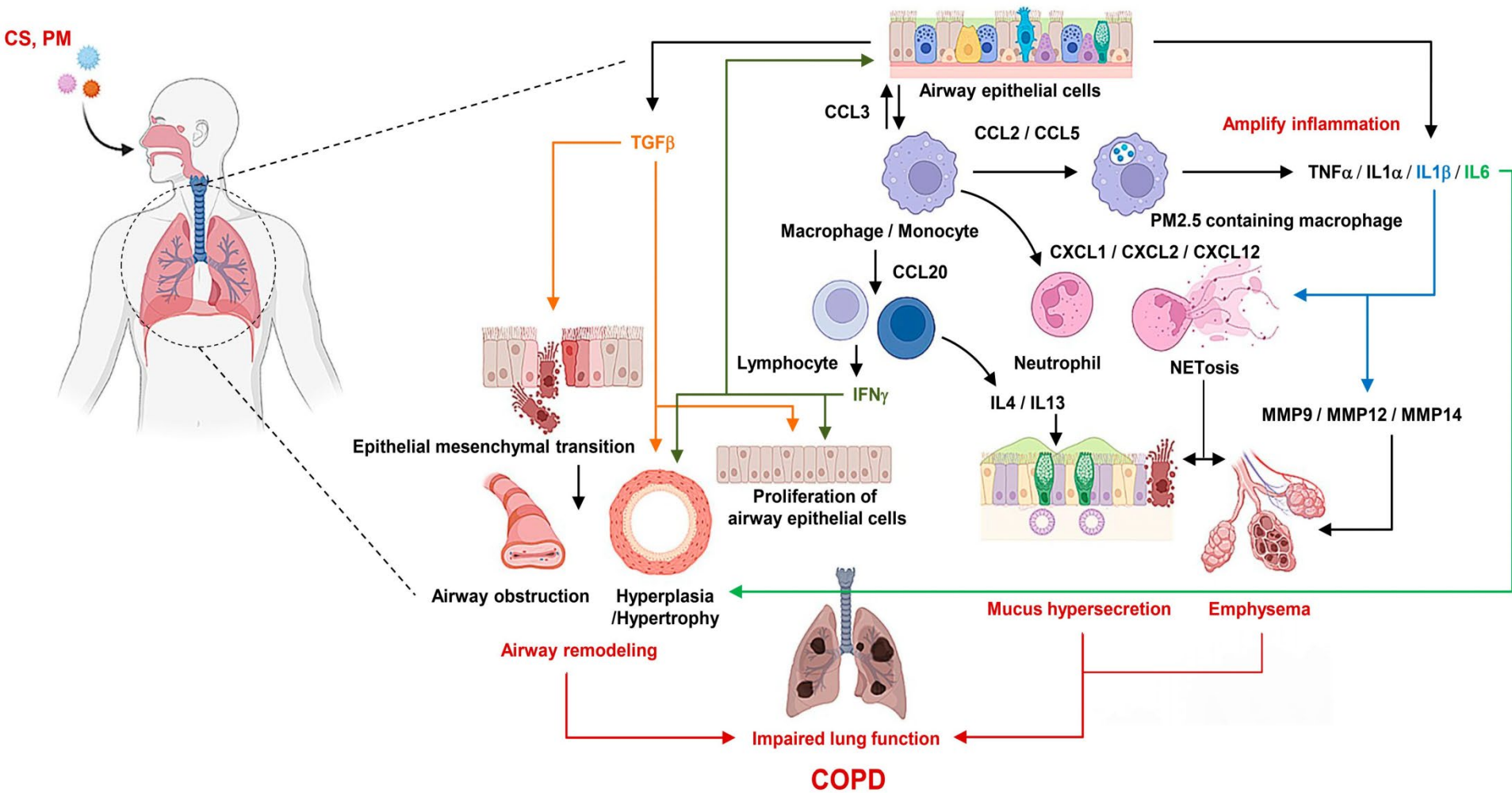


**Stabil ve Alevlenmeye Yatkın
KOAHA'nın Tanımlanması, İzlenmesi ve Yönetimi
GOLD 2026 Önerileri**

Dr.Özlem Şengören Dikiş

13.06.2026



KOAH

Kronik inflamasyon ve yapısal remodeling ile progresif hava akımı kısıtlanmasına yol açan, stabil dönemler ve alevlenmelerle seyreden sistemik, heterojen bir hastalıktır.

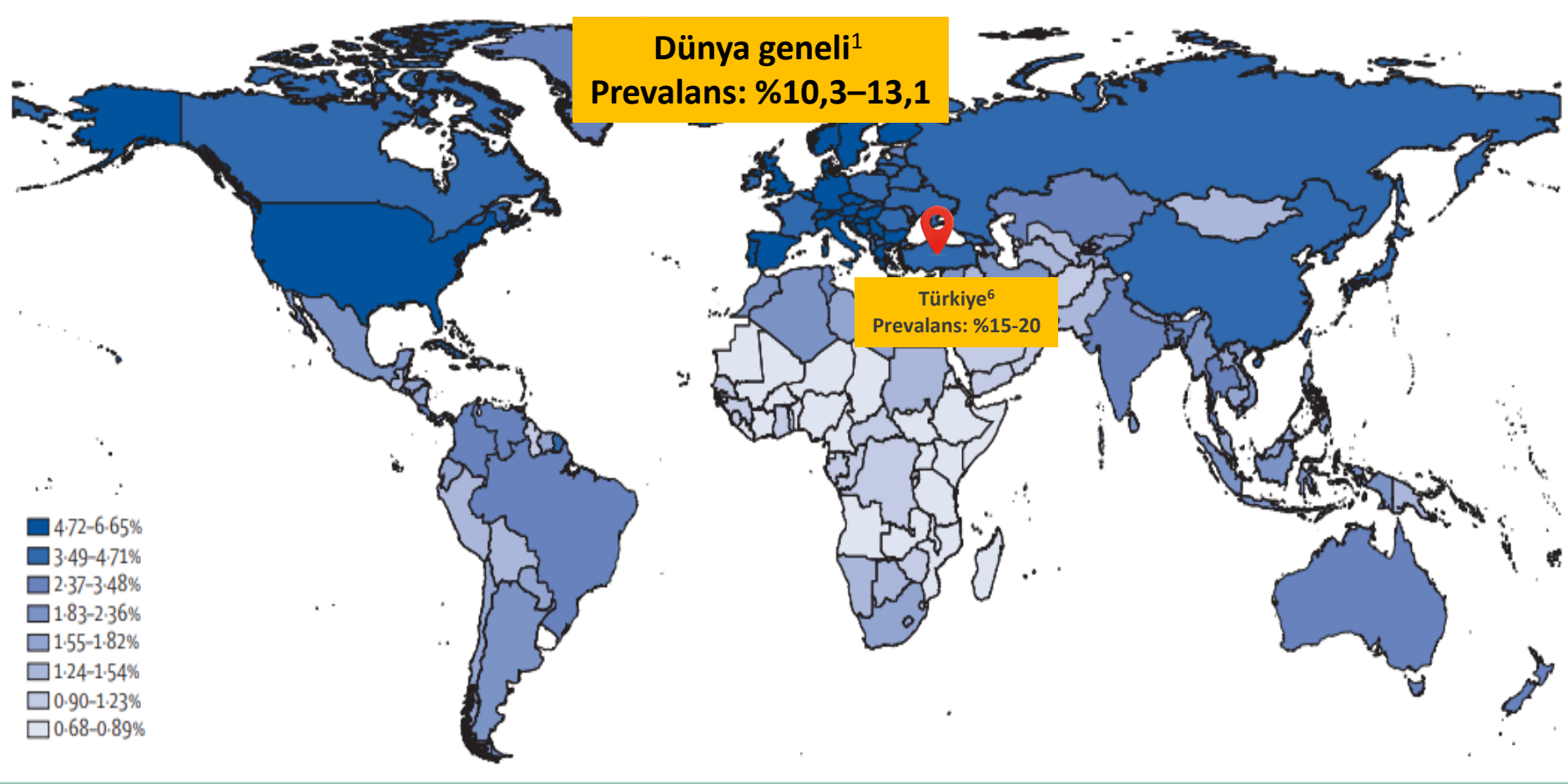


Figure 2: Global crude percentage prevalence of COPD by country, according to GBD 2021
COPD=chronic obstructive pulmonary disease. GBD=Global Burden of Diseases, Injuries, and Risk Factors Study.

KOAH'ın Küresel ve Türkiye'deki Prevalans Yüğü

Türkiye'deki yüksek prevalans yüğü
Sigara kullanımı , biyokütle maruziyeti, hava kirliliğı, yaşlanan nüfus

Table 3. Ten Leading Causes of Death, by Income Group, 2030 (Baseline Scenario)

Income Group	Rank	Disease or Injury	Percent of Total Deaths
World	1	Ischaemic heart disease	13.4
	2	Cerebrovascular disease	10.6
	3	HIV/AIDS	8.9
	4	COPD	7.8
	5	Lower respiratory infections	3.5
	6	Trachea, bronchus, lung cancers	3.1
	7	Diabetes mellitus	3.0
	8	Road traffic accidents	2.9
	9	Perinatal conditions	2.2
	10	Stomach cancer	1.9
High-income countries	1	Ischaemic heart disease	15.8
	2	Cerebrovascular disease	9.0
	3	Trachea, bronchus, lung cancers	5.1
	4	Diabetes mellitus	4.8
	5	COPD	4.1
	6	Lower respiratory infections	3.6
	7	Alzheimer and other dementias	3.6
	8	Colon and rectum cancers	3.3
	9	Stomach cancer	1.9
	10	Prostate cancer	1.8
Middle-income countries	1	Cerebrovascular disease	14.4
	2	Ischaemic heart disease	12.7
	3	COPD	12.0
	4	HIV/AIDS	6.2
	5	Trachea, bronchus, lung cancers	4.3
	6	Diabetes mellitus	3.7
	7	Stomach cancer	3.4
	8	Hypertensive heart disease	2.7
	9	Road traffic accidents	2.5
	10	Liver cancer	2.2
Low-income countries	1	Ischaemic heart disease	13.4
	2	HIV/AIDS	13.2
	3	Cerebrovascular disease	8.2
	4	COPD	5.5
	5	Lower respiratory infections	5.1
	6	Perinatal conditions	3.9
	7	Road traffic accidents	3.7
	8	Diarrhoeal diseases	2.3
	9	Diabetes mellitus	2.1

Table 5. Changes in Rankings for 15 Leading Causes of DALYs, 2002 and 2030 (Baseline Scenario)

Category	Disease or Injury	2002 Rank	2030 Ranks	Change in Rank
Within top 15	Perinatal conditions	1	5	-4
	Lower respiratory infections	2	8	-6
	HIV/AIDS	3	1	+2
	Unipolar depressive disorders	4	2	+2
	Diarrhoeal diseases	5	12	-7
	Ischaemic heart disease	6	3	+3
	Cerebrovascular disease	7	6	+1
	Road traffic accidents	8	4	+4
	Malaria	9	15	-6
	Tuberculosis	10	25	-15
	COPD	11	7	+4
	Congenital anomalies	12	20	-8
	Hearing loss, adult onset	13	9	+4
	Cataracts	14	10	+4
	Violence	15	13	+2
Outside top 15	Self-inflicted injuries	17	14	+3
	Diabetes mellitus	20	11	+9

doi: 10.1371/journal.pmed.0030442.t005

KOAH'ta klinik ve yaşam kalitesi yükü
3,23 milyon ölüm/yıl
≈ 9000 ölüm/gün

Klinik Önceliklerin Deęerlendirilmesi



KOAH Alevlenmeler mi?

Geleneksel akut atak odaklı müdahale modeli, hastalığın asıl ilerleme hızını kontrol altına almakta yeterli midir?



Stabil KOAH mı?

Kesitsel bazda dengede görünen hastaların anlık klinik sapmalarını deęişkenlięin ölçmek öncelikli odağı mı olmalı?



KOAH'ta "Hastalık Stabilitesi"ni sağlamak mı?

Semptom, akcięer fonksiyonu ve atak sıklığı birlikte olarak hedefleyen, en az 12 aylık boylamsal istikrarı yakalamak mı?

Alevlenmeler KOAH'ta FEV1 kaybını hızlandıran temel belirleyici

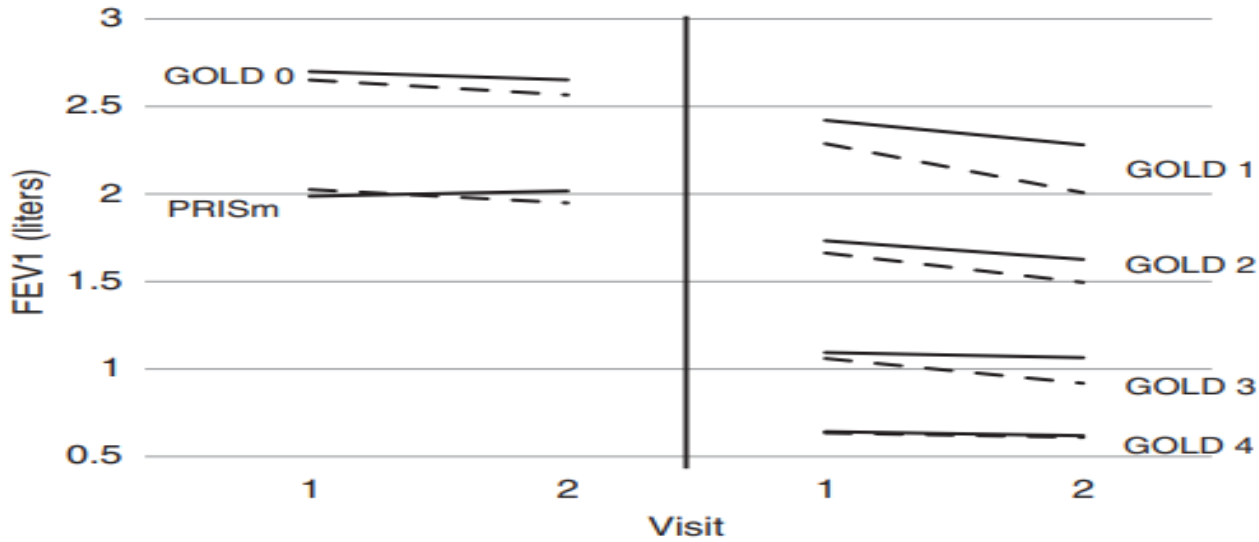


Figure 1. Estimated FEV₁ changes by Global Initiative for Chronic Obstructive Lung Disease (GOLD) group and severe exacerbations status. Estimates were obtained from linear mixed model fits (see STATISTICAL ANALYSES section) for the completer and late subjects. The plot demonstrates that those with at least one severe exacerbation (*dashed lines*) had faster declines in FEV₁, on average, compared with those that did not (*solid lines*), for each GOLD group. PRISm = preserved ratio impaired spirometry.

Etki özellikle erken KOAH evrelerinde belirgin

GOLD 1: Her alevlenme → ek -23 mL/yıl FEV₁ kaybı

Her ağır alevlenme → ek -87 mL/yıl FEV₁ kaybı

GOLD 2-3: FEV₁ kaybı ↓

PRISm: Alevlenme sıklığı yüksek, ek FEV₁ kaybı belirgin değil (PRISm heterojen bir ara fenotip olabilir)

KOAH'ta klinik kontrol (Clinical Control)

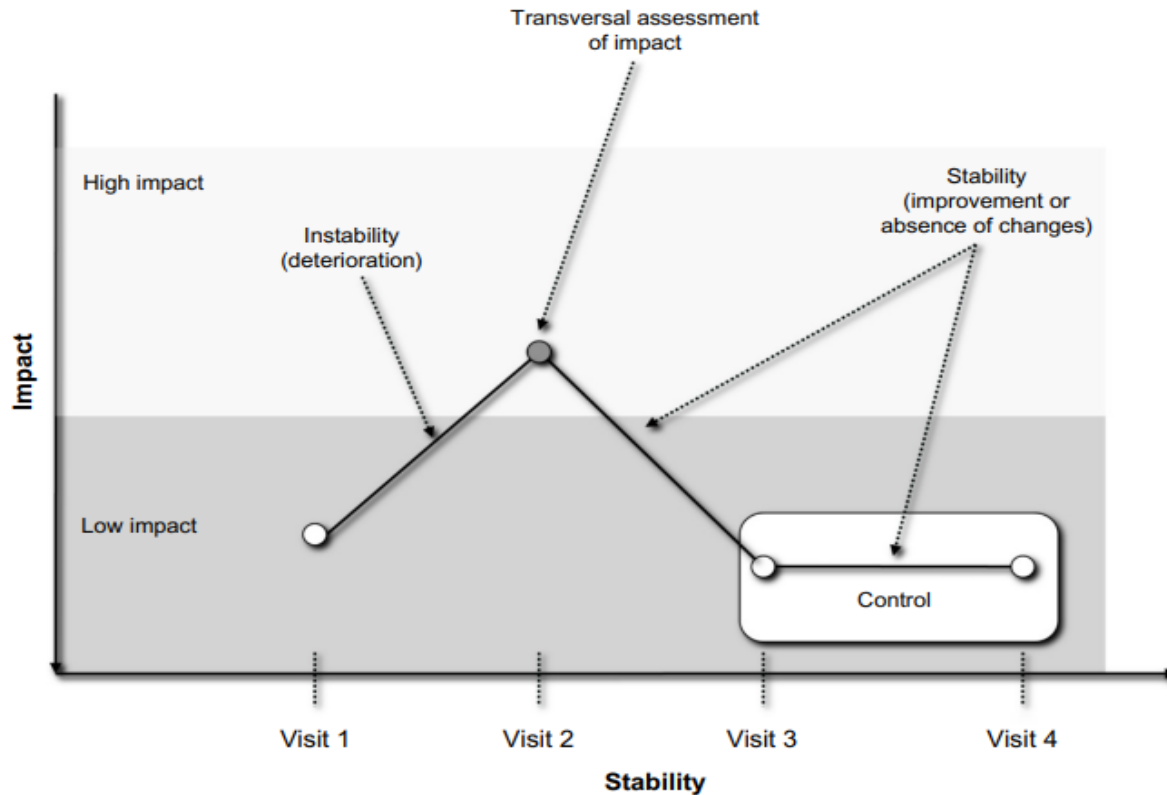


Figure 1 Representation of the concept of impact, stability, and control in COPD.

Notes: The circles represent the transversal measurement of the clinical situation at different times (impact); the lines show the analysis of the changes (degree of stability) and the gray area marks the concept of control understood as the desirable situation in which a condition of low impact is maintained over a long period of time according to the severity of the disease. Copyright © 2014 European Respiratory Society. Reproduced with permission of the European Respiratory Society. *Eur Respir J* erj00644-2014;

Kesitsel etki (cross-sectional impact) ve uzunlamasına /zamansal (longitudinal stability)

Klinik Kontrol Anketi (CC)

Stabilite (Stability)

- Son 3 aydaki alevlenmeler
- Son kontrolden bu yana sağlık algısındaki deęişim

Etki (Impact)

- Kurtarıcı ilaç kullanımı
- Dispne düzeyi
- Günlük yürüme süresi
- Balgam rengi

Questionnaire on clinical control in COPD				
Stability	S ₁	How have you been since your last visit?		
	<input type="checkbox"/> Better	<input type="checkbox"/> The same	<input type="checkbox"/> Worse	
Stability	S ₂	Have you had any exacerbations in the last 3 months?		
	<input type="checkbox"/> No	<input type="checkbox"/> Yes		
Stability	<input type="checkbox"/> Stable (Must meet all criteria)	<input type="checkbox"/> Unstable (If any of the criteria are met)		
	What color has your sputum been in the last few days?			
Impact	<input checked="" type="checkbox"/> White / clear or no sputum	<input type="checkbox"/> Dark		
	How many times have you used the rescue medication in the past week? (Number of occasions rescue medication is required, regardless of the number of inhalations used each time)			
Impact	<input checked="" type="checkbox"/> < 3 times / week	<input type="checkbox"/> ≥ 3 times / week		
	How long (on average) have you walked per day in the last week?			
Impact	<input checked="" type="checkbox"/> ≥ 30 minutes a day	<input type="checkbox"/> < 30 minutes a day		
	What is the current grade of dyspnea (mMRC scale)?			
Impact	<input type="checkbox"/> FEV ₁ ≥ 50% Dyspnea 0 - 1	<input type="checkbox"/> FEV ₁ < 50% Dyspnea 0 - 2	<input type="checkbox"/> FEV ₁ ≥ 50% Dyspnea ≥ 2	<input type="checkbox"/> FEV ₁ < 50% Dyspnea ≥ 3
	<input checked="" type="checkbox"/> Low impact (Must meet 3 of the 4 criteria)	<input type="checkbox"/> High impact (If at least 2 criteria are met)		
Control	<input type="checkbox"/> Grade 0: No dyspnea, except with strenuous exercise			
	<input checked="" type="checkbox"/> Grade 1: Dyspnea when hurrying on a level or when walking up a slight hill			
	<input type="checkbox"/> Grade 2: Dyspnea makes it impossible for them to keep up with other people of the same age on a level, or forces them to stop or rest when walking on level ground at their own pace			
	<input type="checkbox"/> Grade 3: Dyspnea when walking less than 100 meters on level ground			
	<input type="checkbox"/> Grade 4: Dyspnea prevents the patient from leaving home or appears with activities such as dressing or undressing			
Control	Stability <input type="checkbox"/> + <input checked="" type="checkbox"/> Low impact	Instability <input type="checkbox"/> or <input type="checkbox"/> High impact		
	<input type="checkbox"/> Control (Must meet all criteria)	<input type="checkbox"/> No control (If any of the criteria are met)		

Fig. 3. COPD control criteria.

Klinik kontrol kriterlerini karşılayan KOAH...

Table 3 HR of controlled compared to non-controlled patients according to the different criteria used

	HR	95% CI
Control by clinical criteria		
Time to combined event	0.49	0.36–0.66
Time to exacerbation requiring hospitalization	0.27	0.15–0.47
Time to emergency department exacerbation	0.42	0.23–0.77
Time to ambulatory exacerbation	0.53	0.38–0.73
Control by CAT criteria		
Time to combined event	0.82	0.60–1.14
Time to exacerbation requiring hospitalization	0.61	0.33–1.12
Time to emergency department exacerbation	0.72	0.37–1.42
Time to ambulatory exacerbation	0.68	0.48–0.97

Values in bold are statistically significant.

CAT, COPD Assessment Test; COPD, chronic obstructive pulmonary disease; HR, hazard ratio, risk attributable to presenting an event in controlled patients versus non-controlled patients.

- Hastaneye yatış gerektiren alevlenme riski
HR 0,27
- Acil başvuru gerektiren alevlenme riski
HR 0,42
- Ayaktan tedavi edilen alevlenme riski
HR 0,53
- CAT'e göre kontrol sağlanan hastalarda ayaktan alevlenme riski daha düşük
HR 0,68

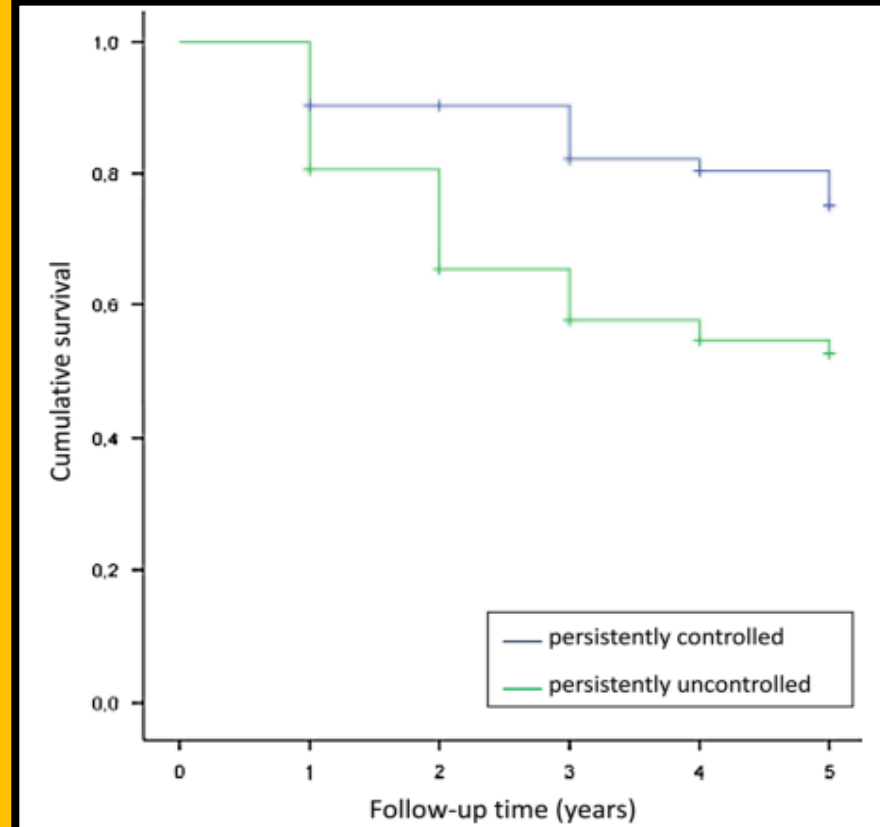


Fig. 3 Kaplan–Meier analysis for all-cause mortality. Persistently controlled patients were associated with a longer survival time than persistently uncontrolled patients

5 yıllık izlemde kontrol kriterlerini karşılayan grupta sağkalım daha yüksek

“Stabil KOAH ” ile “Hastalık stabilitesi ” aynı kavram değil...

2. Materials and Methods

2.1. Study Design and Participants

For this prospective cohort study, patients of the Otto Wagner Hospital, Vienna. A acute exacerbation. Inclusion criteria c least 20 pack-years and spirometrically defined as no acute exacerbation for 8 characterized by a worsening of COPD-

Methods

This cross-sectional comparative in the Department of Pulmonary Institute of Postgraduate Medical Research, Puducherry, India, o months from December 2018 to who were aged ≥ 18 years and clin COPD based on spirometry res Stable COPD patients with medications and not on syste the last three months were patients who presented with ag emergency care were also recruited

Methods

Study Population

We analyzed data from the COPD Phenotyping Study, a collaboration between the University of California Los Angeles (UCLA) and AMGEN (Thousand Oaks, CA).⁶ This prospective, observational cohort study was performed at UCLA from October 2015 to September 2018. The study was approved by the UCLA Institutional Review Board (IRB 14-000748), and written informed consent was obtained from all participants. The participants were 40 to 80 years of age with >10 pack-year smoking history and clinically stable COPD. For subjects with clinically stable COPD, clinical stability was defined as having been on stable medications for COPD for 3 months before Visit 1, not including systemic corticosteroids, and having no history of an exacerbation within the preceding 3 months. The clinically stable subjects were identified at the outpatient clinics of Ronald Reagan University of California and Los Angeles (UCLA) Medical

Stabil KOAH Tanımı

→ Klinik çalışmalarda yaygın giriş kriteri

✓ Alevlenme yok

✓ Tedavi değişikliği yok

✓ Semptomlarda kötüleşme yok

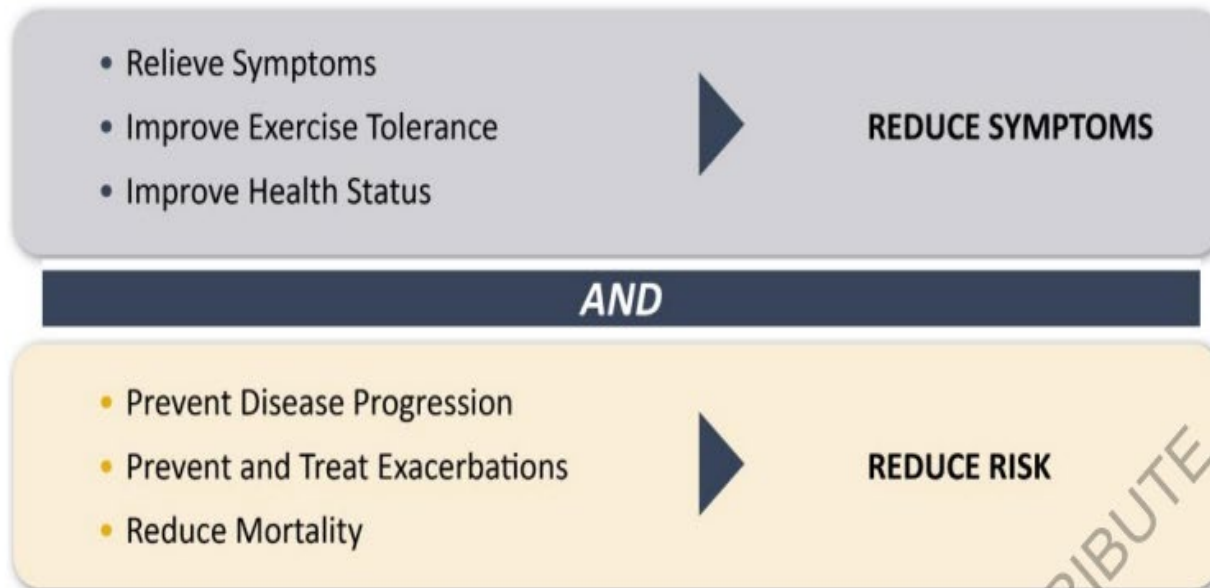
1–4 aylık dönemde klinik stabilite

Hastalık stabilitesi kavramının temel bileşeni



Goals for Treatment of Stable COPD

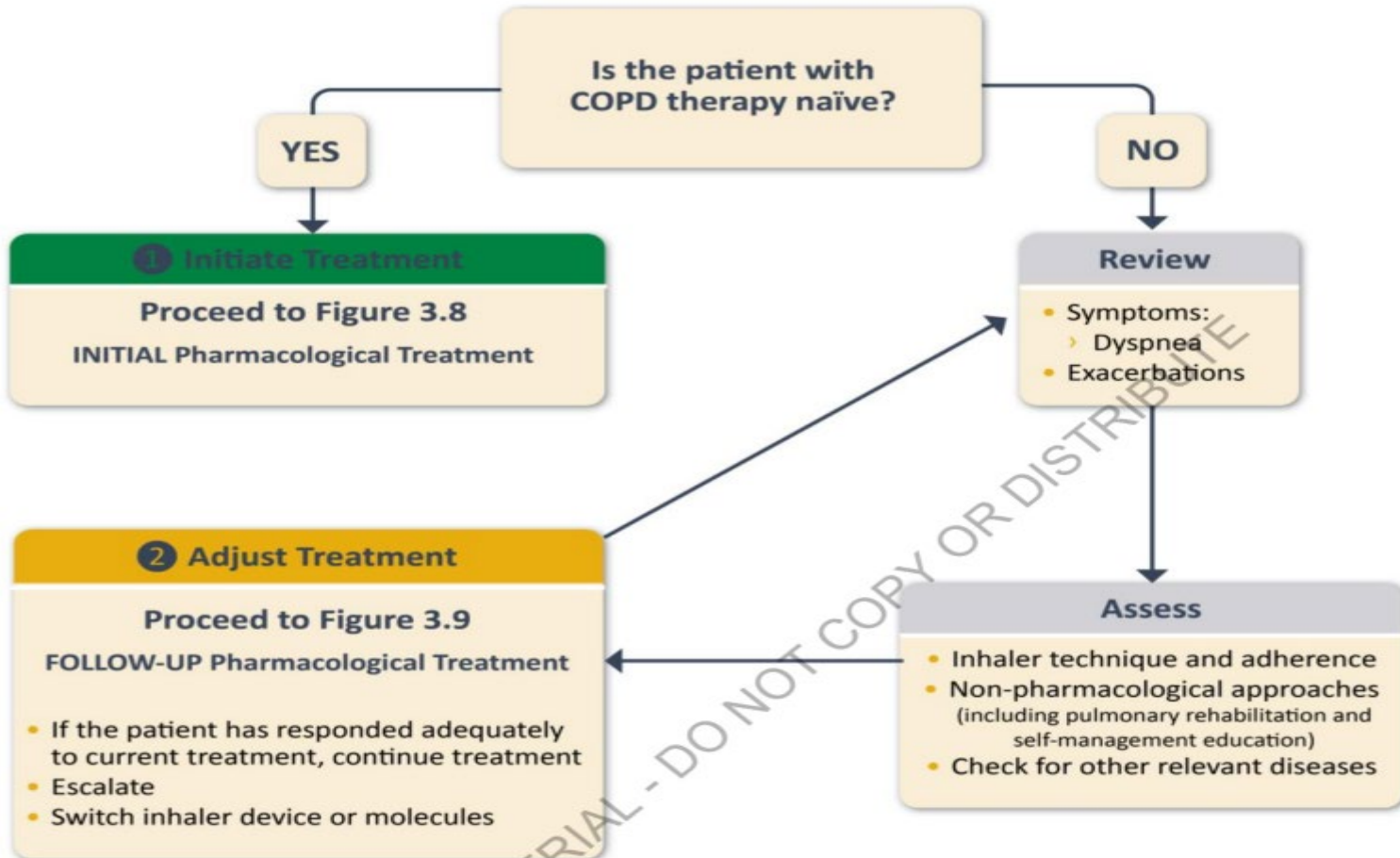
Figure 3.1



Stabil KOAH tedavisinde yalnızca semptom değil, gelecekteki riskin azaltılması

Diagnosis and Management Cycle

Figure 3.7



KOAH yönetimi tek seferlik değil; değerlendir–tedaviyi başlat/optimize et–yanıtı izle

Initial Pharmacological Treatment

Figure 3.8

1 Initiate Treatment

INITIAL treatment - for patients with COPD who are naïve to maintenance pharmacological treatment

EXACERBATION HISTORY (PER YEAR)

One or more (≥ 1)
moderate or severe
exacerbations in the
previous year

Zero (0)
moderate or severe
exacerbations in
the previous year

GROUP E

LABA + LAMA*

consider LABA+LAMA+ICS if blood eos ≥ 300*

GROUP A

A bronchodilator

mMRC 0-1, CAAT < 10

GROUP B

LABA + LAMA*

mMRC ≥ 2 , CAAT ≥ 10

SYMPTOMS

*Single inhaler therapy may be more convenient and effective than multiple inhalers; single inhalers improve adherence to treatment

Exacerbations refers to the number of exacerbations per year; eos: blood eosinophil count in cells per microliter; mMRC: modified Medical Research Council dyspnea questionnaire; CAAT™: Chronic Airways Assessment Test™.

GOLD 2026'da başlangıç tedavi algoritması

Son 1 yılda 1 ve üstü orta veya ağır alevlenme tedavi eskalasyonu

2 Adjust Treatment

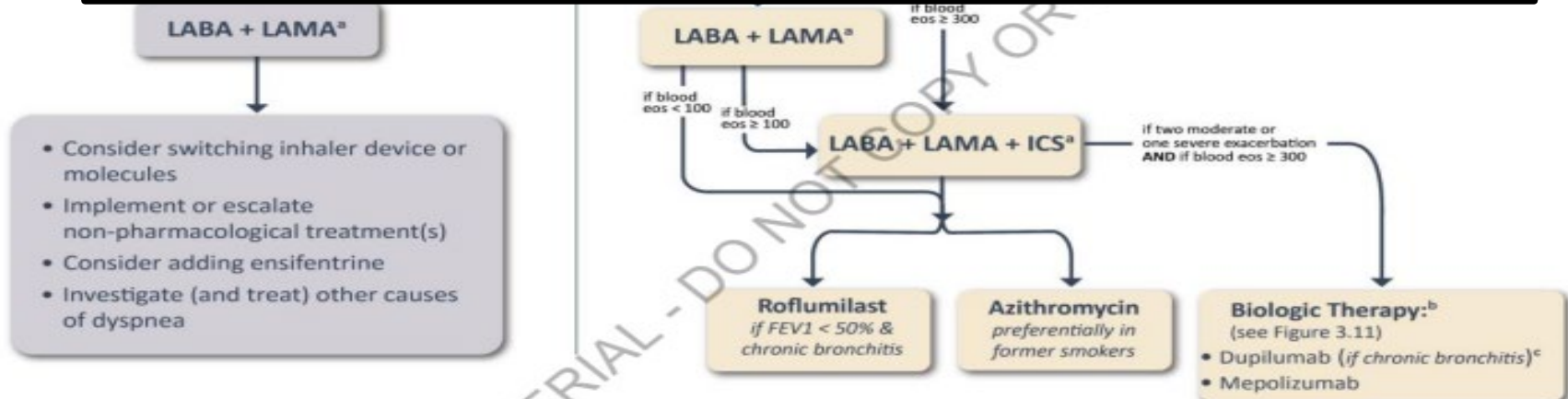
CONTINUE CURRENT TREATMENT

unless dyspnea or exacerbation(s) require optimization

• IF PERSISTENT DYSPNEA

• IF ONE OR MORE MODERATE OR SEVERE EXACERBATION

Treat-to-traits (Tedavi edilebilir özellikler)



Mevcut tedaviye devam et, dispne veya alevlenme varsa tedaviyi optimize et!!!

Abstract

Background: We report the results of the **ETHOS** study with

budesonide/glycopyrrolate/formoterol fumarate (BGF) vs glycopyrrolate/formoterol fumarate (GFF)

and budesonide/formoterol fumarate (BFF) on symptoms and

over 52 weeks in the Phase III ETHOS study of patients with

(UPLIFT) trial (December 2002 - February 2008)

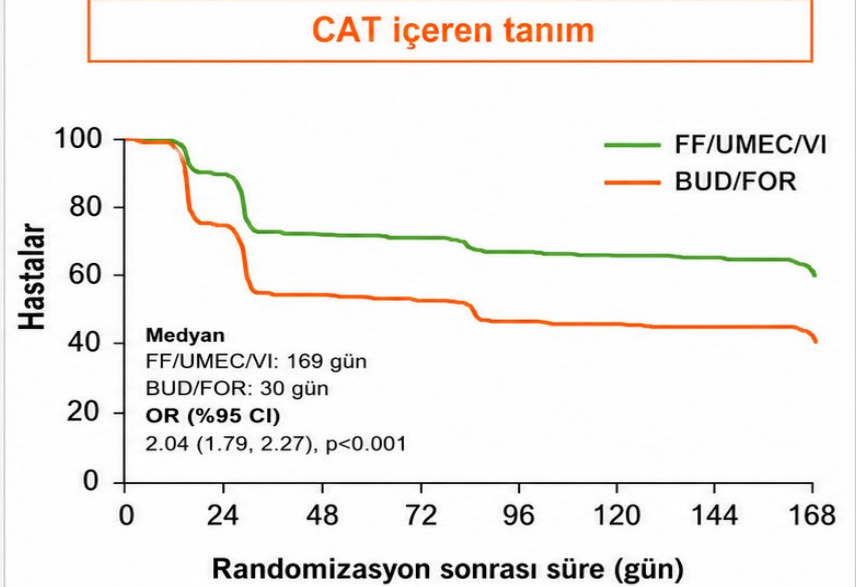
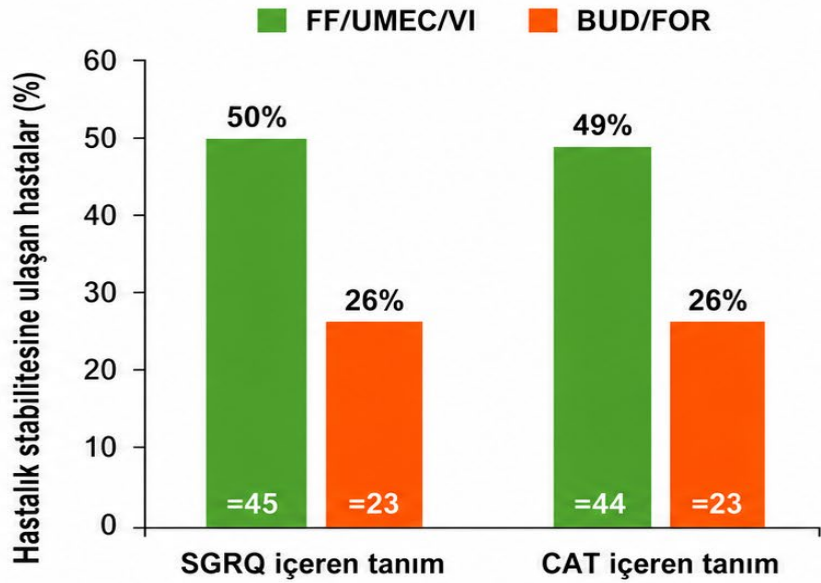
question that this trial was trying to answer is if reducing mortality than either agent alone. The double-blinded trial. The participants were required

The two arms of this trial were tiotropium users and the placebo group which was permitted to use all respiratory medications except inhaled corticosteroids. The primary outcome measured was the rate of decline in the mean FEV1 before and after bronchodilation beginning on day 30. After day 30, the differences between the two groups in the rate of decline in the mean FEV1 before and after bronchodilation were not significant. However, at 4 years and 30 days, tiotropium was associated with a reduction in the risks of exacerbations, related hospitalizations, and respiratory failure. St. George's Respiratory Questionnaire was improved in the tiotropium group, as compared with the placebo group, at each time point throughout the four-year period. This study concluded that Tiotropium treatment in COPD patients was linked to the

Hastalık Stabilitesi Tanımı

- KOAH klinik çalışmalarında yeni bir sonlanım noktası adayı (1-5YIL)
- Mevcut ve yeni tedavilerin etkinliğini değerlendirmeyi kolaylaştırır
- Sadece iyileşmeyi değil, progresyonun önlenmesini de yansıtır

Hastalık stabilitesi ATS ve ERS 2024'te öne çıkan yeni bir tedavi hedefi ...



FF/UMEC/VI ile iki kat fazla hasta 24. haftada hastalık stabilitesine ulaşmış ve stabilitenin korunma süresi 5 kattan daha fazladır.

Singh D ve ark. ATS 2024, A3826.

FULFIL Post Hoc Analizi

Hastalık stabilitesi tanımı

- Orta/ağır alevlenme olmaması
- FEV₁'de <100 mL kötüleşme
- SGRQ'da <4 puan kötüleşme
- CAT'te <2 puan kötüleşme

24. Haftada Hastalık Stabilitesi

SGRQ içeren tanım

FF/UMEC/VI: %50

BUD/FOR: %26

CAT içeren tanım

FF/UMEC/VI: %49

BUD/FOR: %26



- ▶ no accelerated loss of lung function.

While reducing disease activity can prevent symptom worsening and may improve symptoms, additional interventions to further optimize symptom relief may be required. Two terms with similar, but importantly different definitions, have been proposed to describe the clinical state achieved when considering these outcomes over time:

- ▶ disease stability – a low disease activity state with no exacerbations, no worsening of symptoms and no accelerated loss of lung function.⁽⁵⁵⁶⁾
- ▶ disease control – a state of low disease activity, defined by no exacerbations and no worsening of symptoms, plus low impact on the patient defined as symptoms below a threshold value.⁽⁵⁵⁷⁾

Hastalık Stabilitesi

Düşük hastalık aktivitesi

- Alevlenme yok
- Semptom kötüleşmesi yok
- Hızlanmış FEV₁ kaybı yok

Hastalık Kontrol(Klinik kontrol)

Düşük hastalık aktivitesi

- Alevlenme yok
- Semptom kötüleşmesi yok
- Semptomlar belirlenen eşik değerin altında

Sağlık durumu ve semptom stabilitesinin değerlendirilmesi



KOAH özgü anketler

- St. George Solunum Anketi (SGRQ)
- CAAT / CAT
- CCQ = Clinical COPD Questionnaire

Genel anketler

- EuroQoL
- Short-Form Health Survey (SF-36)

CAAT™ Assessment Figure 2.12

For each item below, place a mark (x) in the box that best describes you currently. Be sure to only select one response for each question.

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

Reference: Jones et al. ERJ 2009; 34 (3); 648-54. **TOTAL SCORE:**

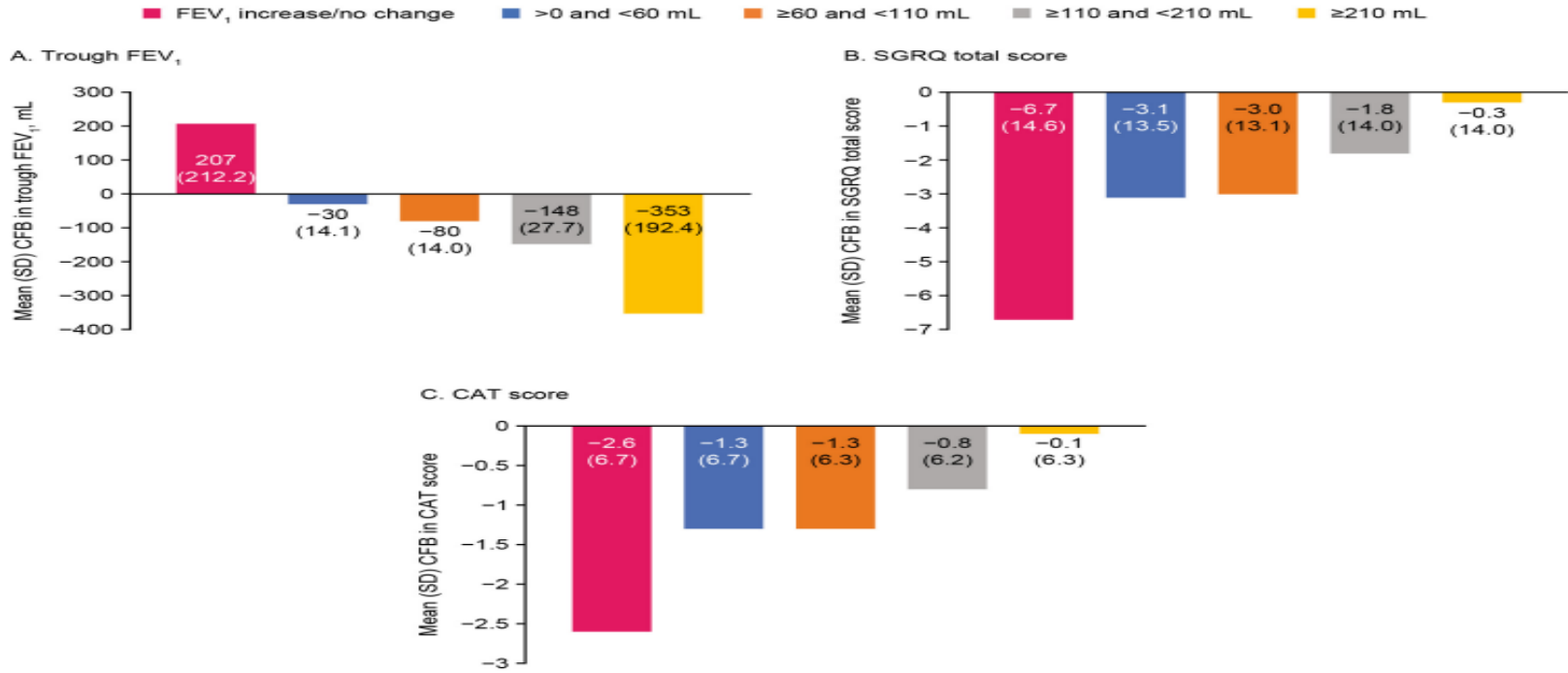
CAT™ has been renamed as the Chronic Airways Assessment Test CAAT™; CAT™ and CAAT™ are equivalent and the scores are interchangeable.

CAAT / CAT Değerlendirmesi

- 8 madde ,Toplam skor: 0–40
- Yüksek skor: Daha fazla semptom yükü
- ≥ 10 puan : Klinik olarak anlamlı semptom düzeyi (SGRQ ≥ 25 'e yaklaşık karşılık gelen pratik eşik değer)

Akciğer fonksiyon kaybının önlenmesi hastalık stabilitesinin temel bileşeni..

Figure 1. Change from Baseline in Trough Forced Expiratory Volume in 1 Second, St George's Respiratory Questionnaire Total Score, and COPD Assessment Test Score at Week 52 Across Forced Expiratory Volume in 1 Second-Decrease Subgroups

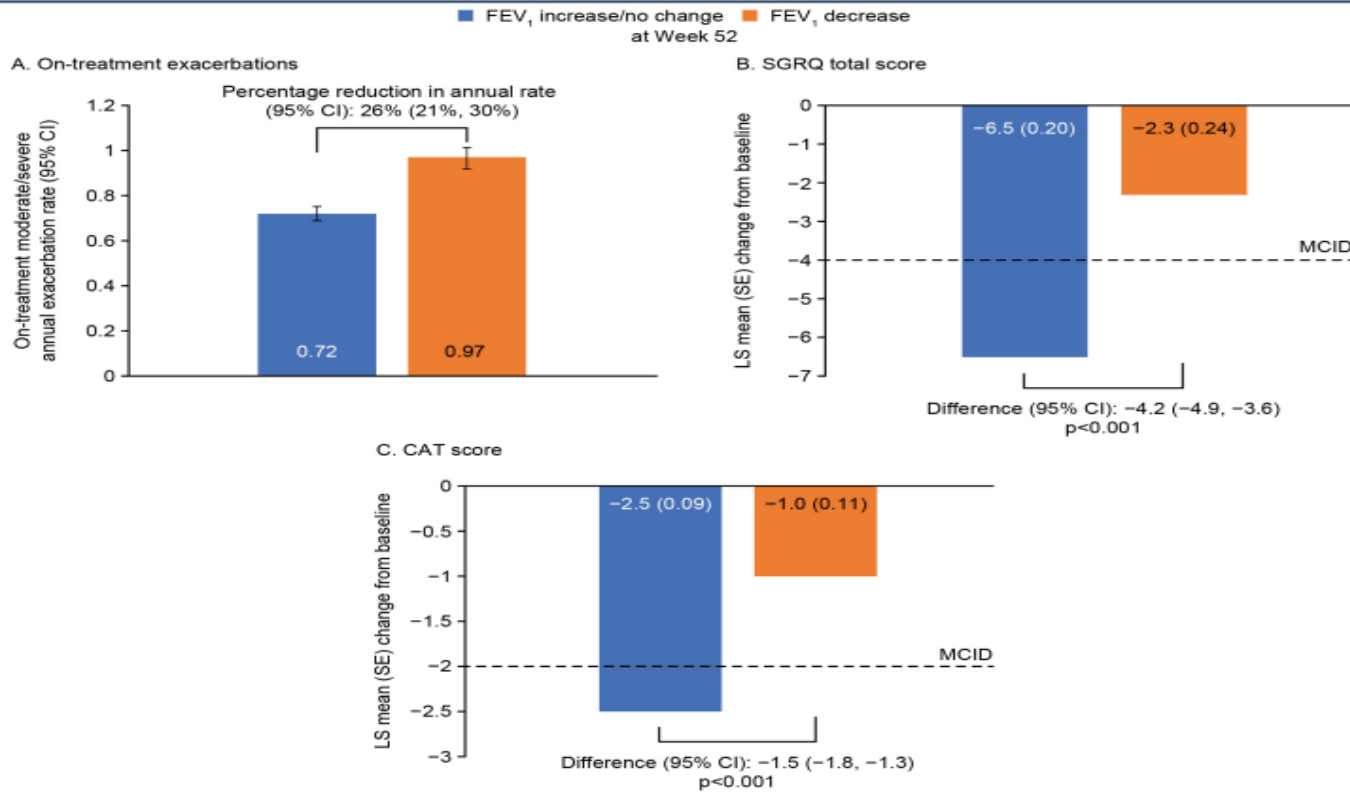


CFB trough FEV₁ at Week 52: >0mL and <60mL, n=795; ≥60mL and <110mL, n=727; ≥110mL and <210mL, n=888; ≥210mL, n=864. CFB SGRQ total score at Week 52: >0mL and <60mL, n=778; ≥60mL and <110mL, n=706; ≥110mL and <210mL, n=865; ≥210mL, n=837. CFB CAT score at Week 52: >0mL and <60mL, n=764; ≥60mL and <110mL, n=692; ≥110mL and <210mL, n=856; ≥210mL, n=826.

IMPACT Çalışması/52 hafta

- FEV₁'i korunan veya artan hastalarda sağlık durumu belirgin olarak daha iyi
SGRQ değişimi: -6,7 puan CAT değişimi: -2,6 puan
- FEV₁ kaybı olan hastalarda
SGRQ değişimi: -0,3 ile -3,1 puan arasında CAT değişimi: -0,1 ile -1,3 puan arasında

Figure 2. Comparison of On-Treatment Moderate/Severe COPD Exacerbations, St George's Respiratory Questionnaire Total Score, and COPD Assessment Test Score, at Week 52 Between Forced Expiratory Volume in 1 Second Decrease and Increase/No Change



Decreases in SGRQ total score and CAT score represent clinical improvement.

FEV₁'de tek ölçüm yerine longitudinal /zamansal FEV₁ eğilimi izlenmeli

IMPACT Çalışması

52. haftada FEV₁'ini koruyan hastalarda alevlenme yükü/semptom yükü düzeyi düşük

Orta/ağır alevlenme oranı: 0,72'ye karşı 0,97

Yıllık alevlenme oranında %26 azalma (%95 GA: %21–30)

Hastalık stabilitesini deęerlendirmede klinik açıdan en anlamlı eşik deęerler nelerdir?

Table 1: Minimal Clinically Important Differences for Commonly Used Outcomes in Chronic Obstructive Pulmonary Disease

Endpoint	MCID (Improvement)	Method of Estimation	Reference
Lung function Trough FEV ₁	100 ml	Anchor-based (exacerbations, patient perception, 2-yr decline in lung function)	9
Exacerbations	No validated MCID	—	—
Dyspnea TDI total score	1 unit	Anchor-based (physician's global evaluation score), distribution-based (SEM, 0.5 SD), expert preference	19
UCSD SOBQ	5 units	Anchor-based (CRQ dyspnea domain, TDI), distribution-based (SEM, Cohen's effect size), estimate by experienced users	20
Health status SGRQ total score	4 units	Anchor-based (MRC dyspnea grade, CRQ dyspnea domain, mortality rate), expert and patient preference	23
CRQ domain scores	0.5 units (average)*	Anchor-based (patient perspectives), distribution-based (SEM, Cohen's effect size), expert panel-based	24
Exercise capacity 6-min walk distance	26 ± 2 m (patients with severe COPD)	Anchor-based (SGRQ, UCSD SOBQ), distribution-based (SEM, Cohen's effect size, empirical rule effect size)	31
Incremental shuttle walking test	47.5 m	Anchor-based (patient perception)	32
Endurance shuttle walking test	45–85 s	Anchor-based (patient perception), distribution-based (0.5 SD)	33
Constant-load cycling endurance tests	46–105 s	Distribution-based (0.5 SD)	8
Dyspnea during exercise tests			
Modified Borg scale	1 unit	Distribution-based (Cohen's effect size)	39
Visual analog scale	10–20 units	Distribution-based (Cohen's effect size)	39

Definition of abbreviations: COPD = chronic obstructive pulmonary disease; CRQ = Chronic Respiratory Questionnaire; MCID = minimal clinically important difference; MRC = Medical Research Council; SGRQ = St George's Respiratory Questionnaire; TDI = Transition Dyspnea Index; UCSD SOBQ = University of California, San Diego Shortness of Breath Questionnaire.

*The MCIDs for the individual domains differ around this mean estimate.

Minimal Klinik Olarak Önemli Farklılık (MCID)

- KOAH'ta akcięer fonksiyonu ve saęlık durumu için tanımlanmış
- Alevlenmeler için tanımlanmış bir MCID yok

	Baseline	Change over 12 months	p value
Mean age (years [SD])	70 (8)
FEV ₁ (% predicted)	47.6 (44.4 to 50.8)	-1.7 (-6.9 to 3.4)	0.51
MRC	3.1 (3.0 to 3.3)	0.1 (-0.1 to 0.2)	0.35
ISW (m)	227 (208 to 247)	-15 (-43 to 12)	0.28
4MGS (m s ⁻¹)	0.92 (0.89 to 0.95)	-0.04 (-0.06 to -0.02)	<0.0001
SGRQ			
Total	50.6 (48.0 to 53.1)	-0.3 (-4.4 to 3.9)	0.91
Symptoms	63.9 (60.9 to 67.0)	-2.5 (-7.4 to 2.4)	0.32
Activities	68.7 (65.4 to 72.0)	1.3 (-3.8 to 6.3)	0.62
Impact	35.9 (33.1 to 38.7)	0.0 (-4.3 to 4.3)	0.99
CAT	20.1 (19.1 to 21.2)	0.6 (-0.4 to 1.5)	0.25

Data are mean (95% CI) unless otherwise specified. FEV₁=forced expiratory volume in 1 s. MRC=Medical Research Council dyspnoea score. ISW=incremental shuttle walk. SGRQ=St George's Respiratory Questionnaire. 4MGS=4 m gait speed. CAT=COPD Assessment Test.

Table 4: Baseline characteristics in patients with stable COPD and change over 12 months (Study 3; n=164)

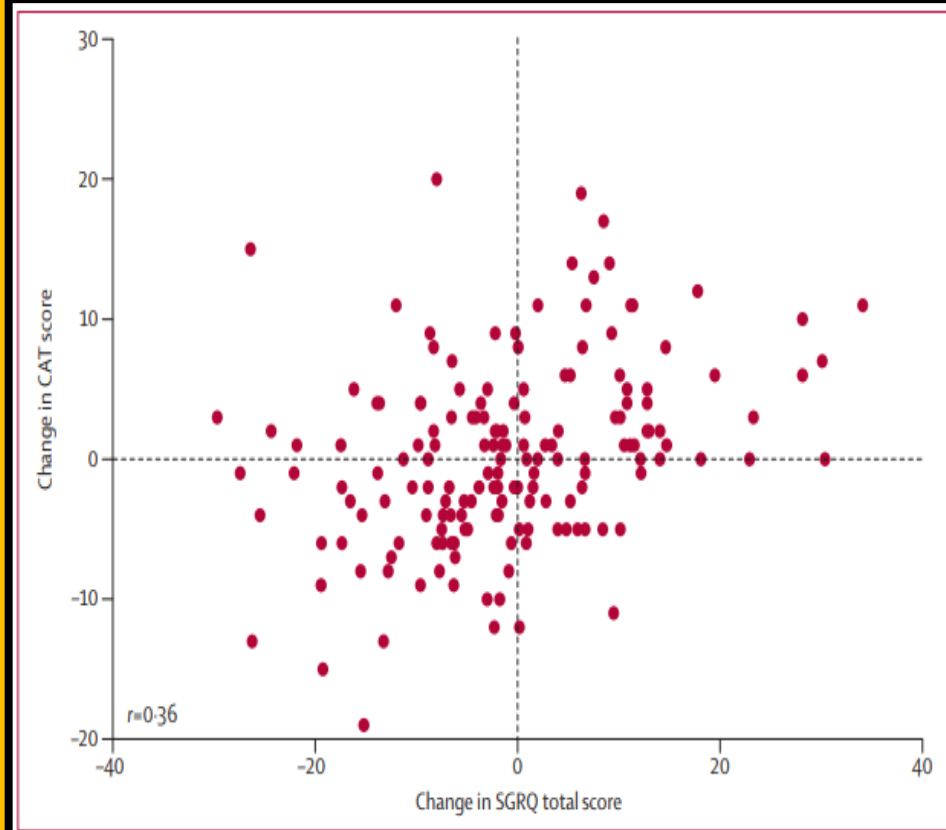


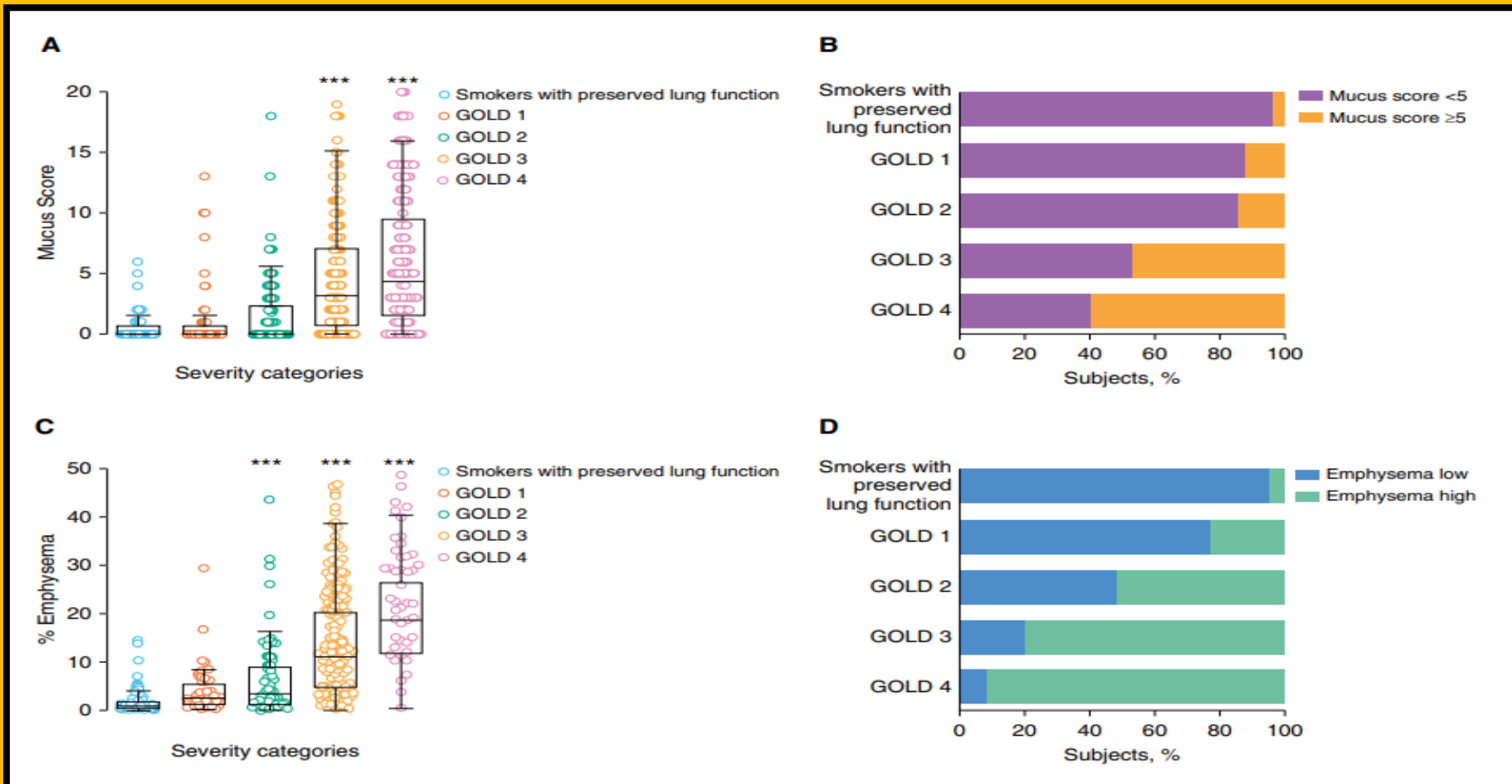
Figure 4: Association between change in CAT score and change in SGRQ score over 12 months in stable patients with COPD (Study 3)

CAT=COPD Assessment Test. SGRQ=St George's Respiratory Questionnaire.

MCID FEV₁ için 100–140 mL mutlak değişim veya %15 değişim

MCID CAT için toplam skorunda >2 puanlık değişim anlamlı

KOAH hastalık stabilitesi ve mukus tıkaç (mucus plug)...



SPIROMICS kohortu, gözlemsel BT-görüntüleme

- 400 smoker / sigarayı bırakmış
- GOLD Evre 3–4'te mukus tıkaçı skoru \geq 5
- Mukus tıkaçı skoru GOLD evresi ile \uparrow

Stabilite /alevlenmede yeni görüntüleme biyobelirteci: Mukus tıkaç (mucus plug) olabilir mi?

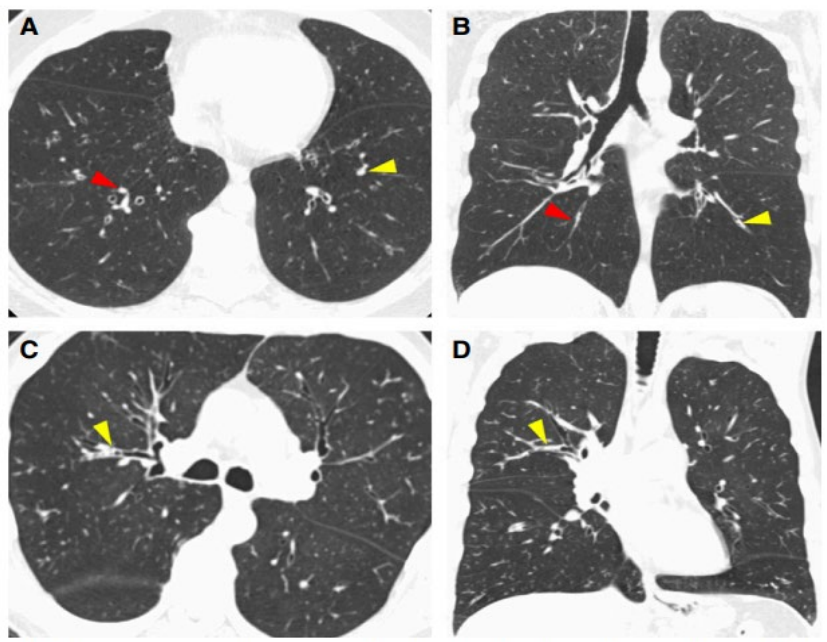
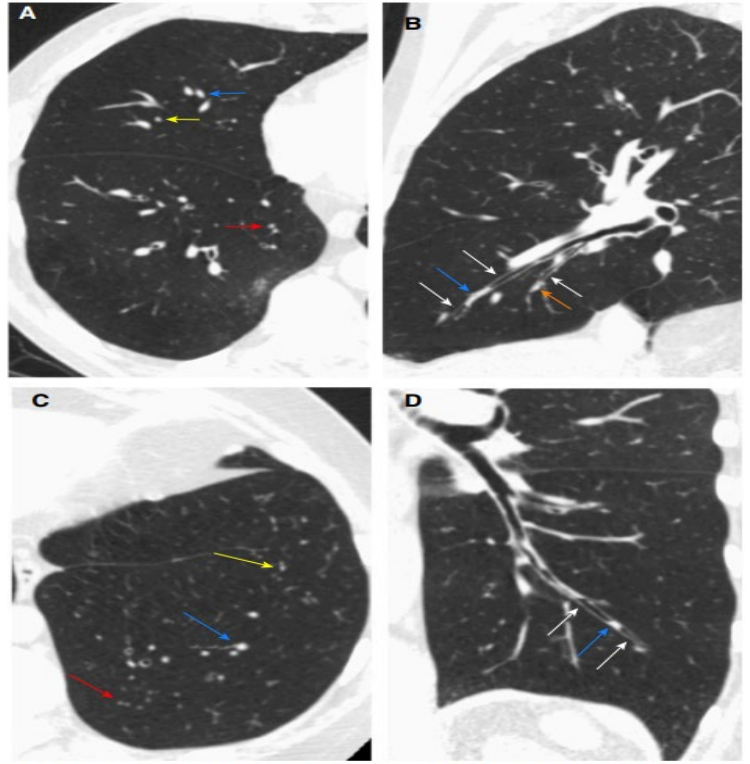


Figure 1. Airway mucus plugs in smokers with chronic obstructive pulmonary disease. (A) An example of two mucus plugs occluding subsegmental airways in the lower lobes identified as tubular opacifications in the axial plane (yellow and red arrowheads). (B) The mucus plugs in the coronal plane (the yellow and red arrowheads indicating the same mucus plugs as in A), revealing that the plugs extend for several millimeters and branch. (C) An example of a branching mucus plug occluding a subsegmental airway. (D) The same plug in the coronal plane.



...s of the lungs.
...ree subsegmental
...lateral segment



Mukus Tıkacı

- BT'de görüntülenebilir ve mukus tıkacı skoru ile değerlendirilebilir
- Yüksek skor, düşük FEV1, yüksek CAAT/CAT skoru, sık alevlenme ve mortalite artışı ile ilişkili
- Kan eozinofil yüksekliği ile ilişkisi, Tip 2 inflamasyon bağlantısı?
- Hastalık stabilitesi ve fenotipleme için aday görüntüleme biyobelirteci

CT an
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bronc
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Hastalık aktivitesi azaltılmasında alevlenme riski ve eozinofil düzeyi temelli ICS kararı

Factors to Consider when Initiating ICS Treatment

Figure 3.10

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 Figures 3.8 & A3.1 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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- KOAH alevlenmesi nedeniyle hastane yatışı öyküsü
- Yılda ≥2 orta şiddette KOAH alevlenmesi
- Kan eozinofil ≥300 hücre/μL
- Astım öyküsü veya eşlik eden astım

Hastalık aktivitesi azaltılmasında alevlenme riski ve eozinofil düzeyi temelli biyolojik tedavi kullanım kararı

Evidence Supporting Use of Biologics in the Treatment of COPD

Figure 3.11

Molecule/RCT*	Key inclusion criteria ^a	Annualized rate of moderate/severe exacerbations	Lung function improvement (pre-BD FEV1) ^d	Quality of life improvement (SGRQ)
Dupilumab (300 mg/2 weeks)				
BOREAS ¹ (n=939)	FEV1 post-BD 30-70% chronic bronchitis ^b eos ≥ 300 (screen)	RR 0.70; P < 0.001	83mL; P < 0.001 (95% CI: 42, 125)	-3.4; P = 0.002 (95% CI: -5.5, -1.3)
NOTUS ² (n=935)	FEV1 post-BD 30-70% chronic bronchitis ^b eos ≥ 300 (screen)	RR 0.66; P < 0.001	62mL; P = 0.02 (95% CI: 11, 113)	-3.4 ^e (95% CI: -5.8, -0.9)
Mepolizumab (100 mg/4 weeks)				
METREO ³ (n=674)	FEV1 post-BD 20-80% eos ≥ 150 (screen) or eos ≥ 300 (previous year)	RR 0.80; NS	19mL; NS (95% CI: -29, 67)	-1.8; NS (95% CI: -4.5, 0.8)
METREX ³ (n=836)	FEV1 post-BD 20-80% eos ≥ 150 (screen) or eos ≥ 300 (previous year) ^c	RR 0.82; P = 0.04	-10mL; NS (95% CI: -54, 33)	0.2; NS (95% CI: -2.8, 3.2)
MATINEE ⁴ (n=804)	FEV1 post-BD 20-80% eos ≥ 300 (screen) and eos ≥ 150 (previous year)	RR 0.79; P = 0.01	-9.0mL; NS (95% CI: -60.1, 42.1)	-2.3; NS (95% CI: -4.6, 0.1)

Boreas, Notus ,Mетро, Metrex, Matinee çalışmaları ve biyolojik ajanlar (DUPİLUMAB,MEPOLİZUMAB)

2 Adjust Treatment

CONTINUE CURRENT TREATMENT

unless dyspnea or exacerbation(s) require optimization

• IF PERSISTENT DYSPNEA

• IF ONE OR MORE MODERATE OR SEVERE EXACERBATION

Treatable- traits



Treat-to-target

- Consider switching inhaler device or molecules
- Implement or escalate non-pharmacological treatment(s)
- Consider adding ensifentrine
- Investigate (and treat) other causes of dyspnea

if blood eos < 100

if blood eos ≥ 100

LABA + LAMA + ICS^a

if two moderate or one severe exacerbation AND if blood eos ≥ 300

Roflumilast

if FEV1 < 50% & chronic bronchitis

Azithromycin

preferentially in former smokers

Biologic Therapy:^b

(see Figure 3.11)

- Dupilumab (if chronic bronchitis)^c
- Mepolizumab

Persistan dispne

- LABA veya LAMA
- LABA + LAMA
- İnhaler cihaz/uyum
- Non-farmakolojik tedaviler gözden geçirilir

Orta/ağır alevlenme

- Kan eozinofil düzeyine göre tedavi yönlendirilir
- LABA + LAMA
- LABA + LAMA + ICS
- Roflumilast, Azitromisin

Biyolojik tedavi

- Dupilumab (Kronik bronşit)
- Mepolizumab

Treat-to- target-Hedefe yönelik tedavi

Belirlenmiş ölçülebilir bir hedefe ulaşana kadar tedaviyi düzenleme/eskale etme yaklaşımı

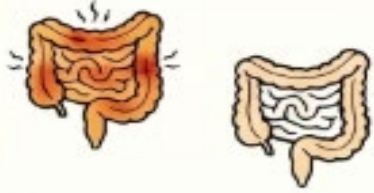
- **Romatoid artrit: Hedef remisyon / düşük hastalık aktivitesi**
- **İBH**
- **Diyabet: Hedef HbA1c düzeyi**
- **Hipertansiyon: Hedef kan basıncı**

KOAH için önerilen hedef: Hastalık stabilitesi

Hastalık kontrolü bozulmaya başladığında ve semptomlar arttığında, henüz akut bir alevlenme gelişmemiş olsa bile, hastanın tedavisini yoğunlaştırabilmesi sağlanmalı

Hastalık stabilitesinin değerlendirilmesinde takip aralıkları ne olmalıdır?

İBH: DİNAMİK TAKİP ARALIĞI



Hastalık aktivitesine göre 3 ile 12 ay arasında değişen periyodik kontroller uygulanır.

İBH modeline benzer şekilde, çoklu kriterler kullanılarak yapılan düzenli hasta değerlendirmesidir.

Kişiselleştirilmiş Takip Faktörleri



Stabilitesi



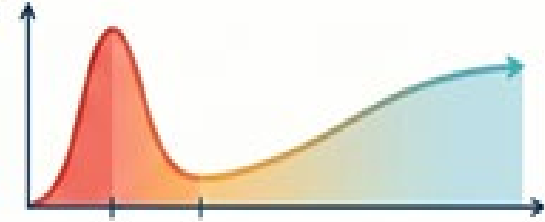
Aktif Kötüleşme Durumu



Alevlenme Geçmişi

Takip süreleri; hastanın stabilitesi, aktif kötüleşme durumu ve alevlenme geçmişine göre belirlenir.

Alevlenme Sonrası İyileşme



Sık ve ağır alevlenme geçiren hastalarda iyileşme süreci daha uzun sürebilmektedir.

Aktif Dönem Takibi



3 - 9 Ay

Önerilen Takip Aralığı



3 - 9 Ay

Önerilen Takip Aralığı



Hastalığın aktif seyrettiği dönemlerde 3 ile 9 ayda bir değerlendirme yapılmalıdır.

Remisyon ve Stabilite Dönemi

Hastalık remisyonunda veya stabil olduğunda takip aralığı aya çıkarılabilir.

STABİL KOAH TAKİBİ



6 - 12 Ay

Önerilen Takip Aralığı



6 - 12 Ay

Önerilen Takip Aralığı





AMAÇ: HASTANIN "KENDİ EN İYİ DURUMUNU- PERSONEL BEST" KORUMAK



Teşekkür ederim...