

**XXVII Congresso Italiano della  
Fibrosi Cistica**

# **Steroide inalatorio : da proscrivere o da prescrivere? PRO**

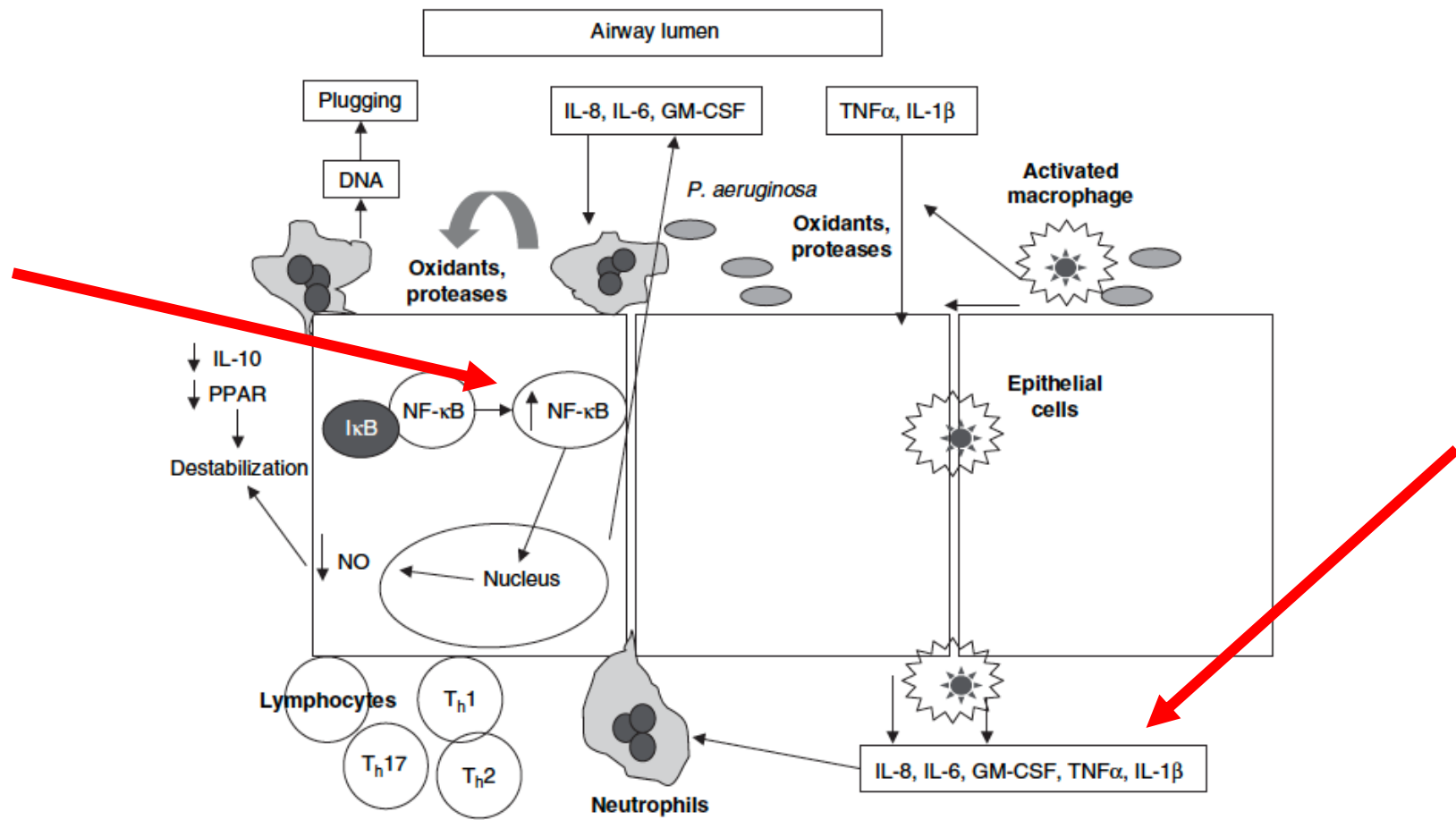
**Francesco Blasi, MD, FERS**

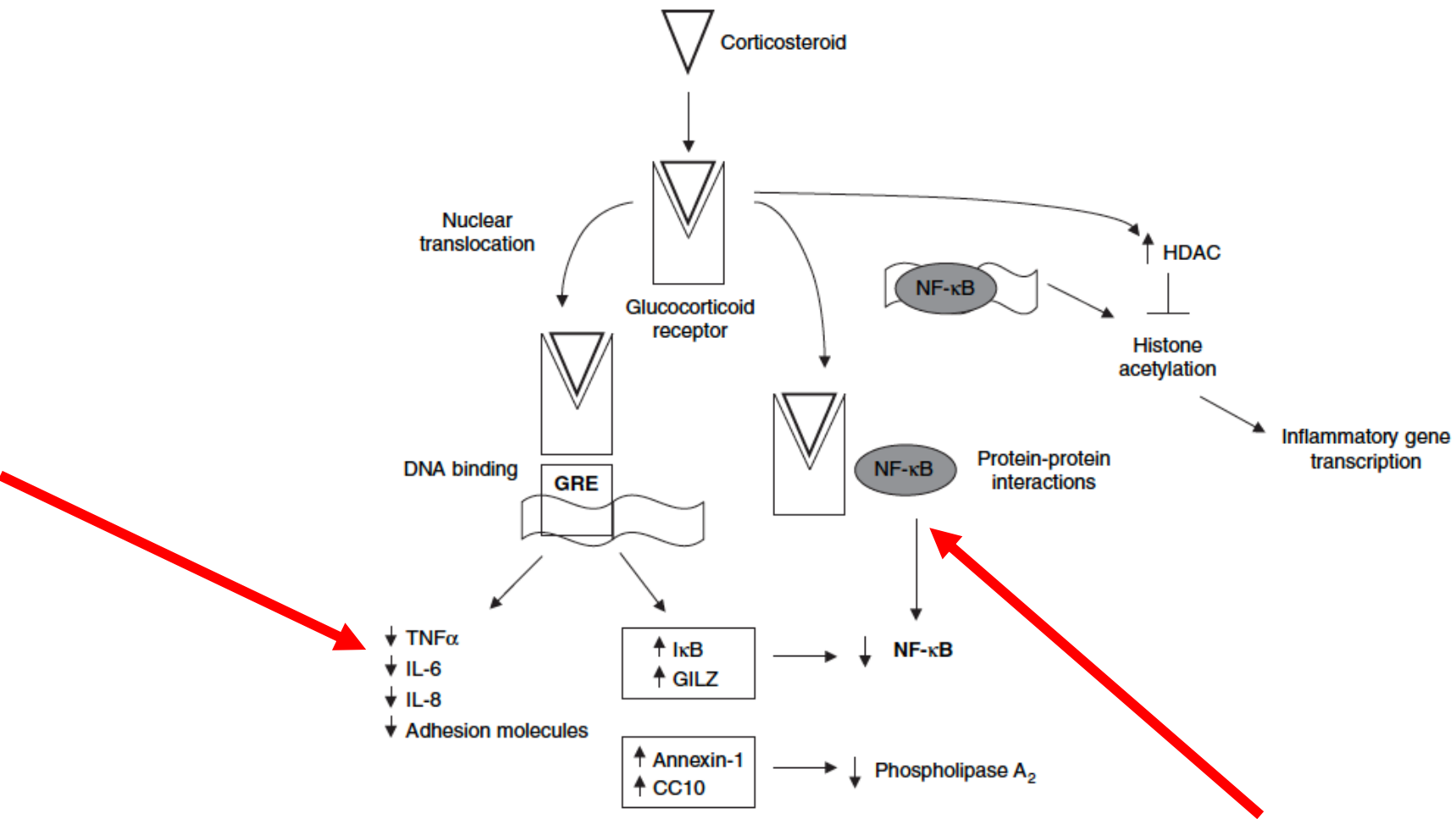
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Fondazione IRCCS Cà Granda Milan, Italy**

## PER COMINCIARE.....

- We conclude that the prescribing practice for an individual with CF should become more like that for an asthmatic and could be restricted to those with symptomatic recurrent wheezing that is not responsive to bronchodilators alone and in whom benefit has been proven.
- Justification is needed to start ICS, reassessment is necessary to see whether they are having an effect (particularly on any tight cough or wheeze), and consideration is always given to reducing the dose or stopping the drugs altogether.
- **It is likely though that the majority of people with CF taking ICS no longer need to do so**
- This view has been reflected in the North American consensus document that states "...for patients with CF, 6 years and older, and without asthma or ABPA, the Cystic Fibrosis Foundation recommends against the routine use of inhaled corticosteroids to improve lung function and to reduce exacerbations





# Inhaled corticosteroids and lower lung function decline in young children with cystic fibrosis

K. De Boeck\*, F. Vermeulen\*, S. Wanyama<sup>#</sup> and M. Thomas<sup>#</sup>, on behalf of the members of the Belgian CF Registry

**TABLE 1** Yearly change in forced expiratory volume in 1 s (FEV<sub>1</sub>) and baseline FEV<sub>1</sub> in years with and without inhaled corticosteroid (ICS) use

Age range	ICS use	Subjects n	Years with ICS use %	Yearly FEV <sub>1</sub> change % pred			Baseline FEV <sub>1</sub> % pred		
				Change	Difference	p-value	Baseline	Difference	p-value
<b>6–12 yrs</b>	No ICS	549	37.6	-1.37±10.2	2.56	0.0003	93.1±15.9	0.01	0.9921
	ICS	331		1.19±10.0			93.1±15.4		
<b>13–17 yrs</b>	No ICS	331	45.5	-1.71±9.34	0.69	0.3624	84.1±19.5	4.43	0.0152
	ICS	276		-1.02±9.39			79.6±21.5		
<b>≥18 yrs</b>	No ICS	758	49.1	-0.90±8.21	0.46	0.2487	66.7±21.2	5.05	<0.0001
	ICS	731		-0.43±7.26			61.6±21.1		
<b>Overall</b>	No ICS	1638	44.9	-1.22±9.17	1.07	0.001	79.1±22.6	5.89	<0.0001
	ICS	1338		-0.15±8.50			73.2±24.0		

Data are presented as mean±SD, unless otherwise stated. Global analysis and analysis in age categories 6–12, 13–17 and ≥18 yrs. % pred: % predicted.

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**In conclusion, analysis of Belgian CF registry data confirms the findings in the American Registry that ICS use is associated with a decrease in lung function decline in children 6–12 yrs of age.**

**A positive effect from ICS treatment in these children is possible, especially in the face of the knowledge that inflammation in the lung starts early**

## Original Articles —

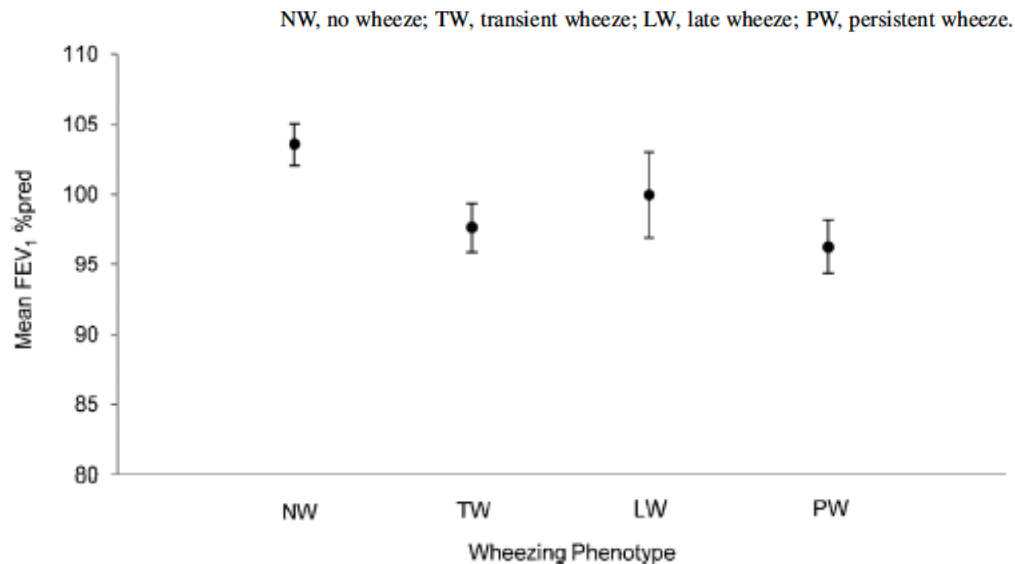
**Epidemiologic Study of Cystic Fibrosis:** Design and  
Implementation of a Prospective, Multicenter, Observational  
Study of Patients With Cystic Fibrosis in the U.S.  
and Canada

**TABLE 2—Common Medical Conditions in All Subjects, in Subjects Less Than 18 Years of Age, and in Subjects 18 Years of Age or Older<sup>1</sup>**

Condition	All ages		<18 years		18+ years	
	N	%	N	%	N	%
Asthma	3,416	19	2,403	19	1,013	18
Sinusitis	3,084	17	1,922	15	1,162	21
Nasal polypsis	1,742	10	1,178	9	564	10
Pharyngitis	677	4	429	3	248	5
Elevated liver enzymes	558	3	397	3	161	3
Cirrhosis	209	1	129	1	80	1
ABPA <sup>2</sup>	285	2	169	1	116	2

## Early Childhood Wheezing Is Associated With Lower Lung Function in Cystic Fibrosis

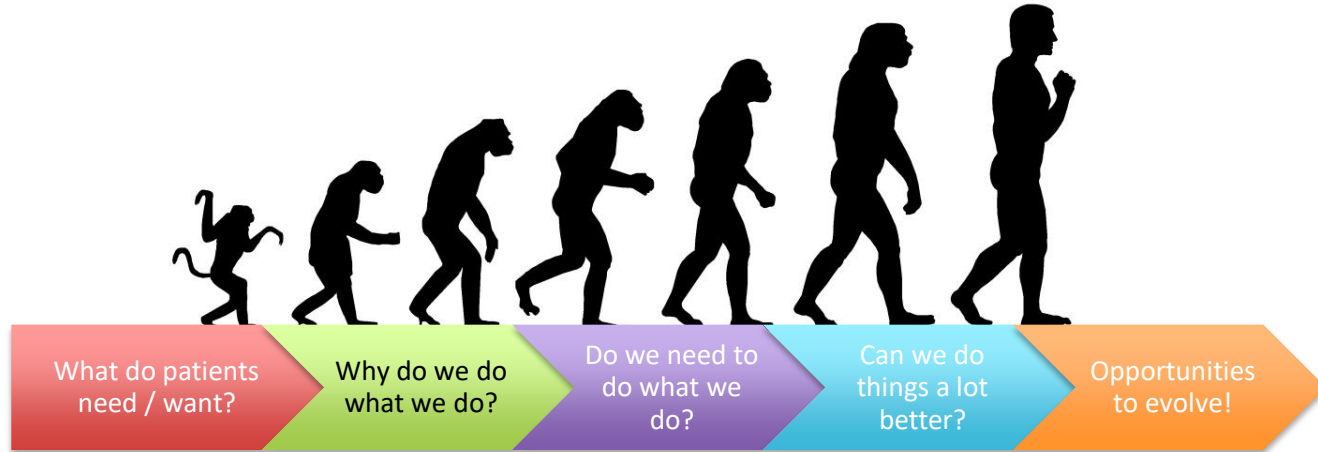
Clement L. Ren, MD,<sup>1\*</sup> Michael W. Konstan, MD,<sup>2</sup> Margaret Rosenfeld, MD, MPH,<sup>3</sup> David J. Pasta, MS,<sup>4</sup> Stefanie J. Millar, MS,<sup>4</sup> Wayne J. Morgan, MD, CM,<sup>5</sup> for the Investigators and Coordinators of the Epidemiologic Study of Cystic Fibrosis



**Fig. 1. Adjusted mean FEV1 % predicted at age 6 to <8 years of age by wheezing phenotype (N = 1,302). Error bars indicate 95% confidence intervals.**



# The COVID Questions





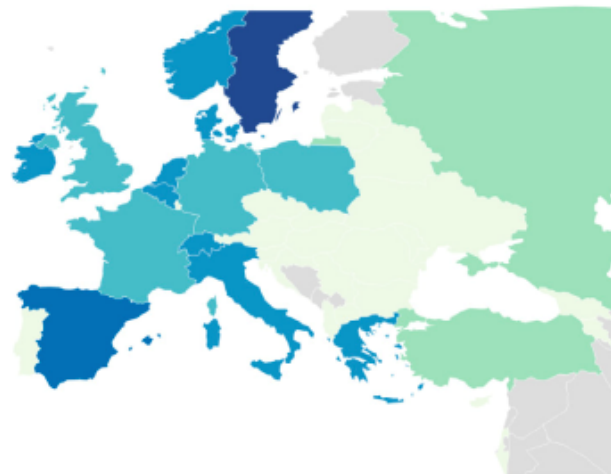
Incidence of SARS-CoV-2 in people with cystic fibrosis in Europe between February and June 2020



## SARS-CoV-2 incidence (per 10000) in the 38 ECFSPR countries

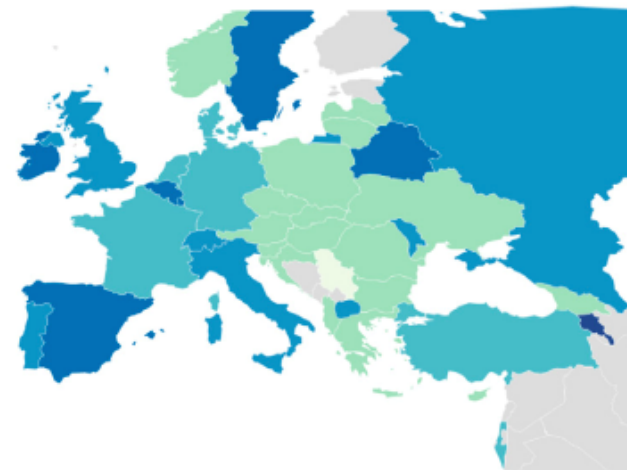
### Cystic fibrosis population

Incidence per 1000



### General population

Incidence per 1000

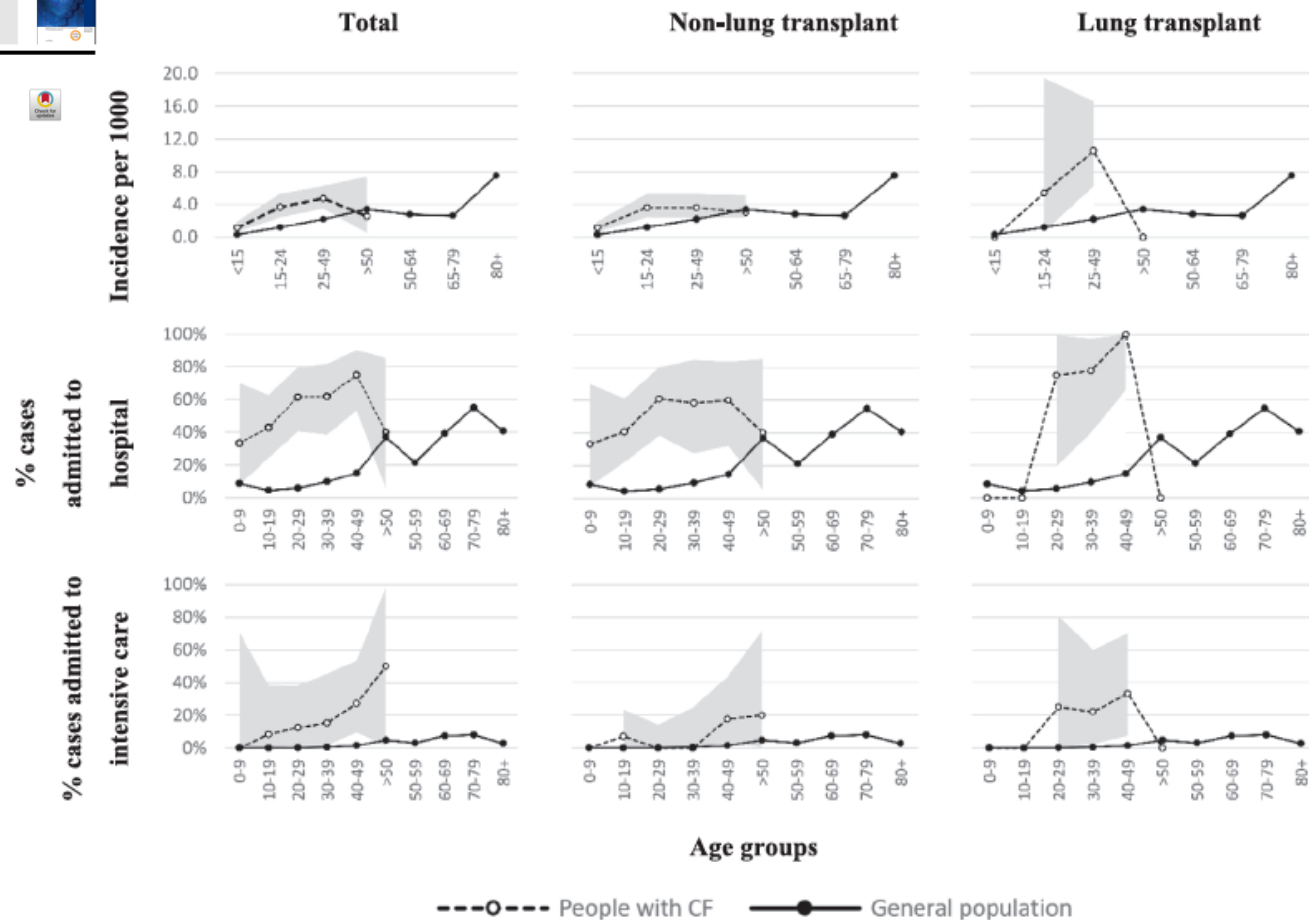


Cases	Population	Incidence (95% CI)
130	48211	2.70 (2.25 - 3.20)

Cases	Population	Incidence (95% CI)
2582924	832750755	3.10 (3.10 – 3.11)



Incidence of SARS-CoV-2 in people with cystic fibrosis in Europe between February and June 2020





# Outcomes of SARS-CoV-2 infection in patients with cystic fibrosis: A multicenter retrospective research network study

Yousaf B. Hadi <sup>a,1</sup>, Dhairya A. Lakhani <sup>b,1</sup>, Syeda F. Naqvi <sup>a</sup>, Nida Ul Fatima <sup>a</sup>, Arif R. Sarwari <sup>a,c,\*</sup>

## Outcomes in the two cohorts of COVID-19 patients with and without cystic fibrosis before and after propensity score matching.

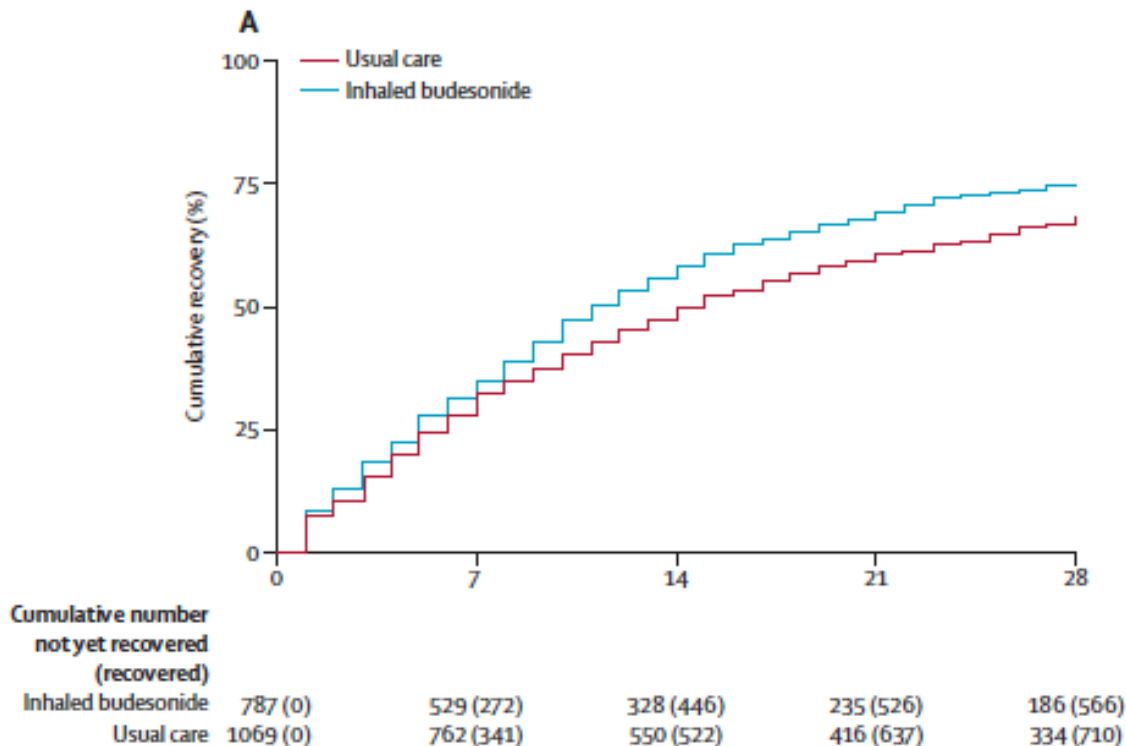
Outcome	Cystic Fibrosis cohort (n = 422)	Percentage	Non-CF cohort (n = 507,388)	Percentage	Risk Ratio
<b>Before propensity score matching</b>					
30-day Mortality	22	5.21	8,705	1.72	3.74
Inpatient services	117	27.73	39,471	7.78	3.56
Critical Care	49	11.61	12,953	2.55	4.55
Mechanical ventilation	26	6.16	7,842	1.55	3.99
30-day composite outcome	37	8.77	13,288	2.62	3.35
Acute renal injury	60	14.22	19,646	3.87	3.67
Outcome	Cystic Fibrosis cohort (n=413)	Percentage	Non-CF cohort (n=413)	Percentage	Risk Ratio
<b>After propensity score matching</b>					
30-day Mortality	22	5.33	12	2.91	1.83
Inpatient services	111	26.88	71	17.19	1.56
Critical Care	48	11.62	27	6.54	1.78
Mechanical ventilation	26	6.30	17	4.12	1.53
30-day composite outcome	37	8.96	23	5.57	1.61
Acute renal injury	56	13.56	35	8.48	1.60

# Inhaled budesonide for COVID-19 in people at high risk of complications in the community in the UK (PRINCIPLE): a randomised, controlled, open-label, adaptive platform trial

Ly-Mee Yu\*, Mona Bafadhel\*, Jienchi Dorward\*, Gail Hayward, Benjamin R Saville, Oghenekome Gbiniigie, Oliver Van Hedde, Emma Ogburn, Philip H Evans, Nicholas P B Thomas, Mahendra G Patel, Duncan Richards, Nicholas Berry, Michelle A Detry, Christina Saunders, Mark Fitzgerald, Victoria Harris, Milena Shanyinde, Simon de Lusignan, Monique J Andersson, Peter J Barnes, Richard E K Russell, Dan V Nicolau Jr, Sanjay Ramakrishnan, F D Richard Hobbs†, Christopher C Butler†, on behalf of the PRINCIPLE Trial Collaborative Group



Lancet 2021; 398: 843-55

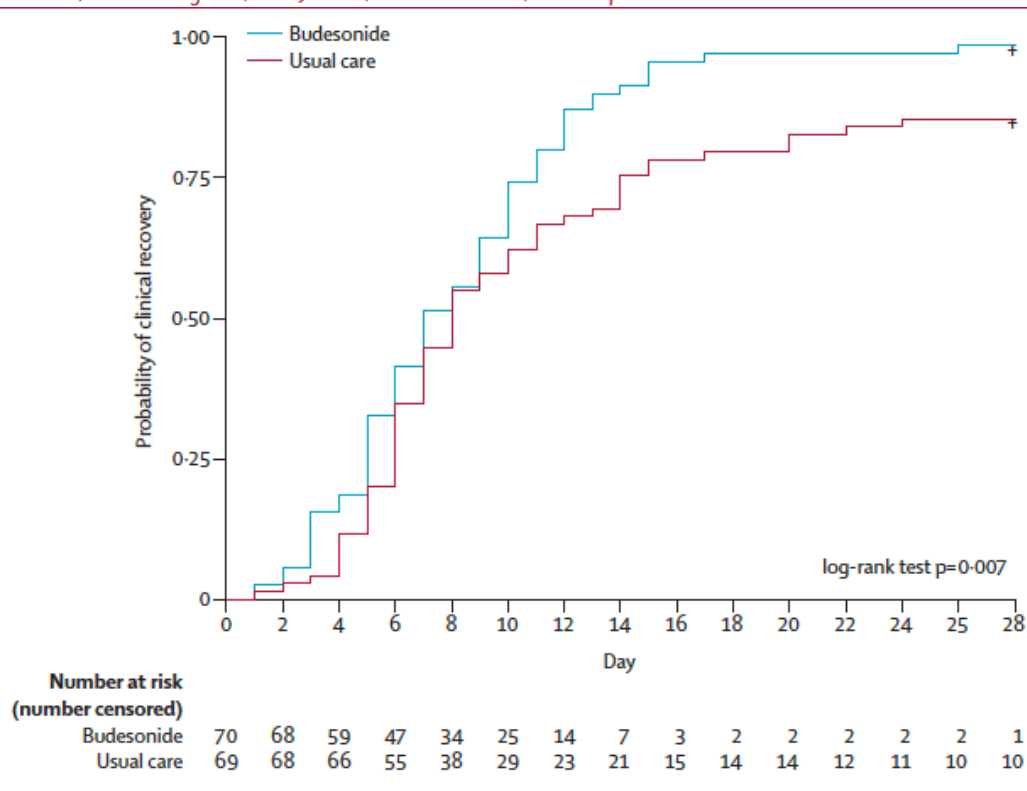


**Interpretation** Inhaled budesonide improves time to recovery, with a chance of also reducing hospital admissions or deaths (although our results did not meet the superiority threshold), in people with COVID-19 in the community who are at higher risk of complications.

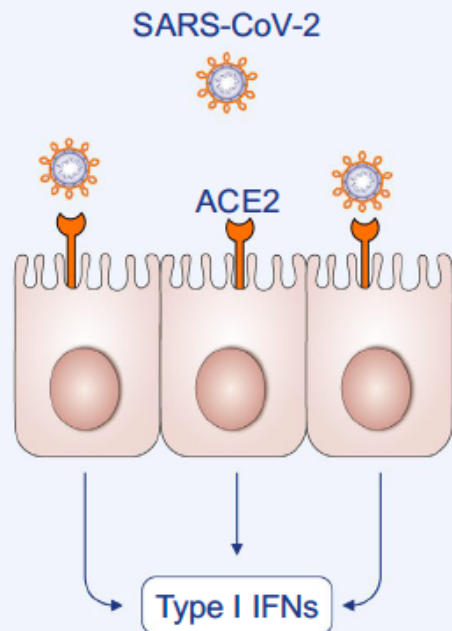
# Inhaled budesonide in the treatment of early COVID-19 (STOIC): a phase 2, open-label, randomised controlled trial

Sanjay Ramakrishnan\*, Dan V Nicolau Jr\*, Beverly Langford, Mahdi Mahdi, Helen Jeffers, Christine Mwasuku, Karolina Krassowska, Robin Fox, Ian Binnian, Victoria Glover, Stephen Bright, Christopher Butler, Jennifer L Cane, Andreas Halner, Philippa C Matthews, Louise E Donnelly, Jodie L Simpson, Jonathan R Baker, Nabil T Fadai, Stefan Peterson, Thomas Bengtsson, Peter J Barnes, Richard E K Russell, Mona Bafadhel

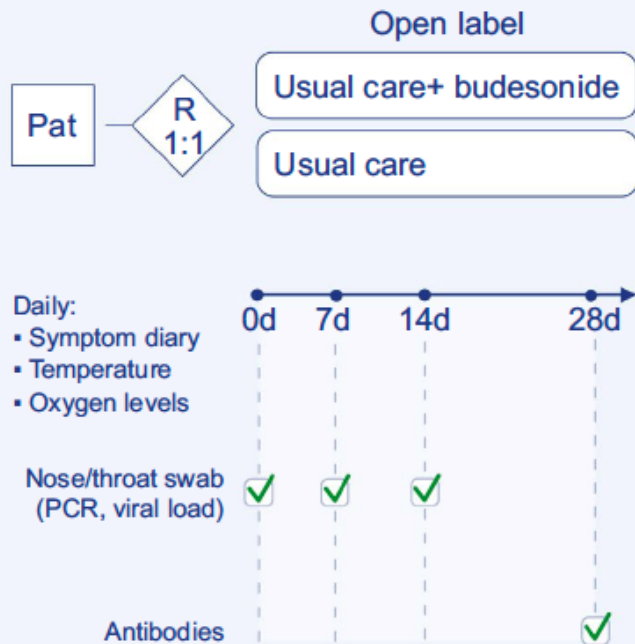
*Lancet Respir Med* 2021



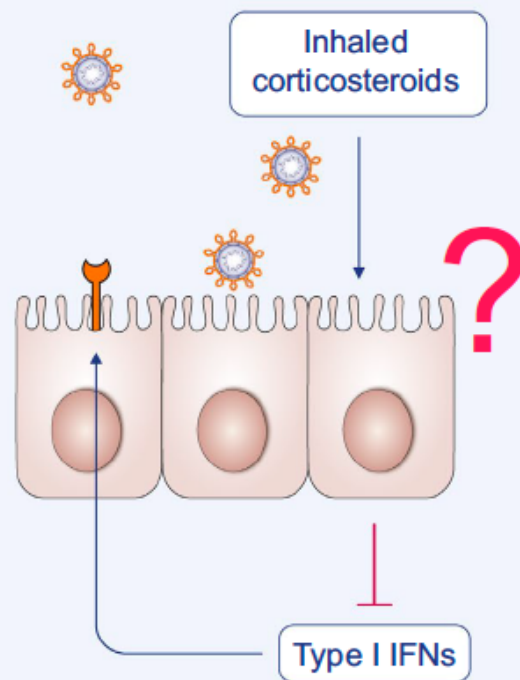
## Viral infection in healthy airways



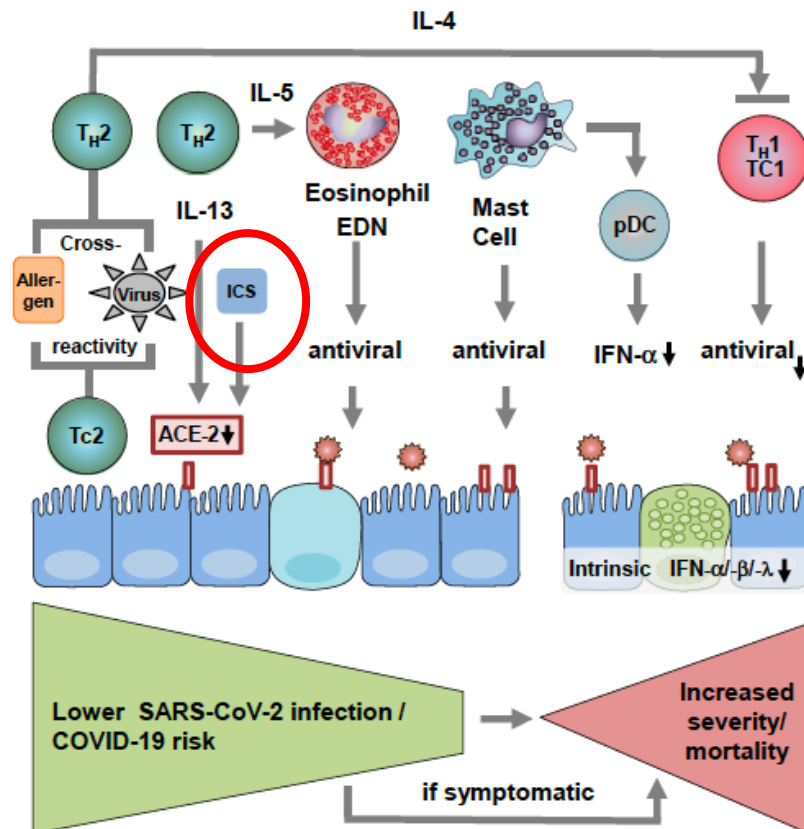
## STOIC Study design



## Reduced ACE2 expression Less severe courses?



## T2 Asthma

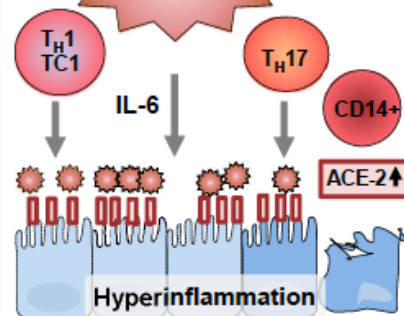


## Non-T2 Asthma

Older age  
Metabolic comorbidities

- Obesity
- T2D
- Metabolic syndrome

chronic (subclinical) inflammation



High-risk COVID-19  
(severity, lung destruction)



