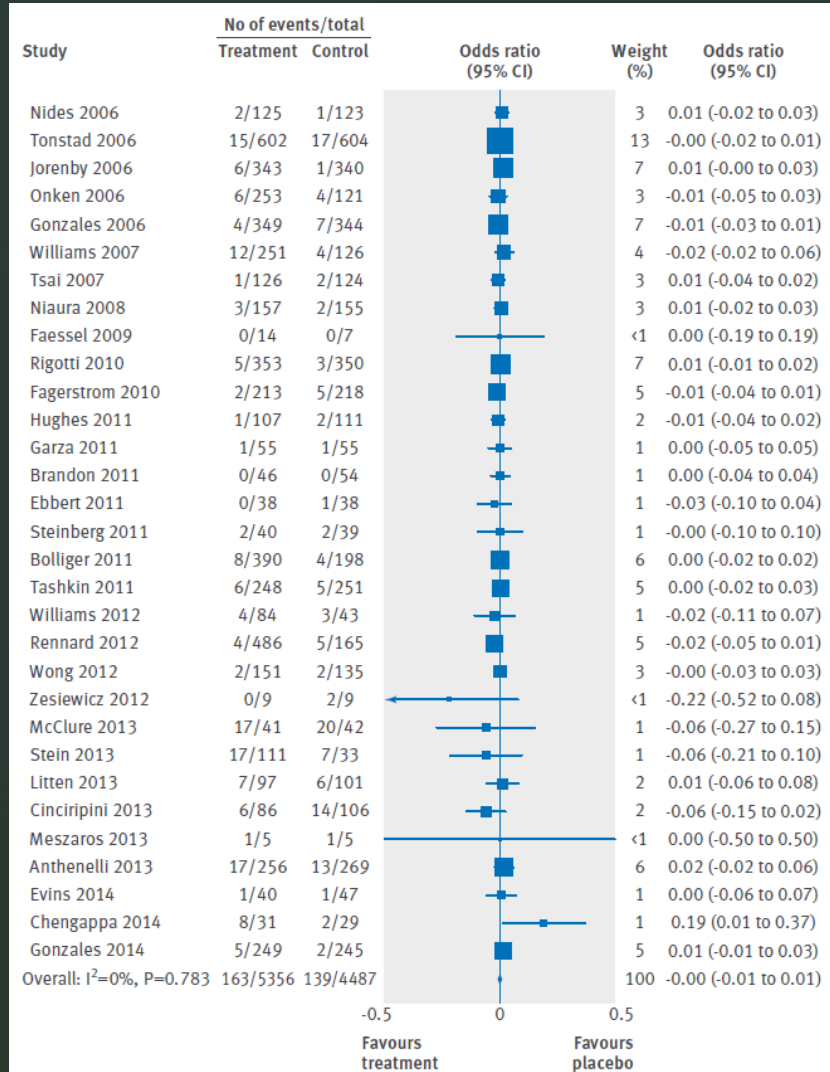
- 92 RCTs involving 32,185 participants

- 28 observational studies involving 145,205 participants



Risk of neuropsychiatric adverse events with varenicline

Risk of depression events associated with varenicline use in 31 placebo RCT



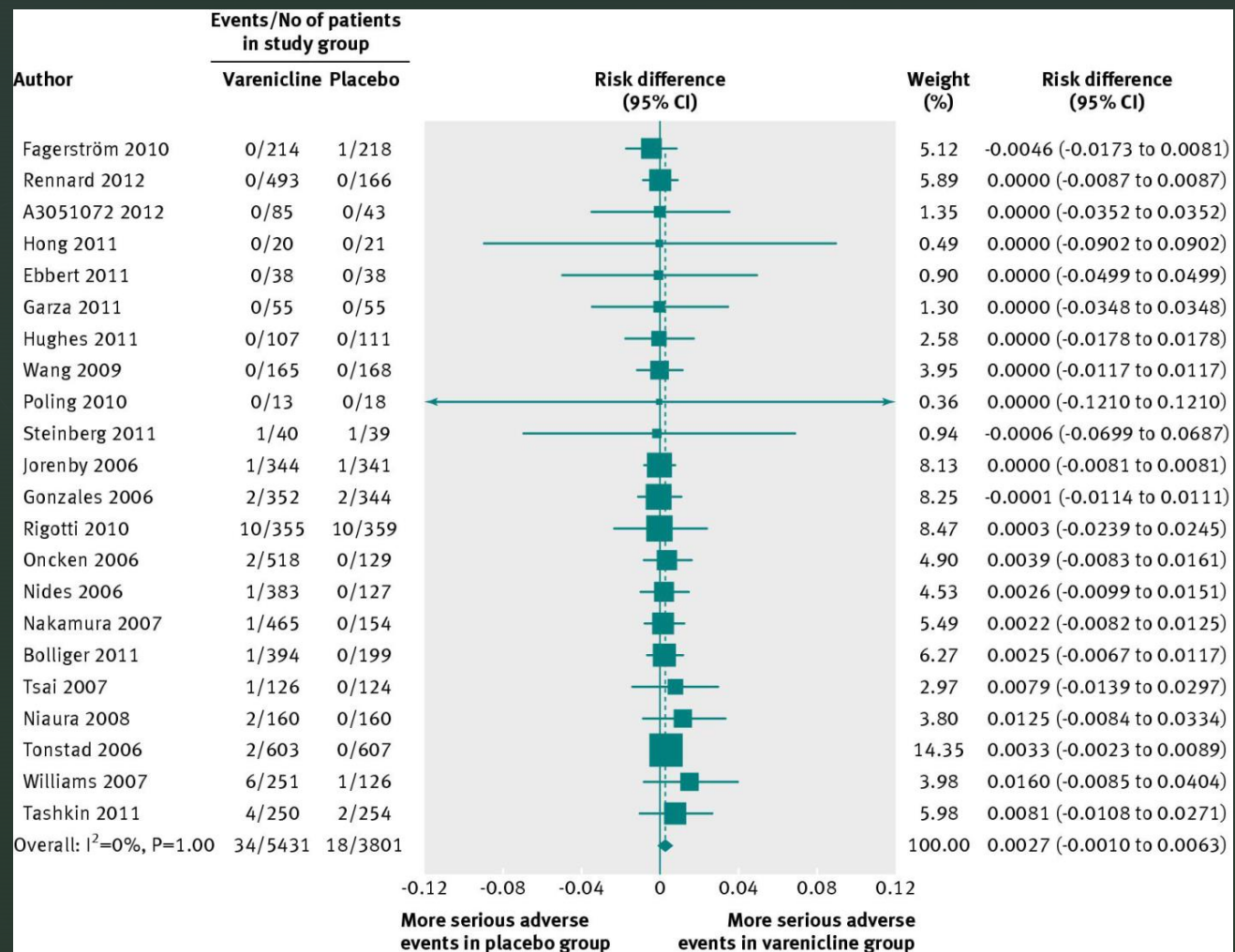
Increased risk:

- sleep disorders
- insomnia
- abnormal dreams
- Fatigue

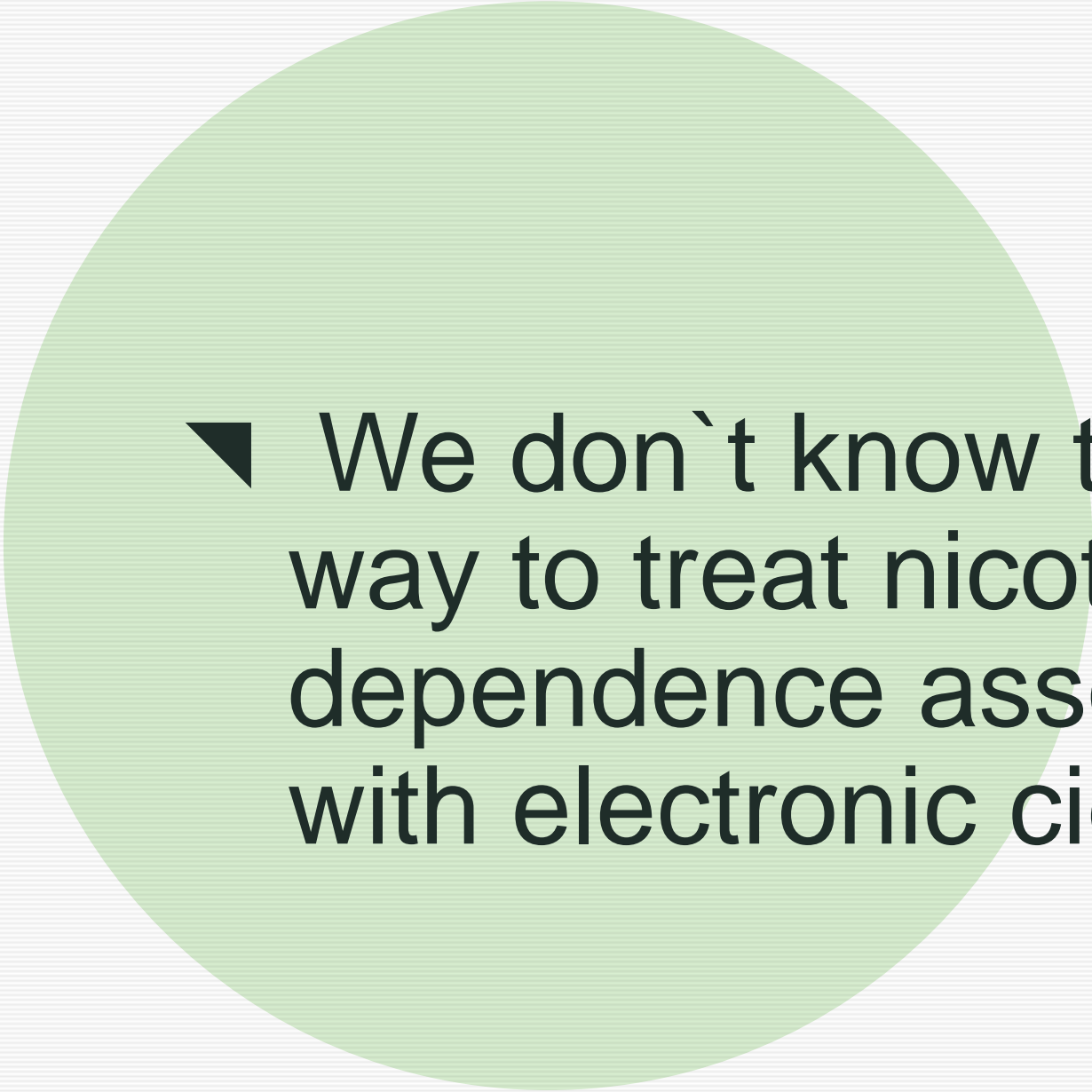
No increased risk:

- Depression
- Suicidal ideation
- Aggression
- Death

No significant increased risk of adverse cardiovascular events associated with varenicline



Examples: myocardial infarction, unstable angina, coronary revascularisation, coronary artery disease, arrhythmias, transient ischaemic attacks, stroke, sudden death or cardiovascular related death, or congestive heart failure.

- 
- ▼ We don't know the best way to treat nicotine dependence associated with electronic cigarettes

Electronic cigarettes

There are different types of e-cigarette devices.



E-liquids

There are even more different types of e-liquids



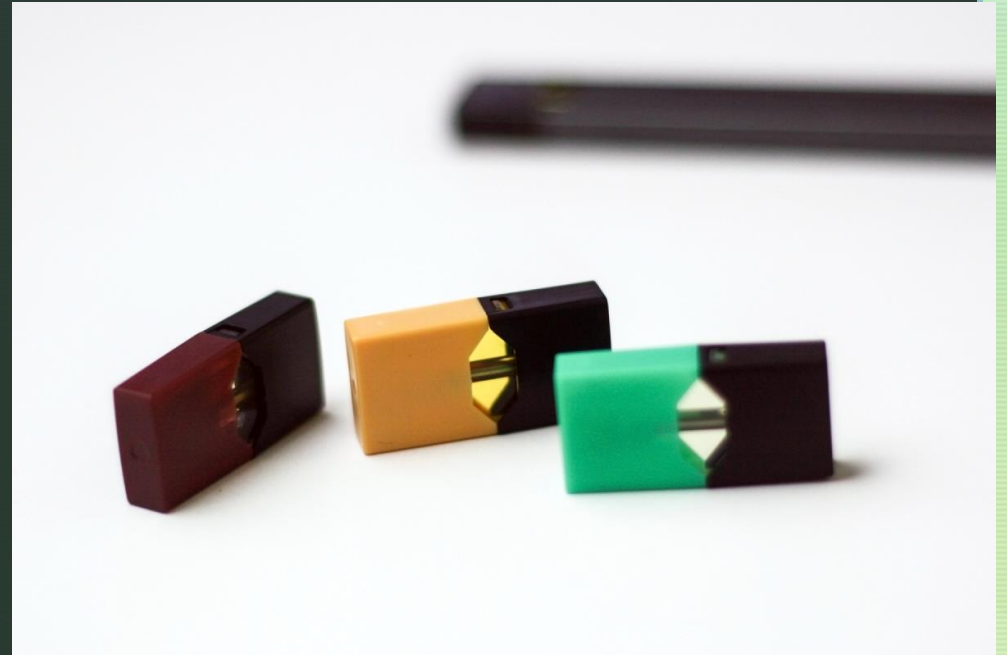
Nicotine content in cigarettes vs e-cigs



1 cigarette = 10-12 mg nicotine

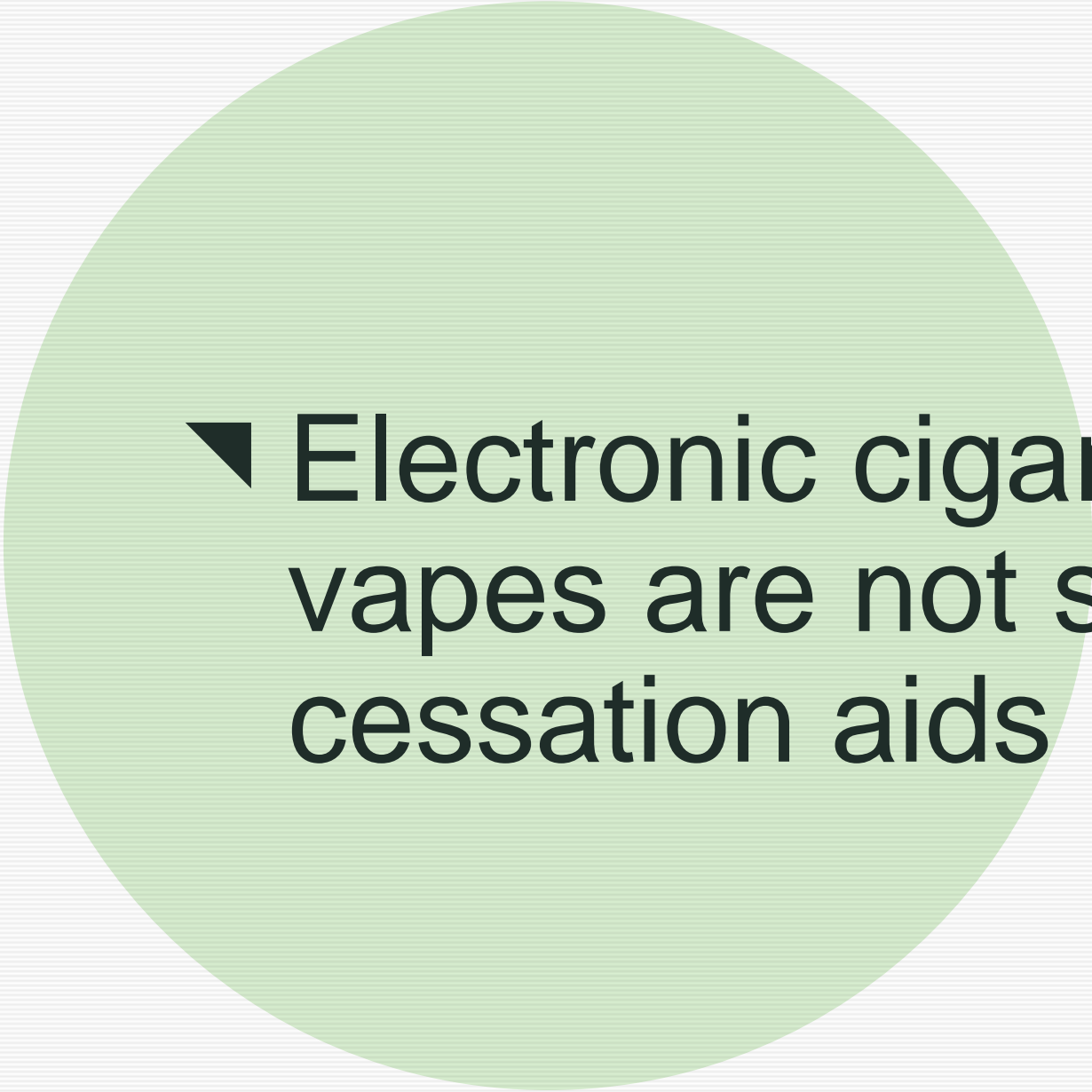
1 cigarette = inhale 1-1.8mg nicotine

1 pack = inhale 20-36 mg nicotine



1 JUUL pod 3% = 23mg nicotine

1 JUUL pod 5% = 40 mg nicotine

- 
- ▼ Electronic cigarettes or vapes are not smoking cessation aids

NRT vs e-cigarette with nicotine 18mg

Table 2. Abstinence Rates at Different Time Points and Smoking Reduction at 52 Weeks.*

Outcome	E-Cigarettes (N = 438)	Nicotine Replacement (N = 446)	Primary Analysis: Relative Risk (95% CI)†	Sensitivity Analysis: Adjusted Relative Risk (95% CI)
Primary outcome: abstinence at 52 wk — no. (%)	79 (18.0)	44 (9.9)	1.83 (1.30–2.58)	1.75 (1.24–2.46)‡
Secondary outcomes				
Abstinence between wk 26 and wk 52 — no. (%)	93 (21.2)	53 (11.9)	1.79 (1.32–2.44)	1.82 (1.34–2.47)§
Abstinence at 4 wk after target quit date — no. (%)	192 (43.8)	134 (30.0)	1.45 (1.22–1.74)	1.43 (1.20–1.71)¶
Abstinence at 26 wk after target quit date — no. (%)	155 (35.4)	112 (25.1)	1.40 (1.14–1.72)	1.36 (1.15–1.67)‡
Carbon monoxide–validated reduction in smoking of ≥50% in participants without abstinence between wk 26 and wk 52 — no./total no. (%)	44/345 (12.8)	29/393 (7.4)	1.75 (1.12–2.72)	1.73 (1.11–2.69)

80% of those in the e-cigarette group were still vaping at 12 months
9% of the NRT group were still using NRT

Electronic cigarettes or vapes are not smoking cessation aids

2021 Cochrane systematic review with 16,759 participants

- E-cigarettes more effective than NRT or non-nicotine devices (moderate certainty)

USPSTF

- Evidence is insufficient to evaluate balance of risk and benefits

General Recommendation

- Use FDA approved smoking cessation aids

COVID-19

Post-COVID
Syndrome (PCS)



- **The most commonly reported symptoms are fatigue, cough, shortness of breath, chest, difficulty concentrating, arthralgia, low-grade fever, and headache**

Cleveland Clinic Journal of Medicine May 2021, 88 (5) 267-272; DOI: <https://doi.org/10.3949/ccjm.88a.21010>

COVID-19 diagnosis

Arrange for multidisciplinary follow-up

Inpatients: ambulatory oximetry, physical therapy, occupational therapy, care coordination, home healthcare

Outpatients: primary care follow-up, referral to local COVID-19 hotline

4 weeks after diagnosis or hospital discharge

Virtual screening^a

No persistent symptoms

Resume routine outpatient care

Persistent symptoms

In-person visit with primary care physician or post-COVID-19 clinic: chest x-ray, spirometry, diffusing capacity for carbon monoxide, psychiatric screening,^b neurocognitive screening^c

If pulmonary embolism diagnosed with COVID-19: echocardiogram, electrocardiogram, ventilation-perfusion scan

For all patients, also consider echocardiogram, electrocardiogram

Normal tests

Consider alternative diagnoses for symptoms
Refer to post-COVID-19 clinic, if available
Optimize comorbid conditions
Supportive symptomatic care

Abnormal neurocognitive screening

Neuropsychiatric referral

Abnormal chest x-ray, spirometry, diffusion capacity, ventilation-perfusion scan

Computed tomography/
computed tomographic pulmonary arteriography
Pulmonary referral

Abnormal echocardiogram and electrocardiogram

Cardiology referral

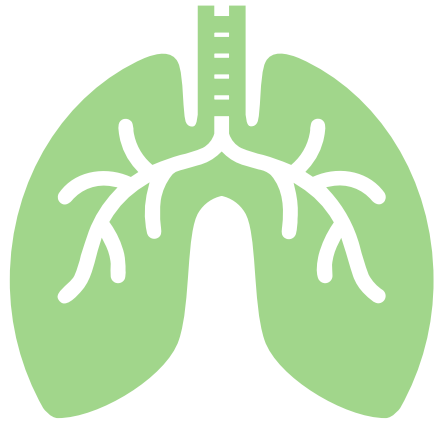
^aScreening tools to consider: Post-COVID-19 Functional Status Scale, COVID-19 Yorkshire Rehabilitation Screen, University of Pennsylvania Post-COVID Screening Measures.

^bAvailable psychiatric screening tools: General Anxiety Disorder-7 (GAD-7), Patient Health Questionnaire-9 (PHQ-9; for depression screening), PTSD Checklist for DSM-5 (PCL-5), Impact of Event Scale-6 (IES-R; for PTSD screening), Hospital Anxiety and Depression Score (HADS).

^cAvailable neurocognitive screening tools: Montreal Cognitive Assessment (MoCA), Mini-Mental State Examination (MMSE), Cognitive Assessment Tool Rapid Version (CAT-rapid).



- Fatigue represents the most common concern in patients with PCS
- Respiratory symptoms are common in PCS patients with PCS, and dyspnea is often the most prevalent



Approximately 5% of patients develop adult respiratory distress syndrome (ARDS). Breathlessness and cough are noted in a substantial proportion of patients with long COVID-19

Batiha GES, Al-Kuraishy HM., Al-Gareeb AI, et al.

In one survivor study, 42% of patients evaluated in clinic three months after hospital discharge had a significant reduction in diffusion capacity of the lung on pulmonary function testing, and this finding is the most commonly reported physiological lung impairment after acute COVID-19.

Van den Borst B, et al.

Roughly half of COVID-19 survivors have persistent and pulmonary radiological changes for up to six months following the acute illness.

Testing

All patients with persistent dyspnea following a COVID-19 illness should undergo **pulmonary function testing and high-resolution chest imaging.**

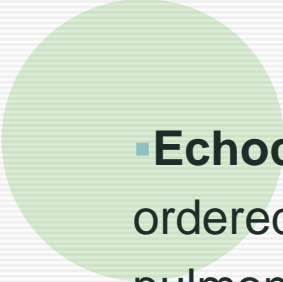
Treatment

Evidence of variable or fixed airflow obstruction can be managed with a trial of **inhaled steroid and long-acting bronchodilator therapy.** If patients do not have a clinical response to inhaled controller therapy, systemic steroids can be prescribed.

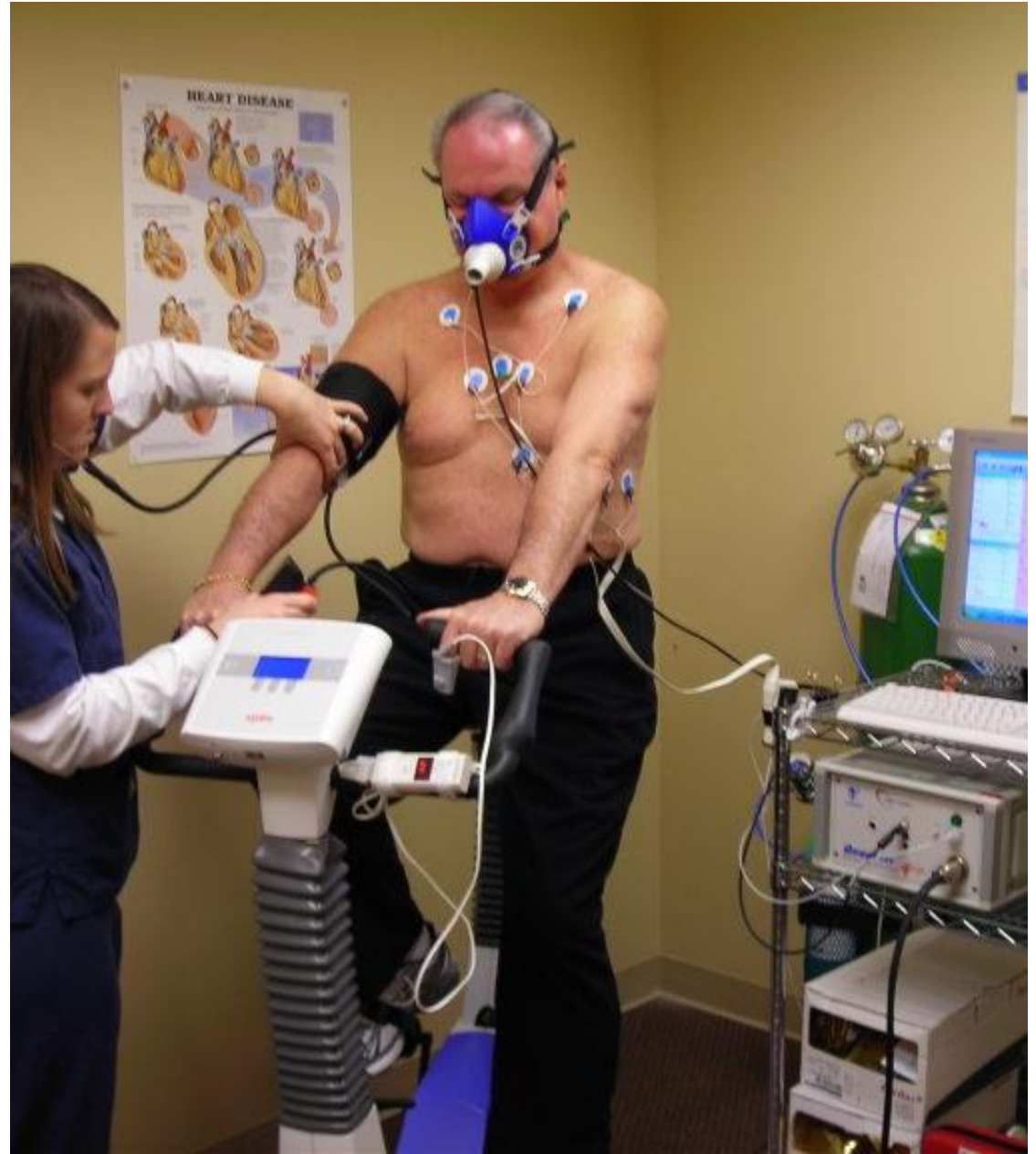
► Persistent Post-COVID-19 Interstitial Lung Disease. An Observational Study of Corticosteroid Treatment

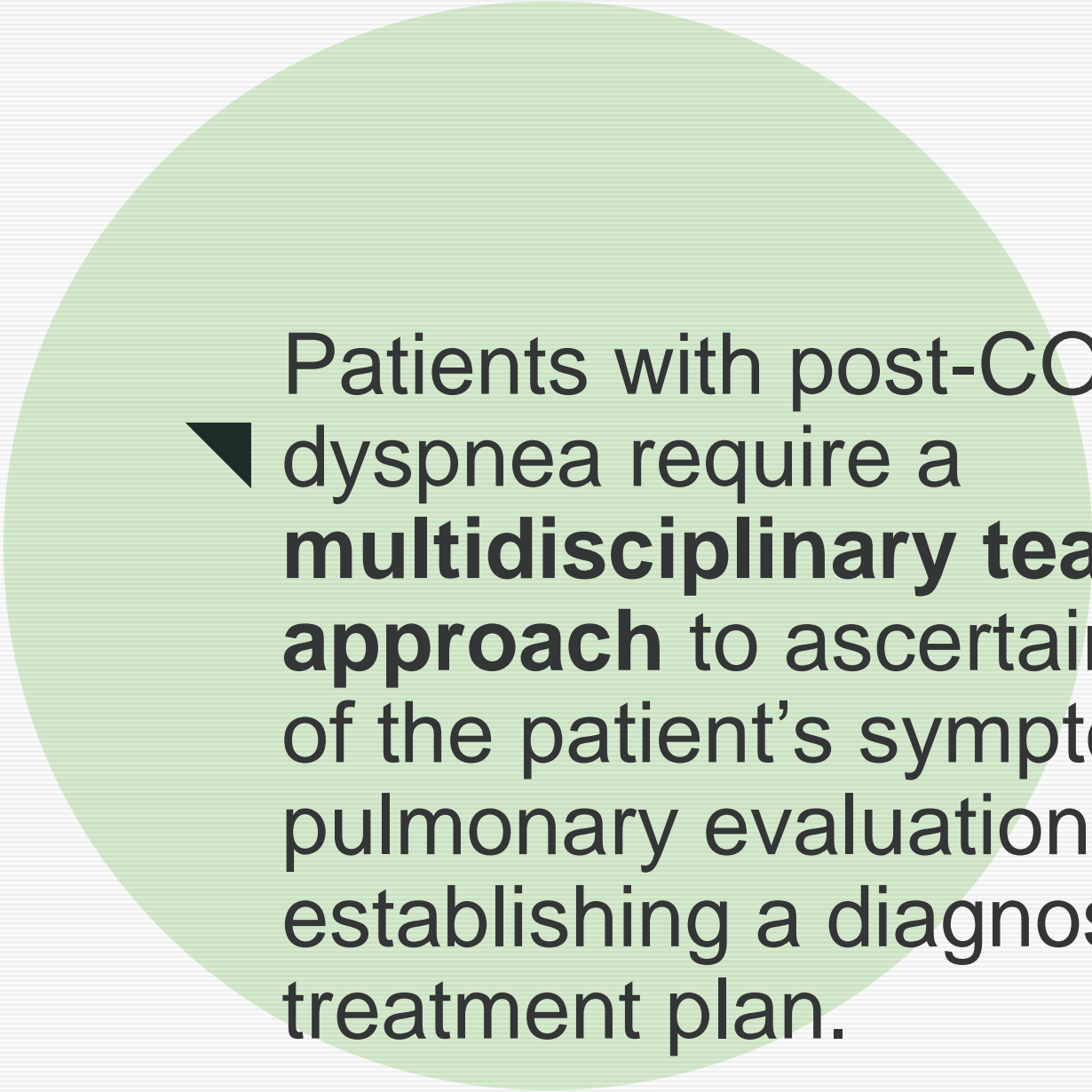
- Annals of American Thoracic Society- 2021
- Observational Study of Corticosteroid Treatment
- 837 patients , 325 had on going symptoms (39%)
- ILD, PNA, functional deficit in 35 (4.8%)
- Given corticosteroids- significant improvement in radiological, and mean of 9.6% increase FVC

doi: [10.1513/AnnalsATS.202008-1002OC](https://doi.org/10.1513/AnnalsATS.202008-1002OC).

- 
- **Echocardiogram** and **ventilation/perfusion lung scanning** can be ordered in the evaluation of persistent dyspnea, particularly if the pulmonary evaluation is unrevealing.
 - **Invasive cardiopulmonary exercise testing** can be performed if the pulmonary and cardiac evaluations are unremarkable.
 - For unexplained dyspnea following COVID-19 illness, we recommend referral to **pulmonary rehabilitation**.
 - Patients can also be referred to **speech-language pathologists** for the evaluation of dysfunctional breathing patterns. This condition can be successfully managed with **respiratory retraining therapy**.

- Cardiopulmonary exercise testing (CPET)



- 
- ▶ Patients with post-COVID-19 dyspnea require a **multidisciplinary team approach** to ascertain the cause of the patient's symptoms, and the pulmonary evaluation is critical to establishing a diagnosis and treatment plan.



<https://recovercovid.org/>

RECOVER: Researching COVID to Enhance Recovery
The National Institutes of Health (NIH) created the RECOVER Initiative to learn about the long-term effects of COVID.

The goal of RECOVER is to rapidly improve our understanding of and ability to predict, treat, and prevent PASC (post-acute sequelae of SARS-CoV-2), including Long COVID.



Questions!

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THANK YOU!



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Cleveland Clinic

Every life deserves world class care.