

**COPD: What does it stand for?**  
**Chronic Obstructive Pulmonary**  
**Disease**

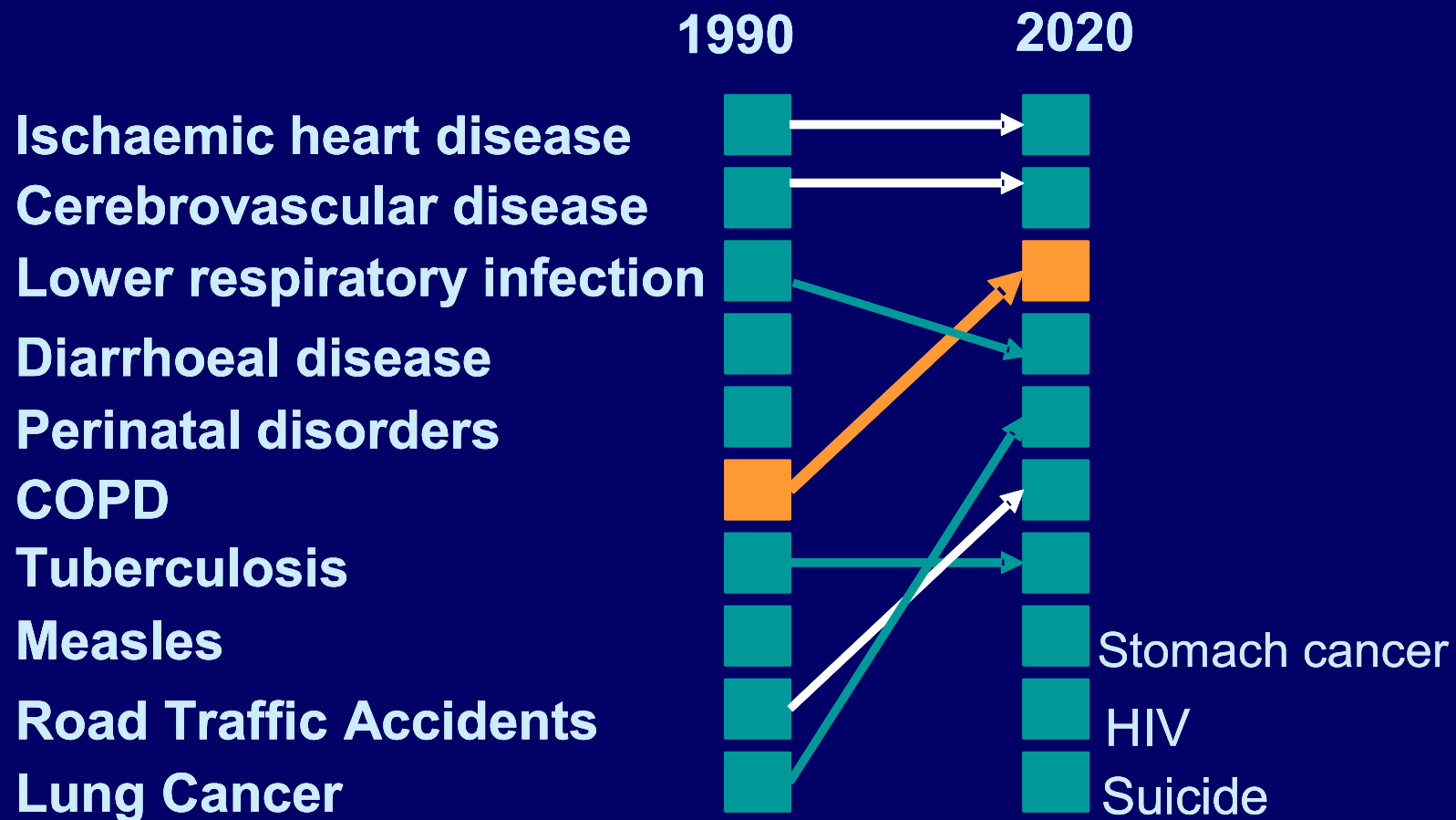
**Professor William MacNee**  
**Past President European Respiratory**  
**Society**

# Definition of COPD

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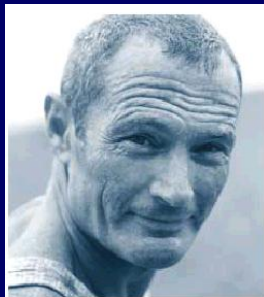
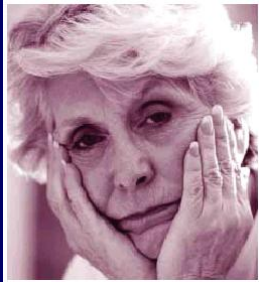
- COPD is a preventable and treatable disease. In the lungs it is characterized by airflow limitation (or obstruction to the flow of air) that is not fully reversible.
- The airflow limitation is usually progressive and associated with an abnormal inflammation in the lungs in response to noxious particles or gases.
- COPD has significant effects out with the lungs that may contribute to the severity in individual patients.

# Global Burden Of Disease Top 10 Causes Of Death 1990 and 2020



# The burden of COPD in Europe

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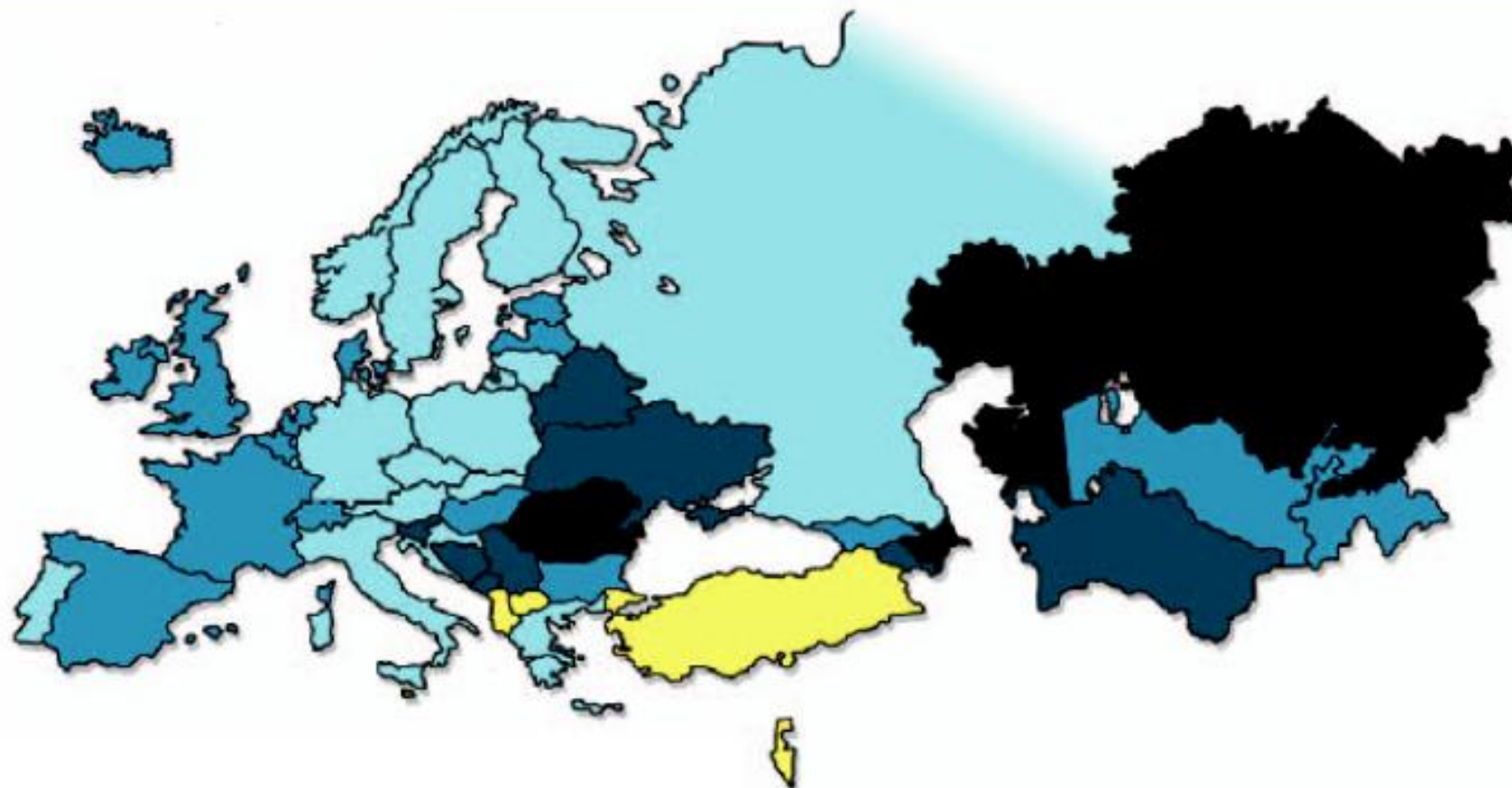
- The frequency of COPD varies in European countries from 4-10% of the adult population, 50/100,000 males ; 20/100,000 females
  - Many more are undiagnosed
  - 2- 300,000 people die in Europe each year because of COPD
  - Estimated direct cost in EU –
    - Out - patient € 4.7 billion
    - In-patient € 2.9 billion
    - Drugs € 2.9 billion
- 41,300 Lost work days/ 100,000 pop<sup>n</sup>
- Productivity losses € 28.5 billion





European Respiratory  
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# Burden of mortality from COPD in Europe



Death rates due to COPD per 100,000    ● >75    ● 51-75    ● 25-50    ● <25    ● No Data

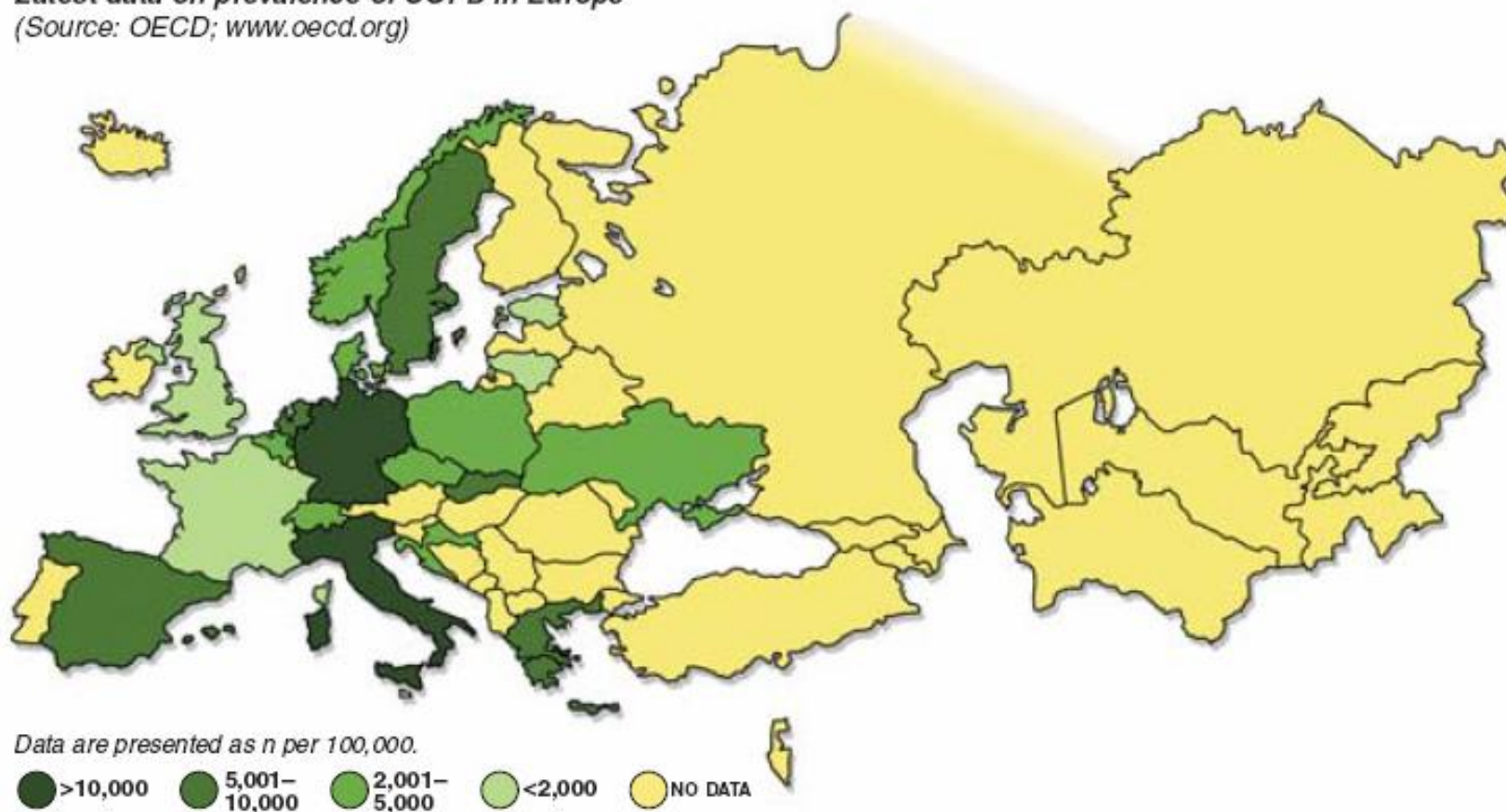
By 2010, COPD will be the fourth leading cause of death in Europe



European Respiratory  
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# Prevalence of COPD in Europe

*Latest data on prevalence of COPD in Europe*  
(Source: OECD; [www.oecd.org](http://www.oecd.org))



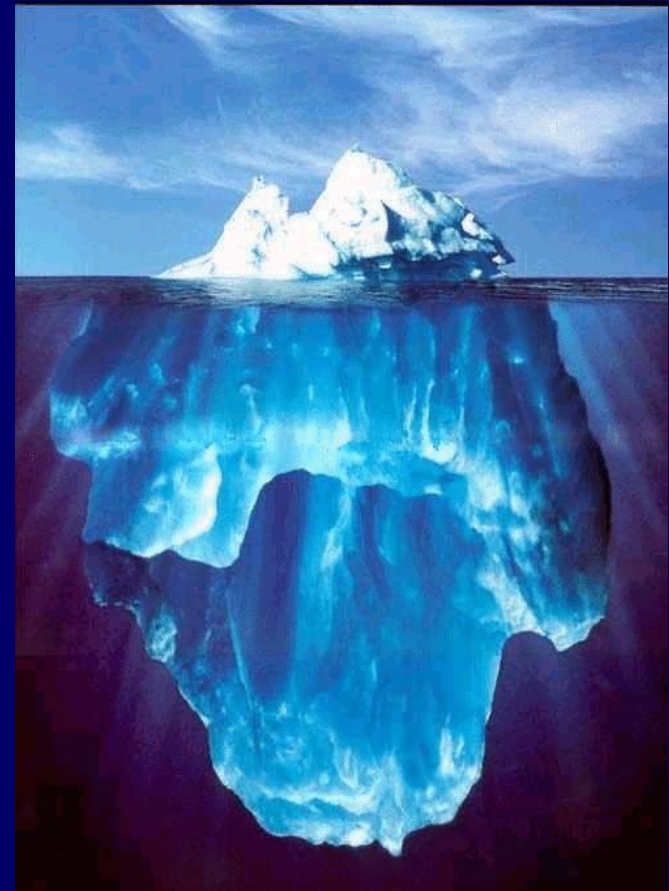
**Better data is urgently needed**

ERS/ELF European White Book 2003

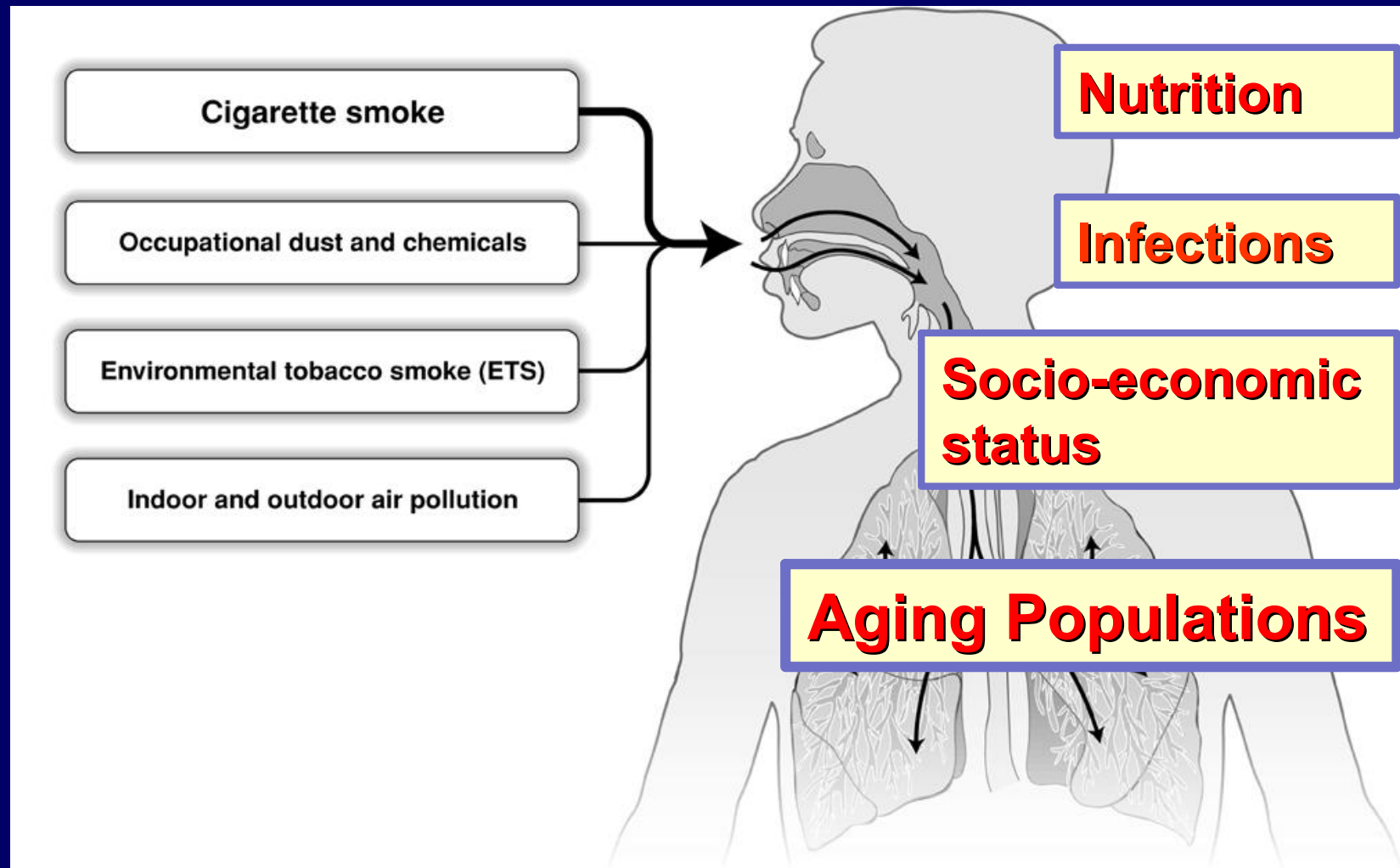
# **COPD is under-recognised and under- diagnosed**

**At least half of those affected with COPD may be undiagnosed**

- **Disease with insidious onset**
- **Often treated as asthma**
- **Smokers don't seek treatment**
- **Can present late with more advanced disease**

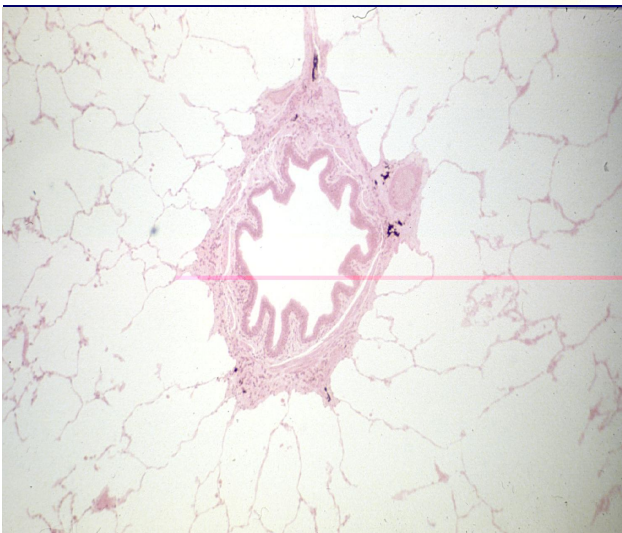
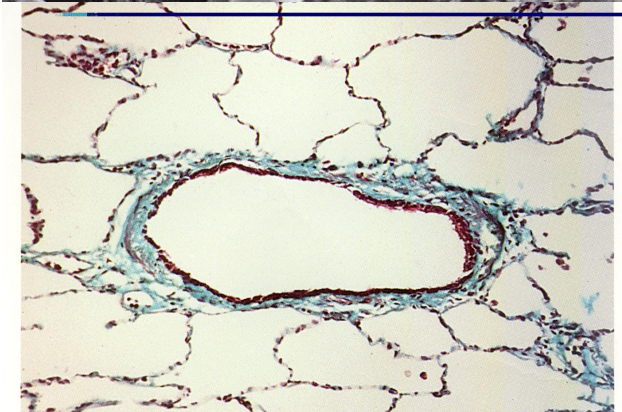
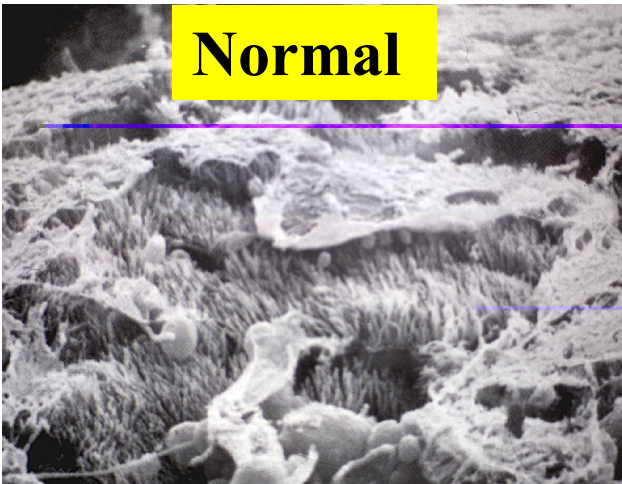


# Risk Factors for COPD





**Normal**

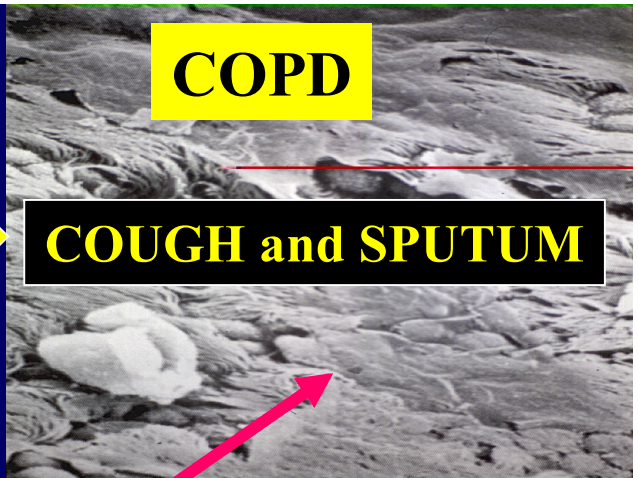


**Chronic  
Bronchitis**

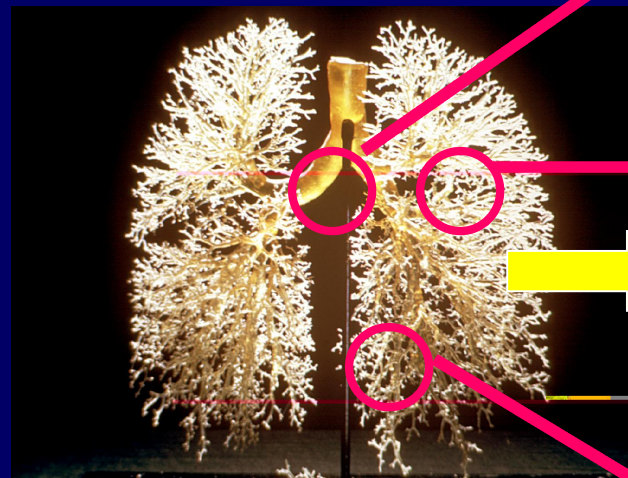


**COPD**

**COUGH and SPUTUM**



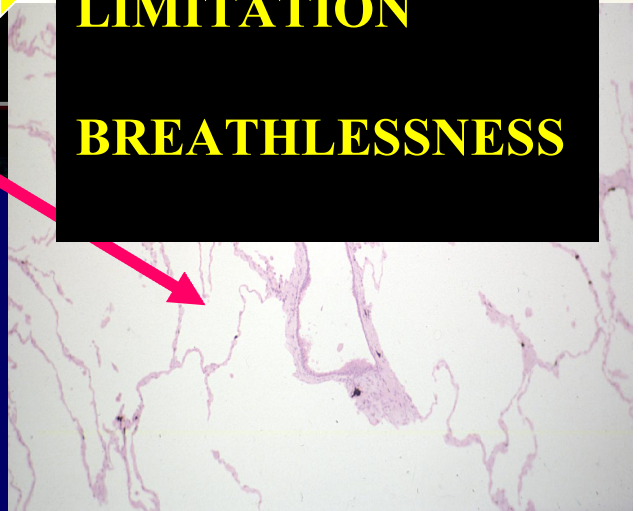
**Bronchiolitis  
Small airways disease**



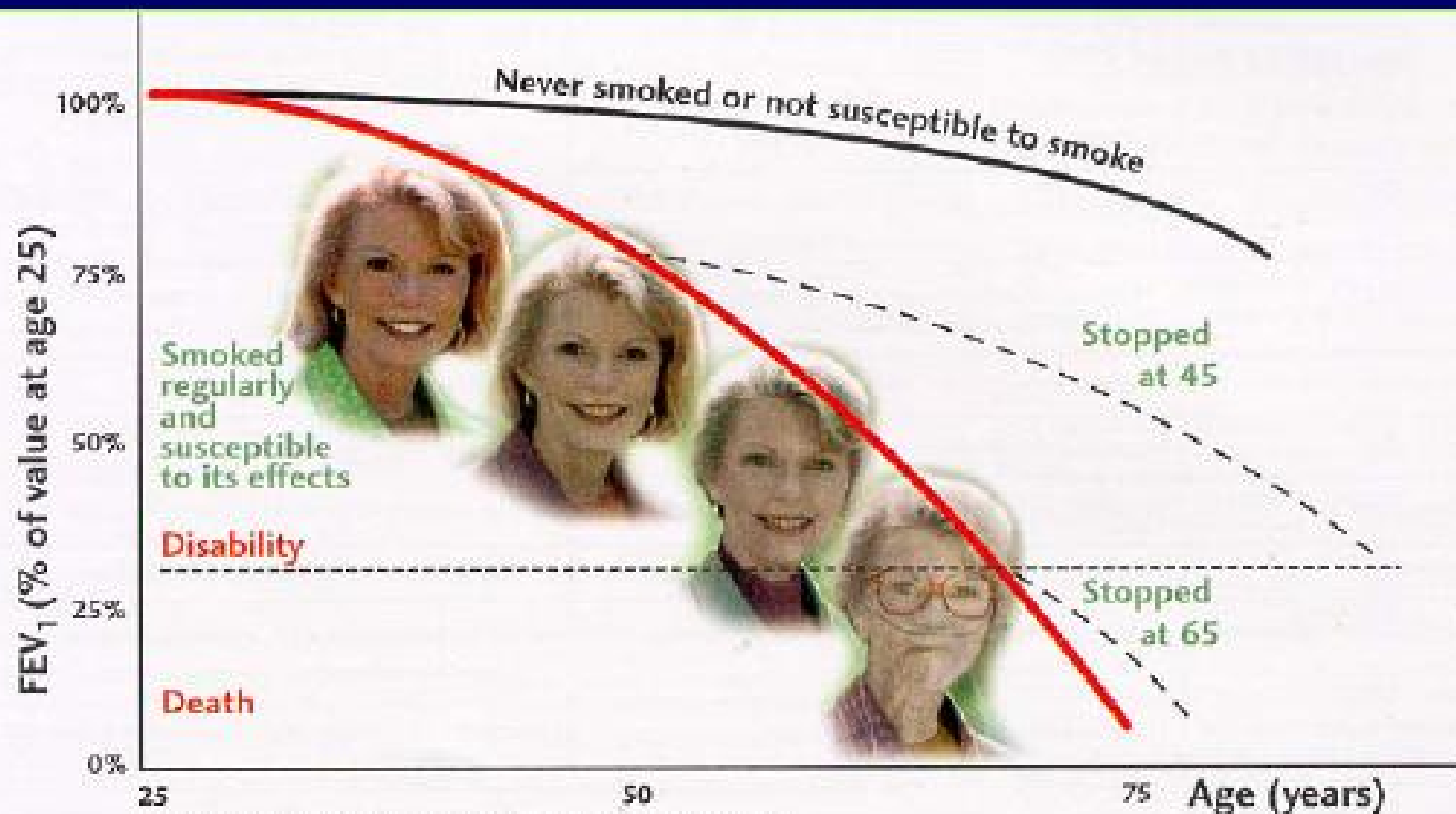
**AIRFLOW  
LIMITATION**

**BREATHLESSNESS**

**Emphysema**



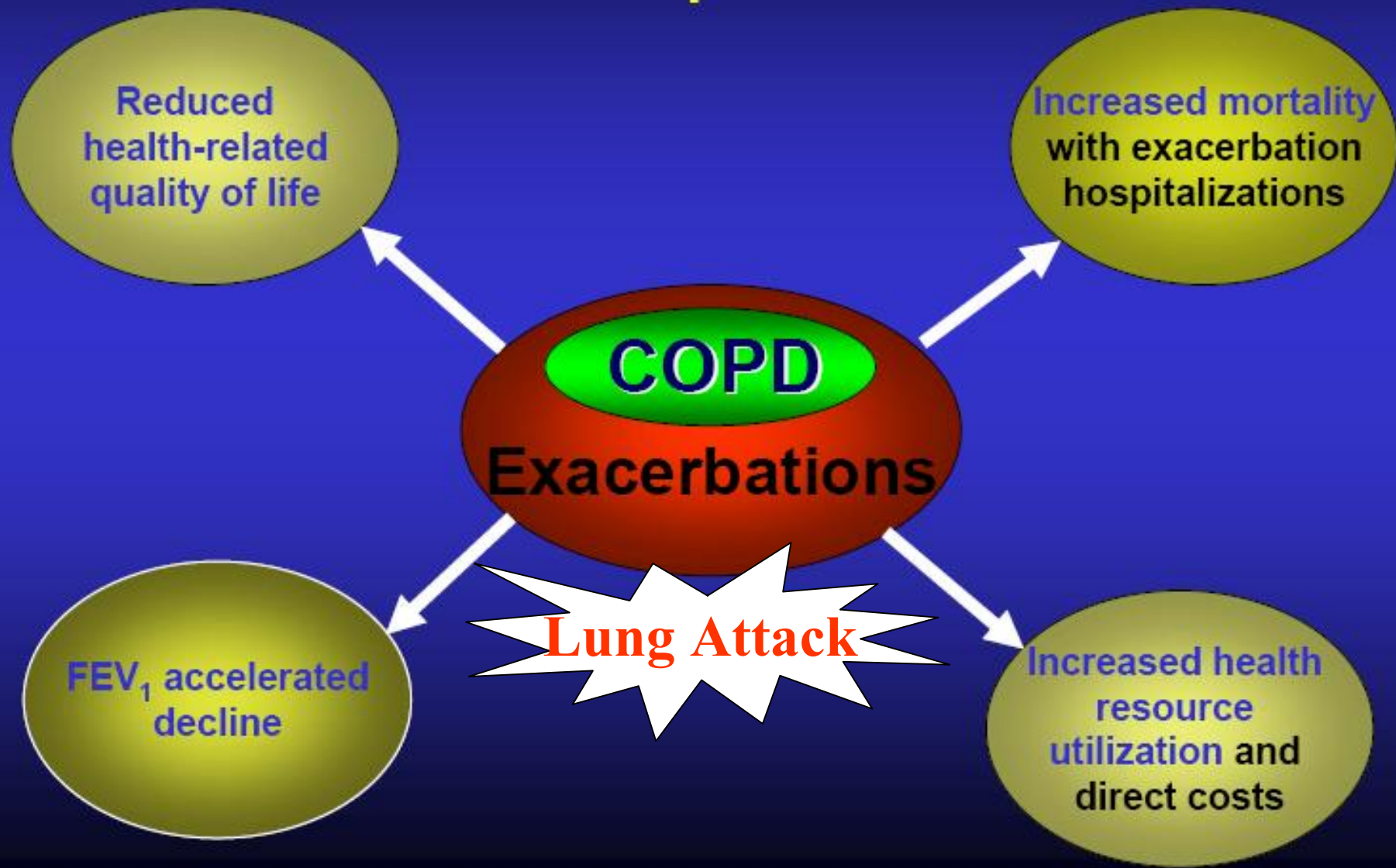
# COPD Progression



Adapted from Fletcher C, Peto R. The natural history of chronic airflow obstruction. *Br Med J* 1975; 1: 1695-1698.



# COPD Exacerbations: Clinical Consequences



# **SYSTEMIC EFFECTS OF COPD - COMORBIDITIES** Target organs

## **Respiratory system**

**Lung infections**

**Lung Cancer**



**Muscle weakness**

**Weight loss**

**Osteoporosis  
( thin bones )**



**Systemic  
inflammation**



## **Other co-morbidities**

- **Depression**
- **Diabetes**
- **Peptic Ulcer**

**Angina**  
**Heart attacks**





# Diagnosis of COPD

## SYMPTOMS

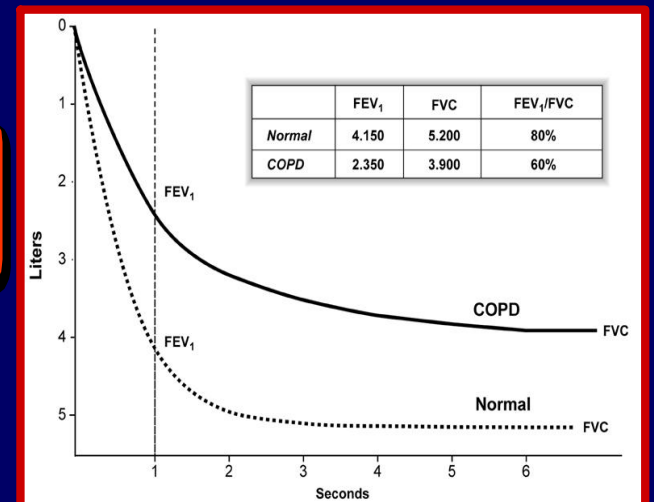
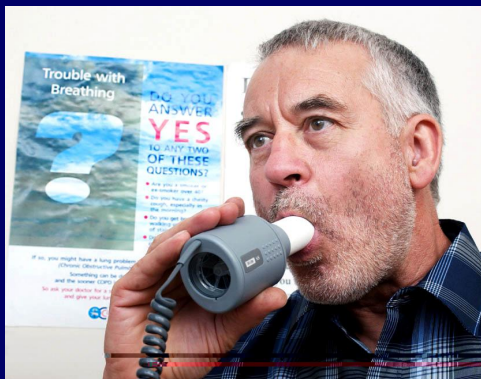
cough  
sputum  
shortness of breath

## EXPOSURE TO RISK FACTORS

tobacco  
occupation  
indoor/outdoor pollution



## SPIROMETRY



# Preventive treatment in COPD

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- Avoidance of risk factors
  - smoking cessation
  - reduction of indoor pollution
  - reduction of occupational exposure
- Influenza vaccination

# GOALS of COPD MANAGEMENT

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- Relieve symptoms
- Prevent disease progression
- Improve exercise tolerance
- Improve health status
- Prevent and treat complications
- Prevent and treat exacerbations
- Reduce mortality

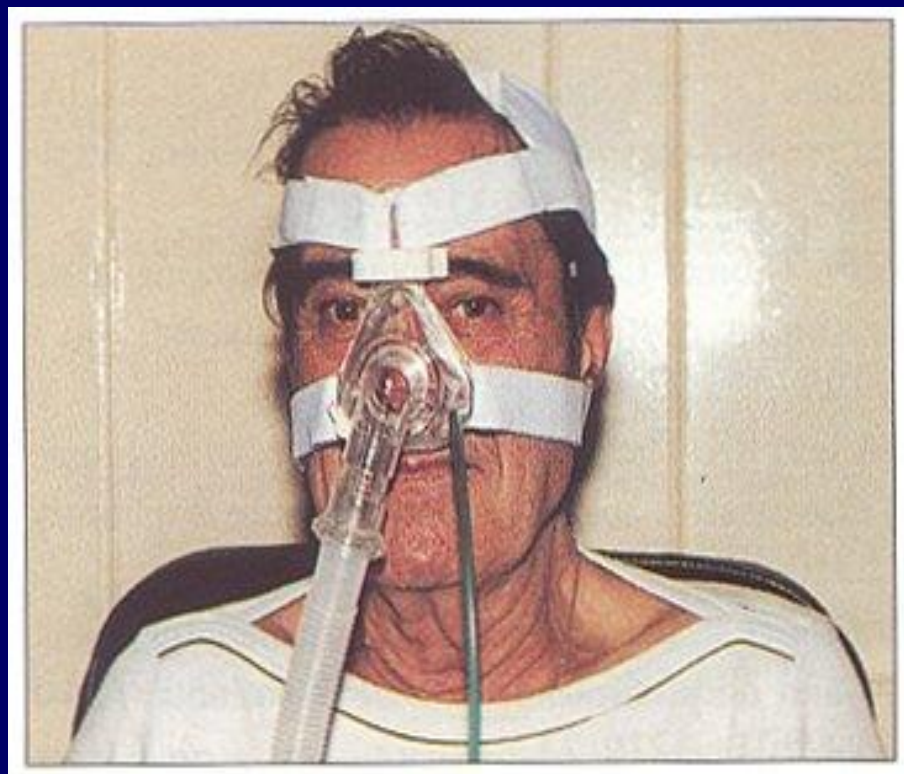
# Management of Stable COPD

## Non-Drug Treatments

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- **Rehabilitation:** All COPD patients benefit from exercise training programs, improving with respect to both exercise tolerance and symptoms of dyspnea and fatigue (**Evidence A**).
- **Provision** of Pulmonary rehabilitation services is inadequate.

# Management of Exacerbations ( Lung Attack )



- Noninvasive ventilation (NIV):-  
improves blood gases , reduces in-hospital mortality,  
decreases the need for invasive mechanical ventilation  
and decreases the length of hospital stay (**Evidence A**).

# WORLD COPD DAY

## November 14, 2007



Raising COPD Awareness Worldwide

# COPD: What do we need?

## *Health care provision*

- Recognition of COPD as a health care priority.
- Spirometry available for screening and diagnosis.
- Vaccination programme specifically targeting COPD patients.
- Smoking cessation services.
- Health care professionals with specific training in COPD and smoking cessation.
- Access to specialist respiratory services for diagnosis and assessment (oxygen therapy, surgical intervention, exacerbations).
- Pulmonary rehabilitation services.
- Access to non-invasive ventilation.
- Hospital at home services to manage non life-threatening exacerbations.
- palliative care services.

# COPD: What do we need?

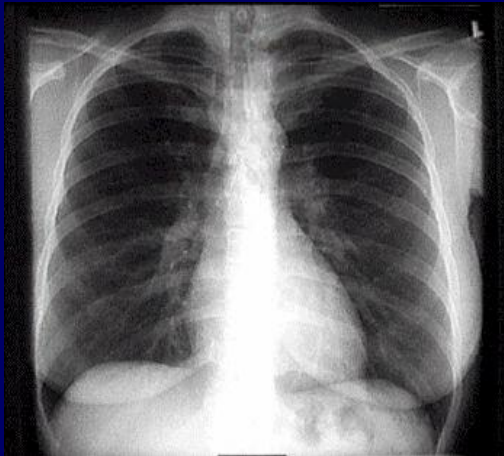
## *Research priorities*

- To create a stronger foundation for fighting COPD by acquiring accurate data on illness, exacerbations, natural history, deaths and costs.
- To perform scientific surveys with spirometry in population samples of all European countries to improve knowledge of the distribution of COPD.
- To establish higher standards of care through studies on the effectiveness of current prevention, education, medication and rehabilitation.
- To develop new therapies to inhibit the progression of the disease.
- To study the most effective smoking cessation intervention techniques to prevent people from starting smoking.
- To guide caregivers and care payers in the most efficient and effective ways to manage this disease.



# Impact of air trapping (hyperinflation)

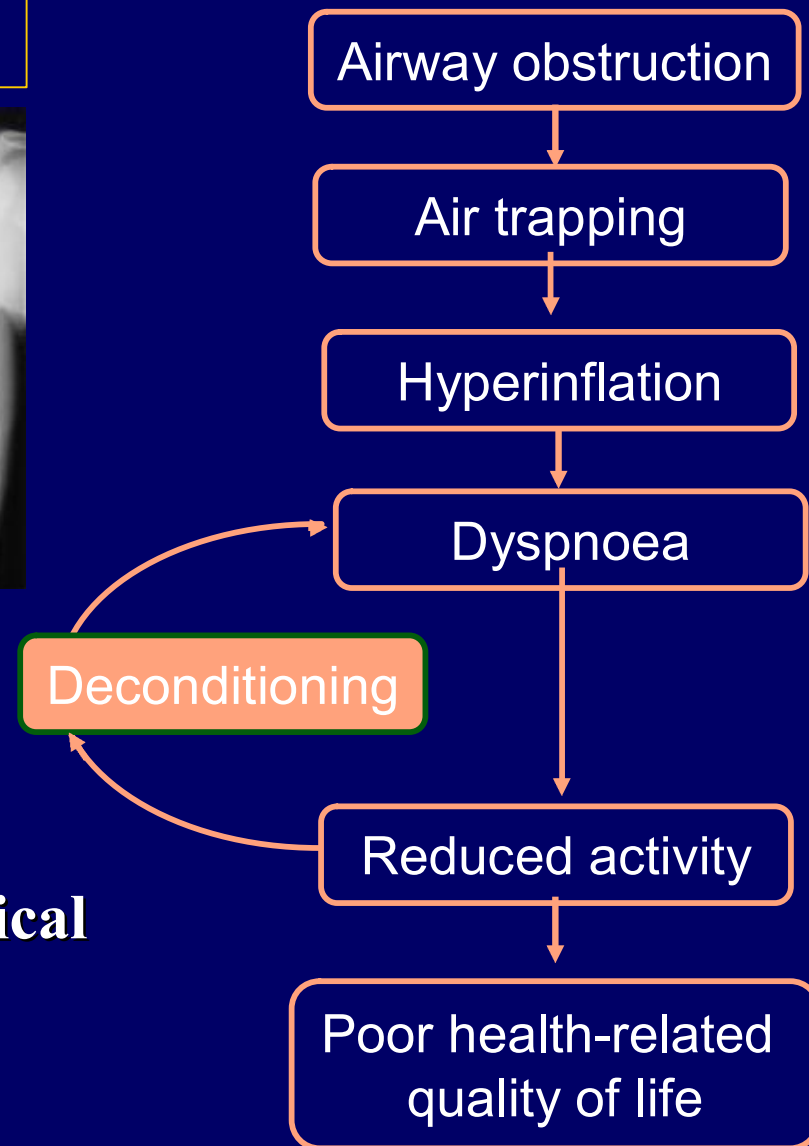
**Normal**



**Hyperinflation**

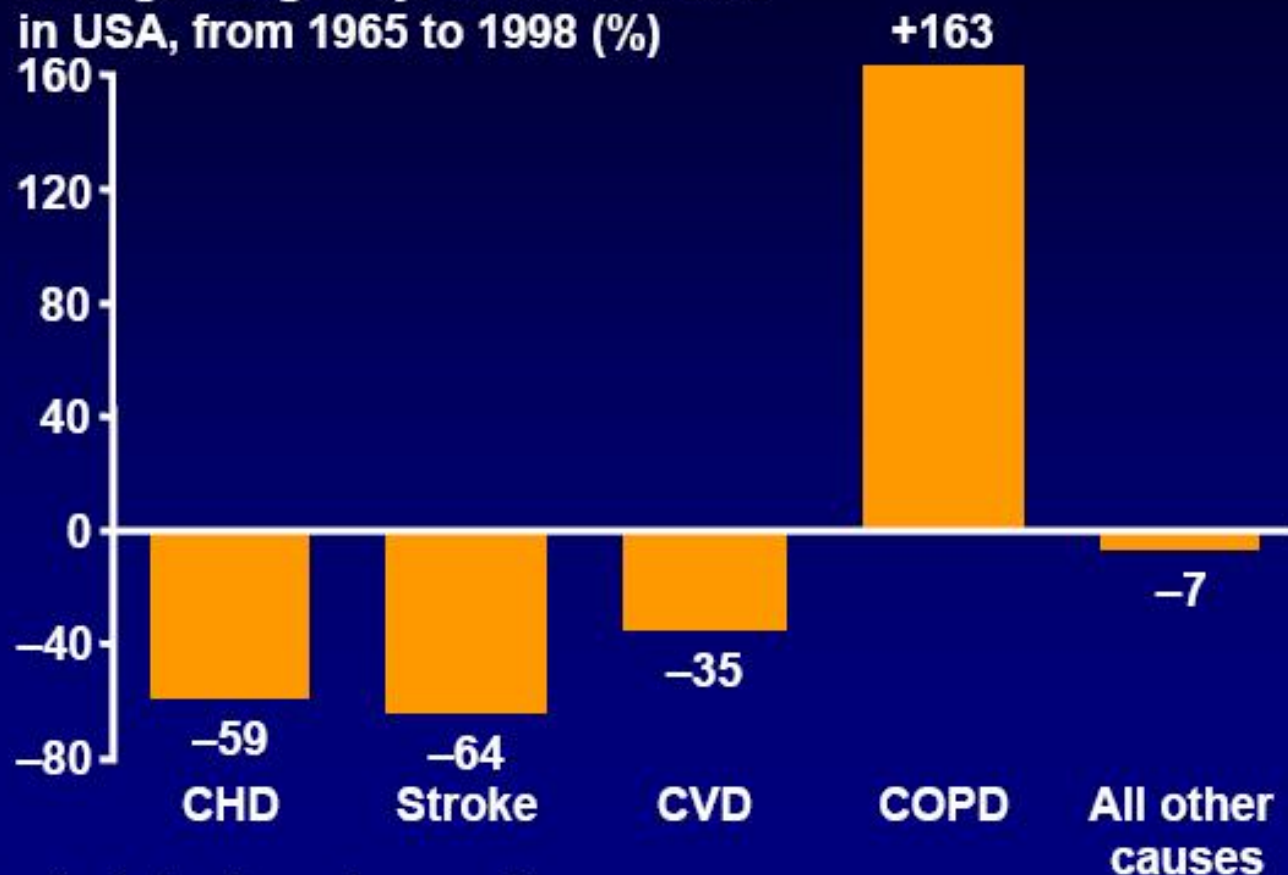


- ❑ Occurs in all patients with COPD
- ❑ Increases the work of breathing
- ❑ Respiratory muscles at a mechanical disadvantage
- ❑ Contributes to breathlessness



# **COPD is the only major cause of death to increase significantly in recent years**

Change in age-adjusted death rate  
in USA, from 1965 to 1998 (%)



COPD = chronic obstructive pulmonary disease

CHD = coronary heart disease

CVD = cerebrovascular disease

Adapted from: [www.copdgold.com](http://www.copdgold.com)

# Increasing burden of diseases and Injuries: change in rank order of DALYs

1999

1. Acute lower respiratory infections
2. HIV/AIDS
3. Perinatal conditions
4. Diarrhoeal diseases
5. Unipolar major depression
6. Ischaemic heart disease
7. Cerebrovascular disease
8. Malaria
9. Road traffic injuries
10. COPD
11. Congenital abnormalities
12. Tuberculosis
13. Falls
14. Measles
15. Anaemias

2020

1. Ischaemic heart disease
2. Unipolar major depression
3. Road traffic injuries
4. Cerebrovascular disease
5. COPD
6. Acute lower respiratory infections
7. Tuberculosis
8. War
9. Diarrhoeal diseases
10. HIV
11. Perinatal conditions
12. Violence
13. Congenital abnormalities
14. Self-inflicted injuries
15. Trachea, bronchus and lung cancers

**DALY = Disability-adjusted life year**

Source: WHO  
Evidence, Information and Policy, 2000

**COPD: What does it stand for?**  
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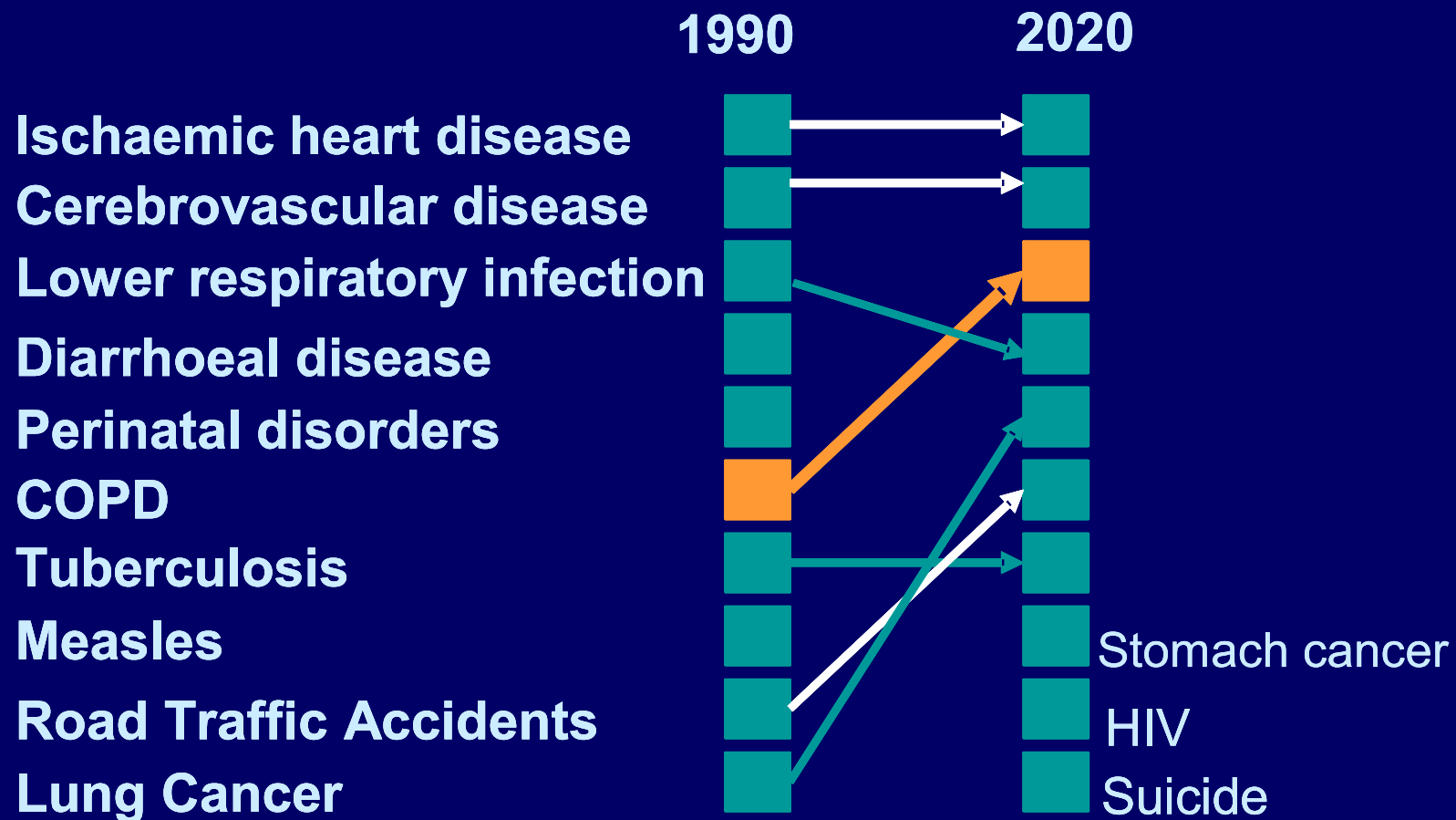
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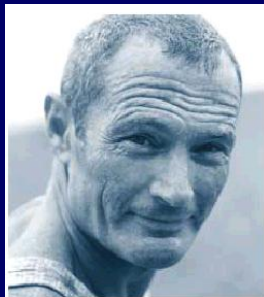
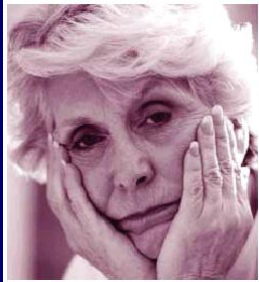
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- COPD is a preventable and treatable disease. Its lung component is characterized by airflow limitation (or obstruction to the flow of air) that is not fully reversible.
- The airflow limitation is usually progressive and associated with an abnormal inflammatory response of the lung to noxious particles or gases.
- COPD has significant extra-pulmonary effects that may contribute to the severity in individual patients.

# Global Burden Of Disease Top 10 Causes Of Death 1990 and 2020



# The burden of COPD in Europe



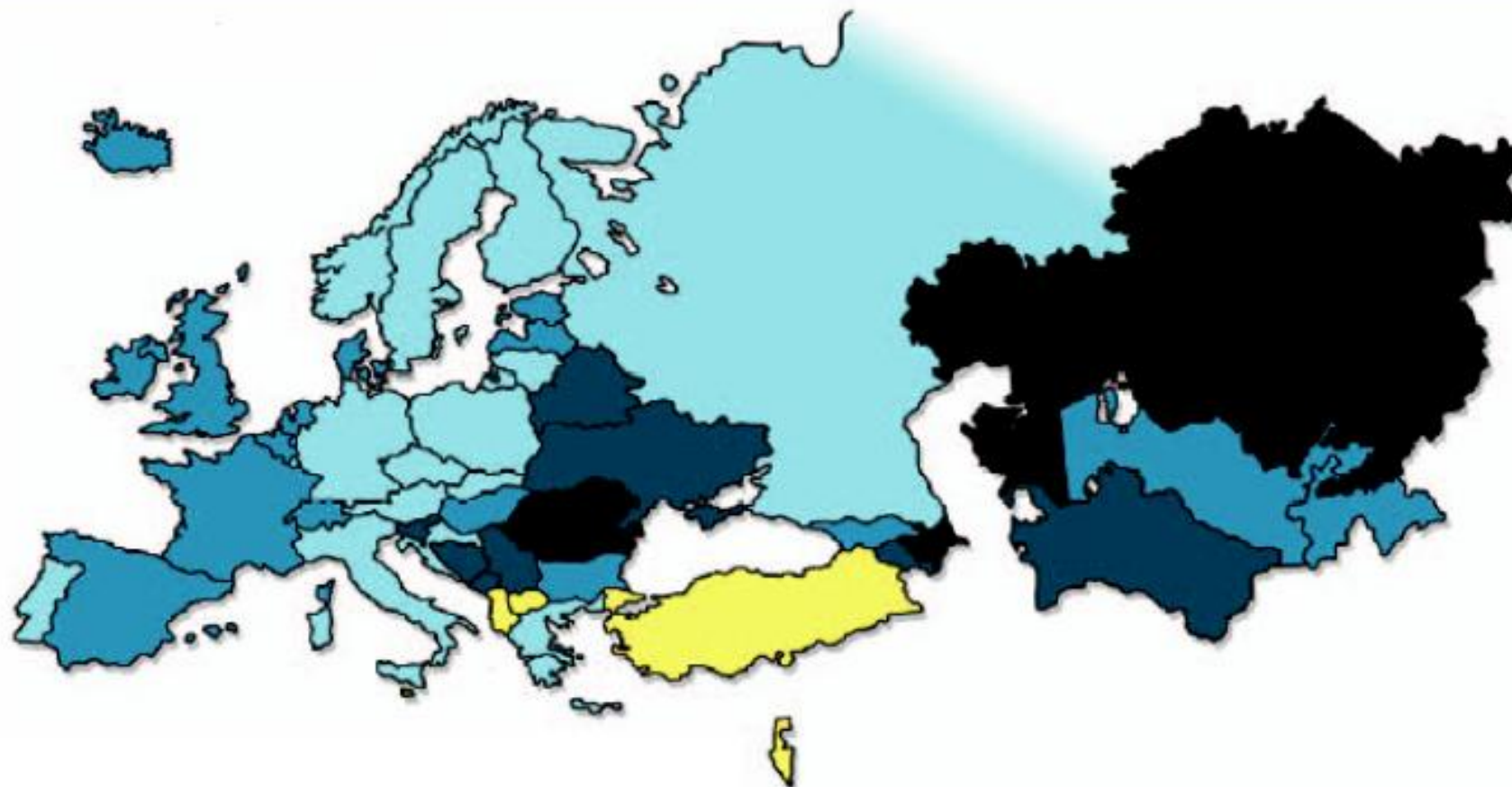
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European Respiratory  
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# Burden of mortality from COPD in Europe

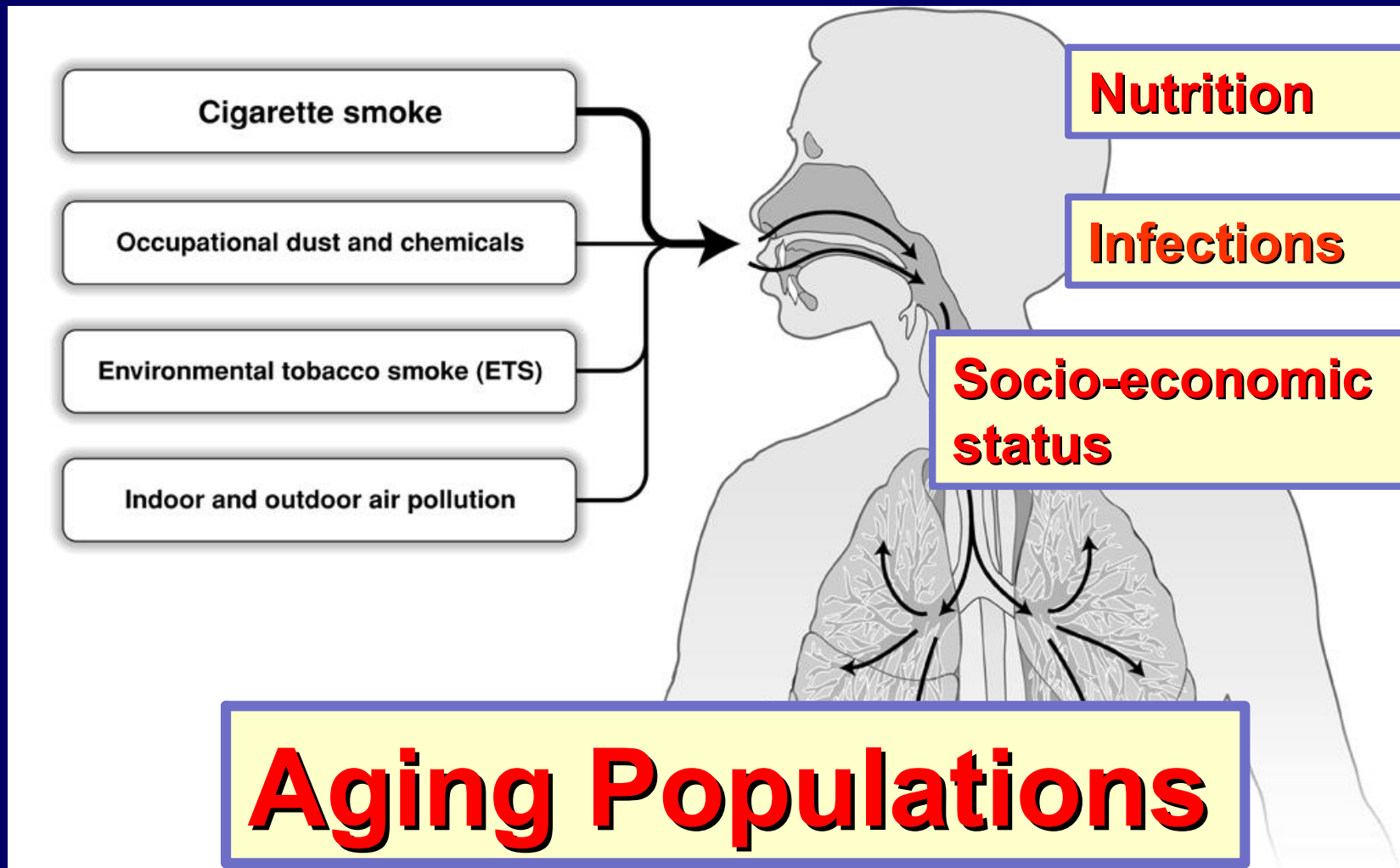


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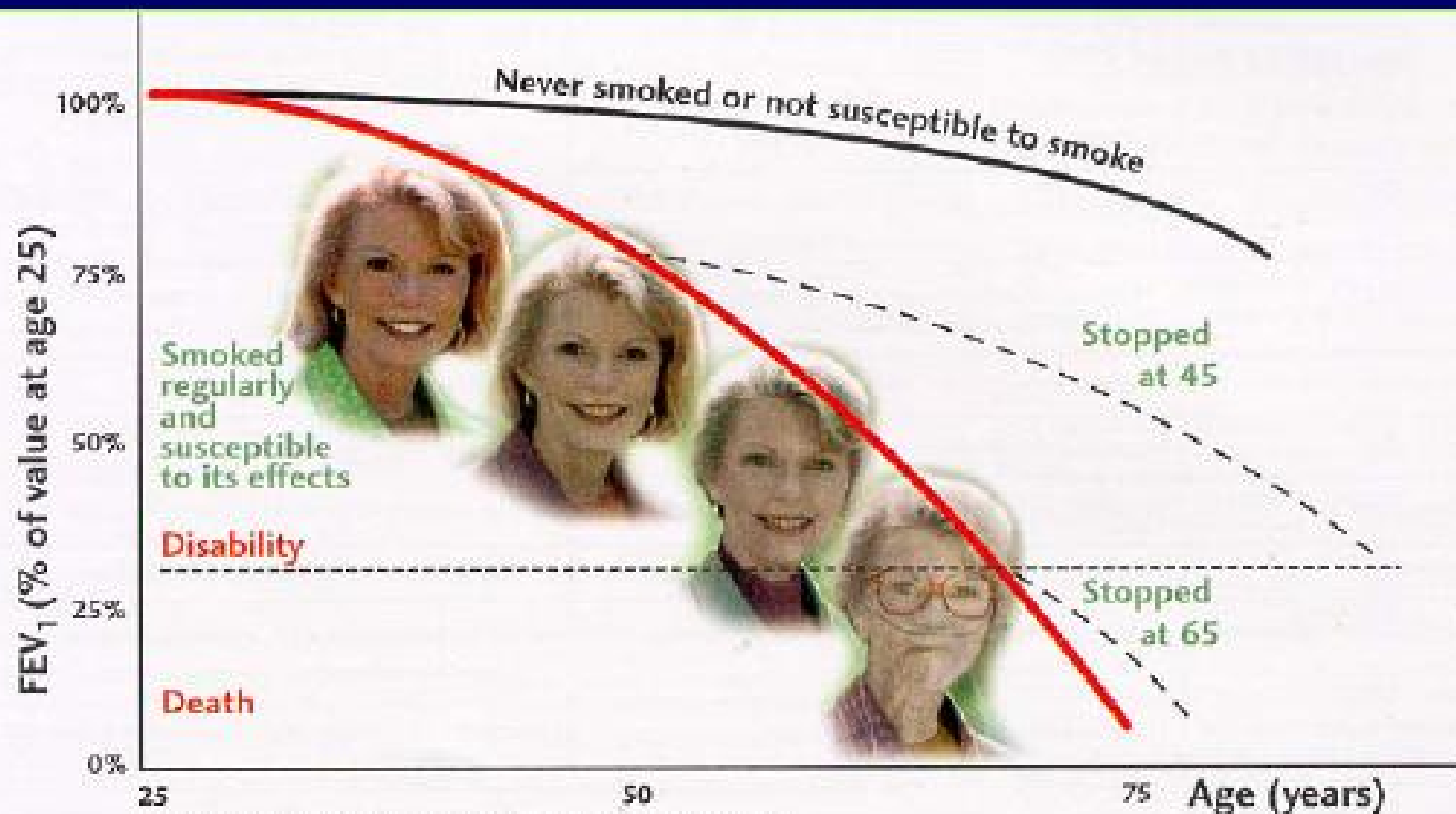
By 2010, COPD will be the fourth leading cause of death in Europe



# Risk Factors for COPD

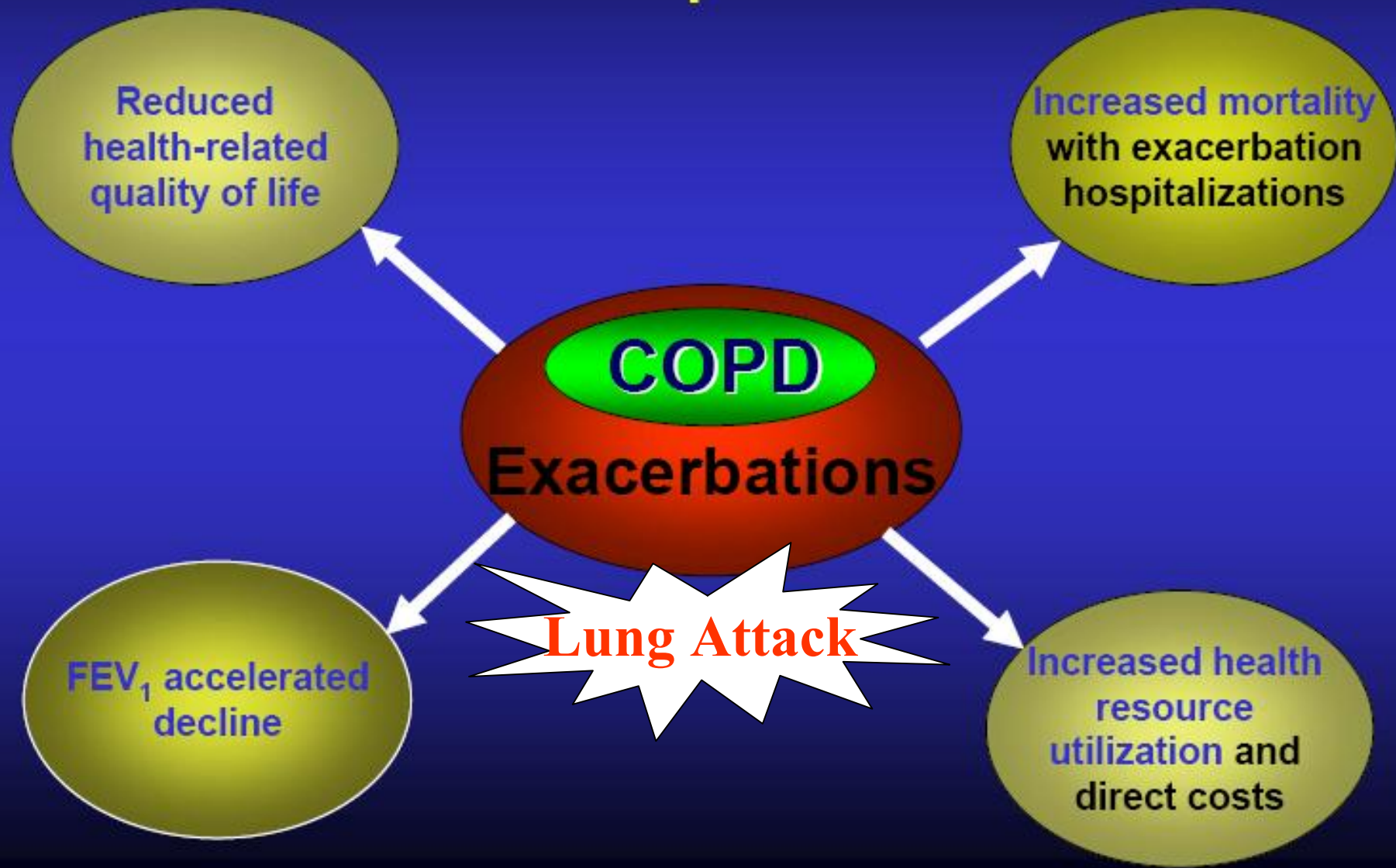


# COPD Progression



Adapted from Fletcher C, Peto R. The natural history of chronic airflow obstruction. *Br Med J* 1975; 1: 1645-1648.

# COPD Exacerbations: Clinical Consequences



# Diagnosis of COPD

## SYMPTOMS

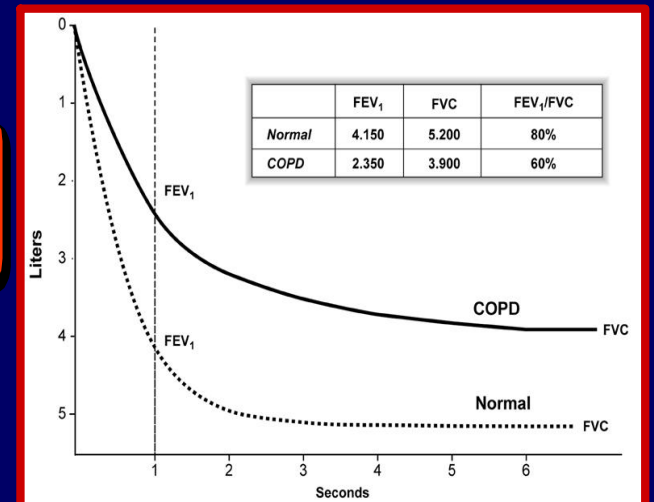
cough  
sputum  
shortness of breath

## EXPOSURE TO RISK FACTORS

tobacco  
occupation  
indoor/outdoor pollution



## SPIROMETRY



# GOALS of COPD MANAGEMENT

- Relieve symptoms
- Prevent disease progression
- Improve exercise tolerance
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Raising COPD Awareness Worldwide

# **COPD: What do we need?**

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- **Recognition of COPD as a health care priority**
- **Spirometry available for screening and diagnosis**
- **Vaccination programme specifically targeting COPD patients**
- **Smoking cessation services**
- **Health care professionals with specific training in COPD and smoking cessation**
- **Access to specialist respiratory services for diagnosis and assessment (oxygen therapy, surgical intervention,exacerbations)**
- **Pulmonary rehabilitation services**
- **Access to non-invasive ventilation**
- **Hospital at home services to manage non life threatening exacerbations**
- **palliative care services**



# COPD: What do we need?

## *Research priorities*

- To create a stronger foundation for fighting COPD by acquiring accurate data on illness, exacerbations, natural history, deaths and costs
- To perform scientific surveys with spirometry in population samples of all European countries to improve knowledge of the distribution of COPD
- To establish higher standards of care through studies on the effectiveness of current prevention, education, medication and rehabilitation
- To develop new therapeutic modalities to inhibit the progression of the disease
- To study the most effective smoking cessation interventions and techniques to prevent people from starting smoking
- To guide caregivers and care payers in the most efficient and effective ways to manage this disease





# **In 2030**

**Plos Med (25 Nov 2006)**

- **COPD: 4th cause of death**
- **Tobacco > 8 million deaths**
- **Biomass fuel combustion > 10 million deaths (Ezzatti, Science 2005)**

# Management of Stable COPD

## Reduce Risk Factors

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- Reduction of total personal exposure to tobacco smoke, occupational dusts and chemicals, and indoor and outdoor air pollutants are important goals to prevent the onset and progression of COPD.
- Smoking cessation is the single most effective — and cost effective — intervention in most people to reduce the risk of developing COPD and stop its progression (Evidence A).



## Translating COPD Guidelines into Primary Care

# KEY POINTS

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- Spirometric confirmation is a key component of the diagnosis of COPD and primary care practitioners should have access to high quality spirometry.
- Older patients frequently have multiple chronic health conditions. Comorbidities can magnify the impact of COPD on a patient's health status, and can complicate the management of COPD.

## Management of Stable COPD

### Non-Pharmacologic Treatments

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- **Rehabilitation:** All COPD patients benefit from exercise training programs, improving with respect to both exercise tolerance and symptoms of dyspnea and fatigue (**Evidence A**).
- **Oxygen Therapy:** The long-term administration of oxygen (> 15 hours per day) to patients with chronic respiratory failure has been shown to increase survival (**Evidence A**).

# **New Advances in COPD**

- **Assessment**
- **Understanding disease processes**
- **Treatment**

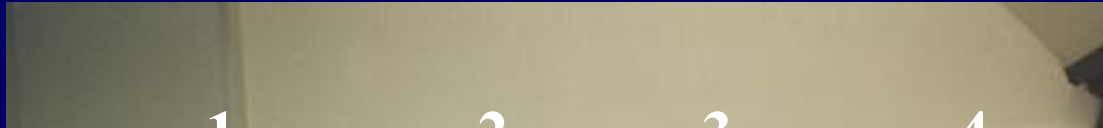


# ***“COPD Lung function Heterogeneity”***

Patient # 1

58 y

FEV1: 28 %



**COPD Sub(Pheno)types  
Important for treatment ?**



Patient # 4

72 y

FEV1: 34%

MRC: 4/4

6MWD: 154 m

BMI: 24

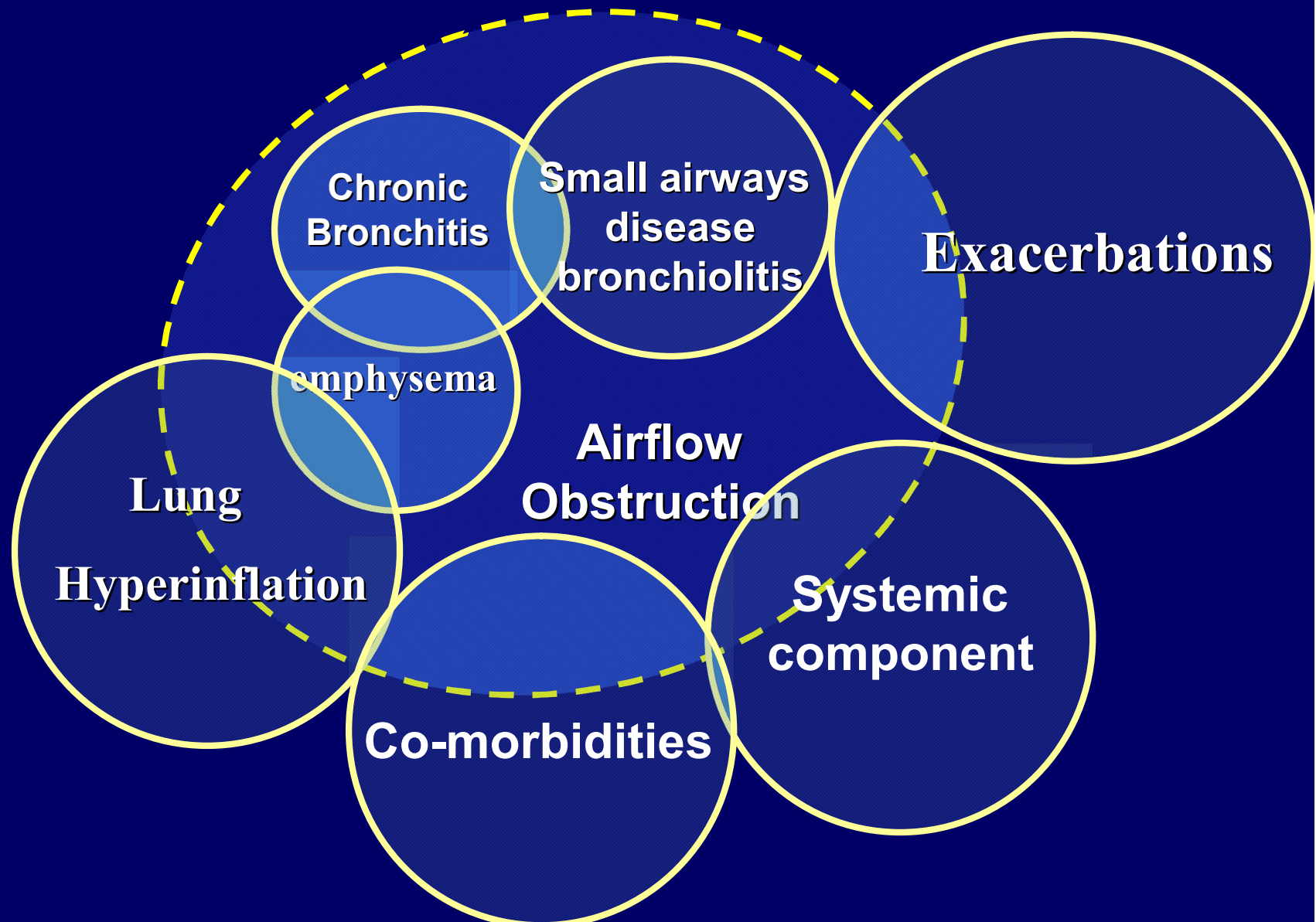
**BODE: 9**

***FEV1 < 35% Predicted***

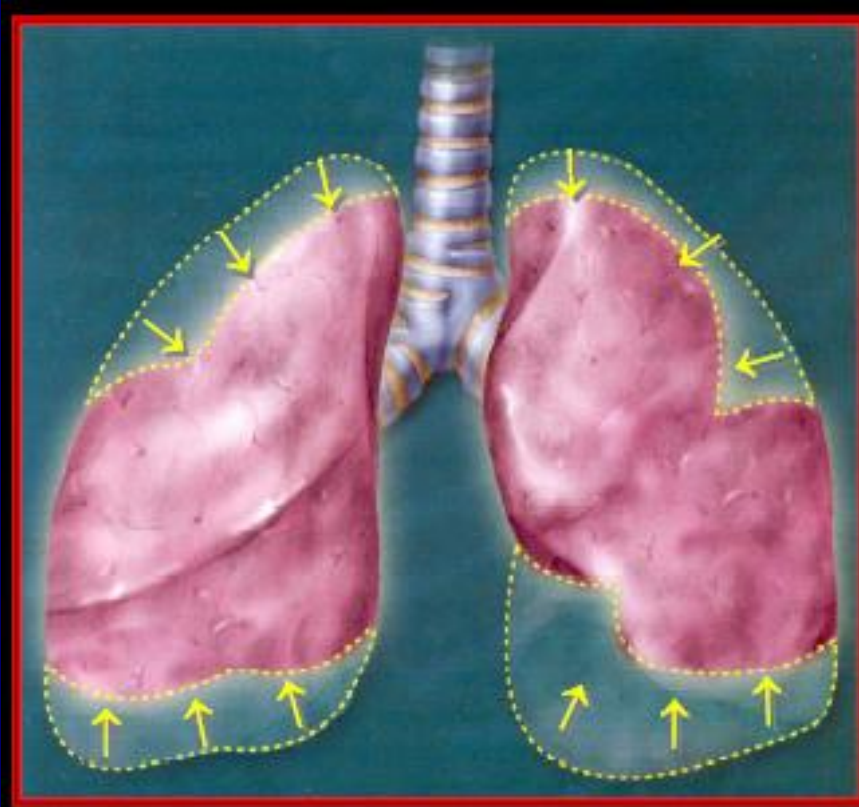
***Severe disease***

Cote et al

# COPD: a multicomponent disease



# Surgical volume reduction for emphysema

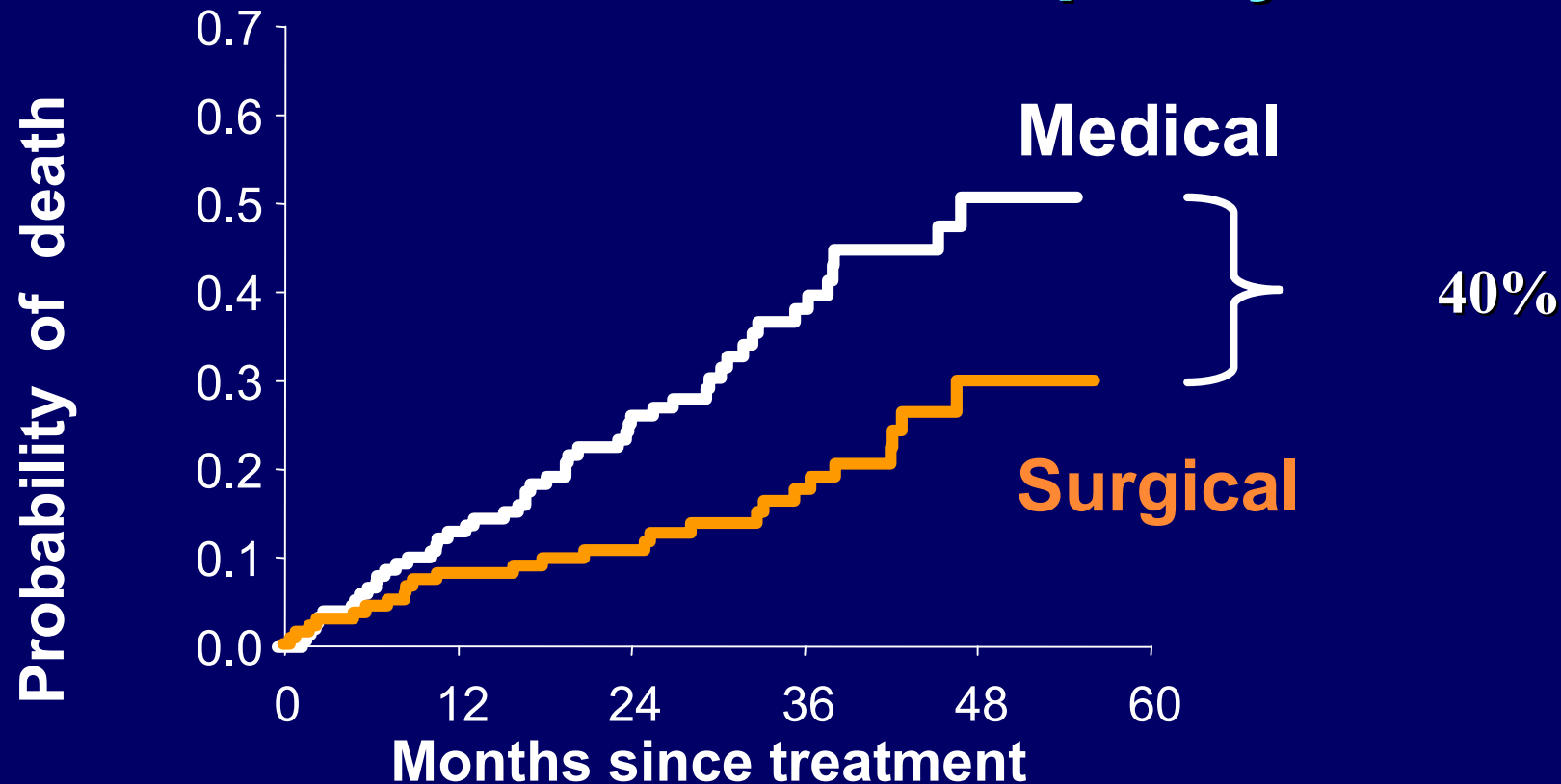


*In selected patients:-*

- Removes hyperinflated areas of lung
- Reduces air trapping
- Improves breathlessness
- Improves health status
- Improves survival

# **LVRS improves mortality:**

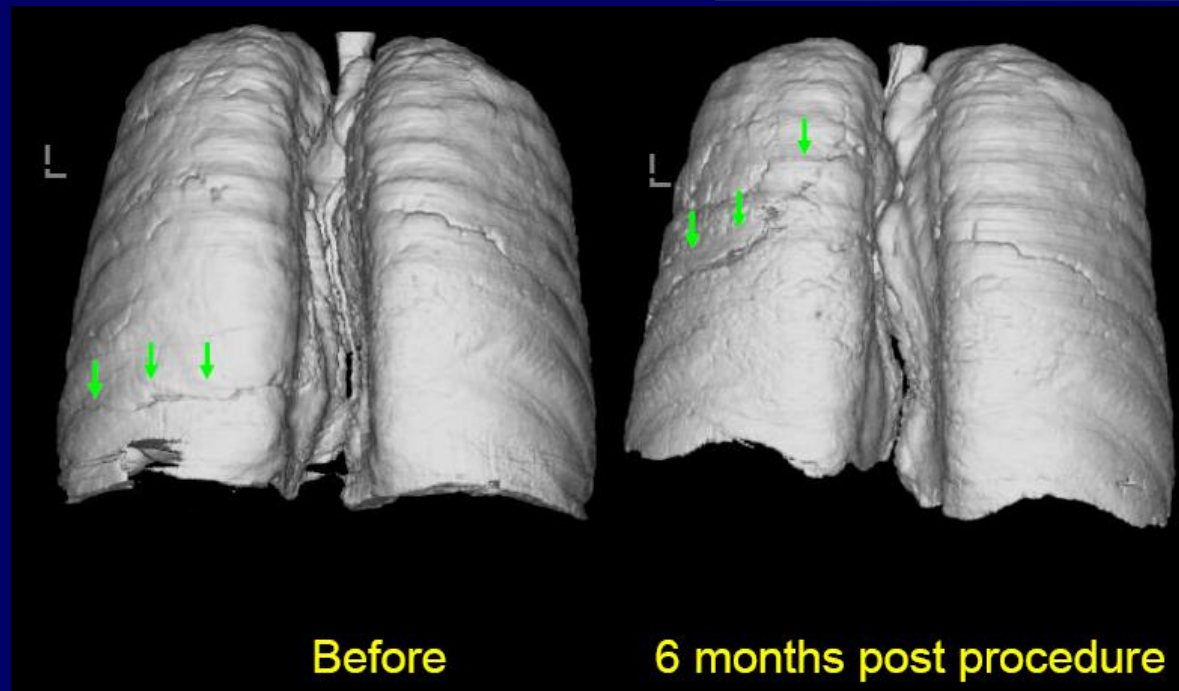
## **Patients Upper-lobe emphysema and low exercise capacity**



**High-risk patients excluded**

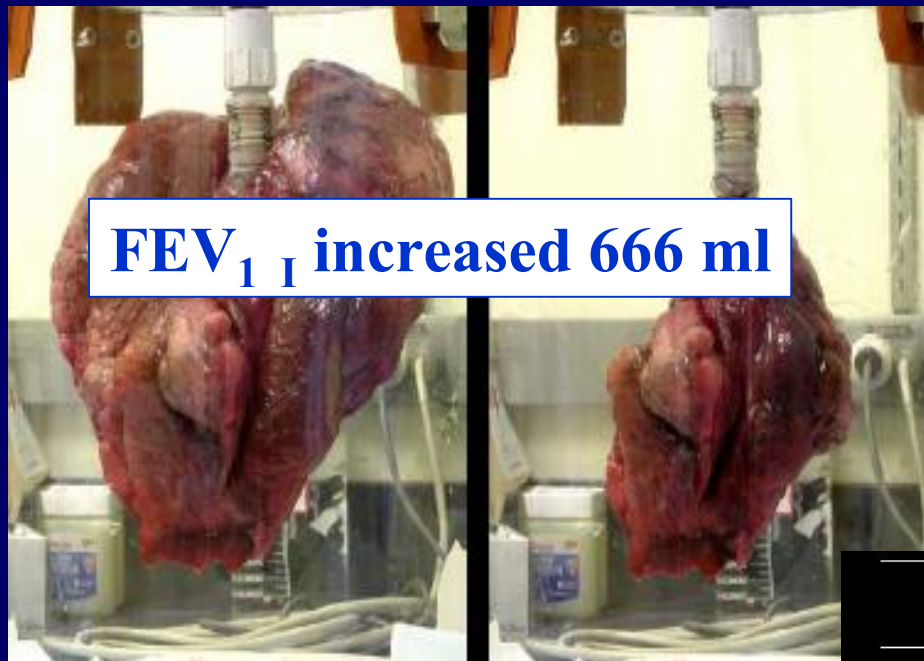


# Bronchoscopic Lung Volume Reduction



# Airway bypass for emphysema

## Proof of concept



**FEV<sub>1</sub> increased 666 ml**

**Before**

**Bypass**

**After**

Designed for patients with diffuse severe emphysema :-

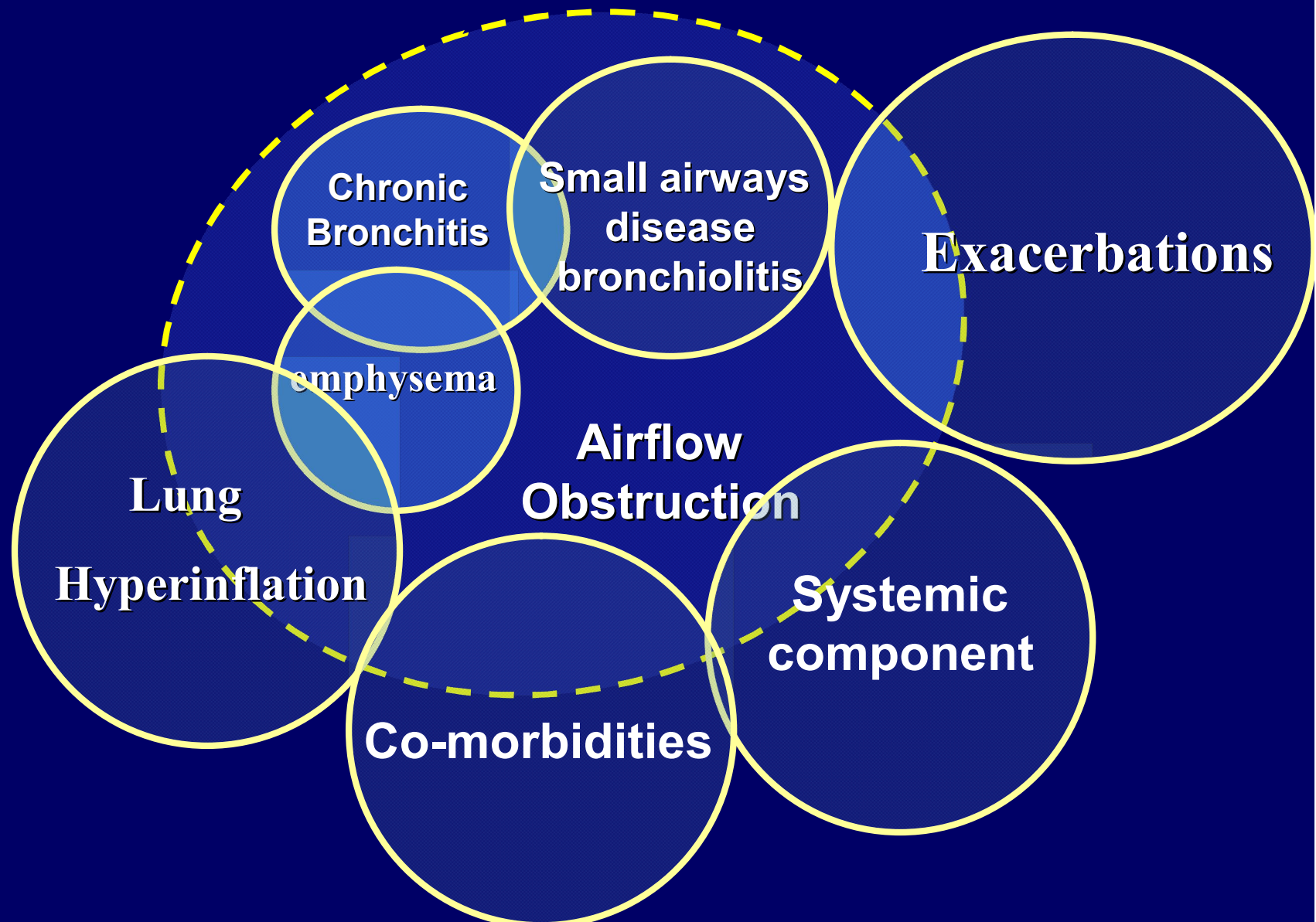
- Creation of new pathways for trapped gas to escape from the lungs
- Reduces hyperinflation
- Improves breathlessness and quality of life

## Preliminary results in emphysema patients

	1 month	6 months
Change in SGRQ scores	-8.1 ± 9.6 (p,0.0001)	-6.8 ± 14.3 (p=0.05)
Proportion with ≥ -4 points	17 (63%)	12 (52%)
Proportion with ≥ -8 points	11 (41%)	9 (39%)
Proportion with FEV <sub>1</sub> change ≥ 15%	3 (10%)	2 (8%)
Proportion with 6MWD change ≥ 15%	8 (29%)	6 (24%)



# COPD: a multicomponent disease



# Small airways disease and emphysema

A

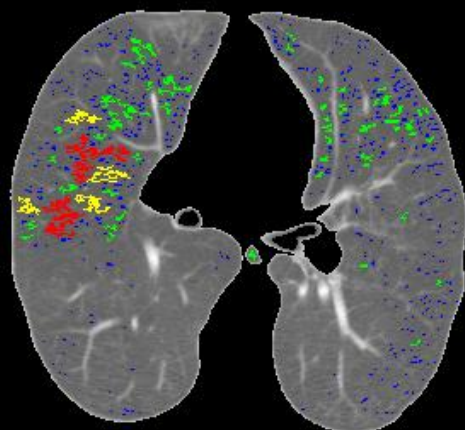
Age 53

44 Pack years

FEV<sub>1</sub> 42.1%

FEV<sub>1</sub>/FVC 0.39

Mild emphysema



B

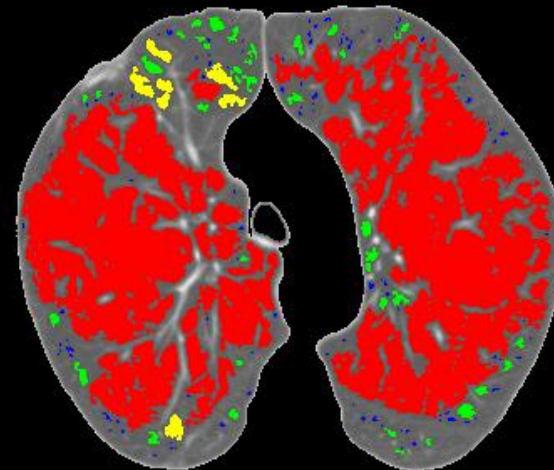
Age 59

48 pack years

FEV<sub>1</sub> 44.2%

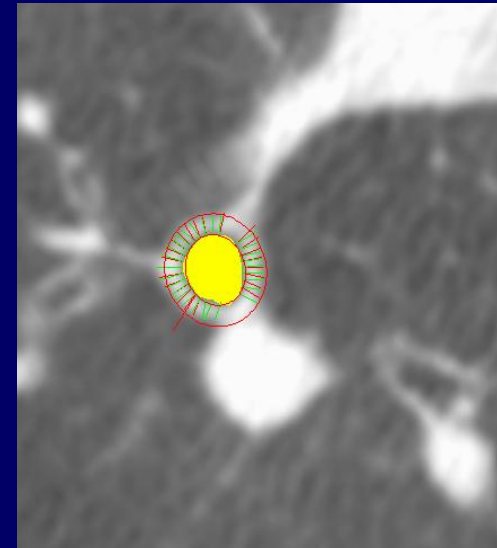
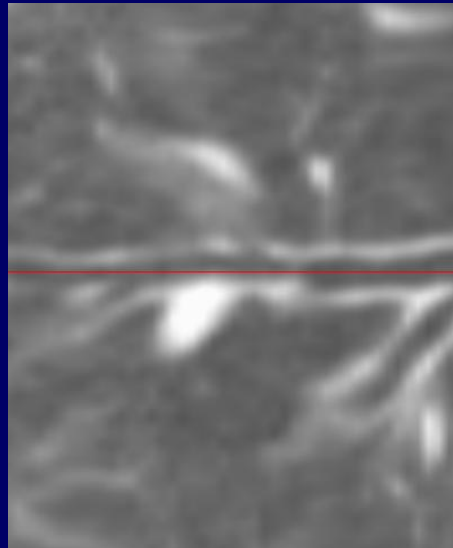
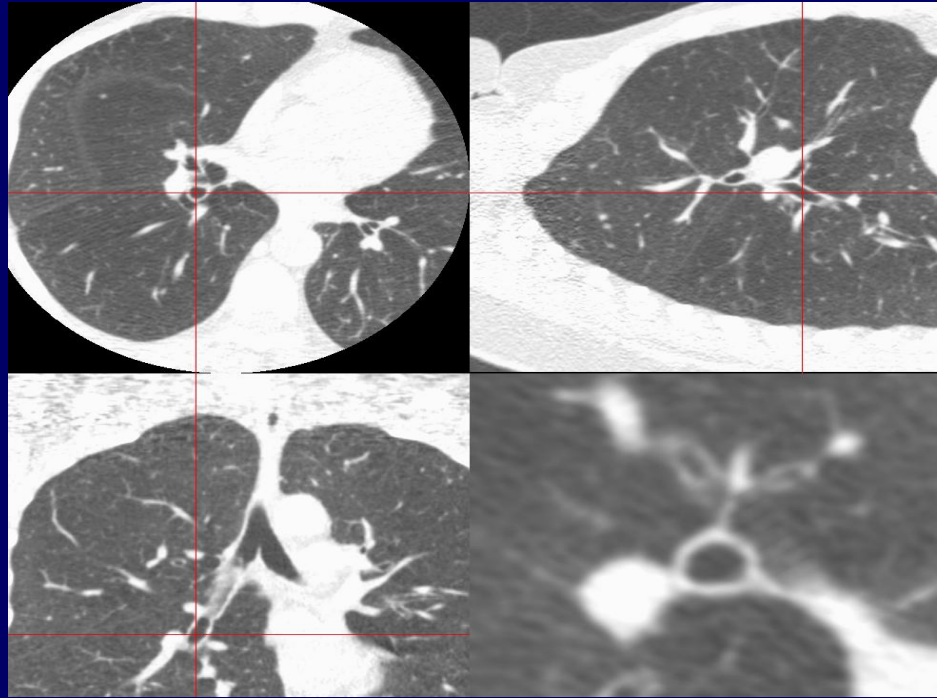
FEV<sub>1</sub>/FVC 0.39

Severe emphysema

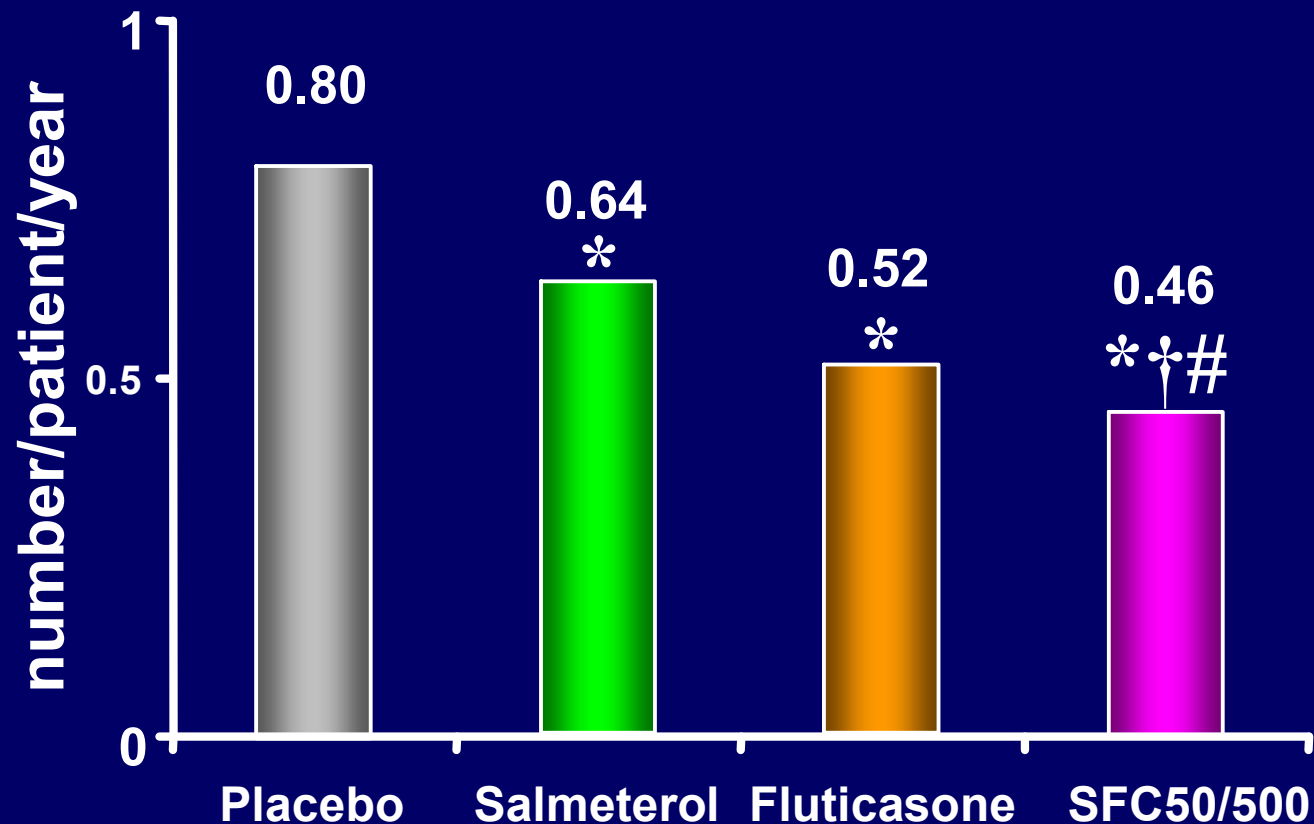


# Airway wall dimensions

- Airway dimensions
  - Lumen diameter
  - Wall thickness
  - Wall area
  - Percentage wall area



# Rate of Exacerbations Requiring Oral Corticosteroids



\*  $p < 0.001$  vs placebo,

†  $p < 0.001$  vs salmeterol,

#  $p = 0.017$  vs fluticasone

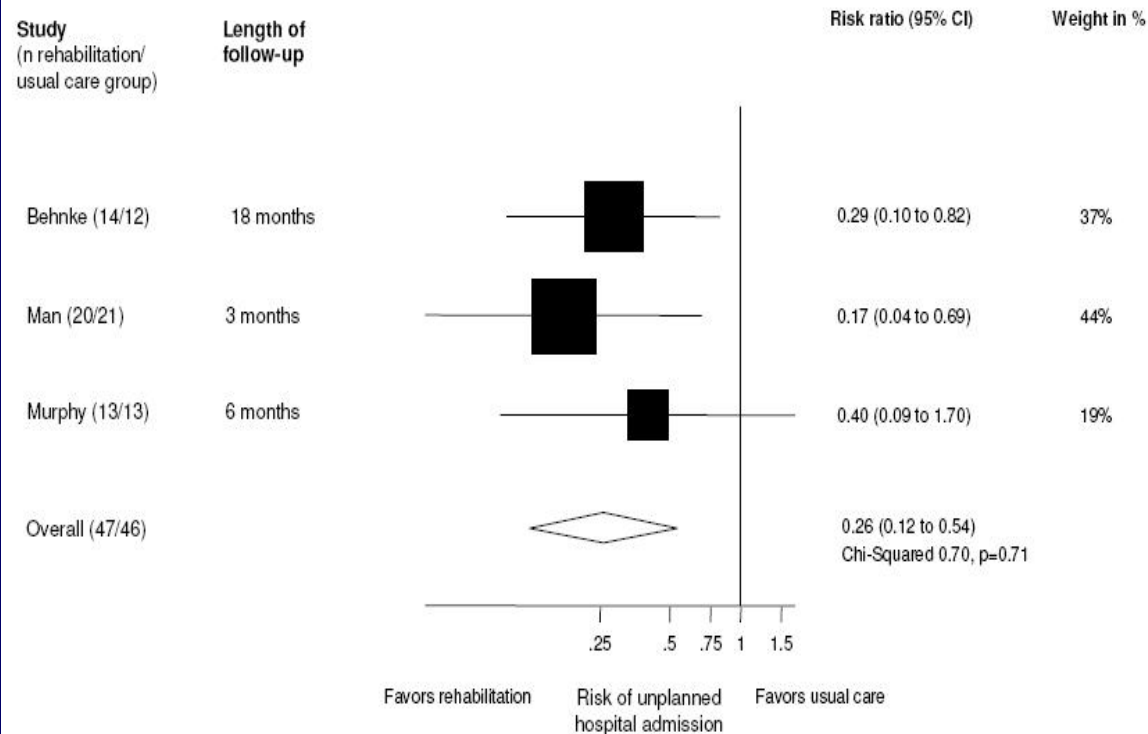
*Calverley NEJM 2007*



# Pulmonary rehabilitation for COPD

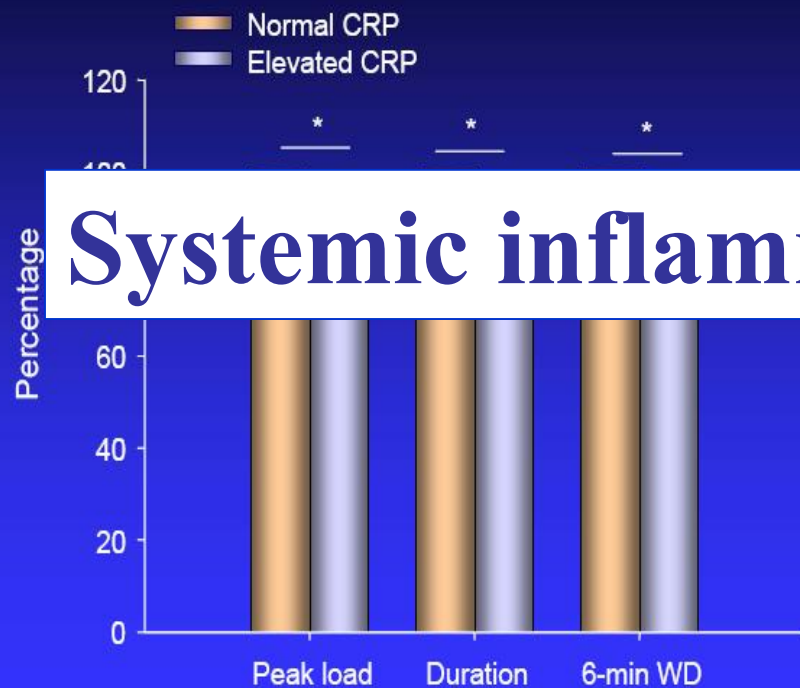


- Improves exercise tolerance
- Improves breathlessness
- Improves quality of life
- Reduces re-admission to hospital with exacerbations of COPD

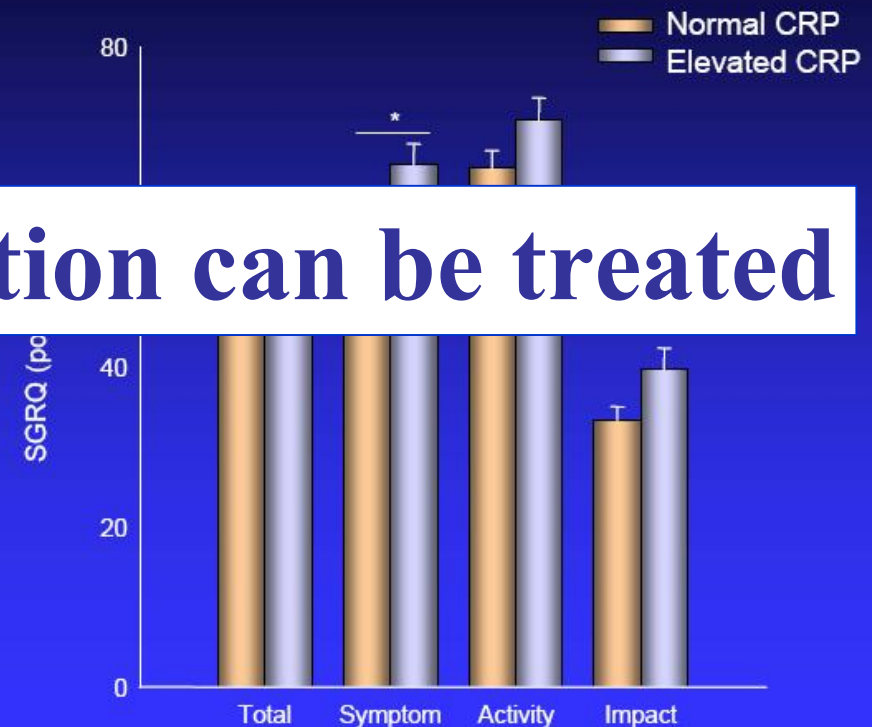


# Systemic inflammation , function and health status in COPD

## Exercise capacity



## Health status

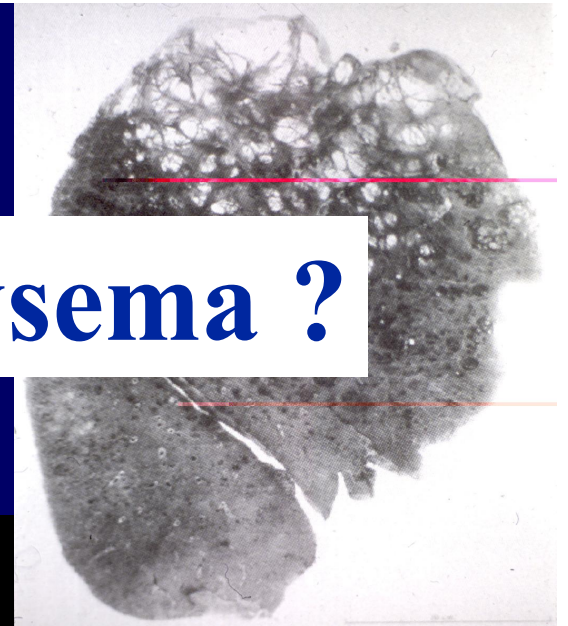
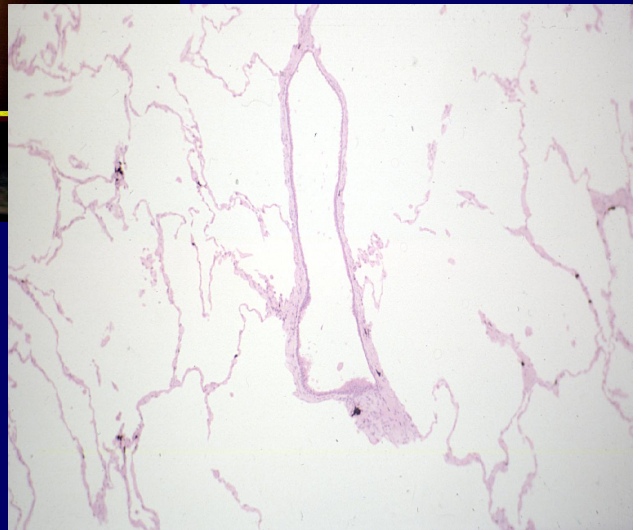
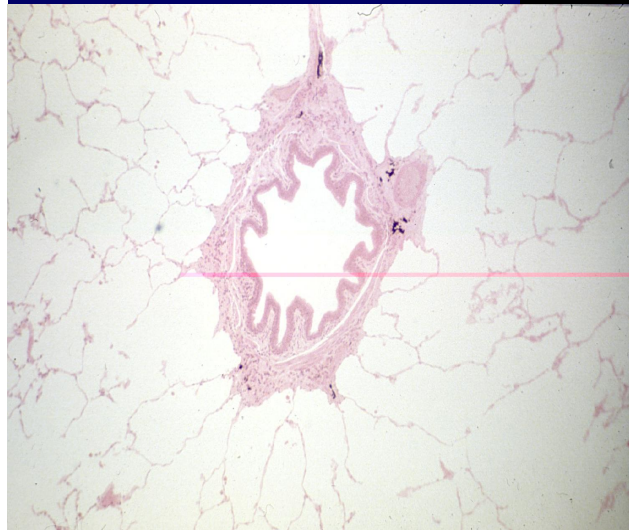
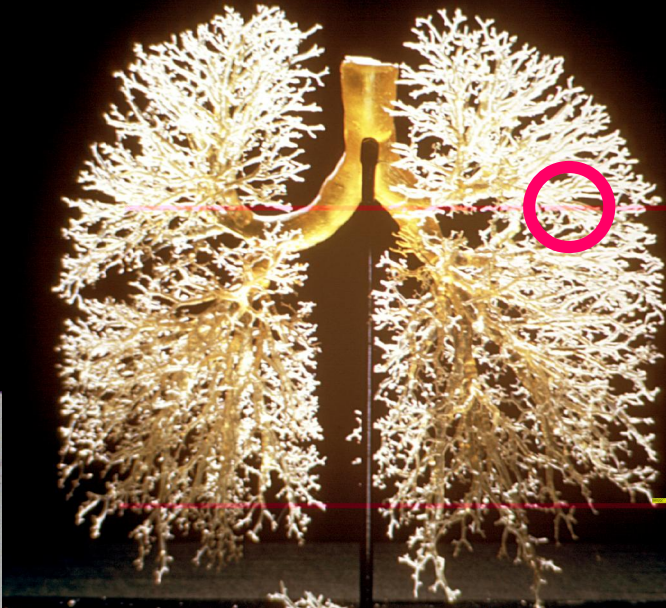


**Systemic inflammation can be treated**

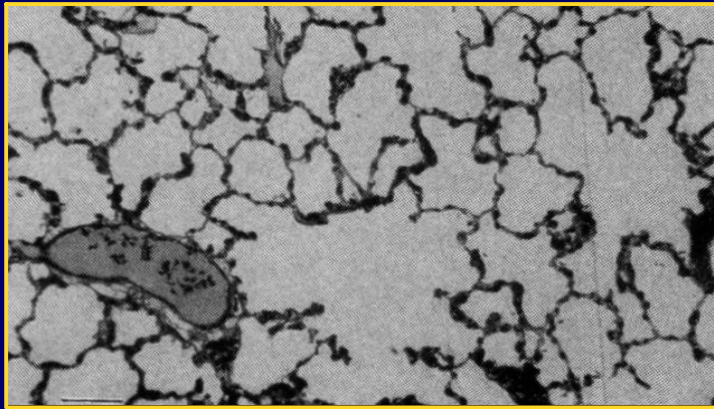


# Can we repair the emphysema ?

**EMPHYSEMA**  
**Macroscopic/  
Microscopic**



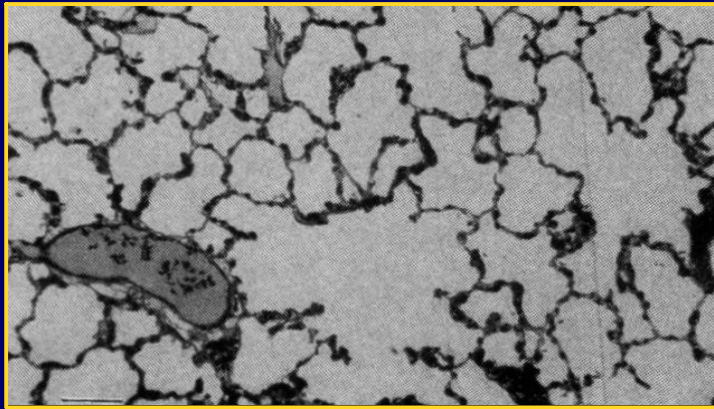
# Retinoic acid reverses elastase-induced pulmonary emphysema in rats



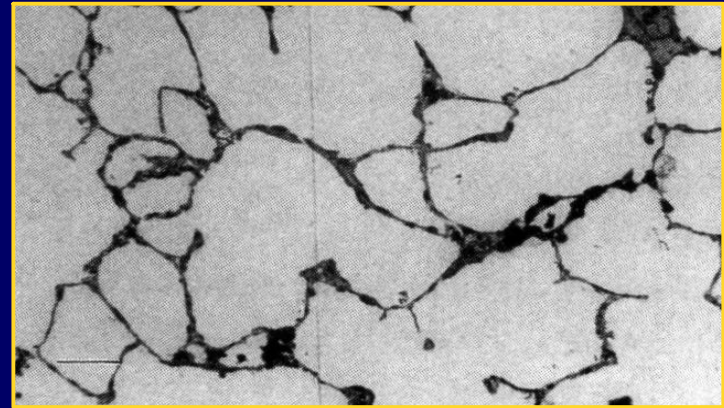
Normal



# Retinoic acid reverses elastase-induced pulmonary emphysema in rats

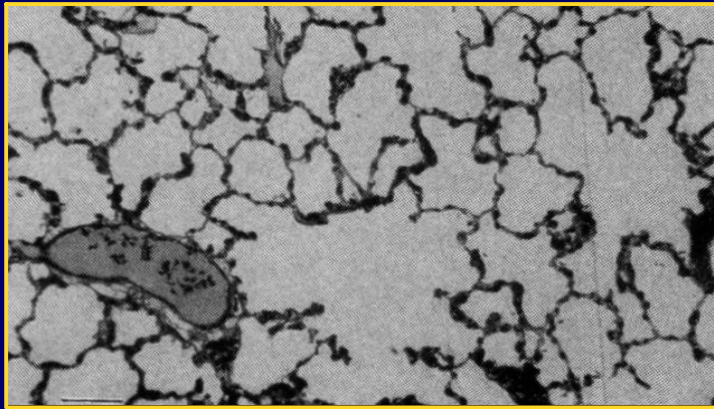


Normal

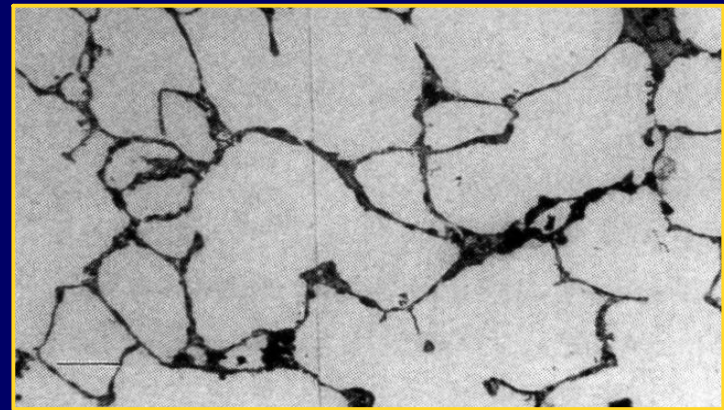


After elastase

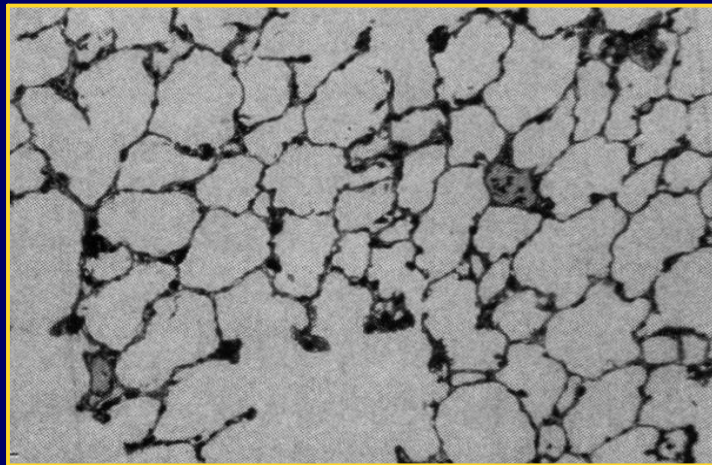
# Retinoic acid reverses elastase-induced pulmonary emphysema in rats



Normal



After elastase



Retinoic acid after elastase

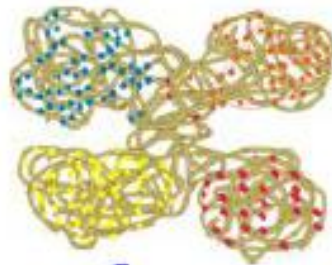
# Tissue Engineering

Short term  
<5 years

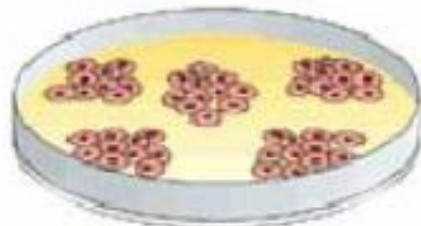
Medium  
5-10 years

Long term  
10-20 years (!)

3D Organoids



Cell therapy  
for local repair



Stem Cells



Lung Implants



# Tissue engineering- the future for chronic diseases ?





# **New Advances in COPD**

- **Assessment**
- **Understanding disease processes**
- **Treatment**




# Website

ELF Home - Microsoft Internet Explorer provided by ISL Computers Ltd. 01484 400979




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Calendar

◀ MAY 2006 ▶

S	M	T	W	T	F	S
	1	2	3	4	5	6
7	8	9	10	11	12	13
14	15	16	17	18	19	20
21	22	23	24	25	26	27
28	29	30	31			

[World No Tobacco Day](#)  
31/5/2006



## European Lung Foundation

Lung diseases are the leading cause of morbidity and mortality in Europe and throughout the world. Lung diseases can affect everyone and they impact on almost every family in Europe.

### Helping Europe to breathe

This website will provide:

- information about [the lungs](#) and how they function
- descriptions of, and information on, [lung diseases](#) and how they impact on individuals and on Europe
- different [risk factors](#) that can cause lung diseases and how they can be avoided


Latest news updates from around the world....

#### Low Levels Of Vitamin D In Teens May Affect Lung Function 23/05/2006

Teenagers who consume low amounts of vitamin D have lower lung function than teens who get the recommended amount of the nutrient, according to a study to be presented at the American Thoracic Society International Conference on May 22nd.

[More...](#)

### Asthma, don't let it slow you down!



Access the [inspirational film](#), commissioned and produced by the ELF, about an olympic swimmer who has suffered with asthma from childhood.

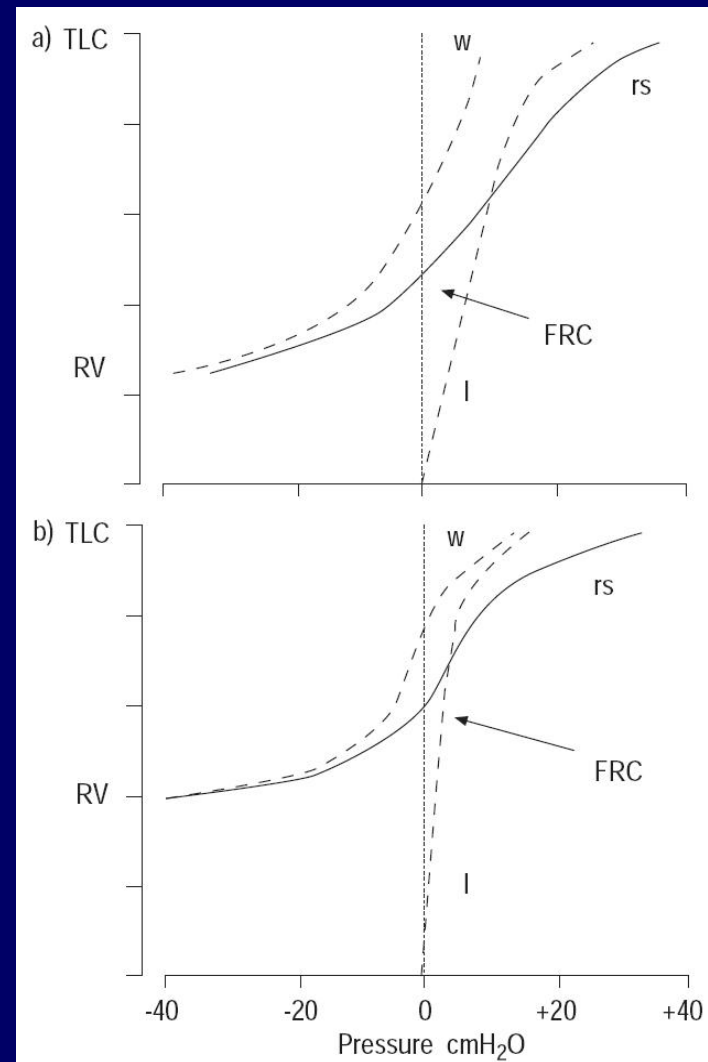
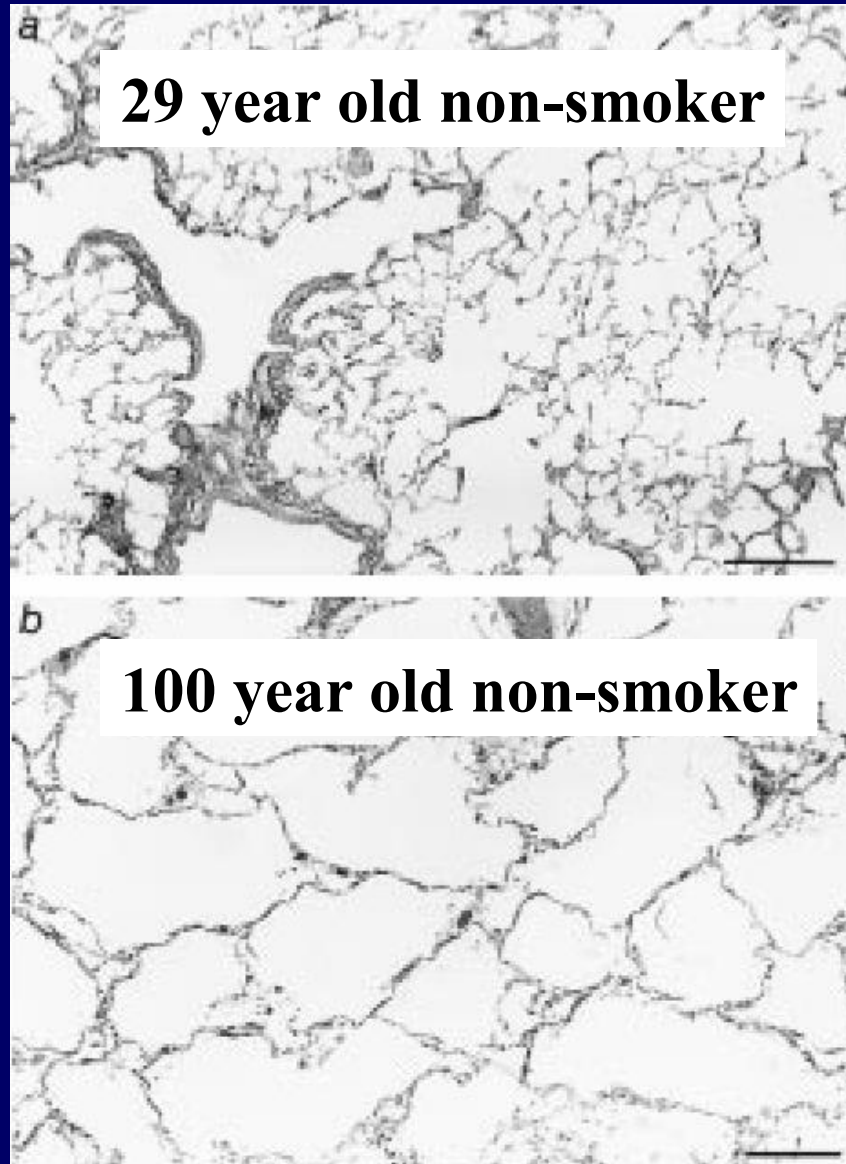
### Bird flu

Access the most up-to-date information on [bird flu](#) from the World Health Organization (WHO).

See how bird flu has spread using this [interactive map](#) from the BBC.

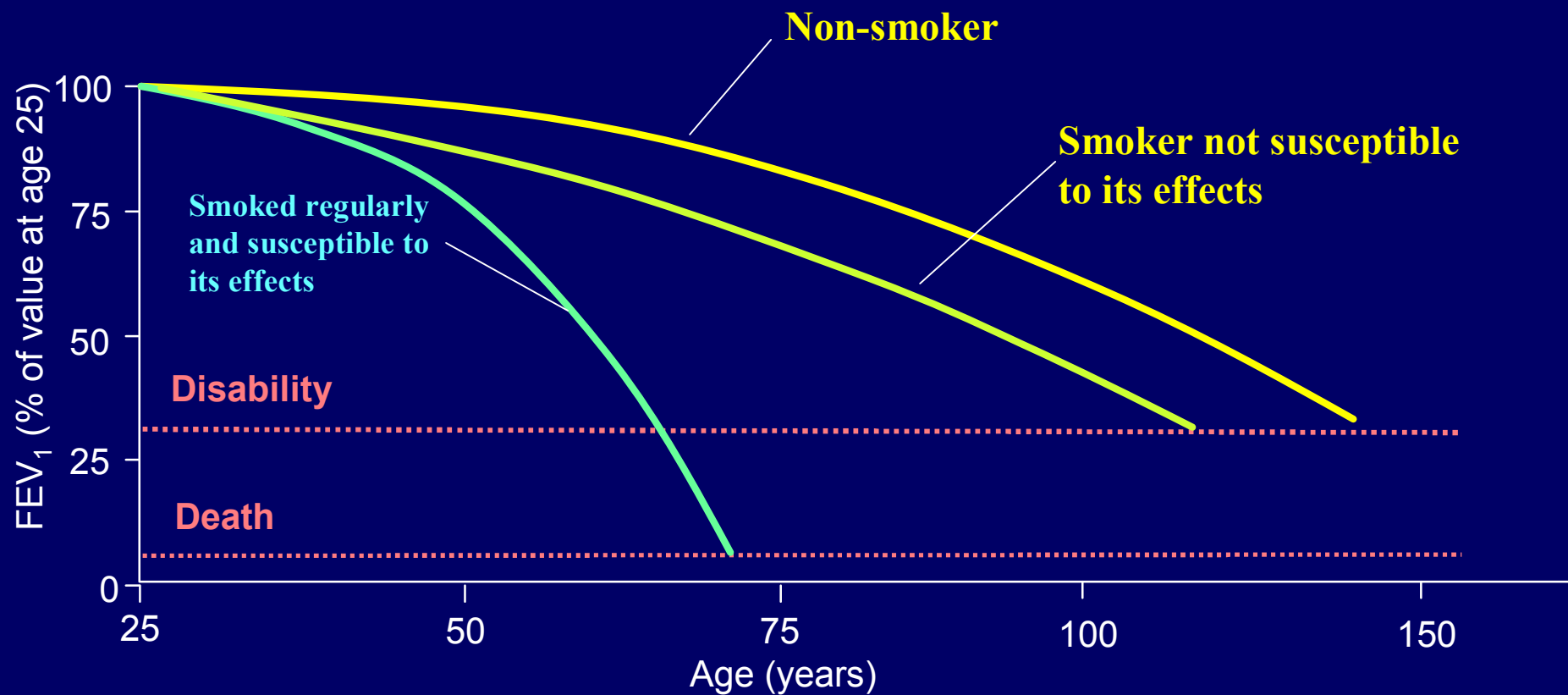
start | ELF Home - Microsoft ... | Internet | 16:06

# Emphysema and aging



*Janssens et al. Eur Respir J 1999; 13: 197*

# Natural history of COPD

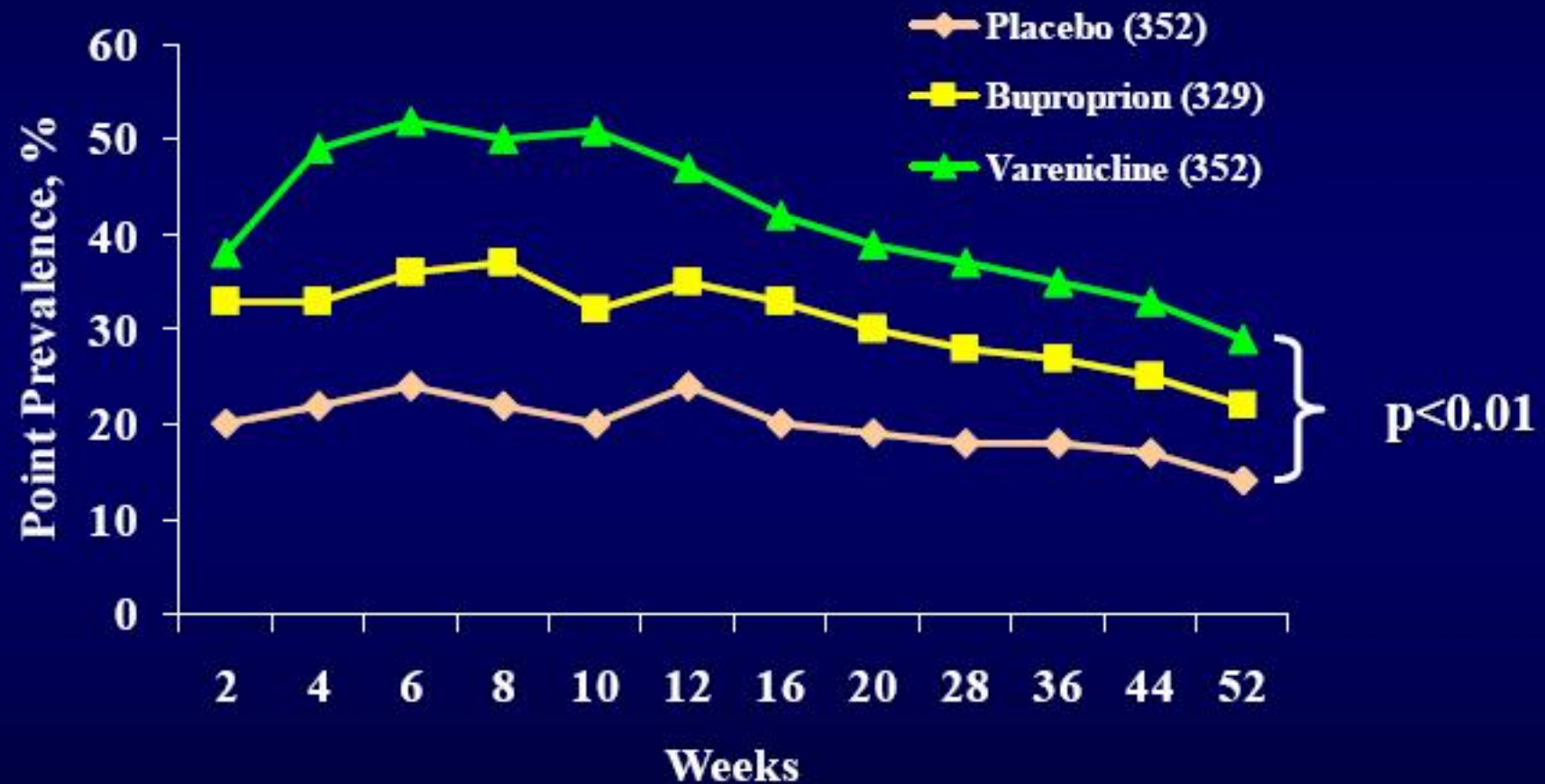


An extrapolation of Fletcher and Peto. 1977



"Are you sure you don't want to add  
something about smoking?"

# Varenicline and Smoking Cessation



Gonzales et al JAMA 2006;296:47



# Factsheets

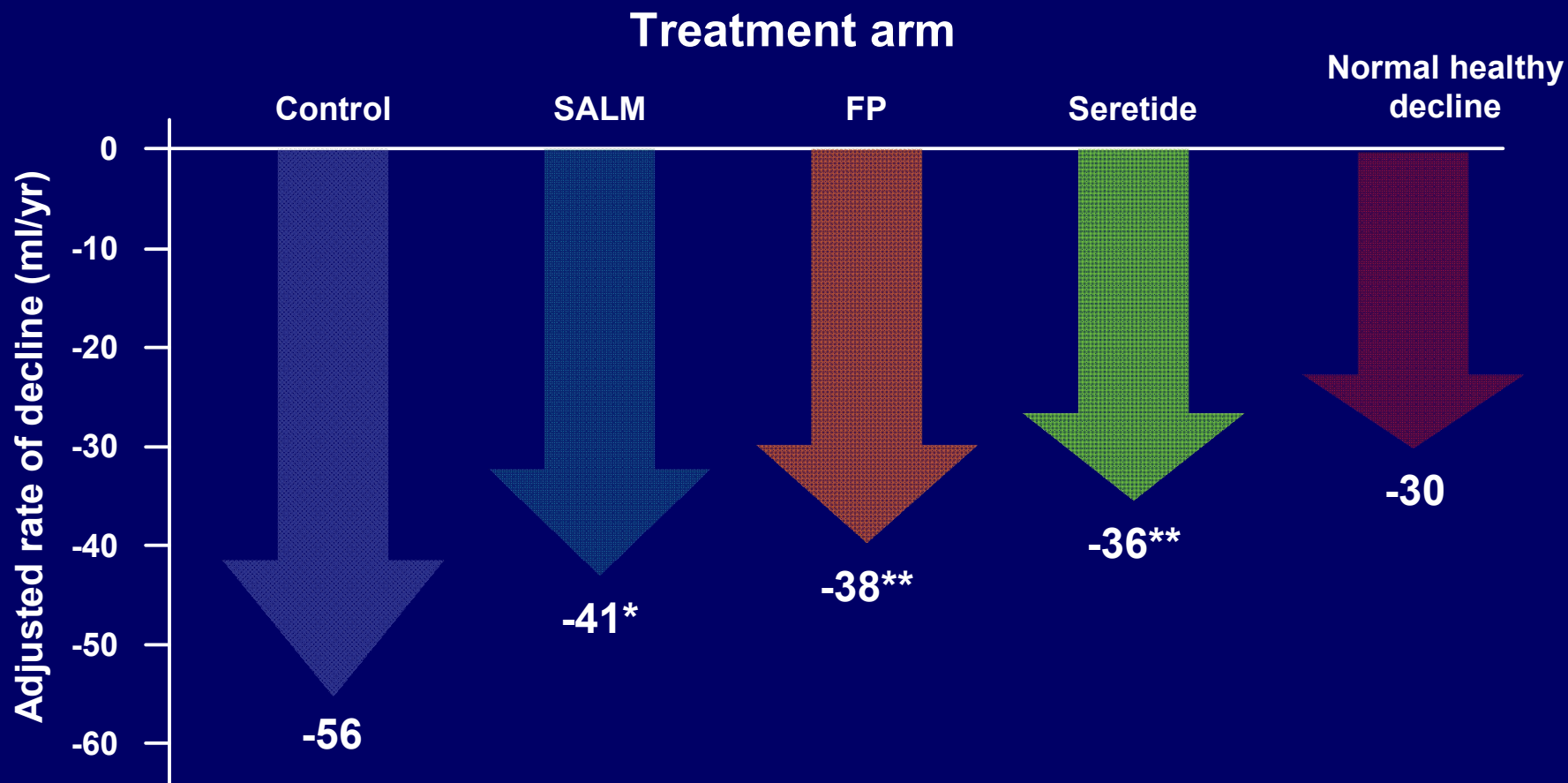
## How can I interpret air pollution levels?

Many countries and international agencies have developed systems to show the different levels of air pollution in different areas each day and this alerts the population when levels are excessive. One of these, from the UK Department of Health, is shown below. This system works by grading each concentration of pollutant, and ranking it at a level between 1 and 10. These levels are then split into four categories: low; moderate; high; and very high.

Band	Index	O <sub>3</sub> µg/m <sup>3</sup>	NO <sub>2</sub> µg/m <sup>3</sup>	PM µg/m <sup>3</sup>	SO <sub>2</sub> µg/m <sup>3</sup>
Low	1	0-32	0-95	0-16	0-88
	2	33-66	96-190	17-32	89-176
	3	67-99	191-286	33-49	177-265
Moderate	4	100-126	287-381	50-57	266-354
	5	127-152	382-476	58-66	355-442
	6	153-179	478-572	67-74	443-531
High	7	180-239	573-635	75-82	532-708
	8	240-299	636-700	83-91	709-886
	9	300-359	701-763	92-99	887-1063
Very high	10	≥360	≥764	≥100	≥1064

# Rate of decline of FEV<sub>1</sub>

## <50% pre-bronchodilator FEV<sub>1</sub>



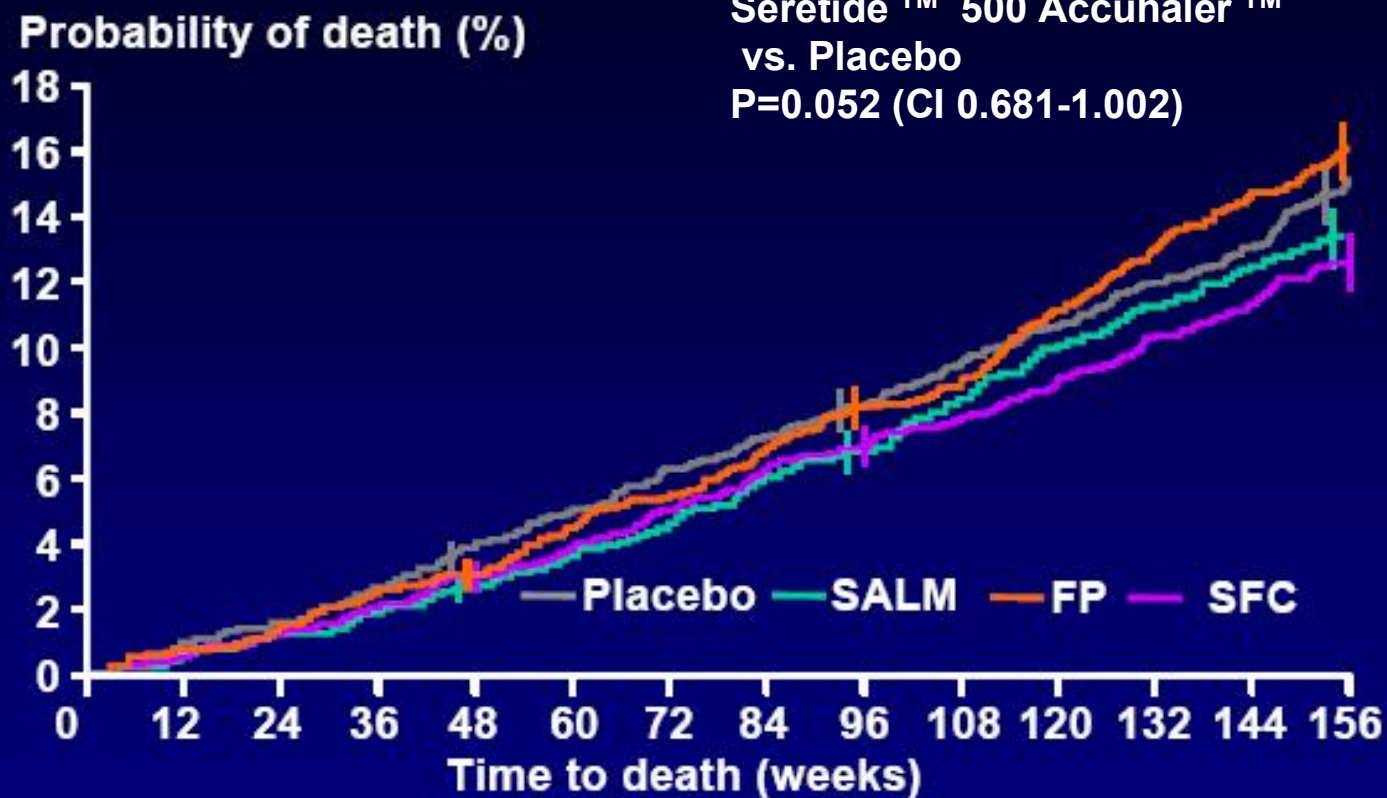
\*p=0.004,

\*\*p<0.001

GSK Data on File SERTCODOF012 <50% FEV<sub>1</sub>

# All-cause mortality at 3 years

RR reduction in mortality of 17.5%  
Absolute reduction 2.6% over 3 years  
Seretide™ 500 Accuhaler™  
vs. Placebo  
P=0.052 (CI 0.681-1.002)



Number	1524	1464	1399	1293
alive	1533	1487	1426	1339
	1521	1481	1417	1316
	1534	1487	1409	1288

Vertical bars are standard errors

Calverley et al. NEJM 2007

# COPD: Life Continues...











## Asthma versus COPD



[www.ginasthma.com](http://www.ginasthma.com)

- different diseases

- different treatments

- different prognoses



[www.goldcopd.org](http://www.goldcopd.org)

## Asthma versus COPD

### Additional investigations

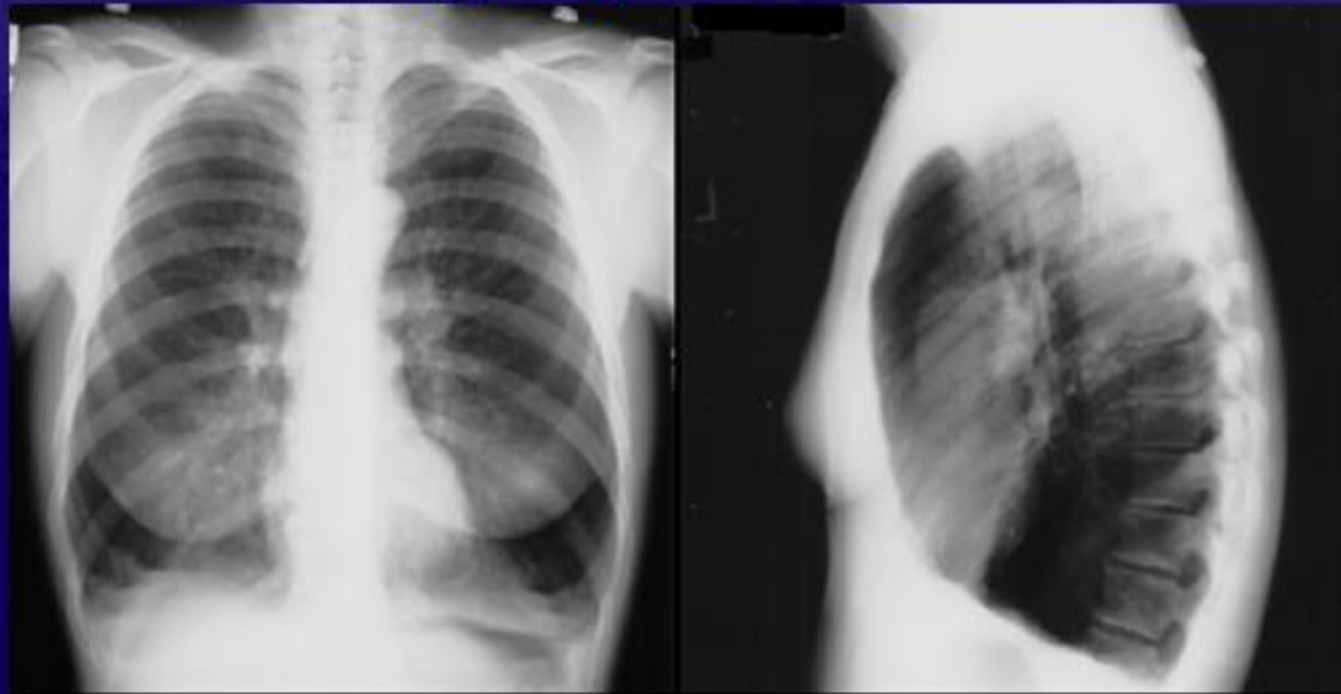


- Imaging



- Allergy testing

# chest radiograph



PA (left) and lateral (right) radiographs of a young woman with alpha-1-antitrypsin deficiency and typical changes of severe obstructive lung disease. There is diminished vascularity (arterial deficiency pattern) along with flattened diaphragms and an increase in the AP dimension of the chest.

## **Bullae**

**Focal absence of pulmonary vessels**

**Reduction in vessel calibre, vessel tapering**

**Signs of overinflation**

*Sensitivity for emphysema ~ 40%*

*Thurlbeck and Simon 1978*

*Sensitive in diagnosis but not quantification in*

*Moderate-severe disease*

*Thurlbeck and Muller 1999*



**Cigarette smoke**  
**Biomass particles**  
**Particulates**



## Pathogenesis of COPD

**Host factors**  
**Amplifying mechanisms**

**LUNG INFLAMMATION**

**Anti-oxidants**

**Anti-proteinases**

**Oxidative stress**

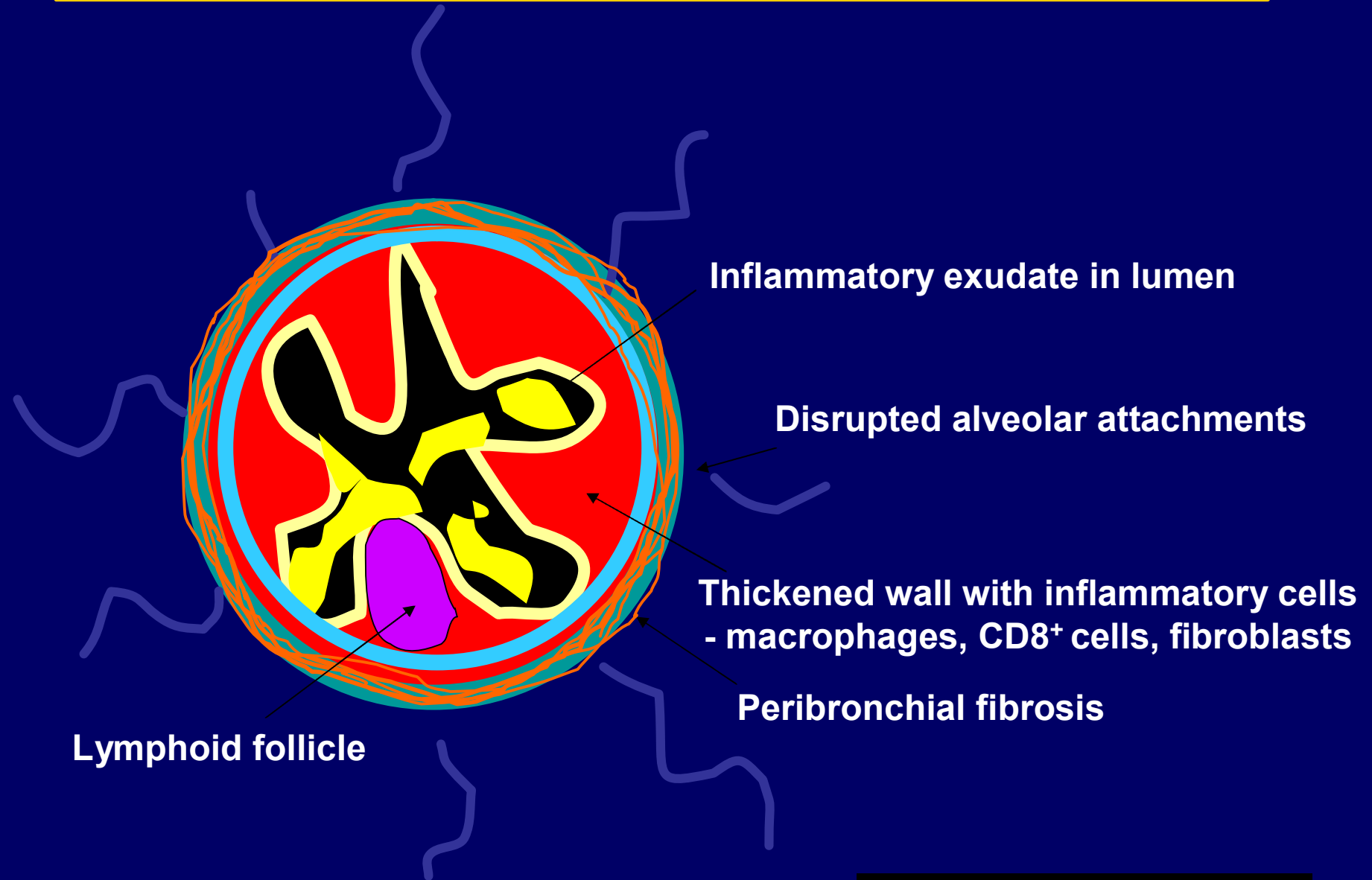
**Proteinases**

**Repair mechanisms**

**COPD PATHOLOGY**

*Source:* Peter J. Barnes, MD

# Changes in Small Airways in COPD Patients



**Source:** Peter J. Barnes, MD





# Risk Factors for COPD

---

## Genes

## Exposure to particles

- Tobacco smoke
- Occupational dusts, organic and inorganic
- Indoor air pollution from heating and cooking with biomass in poorly ventilated dwellings
- Outdoor air pollution

## Lung growth and development

## Oxidative stress

## Gender

## Age

## Respiratory infections

## Socioeconomic status

## Nutrition

## Comorbidities



## Management of Stable COPD

### Assess and Monitor COPD: Key Points

---

- A clinical diagnosis of COPD should be considered in any patient who has dyspnea, chronic cough or sputum production, and/or a history of exposure to risk factors for the disease.
- The diagnosis should be confirmed by spirometry. A post-bronchodilator  $FEV_1/FVC < 0.70$  confirms the presence of airflow limitation that is not fully reversible.
- Comorbidities are common in COPD and should be actively identified.



## Management of Stable COPD

### Assess and Monitor COPD: Spirometry

---

- Spirometry should be performed after the administration of an adequate dose of a short-acting inhaled bronchodilator to minimize variability.
- A post-bronchodilator  $FEV_1/FVC < 0.70$  confirms the presence of airflow limitation that is not fully reversible.
- Where possible, values should be compared to age-related normal values to avoid overdiagnosis of COPD in the elderly.



# COPD and Co-Morbidities

---

COPD has significant extrapulmonary (systemic) effects including:

- Weight loss
- Nutritional abnormalities
- Skeletal muscle dysfunction



# Brief Strategies to Help the Patient Willing to Quit Smoking

---

- **ASK** Systematically identify all tobacco users at every visit.
- **ADVISE** Strongly urge all tobacco users to quit.
- **ASSESS** Determine willingness to make a quit attempt.
- **ASSIST** Aid the patient in quitting.
- **ARRANGE** Schedule follow-up contact.





## Management of Stable COPD

### Reduce Risk Factors: Indoor/Outdoor Air Pollution

---

- Reducing the risk from indoor and outdoor air pollution is feasible and requires a combination of public policy and protective steps taken by individual patients.
- Reduction of exposure to smoke from biomass fuel, particularly among women and children, is a crucial goal to reduce the prevalence of COPD worldwide.



# Therapy at Each Stage of COPD

I: Mild

II: Moderate

III: Severe

IV: Very Severe

- $FEV_1/FVC < 70\%$
- $FEV_1 \geq 80\%$

- $FEV_1/FVC < 70\%$
- $50\% \leq FEV_1 < 80\%$  predicted

- $FEV_1/FVC < 70\%$
- $30\% \leq FEV_1 < 50\%$  predicted

- $FEV_1/FVC < 70\%$
- $FEV_1 < 30\%$  predicted  
or  $FEV_1 < 50\%$  predicted plus chronic respiratory failure

Active reduction of risk factor(s); influenza vaccination

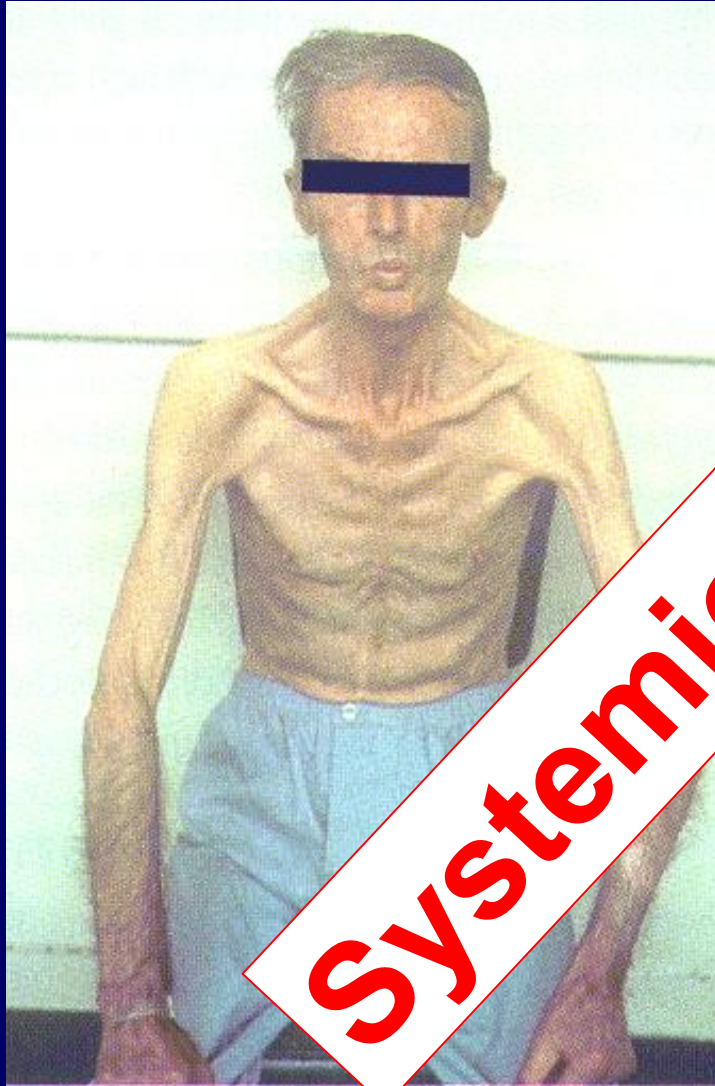
**Add** short-acting bronchodilator (when needed)

**Add** regular treatment with one or more long-acting bronchodilators (when needed); **Add** rehabilitation

**Add** inhaled glucocorticosteroids if repeated exacerbations

**Add** long term oxygen if chronic respiratory failure.  
**Consider** surgical treatments

# Patient with COPD and Cachexia



**Systemic Phenotype**

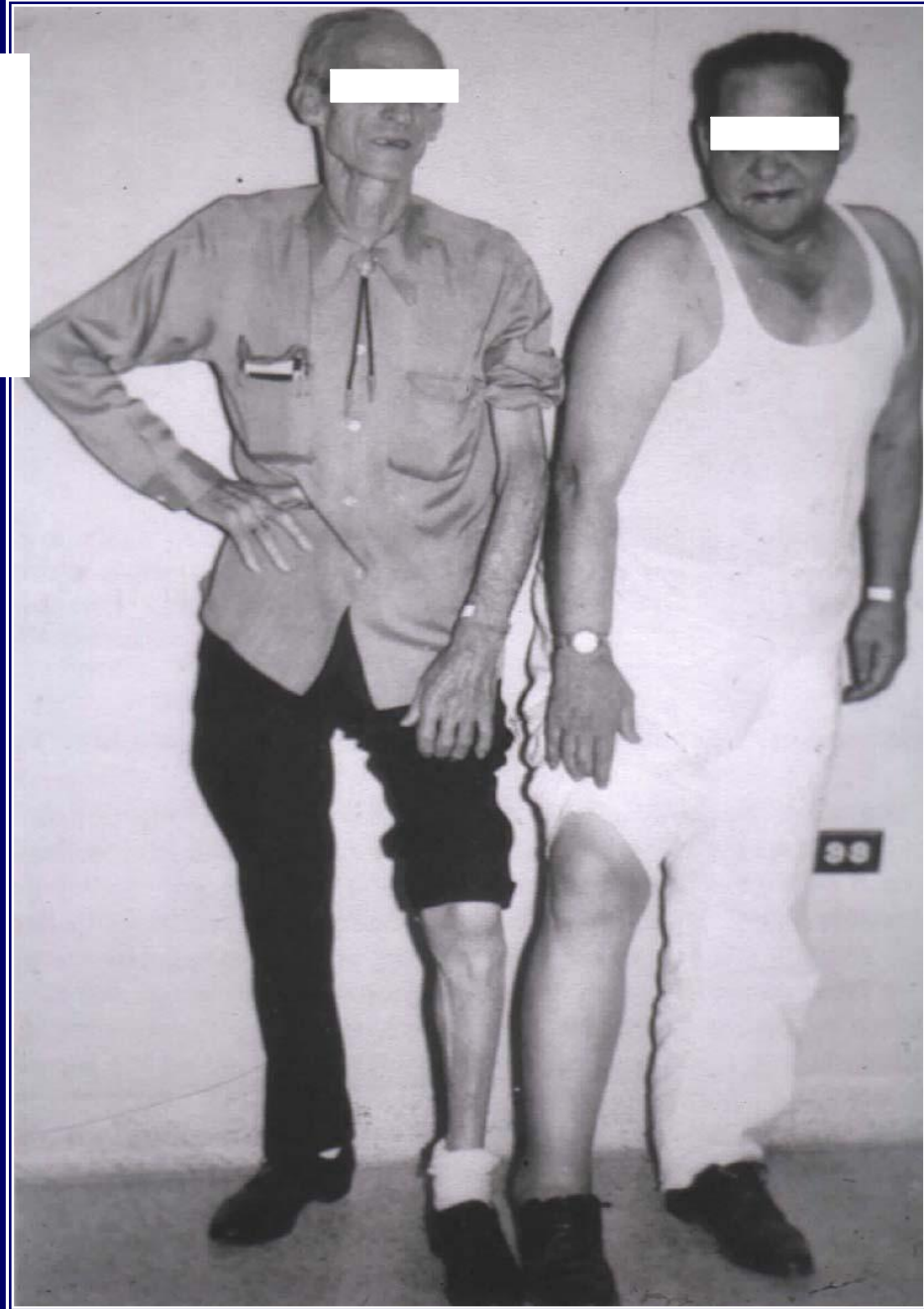
- loss in fat mass (FFM)
- low body mass index (BMI)
- increased mortality
- resistant to diet supplement resist.

- elevated  $\text{TNF}\alpha$
- loss in myosin heavy chain
- Increased oxidative stress
- muscle apoptosis

**These COPD patients  
have different  
phenotypes**

**Should they both  
receive the same  
Treatment ?**

*From T Petty*



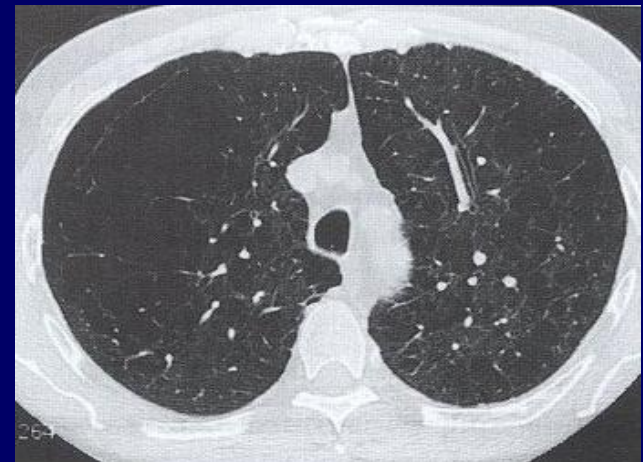
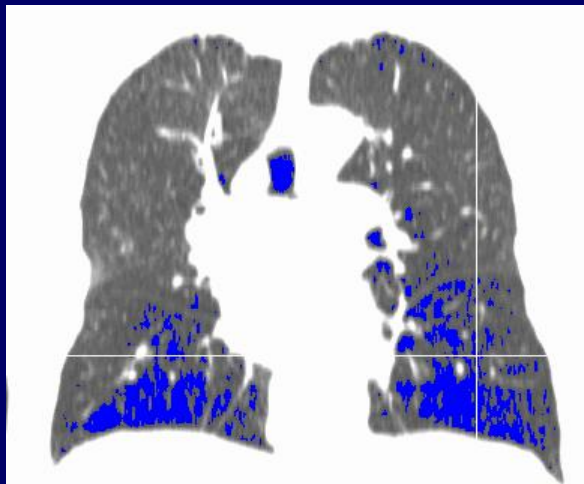
# Facts About COPD

- ♦ COPD is the 4<sup>th</sup> leading cause of death in the United States (behind heart disease, cancer, and cerebrovascular disease).
- ♦ In 2000, the WHO estimated 2.74 million deaths worldwide from COPD.
- ♦ In 1990, COPD was ranked 12<sup>th</sup> as a burden of disease; by 2020 it is projected to rank 5<sup>th</sup>.

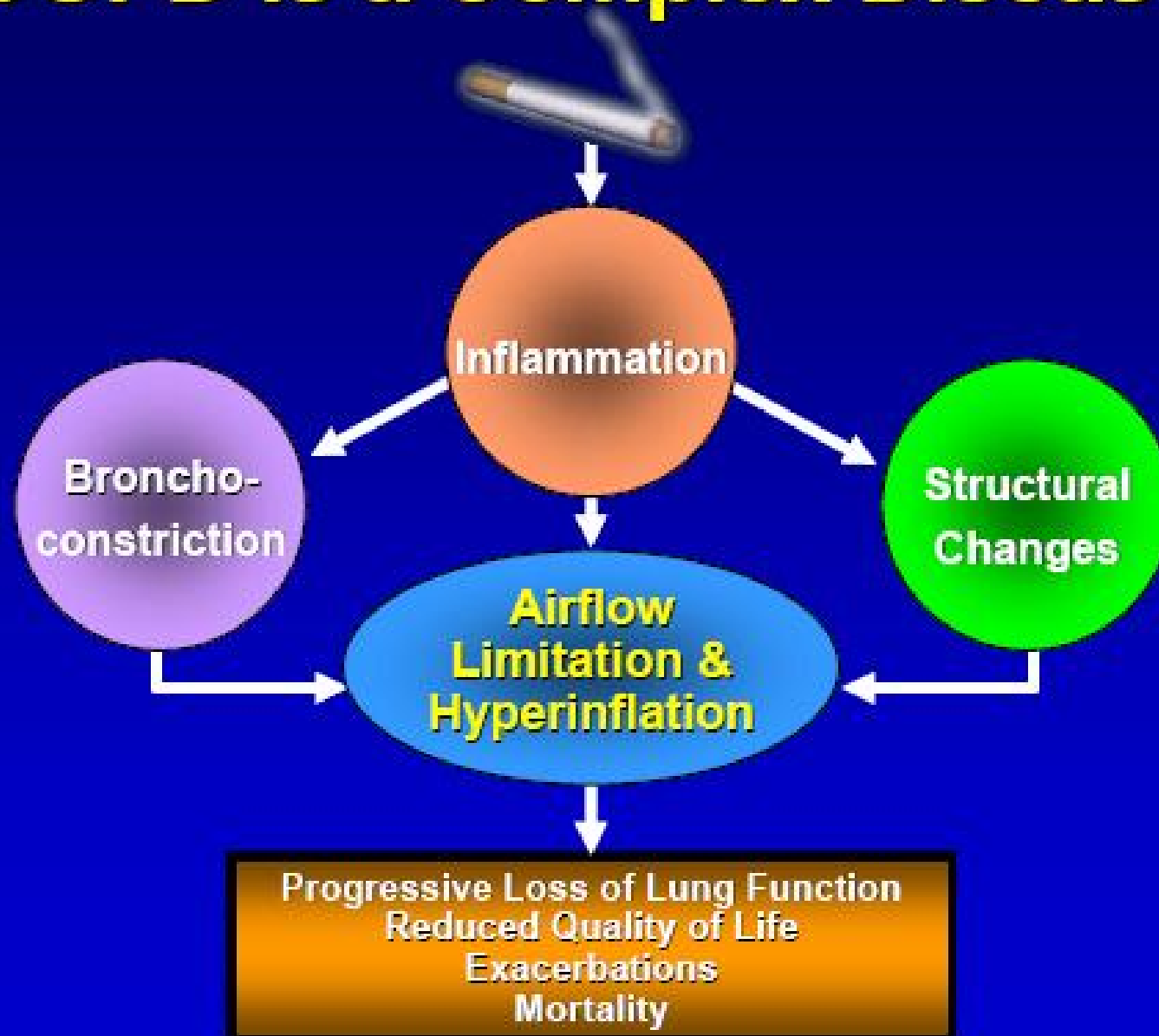


# Alpha –1 antitrypsin deficiency ( $\alpha_1$ proteinase deficiency)

- Pi ZZ 1/5000 Live births in UK
- Liver disease in some cases in childhood
- Emphysema < 50 years of age often in 4<sup>th</sup> decade
- Some individuals lead a normal life
- Panlobular predominantly basal emphysema
- Smoking is a cofactor in the risk of emphysema

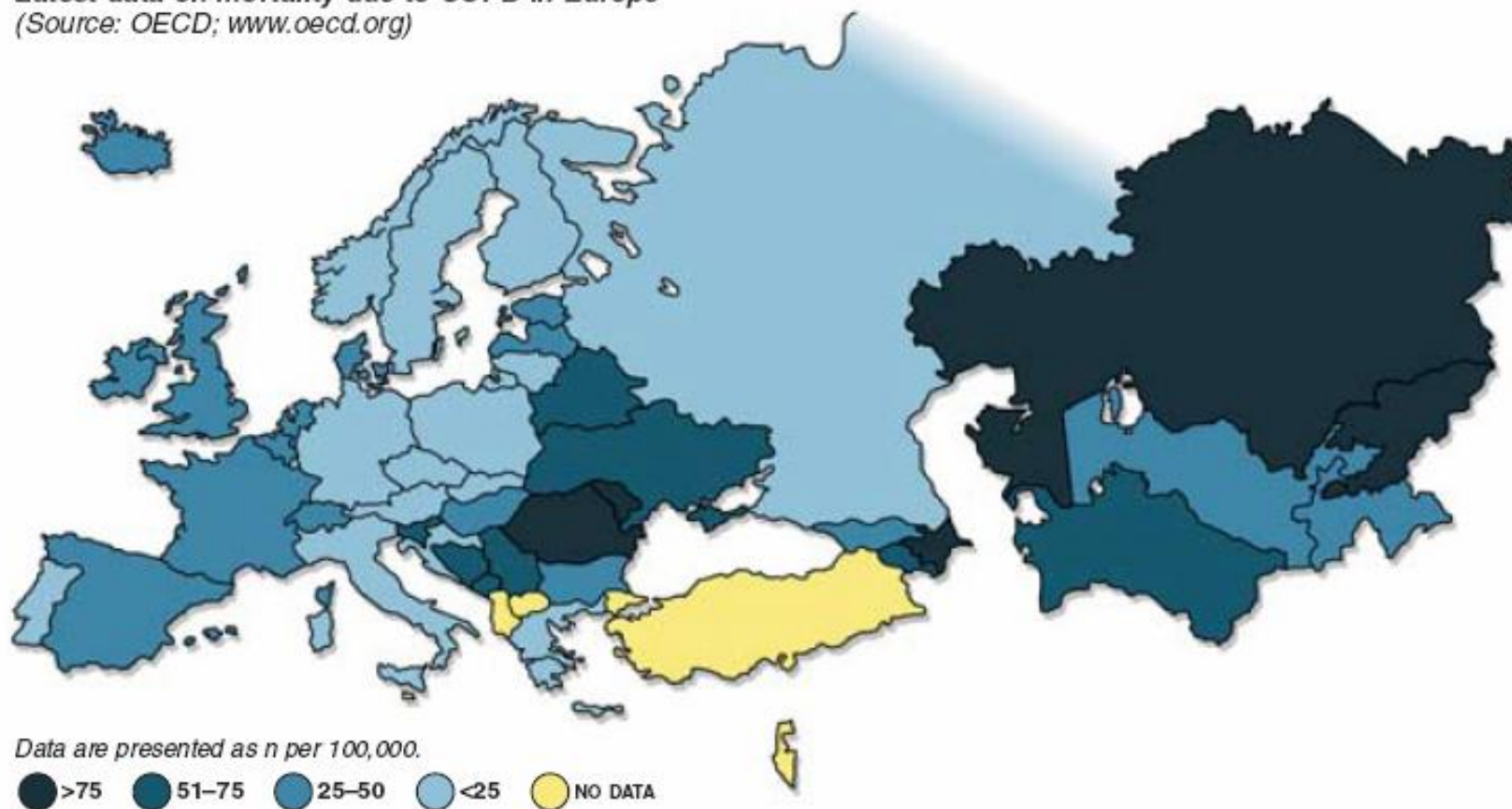


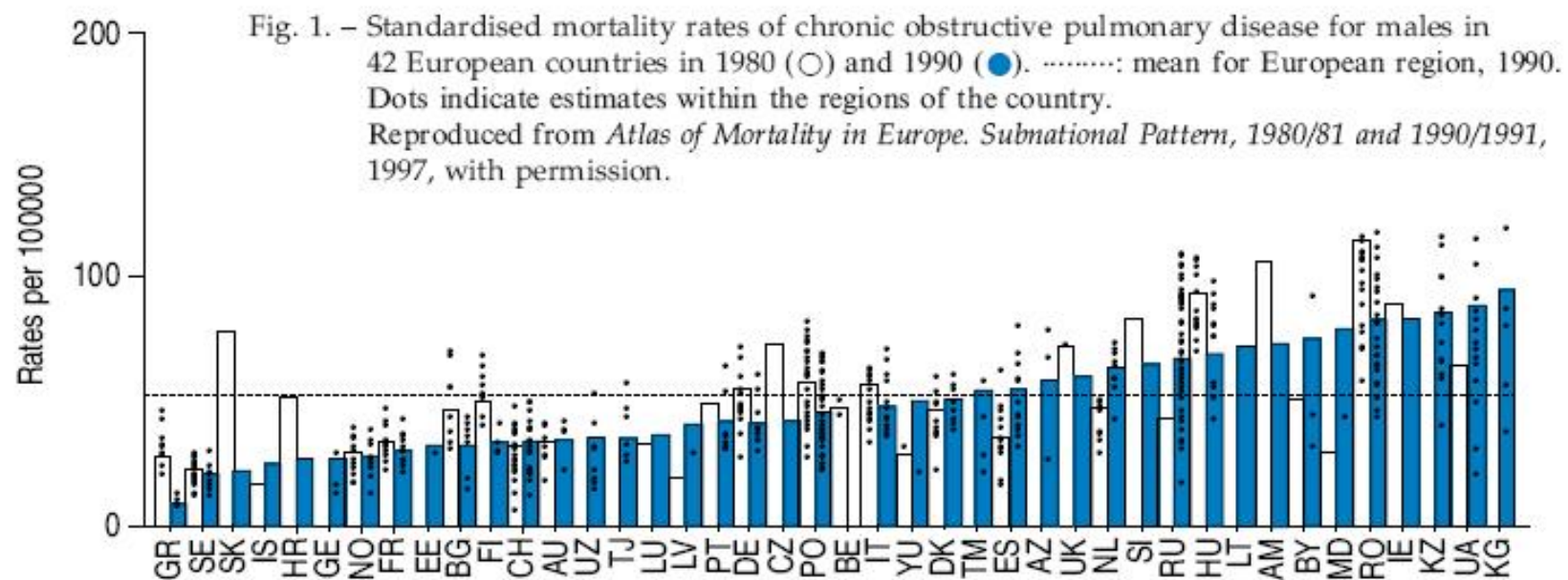
# COPD is a Complex Disease



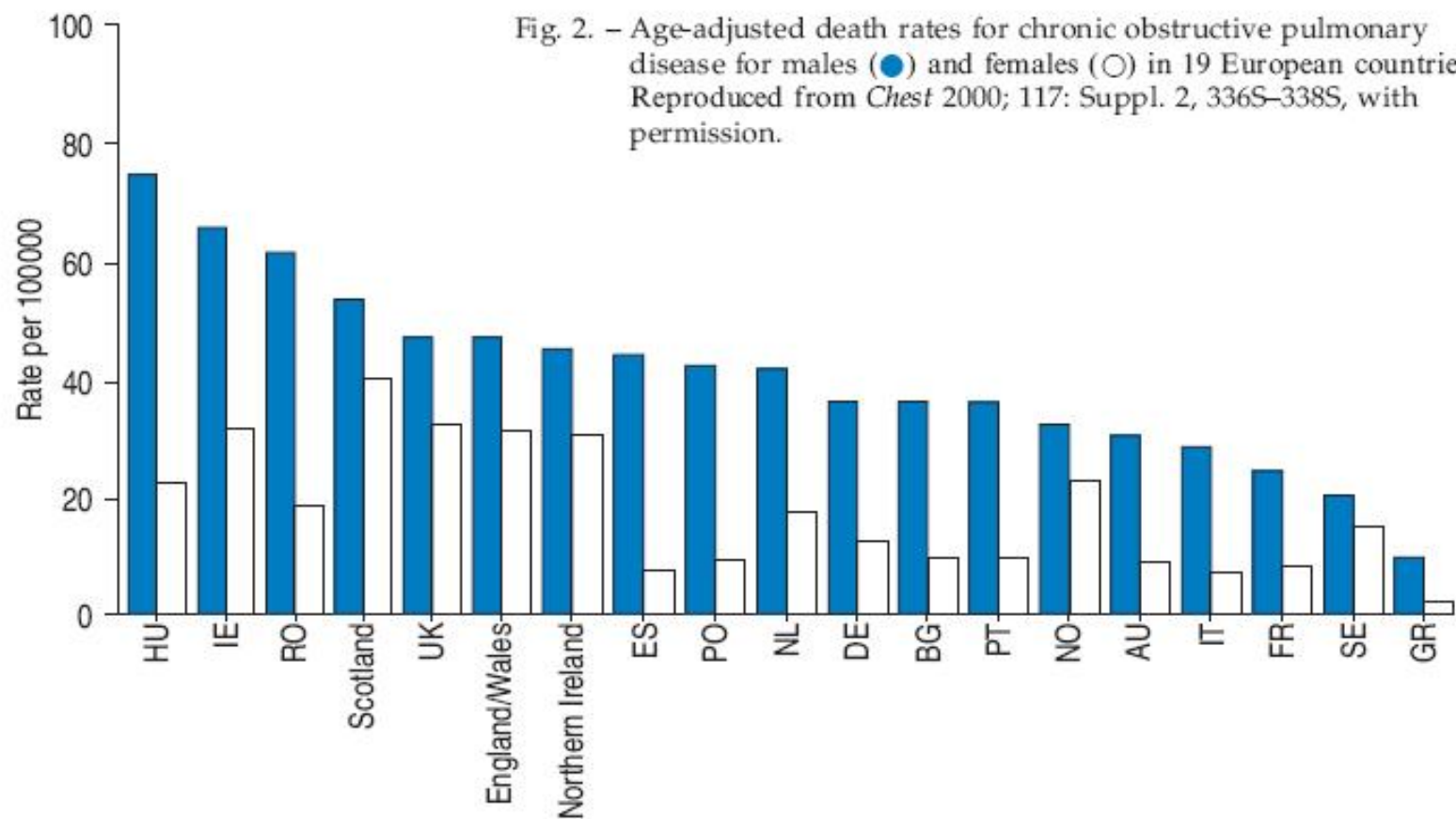


*Latest data on mortality due to COPD in Europe*  
(Source: OECD; [www.oecd.org](http://www.oecd.org))



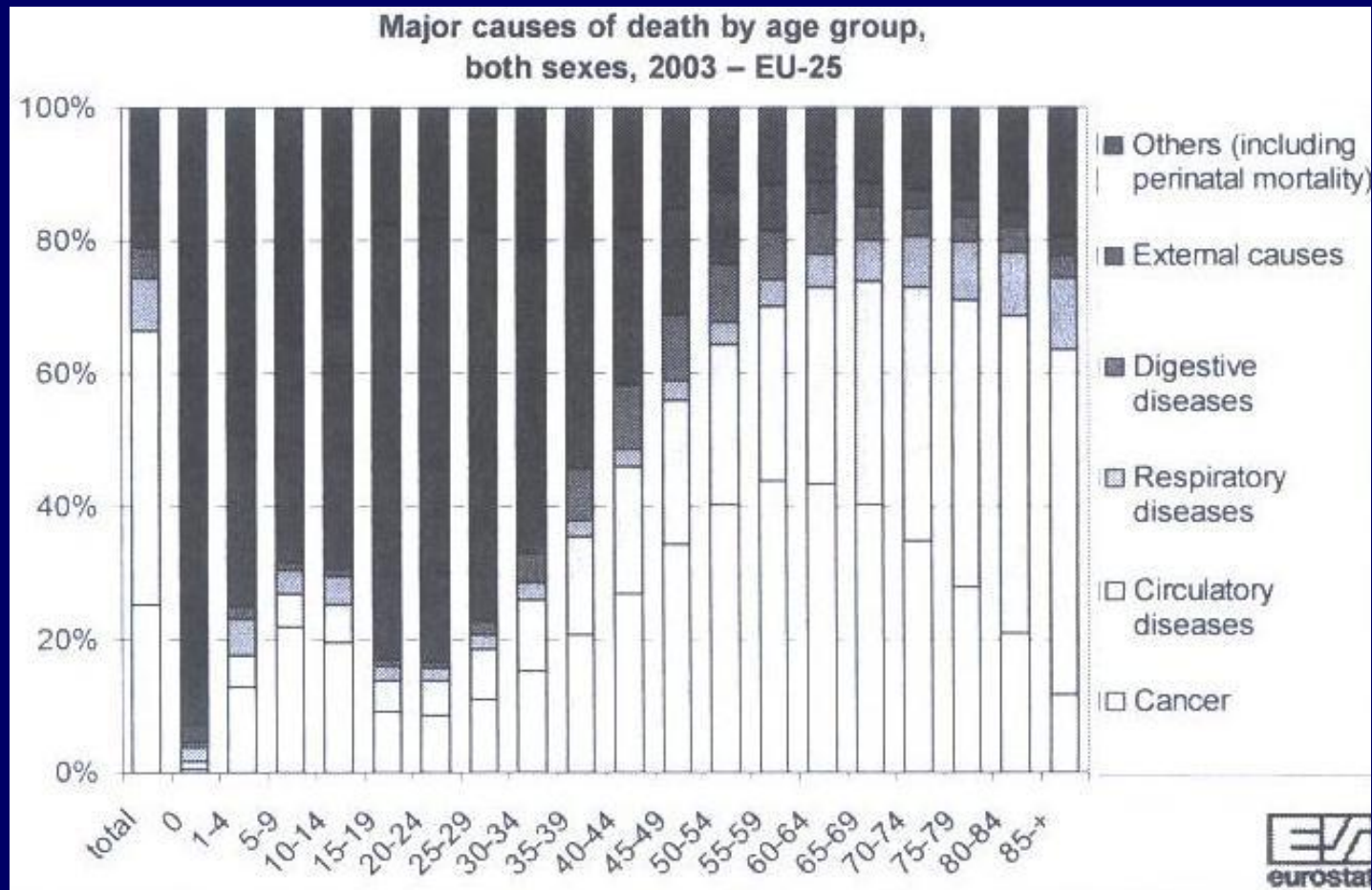






**EU25 - Principal causes of death by age group, in percent**

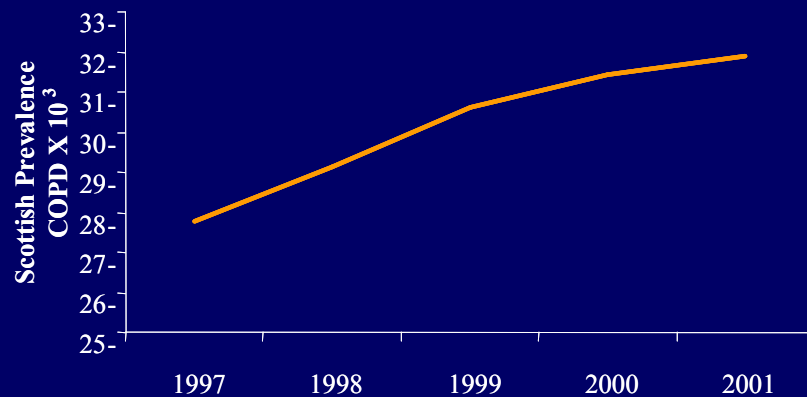
	<b>Cancer</b>	<b>Circulatory diseases</b>	<b>Respiratory diseases</b>	<b>Digestive diseases</b>	<b>External causes<sup>1</sup></b>	<b>Others<sup>1</sup></b>
<b>All age groups</b>	<b>25.3</b>	<b>41.0</b>	<b>8.0</b>	<b>4.6</b>	<b>5.2</b>	<b>15.9</b>
0-19 years old	7.2	3.2	2.8	0.9	26.6	59.3
20-44 years old	19.7	13.8	2.5	6.6	38.0	19.4
45-64 years old	41.4	26.3	4.1	7.6	8.3	12.3
65-84 years old	29.1	42.0	8.2	4.2	2.8	13.7
Over 85 years old	11.6	52.0	10.6	3.5	2.9	19.4



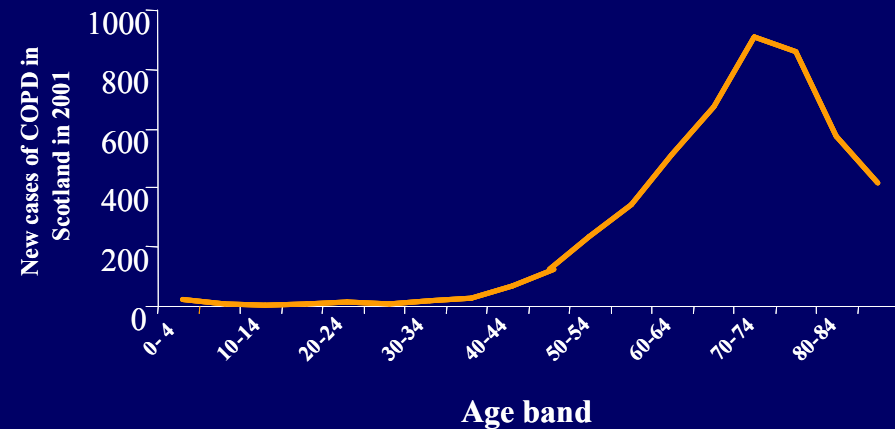
*Figure 1: Main causes of death by 5-year age group*

# COPD epidemiology in Scotland

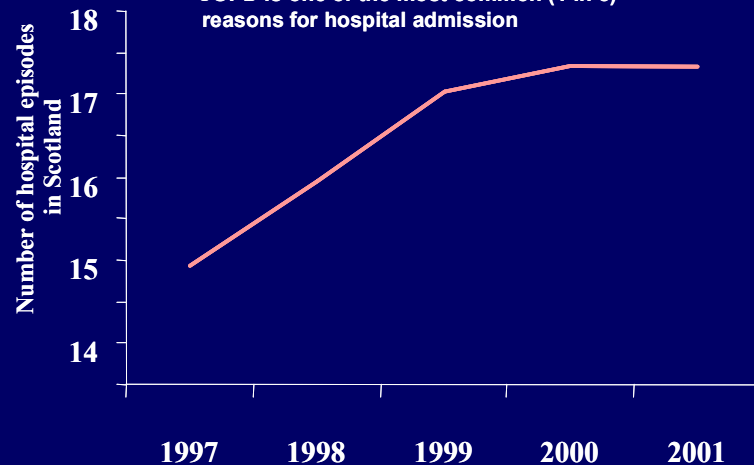
The number of people with COPD is substantial and rising



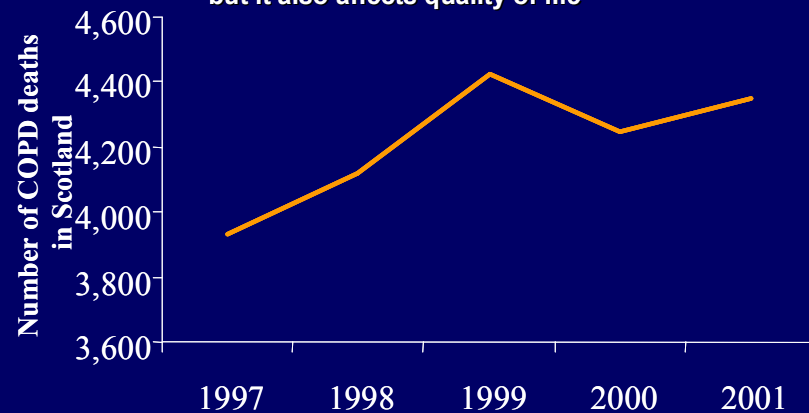
Those most likely to be affected already have the poorest health



COPD is one of the most common (1 in 8) reasons for hospital admission



COPD is a major killer but it also affects quality of life



# Mortality of COPD



- 30,000 in UK <sup>1</sup>
- 5% of all deaths <sup>2</sup>
- 4% of all females <sup>2</sup>
- 6% of all males <sup>2</sup>

Ref. 1. The Burden of Lung Disease BTS, Nov 2001

2. Calverley PMA, Thorax 2002



# Hospital admission for COPD

- 26% of medical admissions are for respiratory disease
- half due to COPD !

**220,000 admissions/year**

- mean length of stay 7.5 days
- over 1 million bed days <sup>1</sup>
- 1 in 8 of all admissions <sup>2</sup>

Ref 1: Crocket A, Men's Health Journal 2003

2: NICE website

# Consultations for COPD



- ♦ Annual GP consultation rates 2 x angina
- ♦ 417/1000 for patients 45-64 years
- ♦ 1032/1000 for patients 75-84 years

Pearson, BTS Guidelines Group / Standards of Care Committee

Thorax 1997

# **COPD and Systemic Effects**

---

**COPD has significant extrapulmonary  
(systemic) effects including:**

- Weight loss**
- Nutritional abnormalities**
- Skeletal muscle dysfunction**

# **COPD and Co-Morbidities**

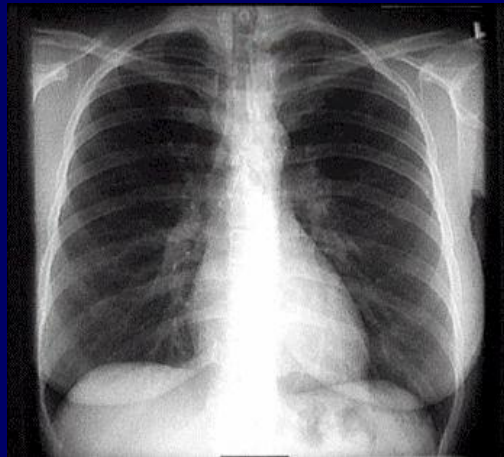
**COPD patients are at increased risk for:**

- **Myocardial infarction, angina**
- **Osteoporosis**
- **Respiratory infection**
- **Depression**
- **Diabetes**
- **Lung cancer**

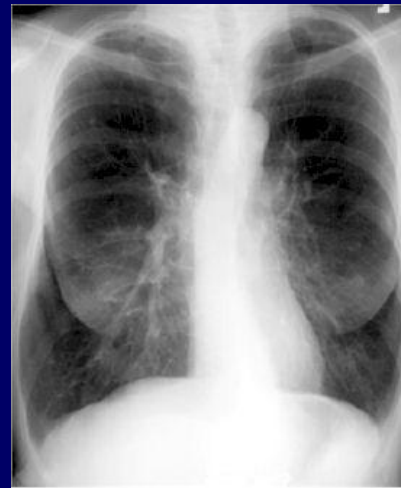
# Air Trapping

- ▲ Occurs in all patients with COPD
- ▲ Results in an increase in the work of breathing
- ▲ Places respiratory muscles at a mechanical disadvantage
- ▲ Contributes to the sensation of breathlessness (dyspnea)

**Normal**



**Hyperinflation**





# Leading causes of death

	1990		2020
Ischemic heart disease	1	→	1
Cerebrovascular disease	2	→	2
COPD	6	→	3
Lower respiratory infection	3	→	4
Lung cancer	10	→	5
Road traffic accidents	9	→	6
Tuberculosis	7	→	7
Stomach cancer	14	→	8

WHO Global Burden of Disease study

# Increasing burden of diseases and Injuries: change in rank order of DALYs

**1999**

1. Acute lower respiratory infections
2. HIV/AIDS
3. Perinatal conditions
4. Diarrhoeal diseases
5. Unipolar major depression
6. Ischaemic heart disease
7. Cerebrovascular disease
8. Malaria
9. Road traffic injuries
10. **COPD**
11. Congenital abnormalities
12. Tuberculosis
13. Falls
14. Measles
15. Anaemias

**2020**

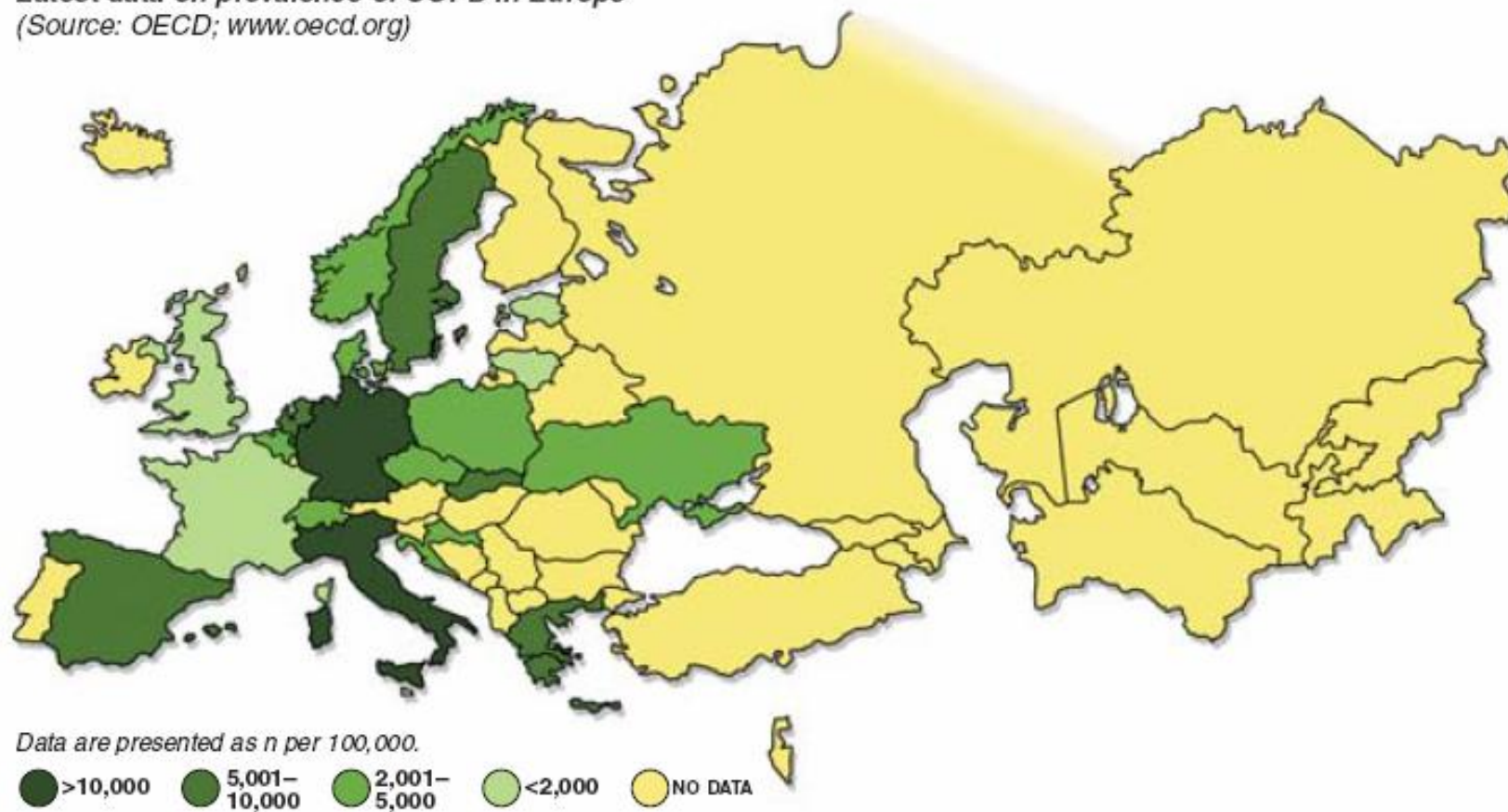
1. Ischaemic heart disease
2. Unipolar major depression
3. Road traffic injuries
4. Cerebrovascular disease
5. **COPD**
6. Acute lower respiratory infections
7. Tuberculosis
8. War
9. Diarrhoeal diseases
10. HIV
11. Perinatal conditions
12. Violence
13. Congenital abnormalities
14. Self-inflicted injuries
15. Trachea, bronchus and lung cancers

**DALY = Disability-adjusted life year**

**Source: WHO  
Evidence, Information and Policy, 2000**



*Latest data on prevalence of COPD in Europe*  
(Source: OECD; [www.oecd.org](http://www.oecd.org))



# Underdiagnosis of COPD

Diagnosed  
COPD  
2.4 - 7 million

Estimated total  
COPD  
16 million

56 - 85%  
Undiagnosed/misdiagnosed

US estimates

Stang, 2000

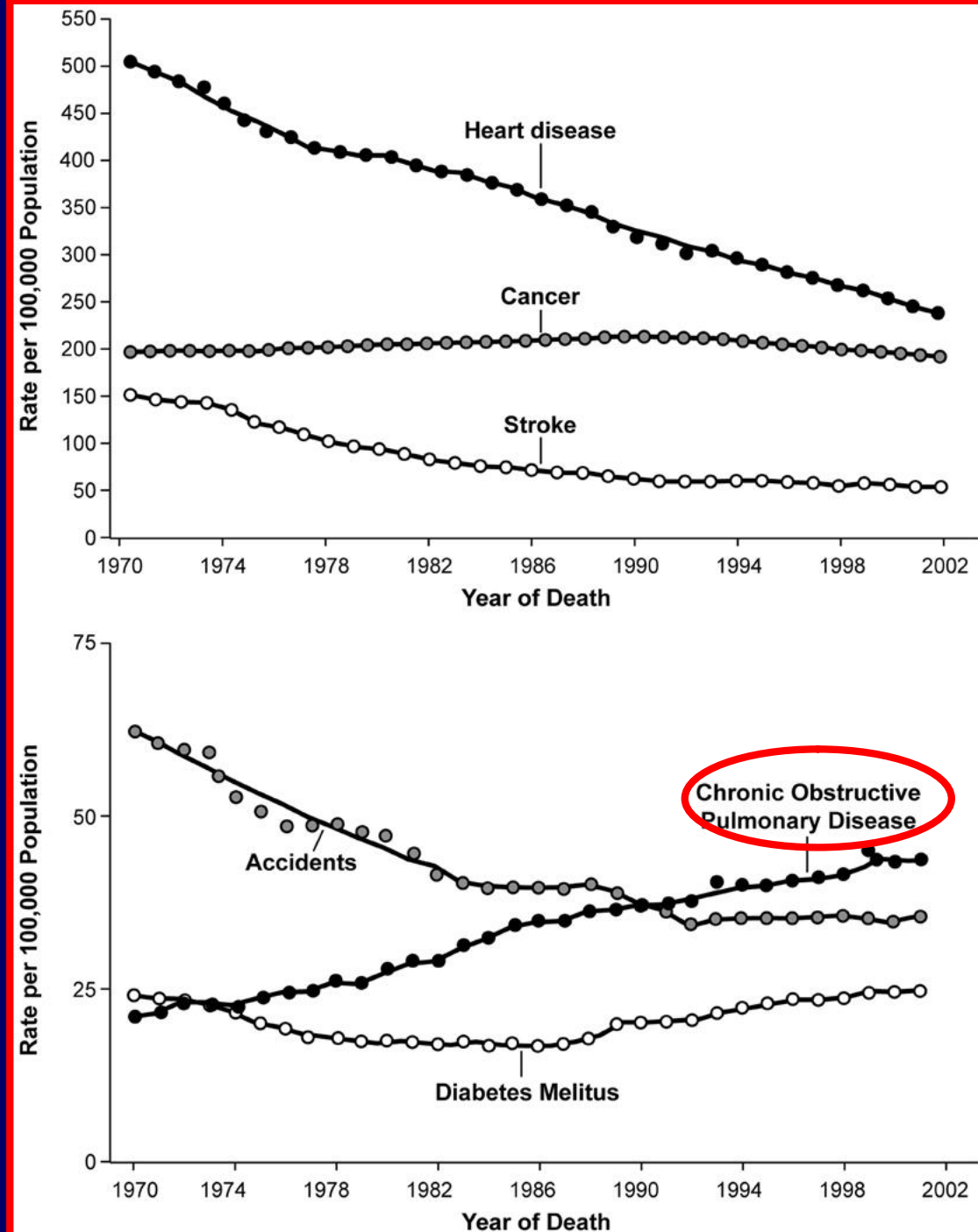
# Burden of COPD: Key Points

- COPD is a leading cause of morbidity and mortality worldwide and results in an economic and social burden that is both substantial and increasing.
- COPD prevalence, morbidity, and mortality vary across countries and across different groups within countries.
- The burden of COPD is projected to increase in the coming decades due to continued exposure to COPD risk factors and the changing age



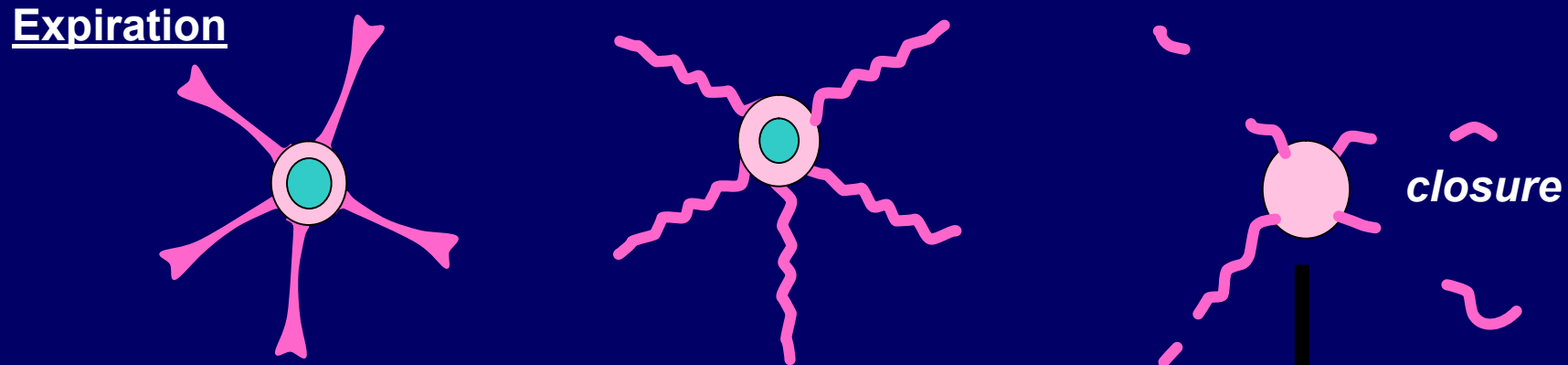
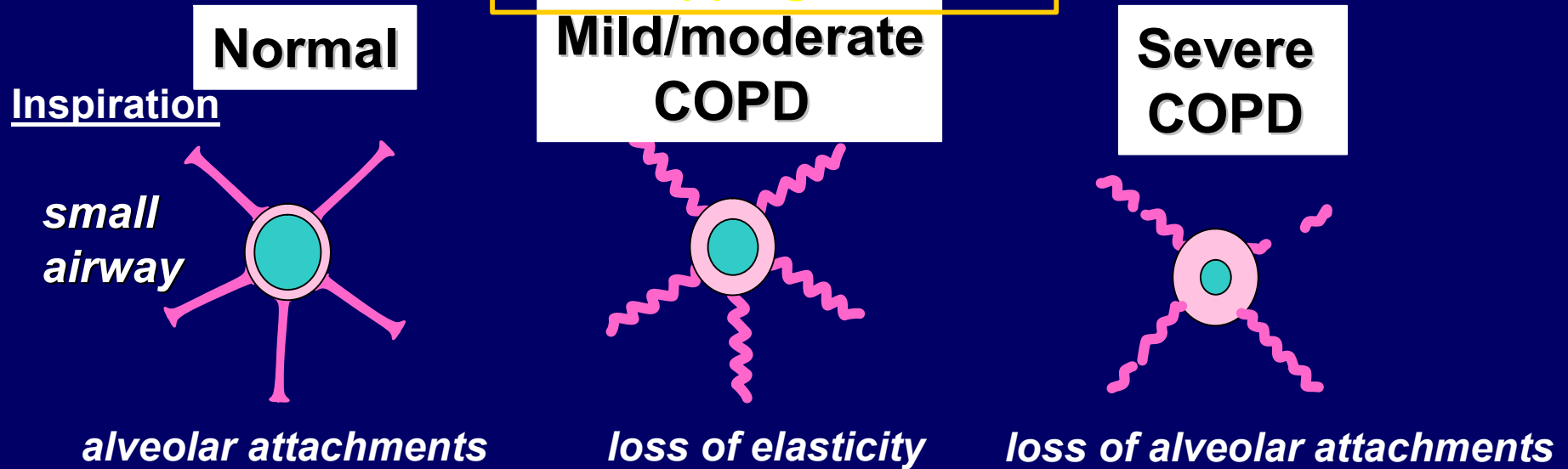
Of the six leading causes of death in the United States, only COPD has been increasing steadily since 1970

Source: Jemal A. et al. *JAMA* 2005





## Air Trapping in COPD



↓ **Health status**

**Dyspnea**  
↓ **Exercise capacity**

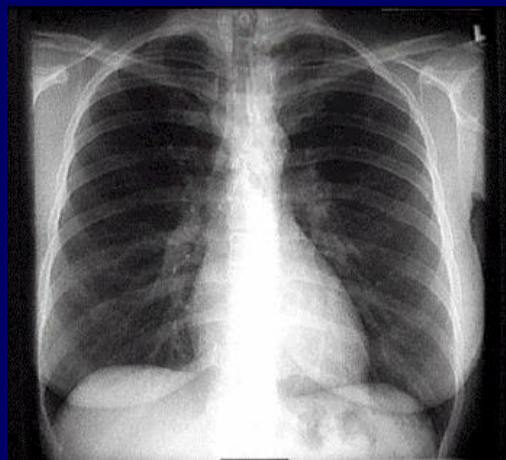
**Air trapping**  
**Hyperinflation**

*Source: Peter J. Barnes, MD*

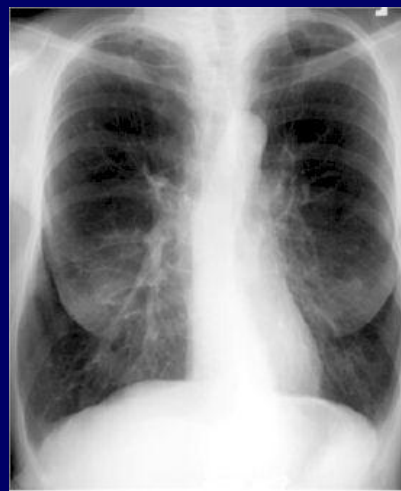
# Air Trapping

- ▲ Occurs in all patients with COPD
- ▲ Results in an increase in the work of breathing
- ▲ Places respiratory muscles at a mechanical disadvantage
- ▲ Contributes to the sensation of breathlessness

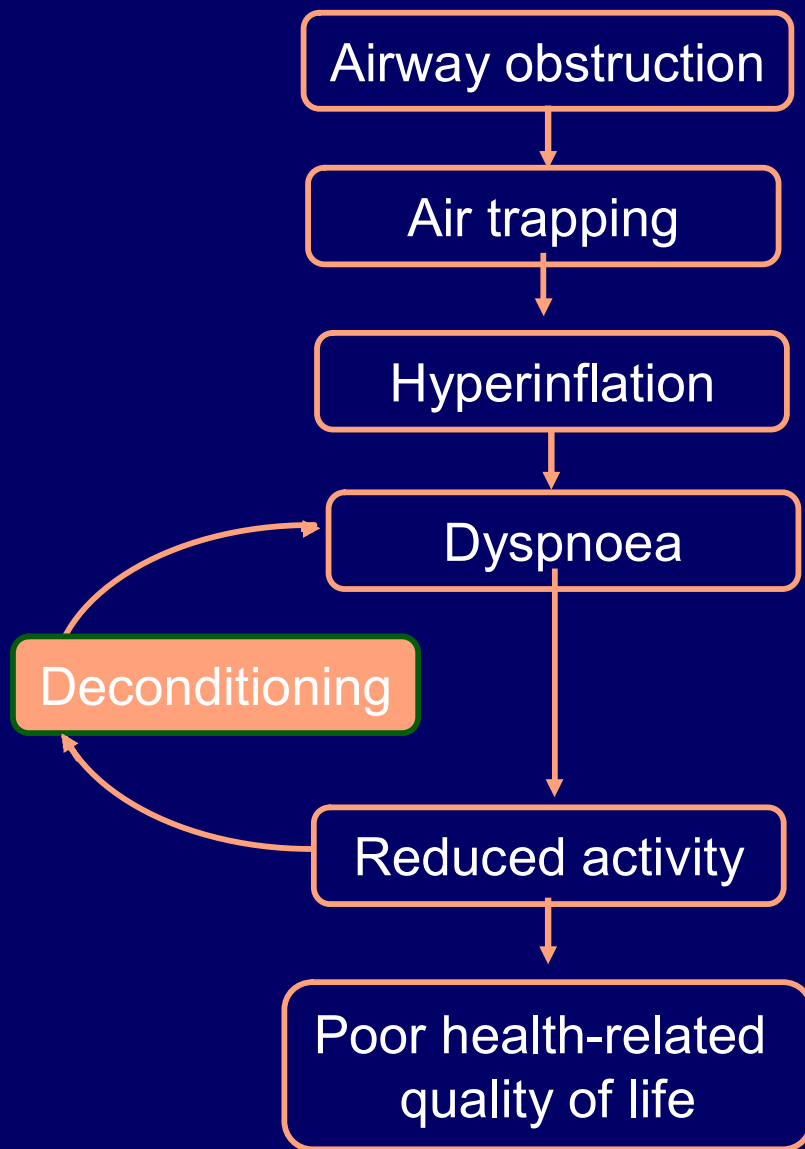
**Normal**



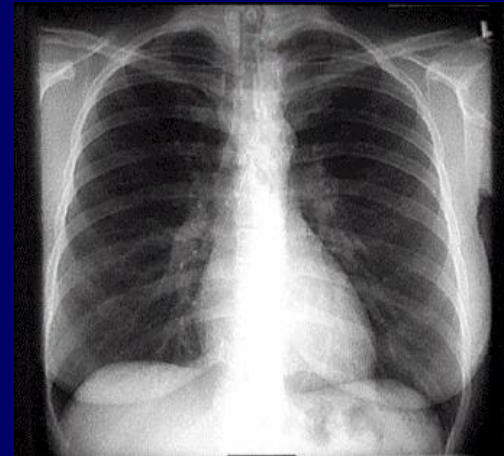
**Hyperinflation**



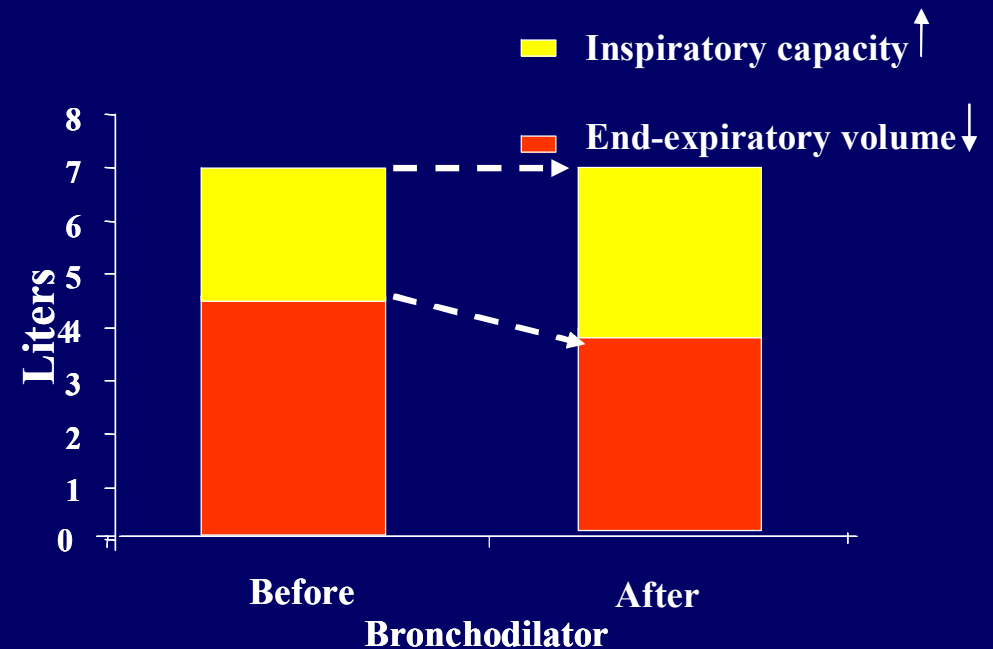
# Impact of air trapping (hyperinflation)



**Normal**



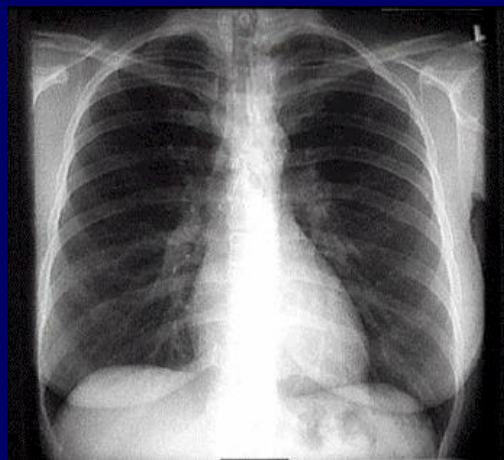
**Hyperinflation**



# Air Trapping

- ▲ Occurs in all patients with COPD
- ▲ Results in an increase in the work of breathing
- ▲ Places respiratory muscles at a mechanical disadvantage
- ▲ Contributes to the sensation of breathlessness

**Normal**



**Hyperinflation**



# Diagnosis of COPD

## SYMPTOMS

cough  
sputum  
shortness of breath

## EXPOSURE TO RISK FACTORS

tobacco  
occupation  
indoor/outdoor pollution



**SPIROMETRY**



# Definition of COPD

- COPD is a **preventable and treatable** disease. Its lung component is characterized by airflow limitation (or obstruction to the flow of air) that is not fully reversible.
- The airflow limitation is usually **progressive** and associated with an **abnormal inflammatory response** of the lung to noxious particles or gases.
- COPD has significant **extra-pulmonary effects** that may contribute to the severity in individual patients.

# Standards for the diagnosis and treatment of patients with COPD : a summary of the ATS/ERS position paper

## Diagnosis

The diagnosis of COPD should be considered in any patient who has the following:  
symptoms of cough; sputum production; or dyspnoea; or history of exposure to risk factors for the disease.

The diagnosis requires spirometry;  
a post-bronchodilator forced expiratory volume in one second (FEV<sub>1</sub>)/forced vital capacity (FVC)  $\leq 0.7$  confirms the presence of airflow limitation that is not fully reversible.

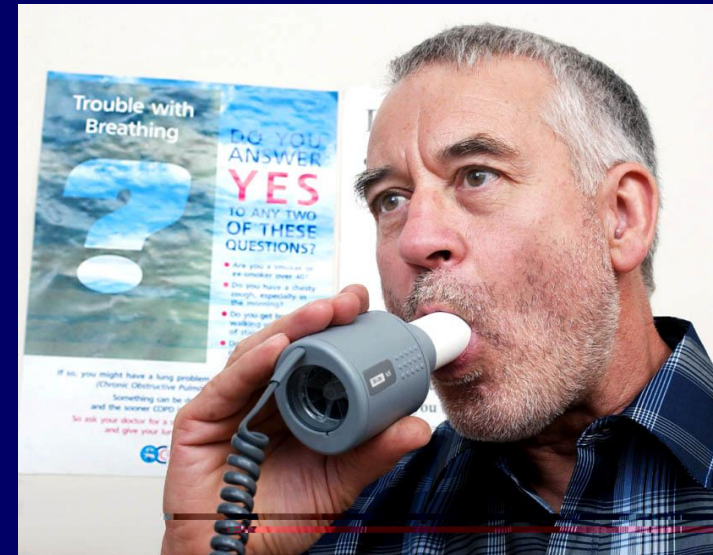
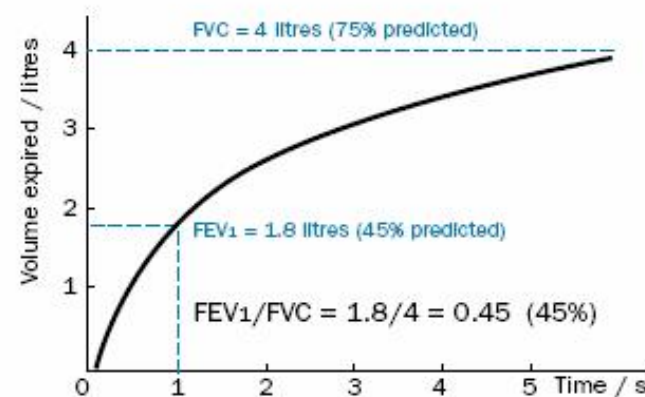
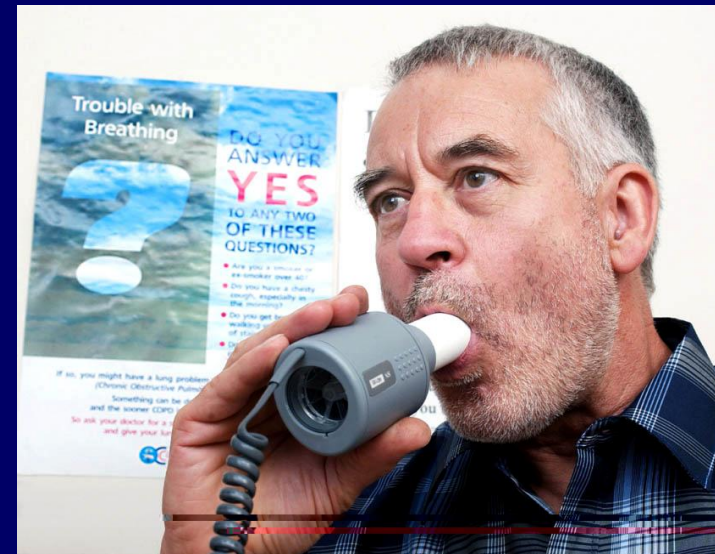


Figure 9a. Spirometry tracing of a patient with COPD – an obstructive picture



## NICE recommendations: Diagnosis (2)

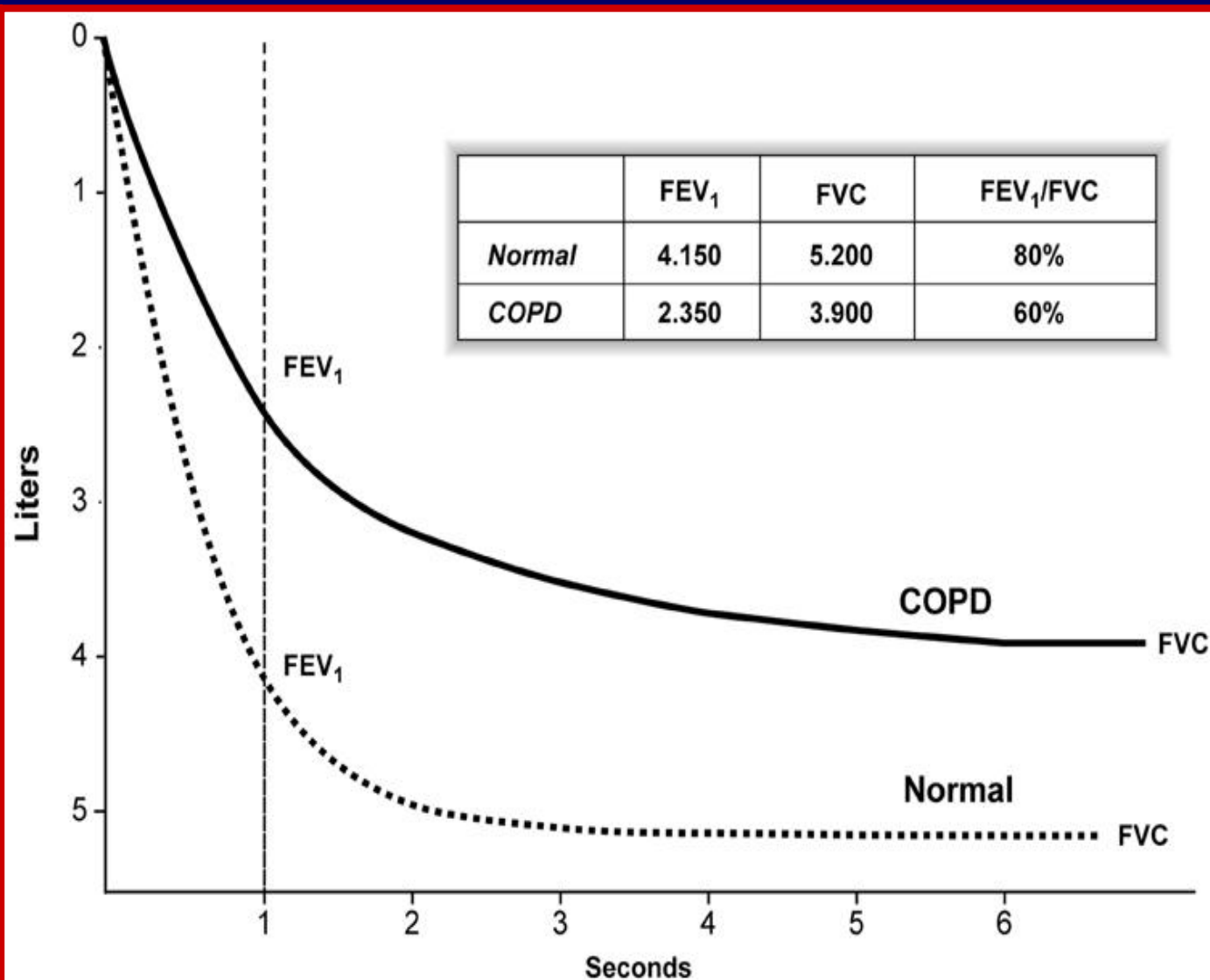
- If a diagnosis of COPD seems likely, perform spirometry
  - A diagnosis of airflow obstruction can be made if:
    - $FEV_1/FVC < 0.7$
    - $FEV_1 < 80\%$  predicted



Diagnosis can be made on patients' symptoms and be confirmed with spirometry



# Spirometry: Normal and Patients with COPD





# COPD and Co-Morbidities

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COPD patients are at increased risk for:

- Myocardial infarction, angina
- Osteoporosis
- Respiratory infection
- Depression
- Diabetes
- Lung cancer