



Simposio: Explorando a la EPOC aguas arriba - Parte II

# ¿Qué nos espera en la EPOC en los próximos 10 años?

Alvar Agusti



# Conflictos de interés

Research grants/clinical trials:	AstraZeneca, GSK, Menarini,
Lector for:	AstraZeneca, Chiesi, Menarini, GSK, Zambon
Member of scientific (advisory) board:	AstraZeneca, Chiesi, Menarini, GSK
Consultant:	None
Employer (including part-time):	None
Tobacco Industry relationship:	None





**¿Qué nos espera en la EPOC en los próximos 10 años?**



Hacer predicciones es muy difícil, especialmente cuando se trata del futuro.

(Niels Bohr)

1885 - 1962



## ¿Qué nos espera en la EPOC en los próximos 10 años?

1. Considerar el eje temporal: la EPOC no empieza a los 65 años
2. Abordar la complejidad biológica de la EPOC en la práctica clínica



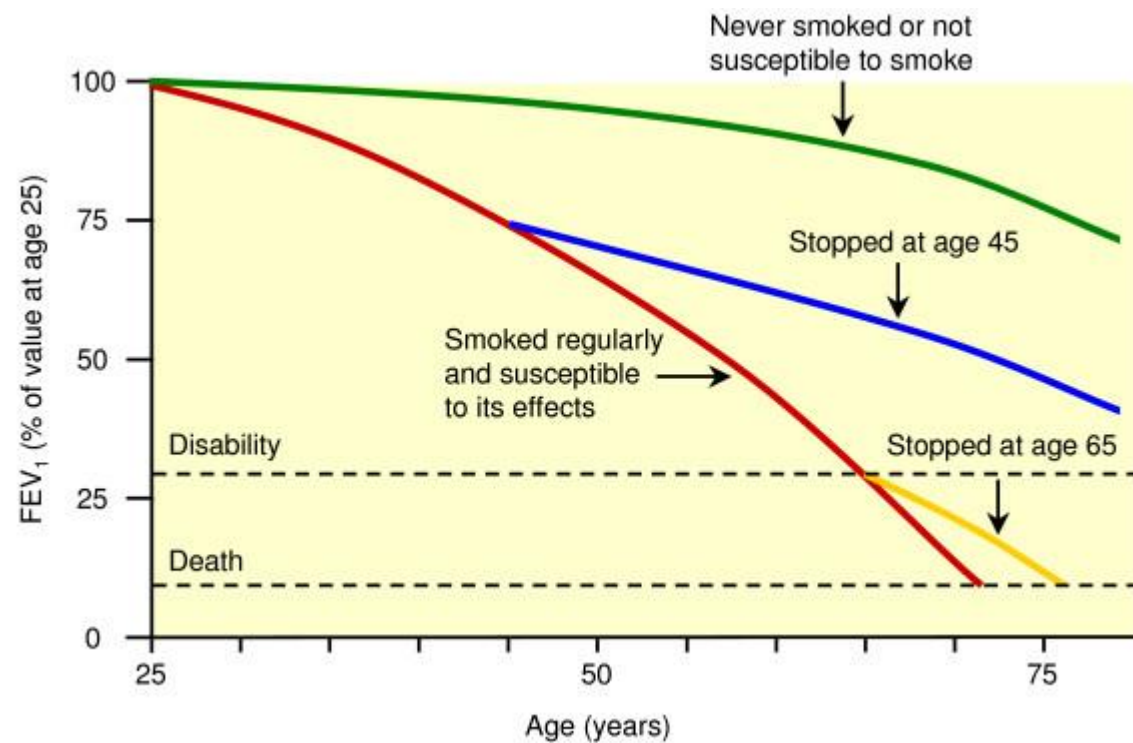
## ¿Qué nos espera en la EPOC en los próximos 10 años?

### 1. Considerar el eje temporal: la EPOC no empieza a los 65 años

- Mas allá del tabaco
- GETomics
- Trayectoma
- Plasticidad individual

### 2. Abordar la complejidad biológica de la EPOC en la práctica clínica

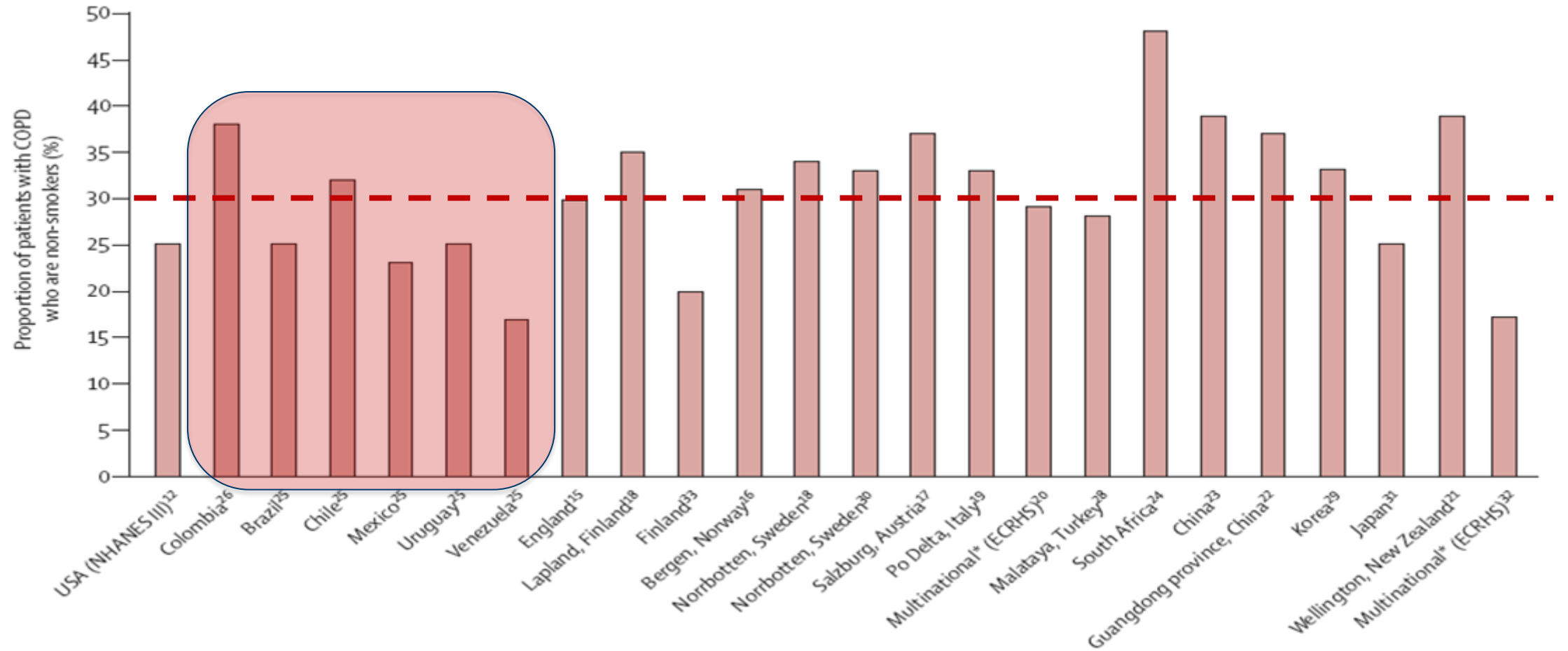
# EPOC: paradigma tradicional



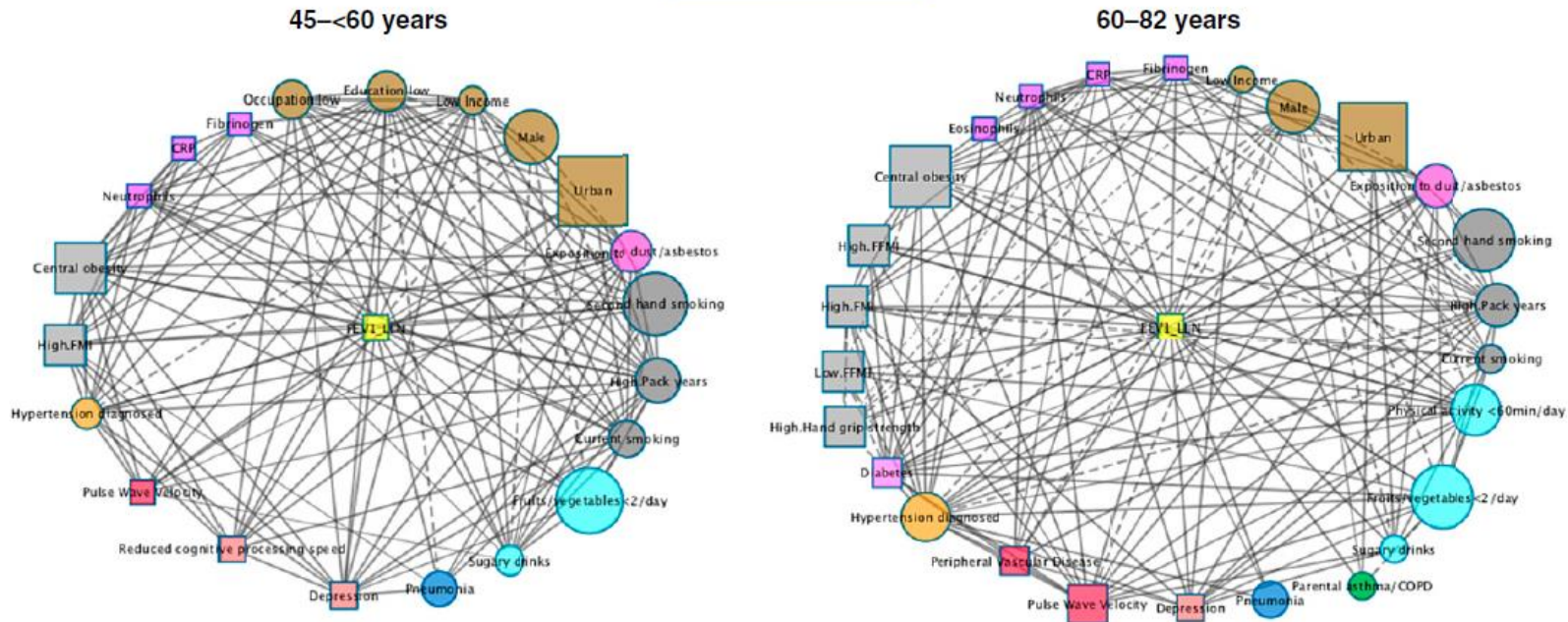
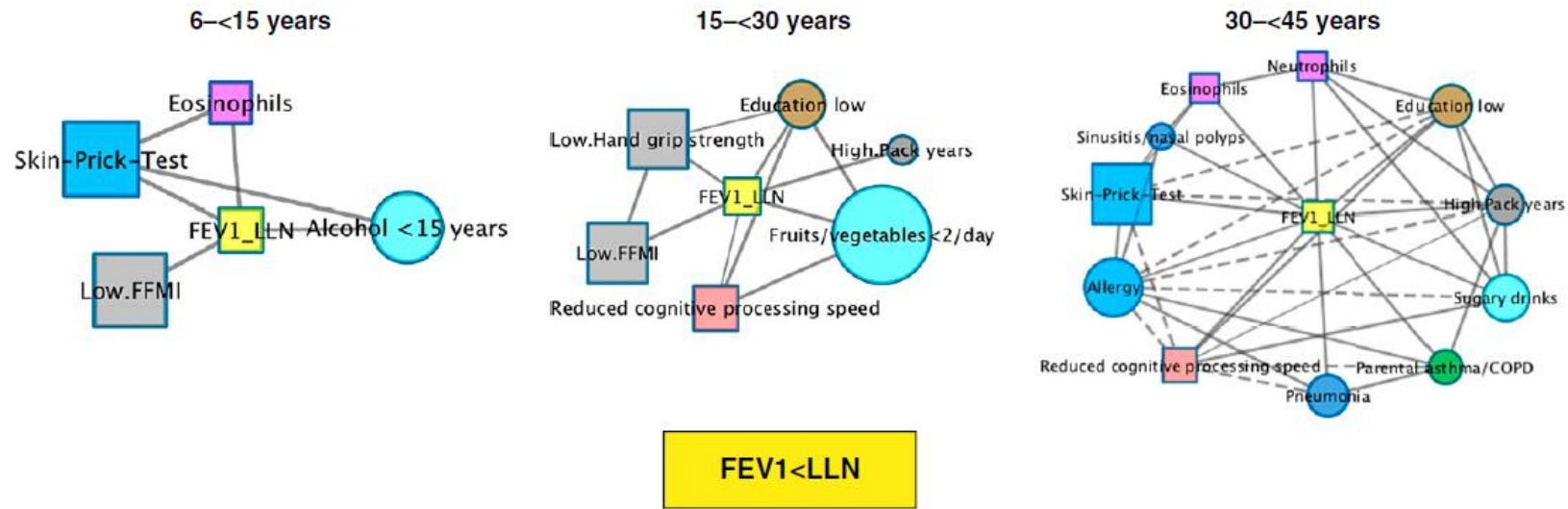
CHARLES FLETCHER, RICHARD PETO

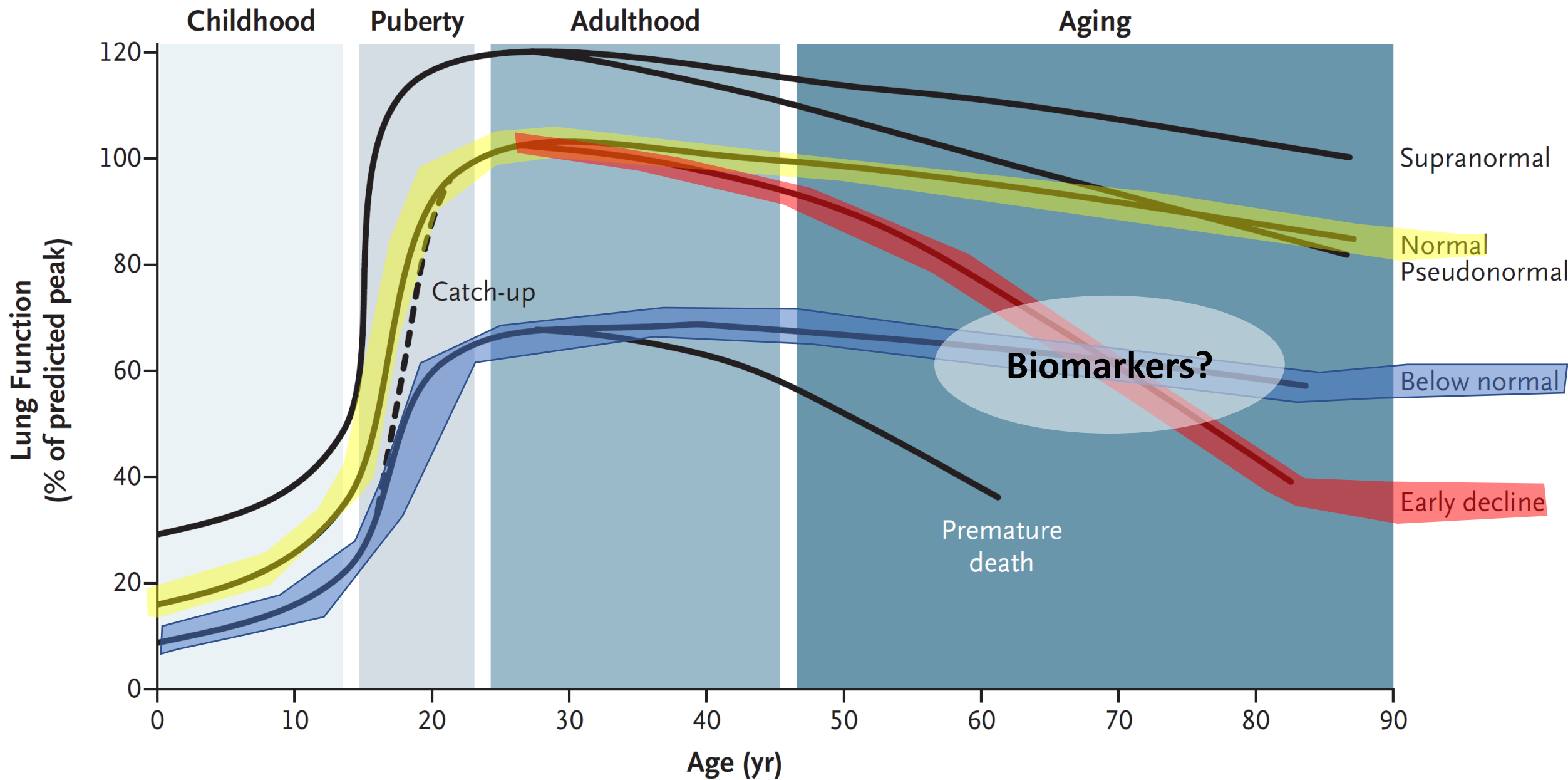
*British Medical Journal*, 1977, 1, 1645-1648

# EPOC: mas allá del tabaco

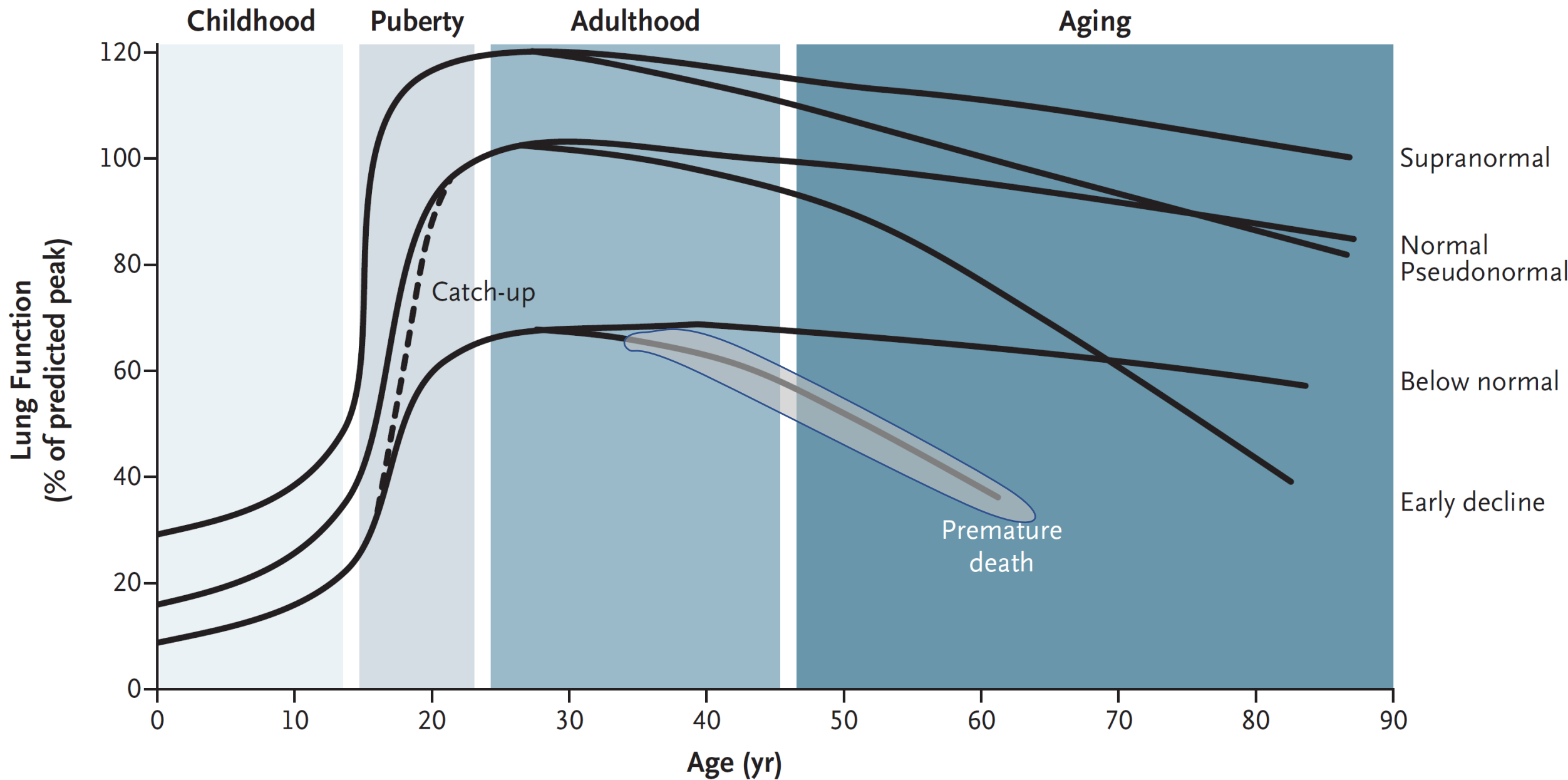








# The trajectome



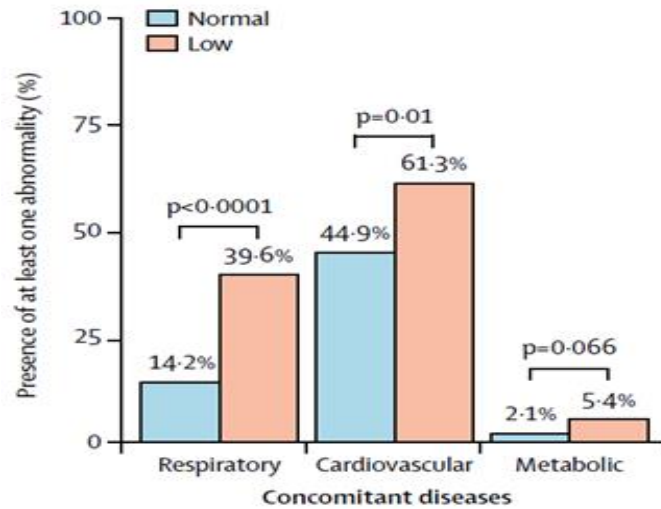
# The trajectome

# Lung function in early adulthood and health in later life: a transgenerational cohort analysis

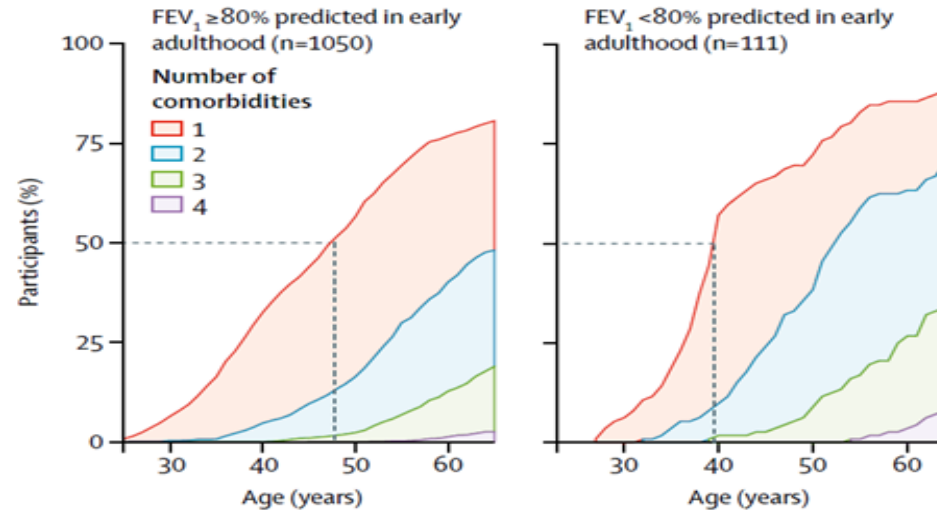


Alvar Agustí\*, Guillaume Noell\*, Josep Brugada, Rosa Faner

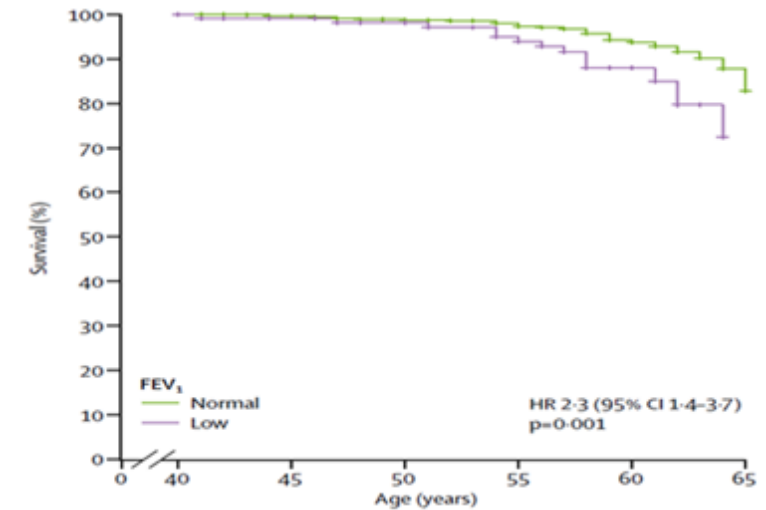
## Prevalence

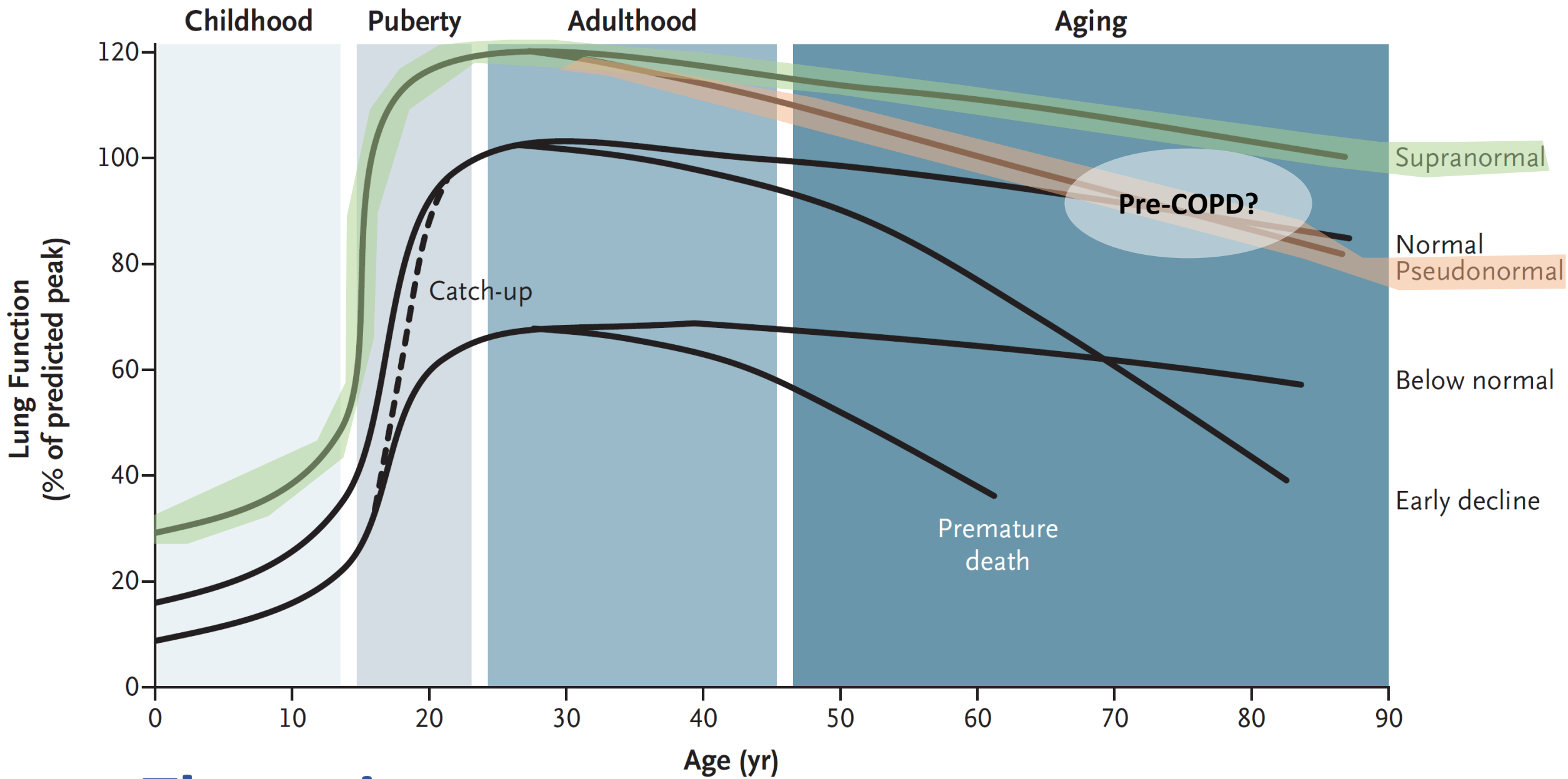


## Incidence

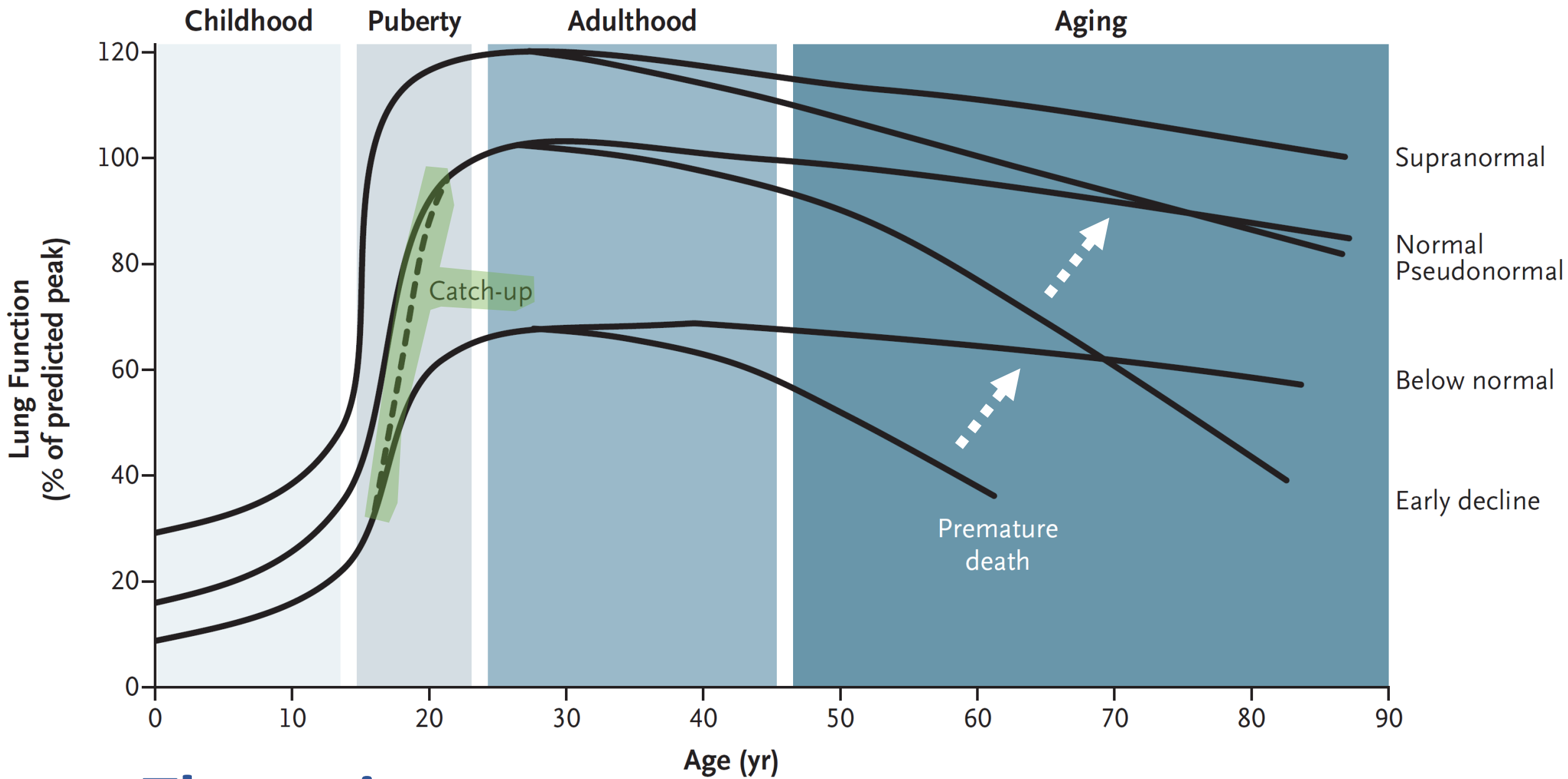


## Mortality

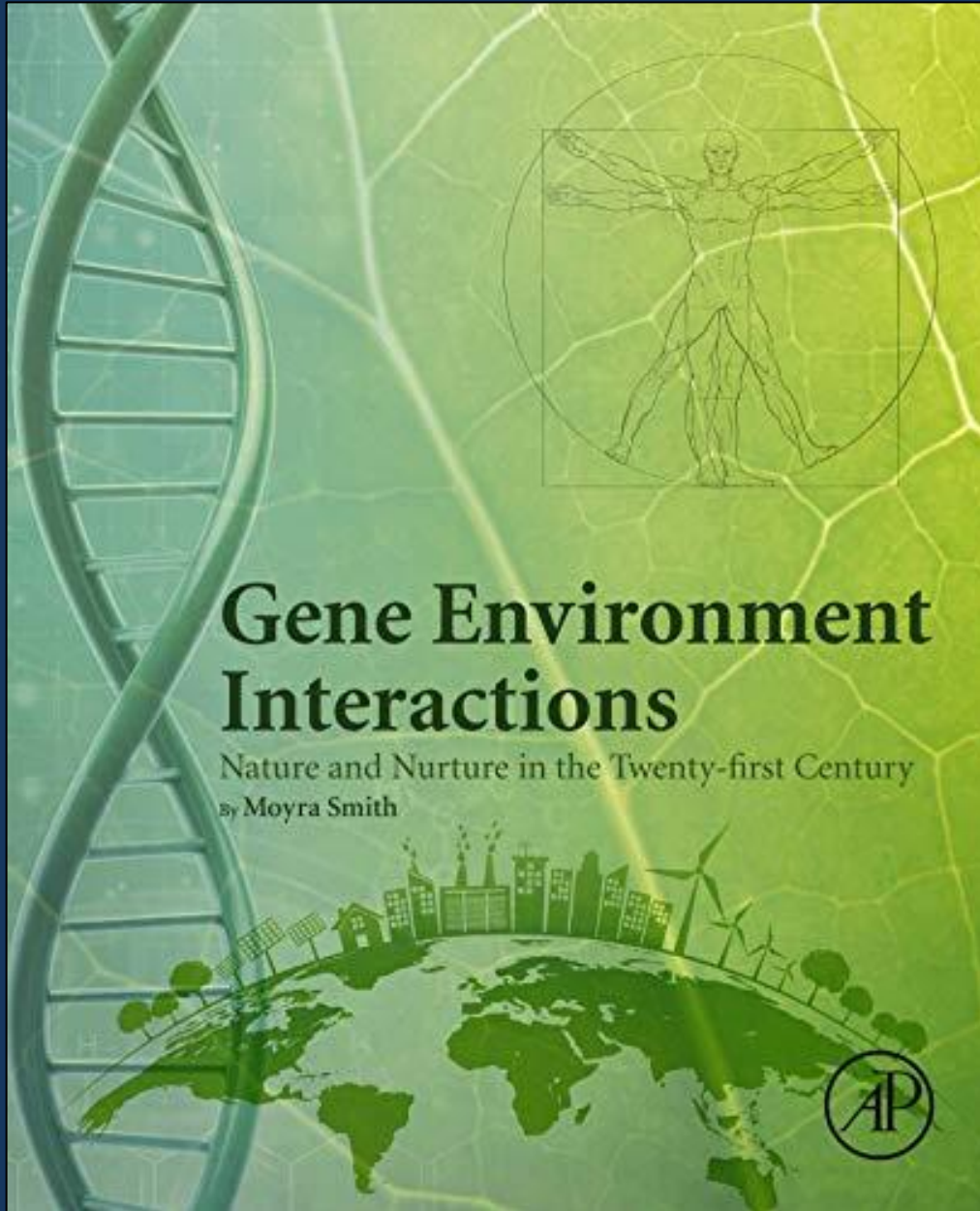




# The trajectome



# The trajectome



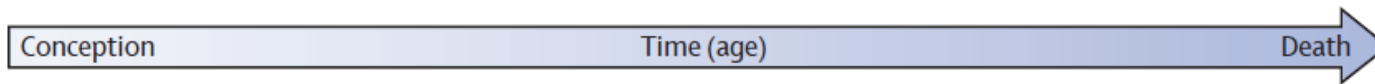
**GxE**



**GxExT**



**GETomics**





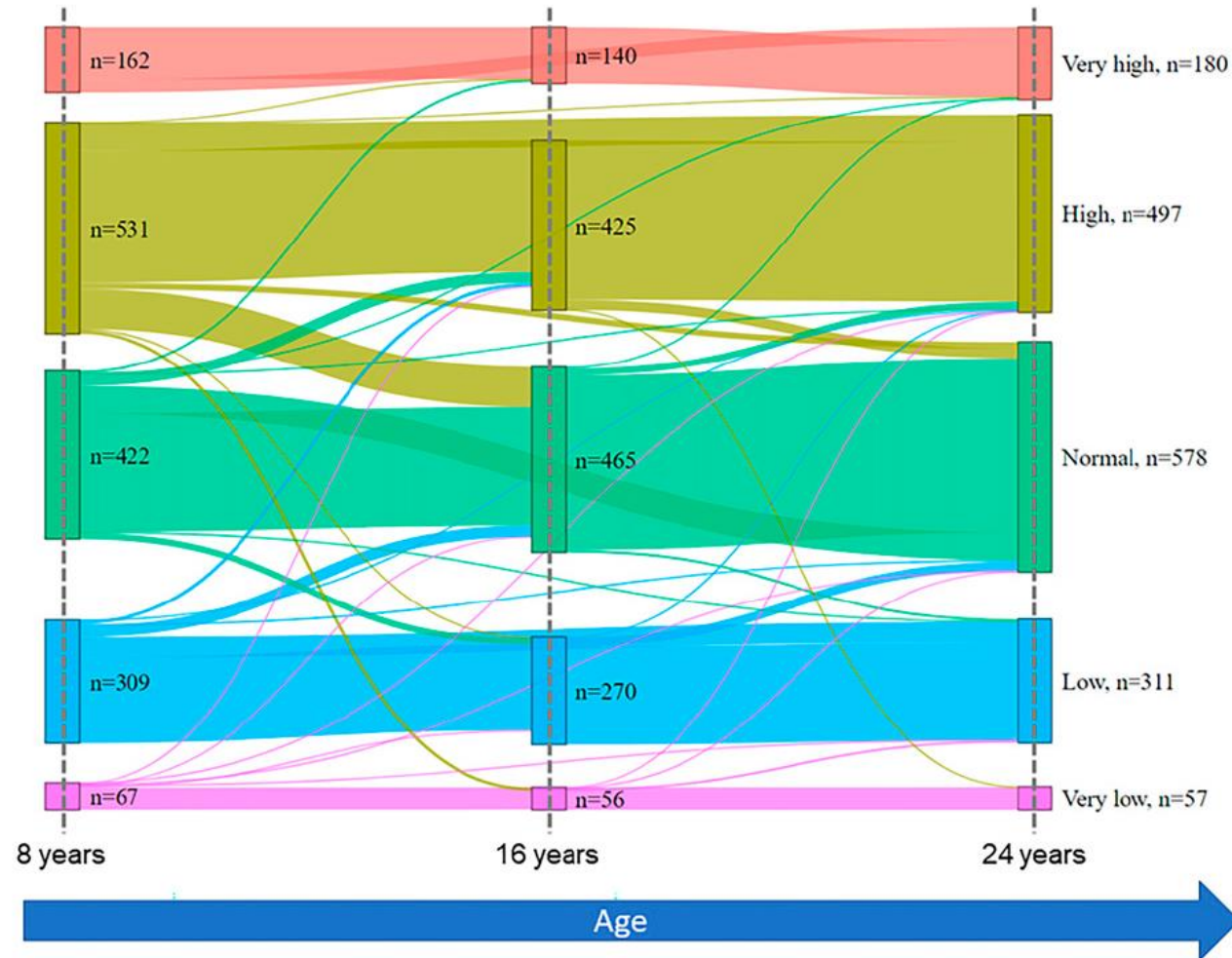
# A GETomics approach to human health and disease

The final outcome (health/disease) of any GxE interaction depends on:

1. The age of the individual at which that interaction occurs (pregnancy, infancy, adolescence, adulthood, elderly) because organs are at different developmental/ageing stages with different potential to respond.
2. The previous history of GxE interactions and the biological response they elicited (**biological memory**): epigenetics & immune response.

# Plasticity of Individual Lung Function States from Childhood to Adulthood

⑧ Gang Wang<sup>1,2,3,4</sup>, Jenny Hallberg<sup>2,5</sup>, Rosa Faner<sup>6,7</sup>, Hans-Jacob Koefoed<sup>8</sup>, Simon Kebede Merid<sup>2</sup>, Susanna Klevebro<sup>2,5</sup>, Sophia Björkander<sup>2</sup>, Olena Gruzieva<sup>3,9</sup>, Göran Pershagen<sup>3,9</sup>, Marianne van Hage<sup>10</sup>, Stefano Guerra<sup>11,12</sup>, Matteo Bottai<sup>13</sup>, Antonios Georgelis<sup>9</sup>, Ulrike Gehring<sup>14</sup>, Anna Bergström<sup>3,9</sup>, Judith M. Vonk<sup>8,15</sup>, Inger Kull<sup>2,5</sup>, Gerard H. Koppelman<sup>8,16</sup>, Alvar Agusti<sup>6,7,17,18\*</sup>, and Erik Melén<sup>2,5\*</sup>



- **Catch-up: 14.5%**
- **Growth failure: 2.4%**
  
- **IL-6 and CXCL10: negatively associated with impaired lung function development.**

TIME  
MATTERS



**THE EARLIER**

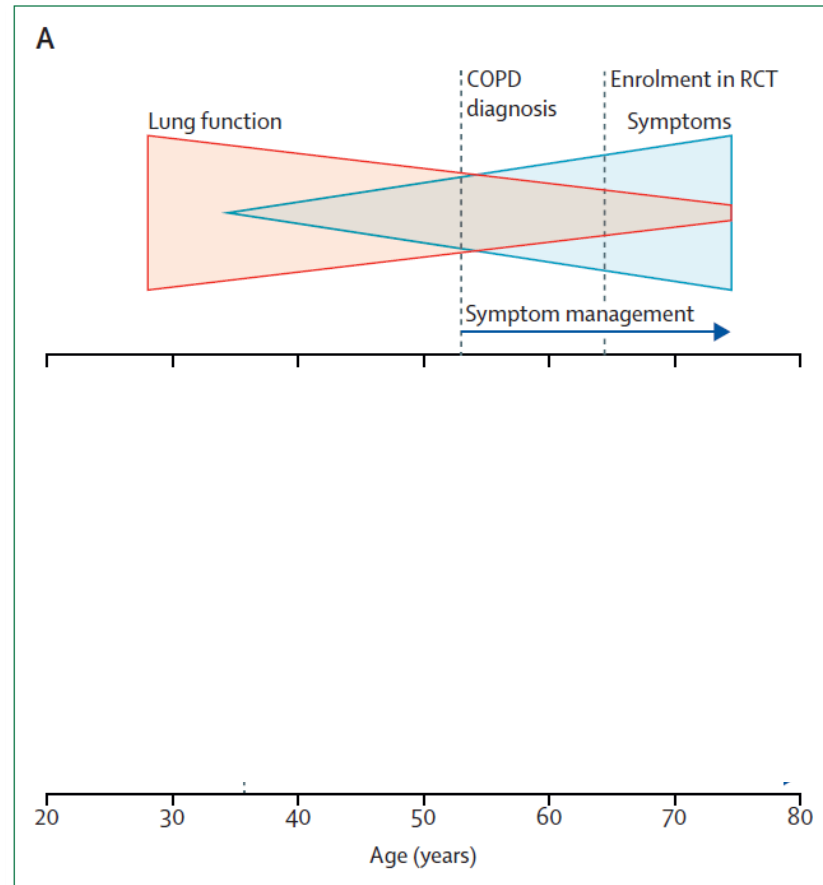
the better

# Towards the elimination of chronic obstructive pulmonary disease: a *Lancet* Commission



*Daiana Stolz, Takudzwa Mkorombindo, Desiree M Schumann, Alvar Agusti, Samuel Y Ash, Mona Bafadhel, Chunxue Bai, James D Chalmers, Gerard J Criner, Shyamali C Dharmage, Frits M E Franssen, Urs Frey, Mei Lan Han, Nadia N Hansel, Nathaniel M Hawkins, Ravi Kalhan, Melanie Konigshoff, Fanny W Ko, Trisha M Parekh, Pippa Powell, Maureen Rutten-van Mölken, Jodie Simpson, Don D Sin, Yuanlin Song, Bela Suki, Thierry Troosters, George R Washko, Tobias Welte, Mark T Dransfield*

Current





# ¿Qué nos espera en la EPOC en los próximos 10 años?

## 1. Considerar el eje temporal: la EPOC no empieza a los 65 años

- Mas allá del tabaco
- GETomics
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## 2. Abordar la complejidad biológica de la EPOC en la práctica clínica

- Rasgos Tratables
- Biomarcadores
- Tratamientos biológicos

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ORIGINAL ARTICLE



## Treatable traits in the NOVELTY study

Alvar Agustí<sup>1,2,3,4</sup>  | Eleni Rapsomaniki<sup>5</sup> | Richard Beasley<sup>6</sup>  | Rod Hughes<sup>7</sup> |  
Hana Müllerová<sup>8</sup> | Alberto Papi<sup>9,10</sup>  | Ian D. Pavord<sup>11</sup>  | Maarten van den Berge<sup>12</sup> |  
Rosa Faner<sup>3,4</sup>  | for the NOVELTY Study Investigators

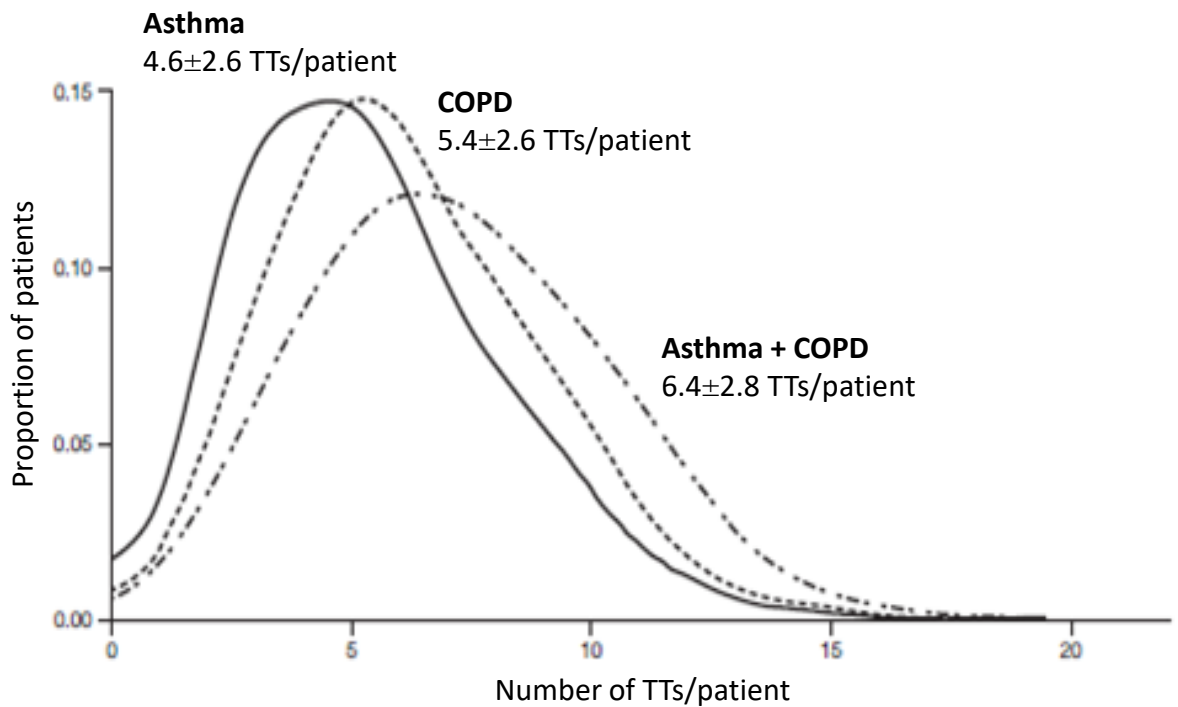
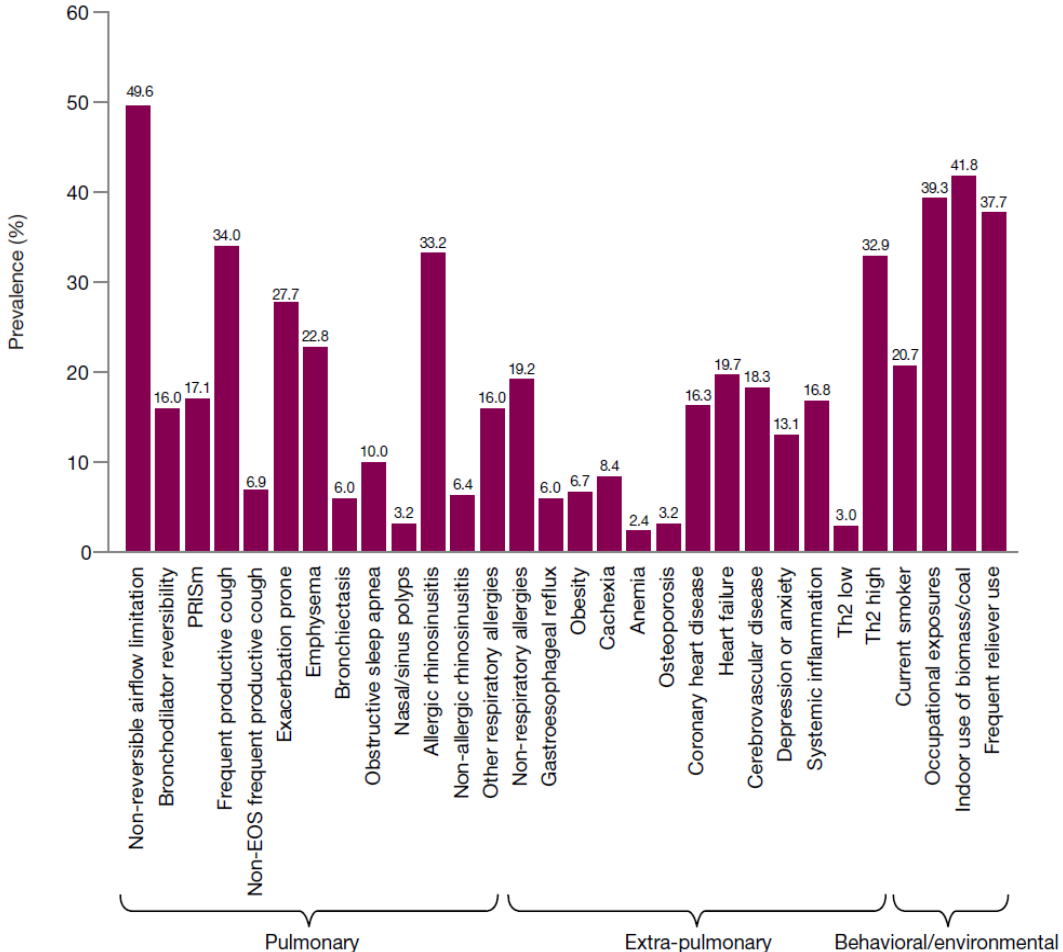
*Respirology*. 2022;1–12.

**11,226 participants**: real-life global study:

- “Asthma” (n=5.932, 52.7%)
- “COPD” (n=3.898, 34.6%)
- “asthma+COPD” (n=1.396, 12.4%)



# Prevalence of the 30 studied TTs in the NOVELTY COHORT



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EDITORIAL



WILEY

# CT in COPD: To be or not to be

Alvar Agusti MD, PhD<sup>1,2,3,4</sup> 

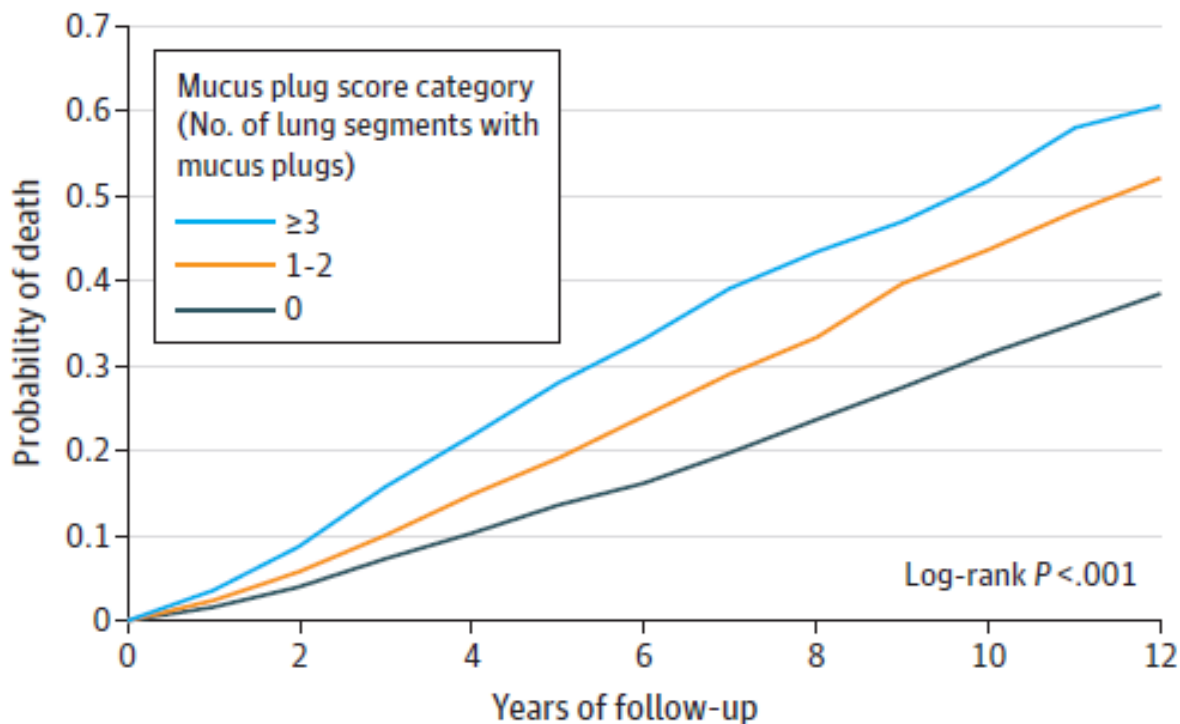
Rosa Faner PhD<sup>2,3,4</sup> 

*Respirology*. 2022;27:258–259.

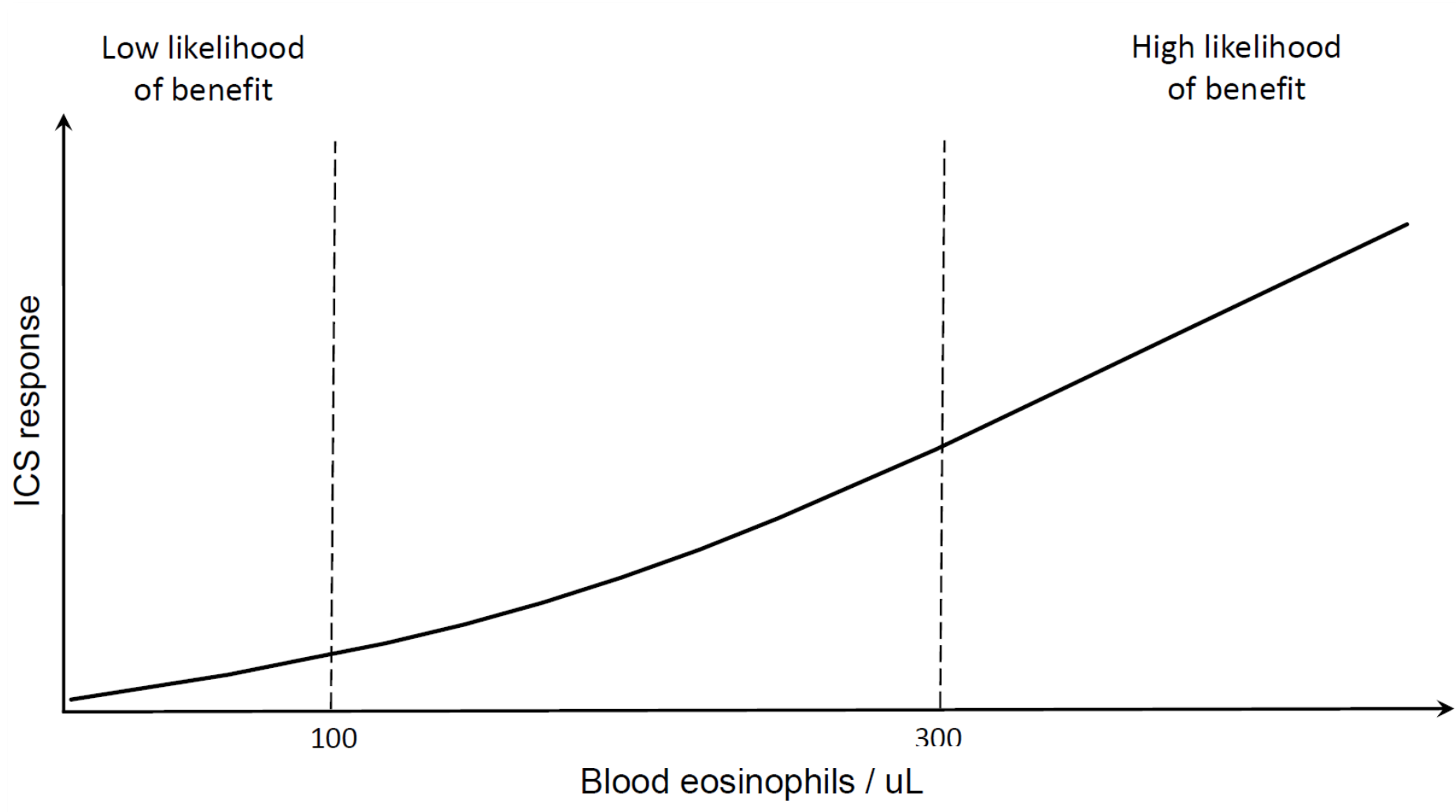
# Airway-Occluding Mucus Plugs and Mortality in Patients With Chronic Obstructive Pulmonary Disease

Alejandro A. Diaz, MD, MPH; José L. Orejas, MD; Scott Grumley, MD; Hrudaya P. Nath, MD; Wei Wang, PhD; Wojciech R. Dolliver, MD; Andrew Yen, MD; Seth J. Kligerman, MD; Kathleen Jacobs, MD; Padma P. Manapragada, MD; Mostafa Abozeed, MD, MSc, PhD; Muhammad Usman Aziz, MD; Mohd Zahid, MD; Asmaa N. Ahmed, MD, MBBCh, MSc; Nina L. Terry, MD; Ruben San José Estépar, MSc; Victor Kim, MD; Barry J. Make, MD; MeiLan K. Han, MD; Sushilkumar Sonavane, MD; George R. Washko, MD, MSc; Michael Cho, MD, MPH; Raúl San José Estépar, PhD

**A** Unadjusted probability of death by mucus plug score



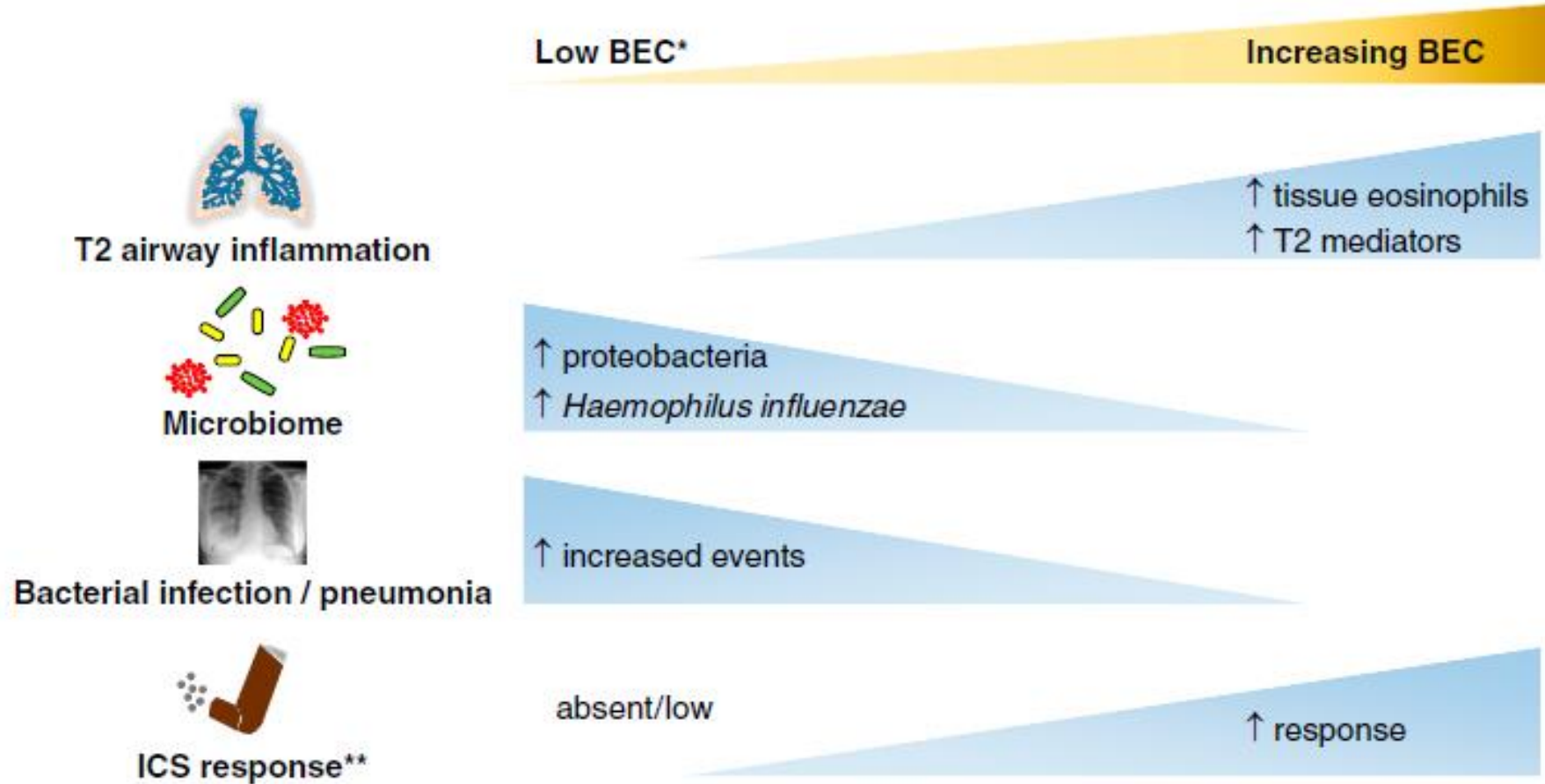
# Relationship between blood eosinophils and the efficacy of ICS in preventing exacerbations



# Blood Eosinophils and Chronic Obstructive Pulmonary Disease

A Global Initiative for Chronic Obstructive Lung Disease Science Committee  
2022 Review

Dave Singh<sup>1</sup>, Alvar Agusti<sup>2</sup>, Fernando J. Martinez<sup>3</sup>, Alberto Papi<sup>4</sup>, Ian D. Pavord<sup>5</sup>, Jadwiga A. Wedzicha<sup>6</sup>,  
Claus F. Vogelmeier<sup>7</sup>, and David M. G. Halpin<sup>8</sup>



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**Biologics for COPD — Finally Here**

Alvar Agusti, M.D., Ph.D.

N ENGL J MED 389;3 NEJM.ORG JULY 20, 2023



**Muchas gracias por su atención**

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