



# Oxygène à haut débit dans l'IRA hypoxémique du patient immunodéprimé

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# Quelle sont les questions ?

- Les immunodéprimés sont-ils différents ?
- Chez les ID, la stratégie d’oxygénation/ventilation a-t-elle un impact sur le pronostic ?
- Faut-il encore tout faire pour éviter l’intubation dans l’IRA hypoxémique du patient immunodéprimé ?

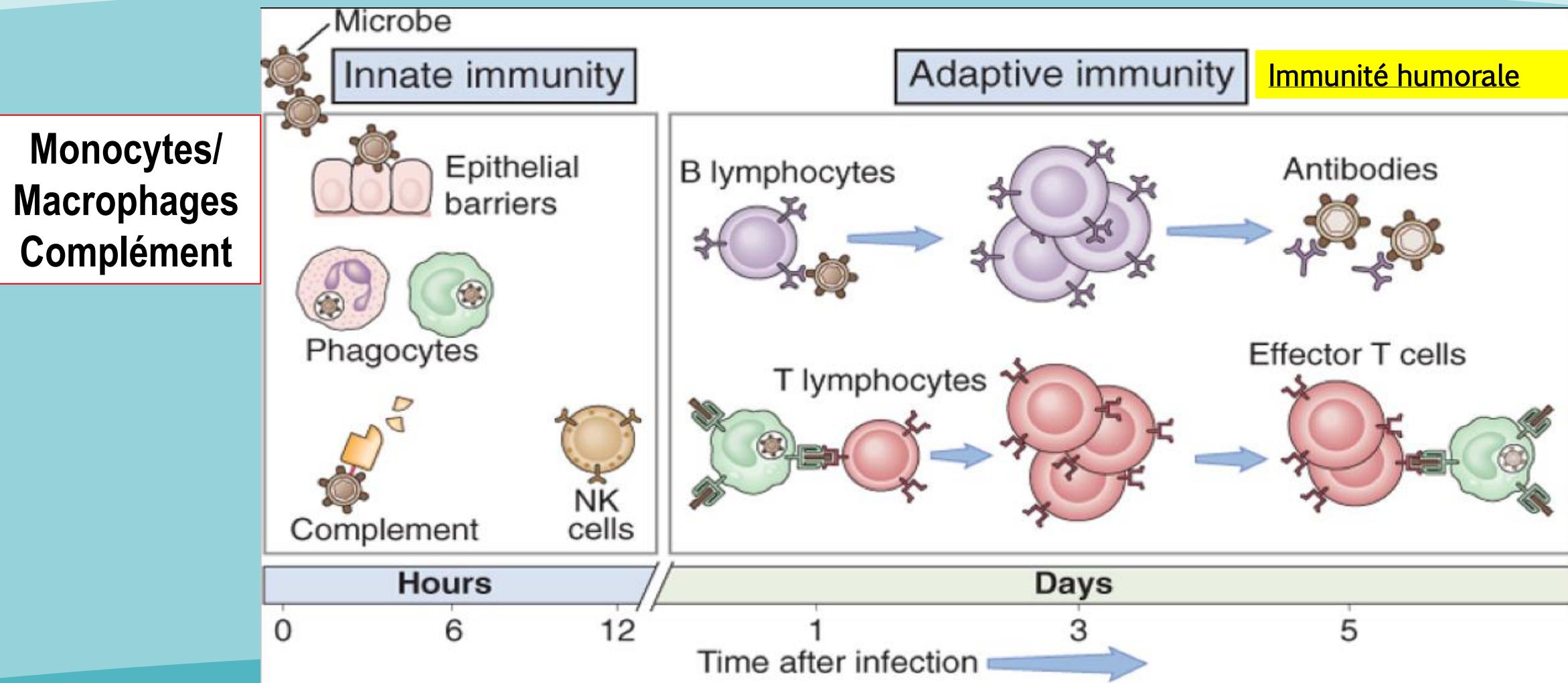
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# Clarifications

# Immunocompromized Patients with ARF: Striking Differences

	Unselected Patients	Immunocompromized patients
Incidence of respiratory events	1-5%	1-50%
Risk for severe hypoxemia	Ref	↑↑
Increased mortality if delayed intubation	Probably	Yes (NIV or HFNO failure)
Complications of MV	Similar	Similar or slightly ↑
Respiratory drive	=	=
VTE under NIV	Similar	Similar
Need for diagnostic strategy	Sometimes	Almost always
Intubation rate (if not intubated at admission)	25-30%	40%
Mortality of patients not intubated at admission	10-15%	25-35%
Reduced mortality over time	Slightly	Yes +++
Conclusive literature	No	No

# Système immunitaire



# Immunodéprimés: quo va dis?

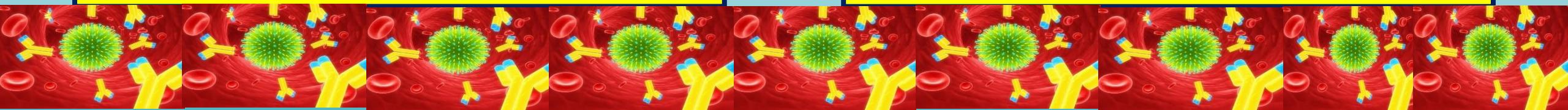
## Déficit immunitaire secondaire

- Immunodépression médicamenteuse
- Hémopathies malignes
- Tumeurs solides?
- Chimiothérapie
- Greffe d'organe
- SIDA
- Dénutrition
- Diabète
- ...

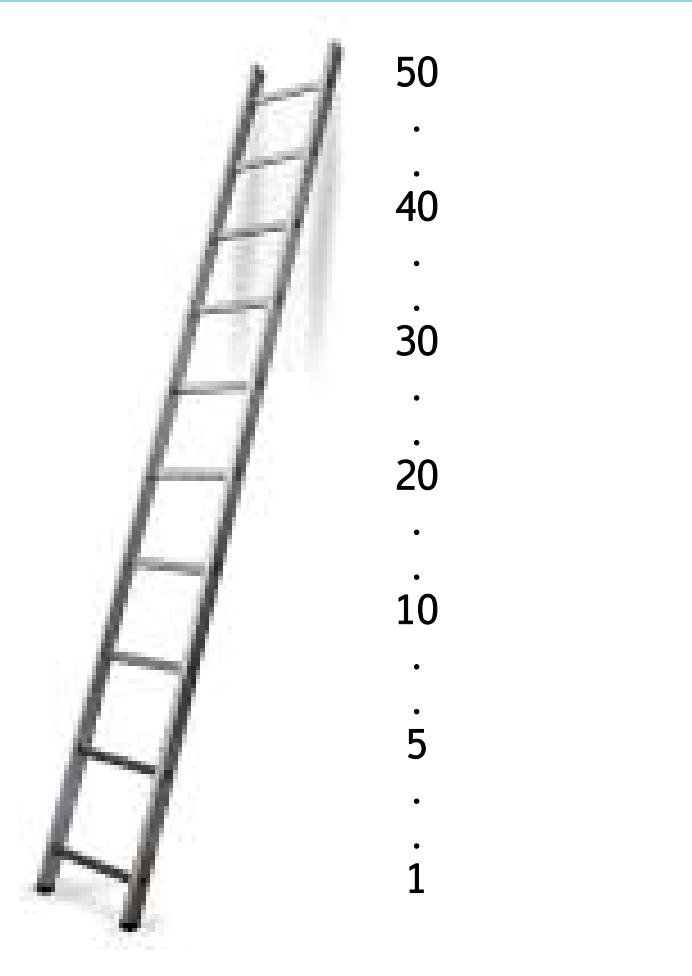
## Déficit immunitaire secondaire

### >200 maladies, 5 grands groupes.

- Les DIP humoraux: DICV les + fréquents.  
Déficit en Ig sériques, manque d'Ac
- Les DIP combinés: DICS: Altération des lymphocytes et des anticorps
- Le déficit de la phagocytose
- Altérations du système du complément
- Les DIP complexes (mdies génétiques / Sd (Wiskott-Aldrich= eczéma, thrombopénie)



# IRA: incidence selon le type d'immunodépression



Allogreffe de moelle, LAM

- .
- .

Neutropénie prolongée

- .
- .

Hémopathie lymphoïde

- .

Greffé pulmonaire

- .

Greffé rénaux, LAL, lymphome

Cancer du poumon,

maladies

systémiques,

greffe cardiaque

Tumeurs solides

# IRA: Intubation et mortalité selon le type d'ID



	IOT	MORTALITE
Allogreffe de moelle	50%	80%
LAM inaugurale	50%	50%
LAM (neutropénie)	40%	40%
Hémopathie lymphoïde	30%	40%
Greffé rénale	30%	30%
Greffé pulmonaire	25-30%	20-25%
LAL	25-40%	40%
Lymphome	25%	40%
Cancer du poumon	50%	60%
Maladies systémiques	25-30%	20%
Greffé cardiaque	10-15%	20%
Tumeurs solides (post-op exclus)	20%	30%

# Insuffisance respiratoire aiguë: quo va dis?

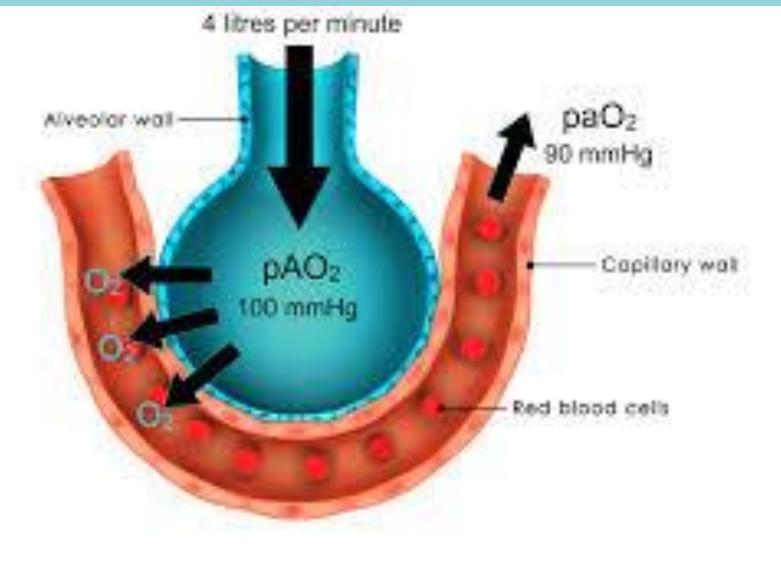
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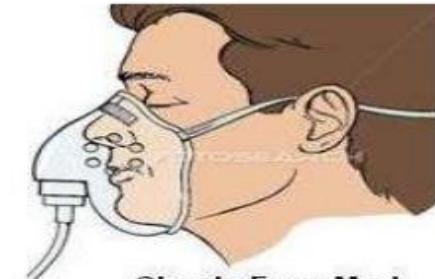


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3





Simple Face Mask



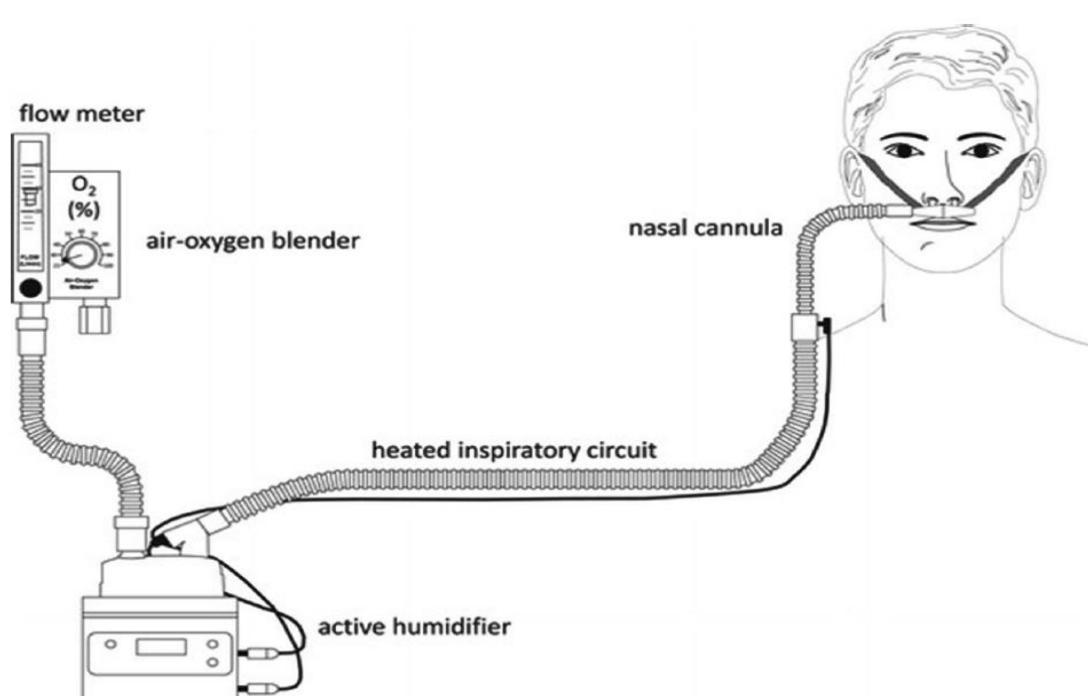
Partial Rebreather Mask



Non Rebreather Mask



Venturi Mask





# Oxygenation strategy in hypo ARF in IC: *The menu is the same*

	Strength	Weakness
Standard oxygen	Easy, can be used everywhere	Low FiO <sub>2</sub> , poor comfort, Lack of monitoring
NIV (PS/CPAP/Helmet)	Has changed the perspective for COPD/CPO..., has been very successful at a time when mortality was high with intubation, allows to monitor VTE, ...	Comfort, increased mortality if delayed intubation, workload, Helmet?
High flow oxygen	Easy, high FiO <sub>2</sub> , humidified and warmed, some positive pressure, comfort, reduces WOB, reduces intubation, reduces mortality vs. NIV	Lack of monitoring Same issues with HFNC failure Reduces intubation rate only in the most hypoxicemic
First line intubation	May control PSILI Allows protective ventilation Is not anymore the “end”	Intubated patients always have higher mortality rates. Marker of severity > a decision
A+B+C+D	...	...

$\text{FiO}_2$  values are higher and more stable

Because the delivered flow is higher than the spontaneous inspiratory demand and because the difference between the delivered flow rate and the patient's inspiratory flow rate is smaller

*The flow must be set to match the patient's inspiratory demand and/or the severity of the respiratory distress*

The anatomical dead space is decreased via washout of the nasopharyngeal space

Consequently, a larger fraction of the minute ventilation participates in gas exchange

Respiratory efforts become more efficient

Thoracoabdominal synchrony improves

The work of breathing is decreased

Because HFNC mechanically stents the airway

Provides flow rates that match the patient's inspiratory flow, and markedly attenuates the inspiratory resistance associated with the nasopharynx, thereby reducing the work of breathing

The gas delivered is heated and humidified

Warm humid gas reduces the work of breathing and improves mucociliary function, thereby facilitating secretion clearance, decreasing the risk of atelectasis, and improving the ventilation/perfusion ratio and oxygenation

The body is spared the energy cost of warming and humidifying the inspired gas (neonates +++)

Warm humid gas is associated with better conductance and pulmonary compliance compared to dry, cooler gas

*HFNC delivers adequately warmed and humidified gas only when the flow is >40 L/min*

Positive airway pressures are increased

The nasal cannula generates continuous positive pressures in the pharynx of up to 8 cmH<sub>2</sub>O, depending on flow and mouth opening

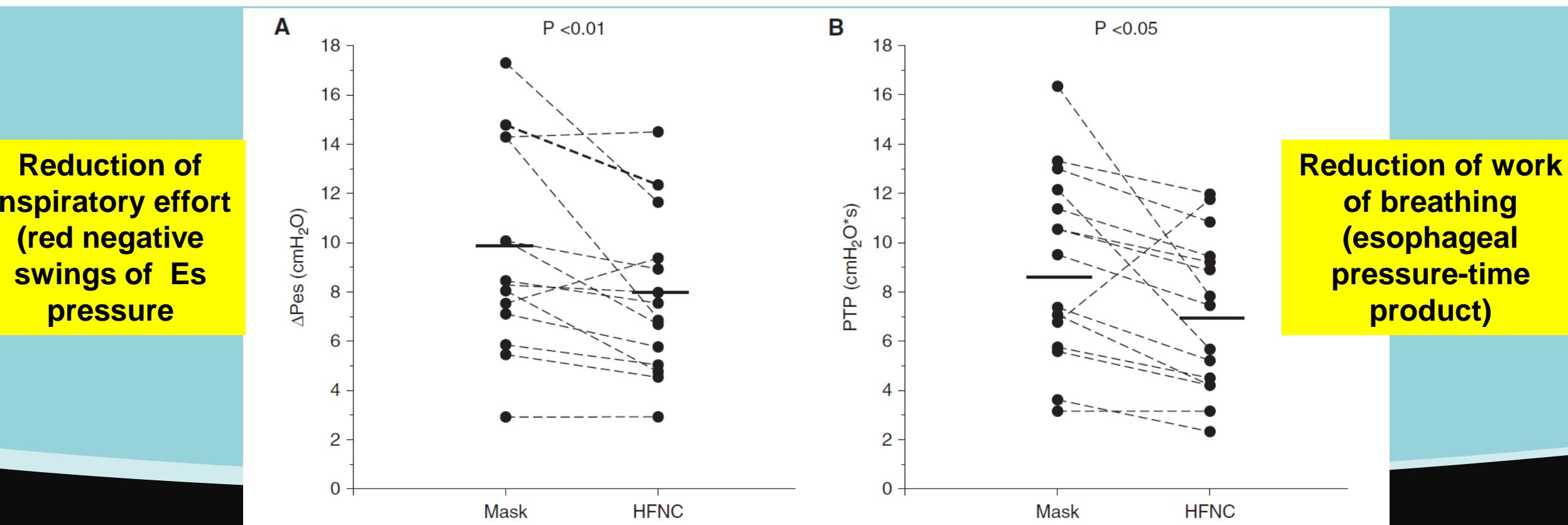
The positive pressure distends the lungs, ensuring lung recruitment and decreasing the ventilation-perfusion mismatch in the lungs

End-expiratory lung volume is greater with HFNO than with low-flow oxygen therapy

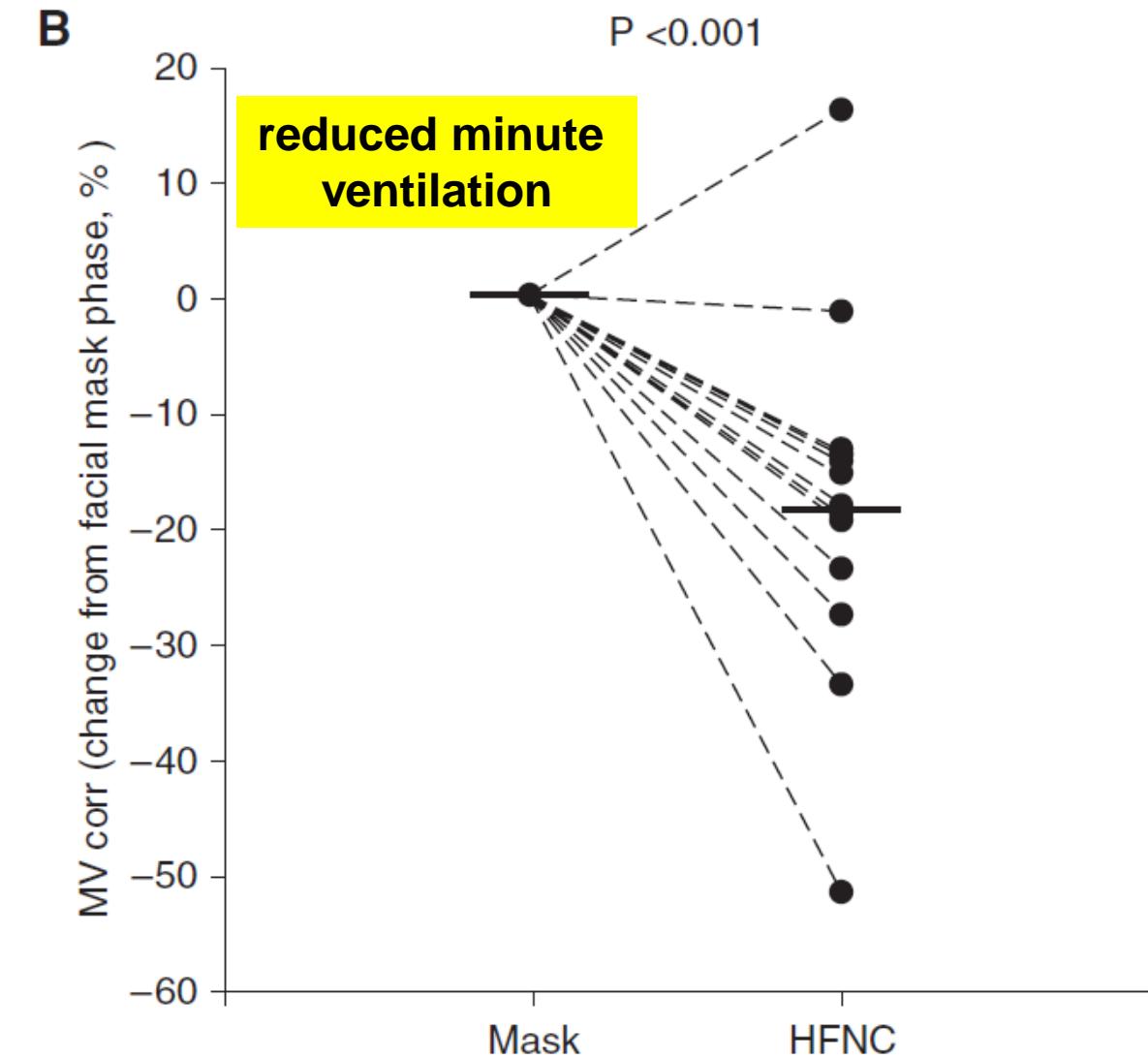
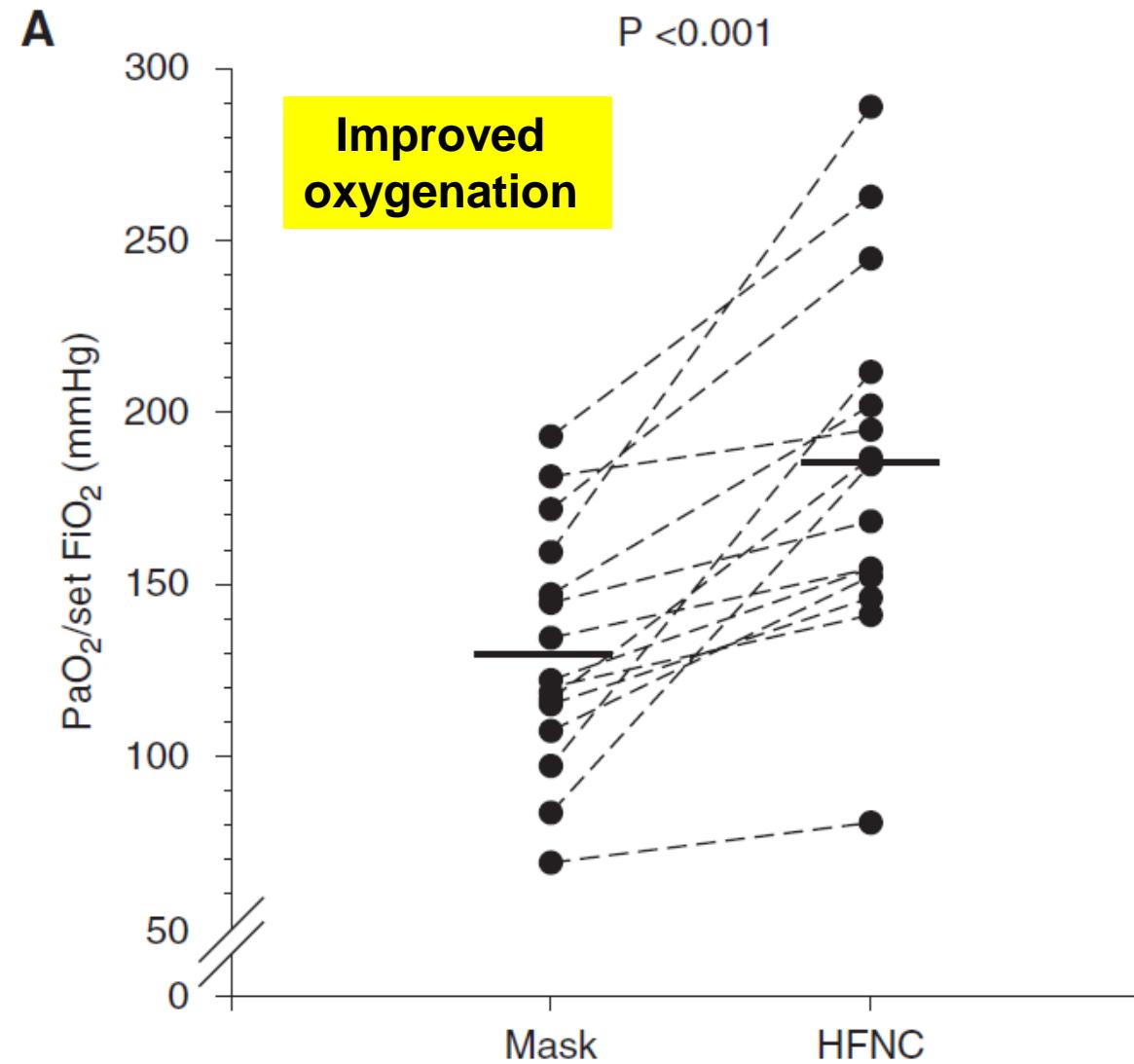
*Minimizing leaks around the cannula prongs is of the utmost importance*

## **HFNC delivered at 40 L/min vs. standard nonocclusive facial mask at 12 L/min Physiologic Effects of High-Flow Nasal Cannula in Acute Hypoxemic Respiratory Failure**

Tommaso Mauri<sup>1,2</sup>, Cecilia Turrini<sup>1,3</sup>, Nilde Eronia<sup>4</sup>, Giacomo Grasselli<sup>1</sup>, Carlo Alberto Volta<sup>3</sup>, Giacomo Bellani<sup>4,5</sup>, and Antonio Pesenti<sup>1,2</sup>



## HFNC delivered at 40 L/min vs. standard nonocclusive facial mask at 12 L/min





# Optimum support by high-flow nasal cannula in acute hypoxemic respiratory failure: effects of increasing flow rates

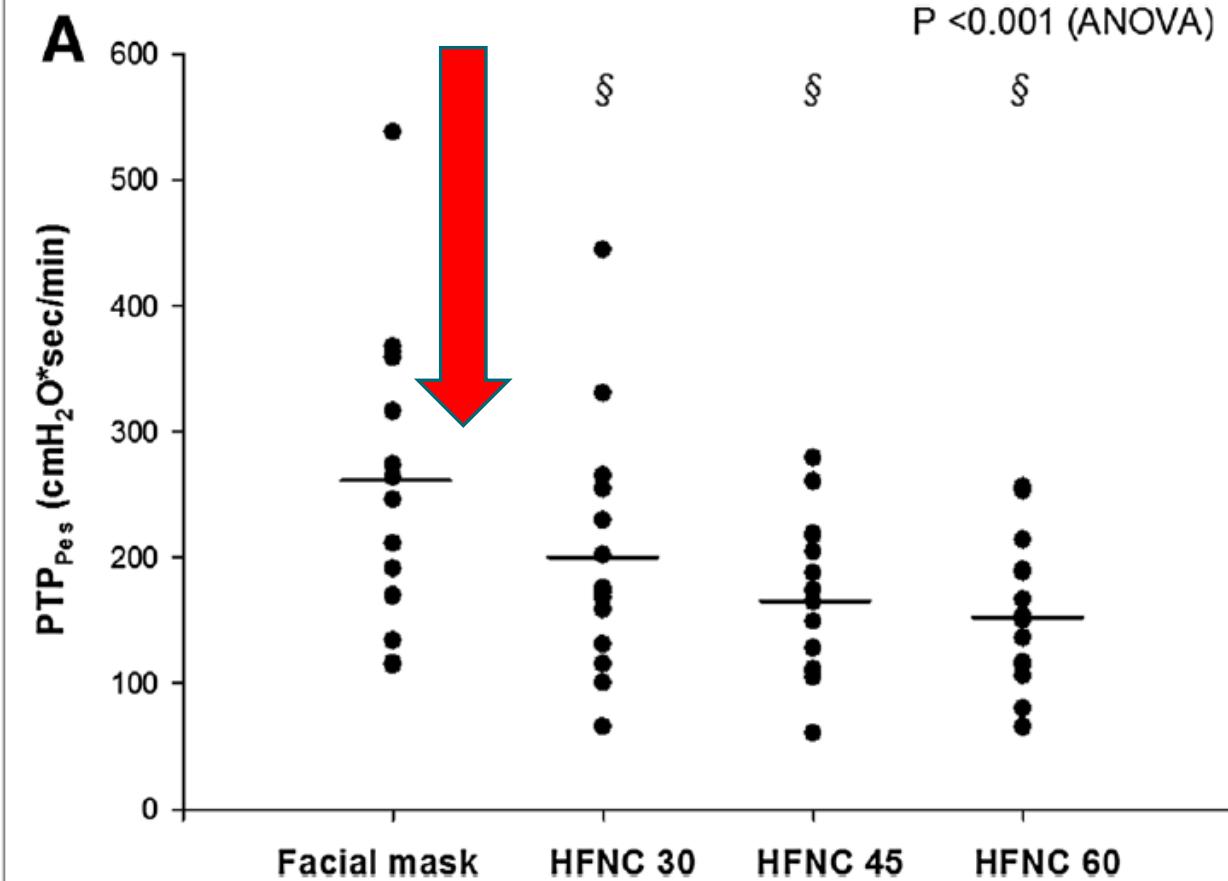
Tommaso Mauri<sup>1,2</sup>, Laura Alban<sup>3</sup>, Cecilia Turrini<sup>3</sup>, Barbara Cambiaghi<sup>4</sup>, Eleonora Carlesso<sup>1</sup>, Paolo Taccone<sup>2</sup>, Nicola Bottino<sup>2</sup>, Alfredo Lissoni<sup>2</sup>, Savino Spadaro<sup>3</sup>, Carlo Alberto Volta<sup>3</sup>, Luciano Gattinoni<sup>5</sup>, Antonio Pesenti<sup>1,2\*</sup> and Giacomo Grasselli<sup>2</sup>

*Intensive Care Med (2017) 43:1453–1463*

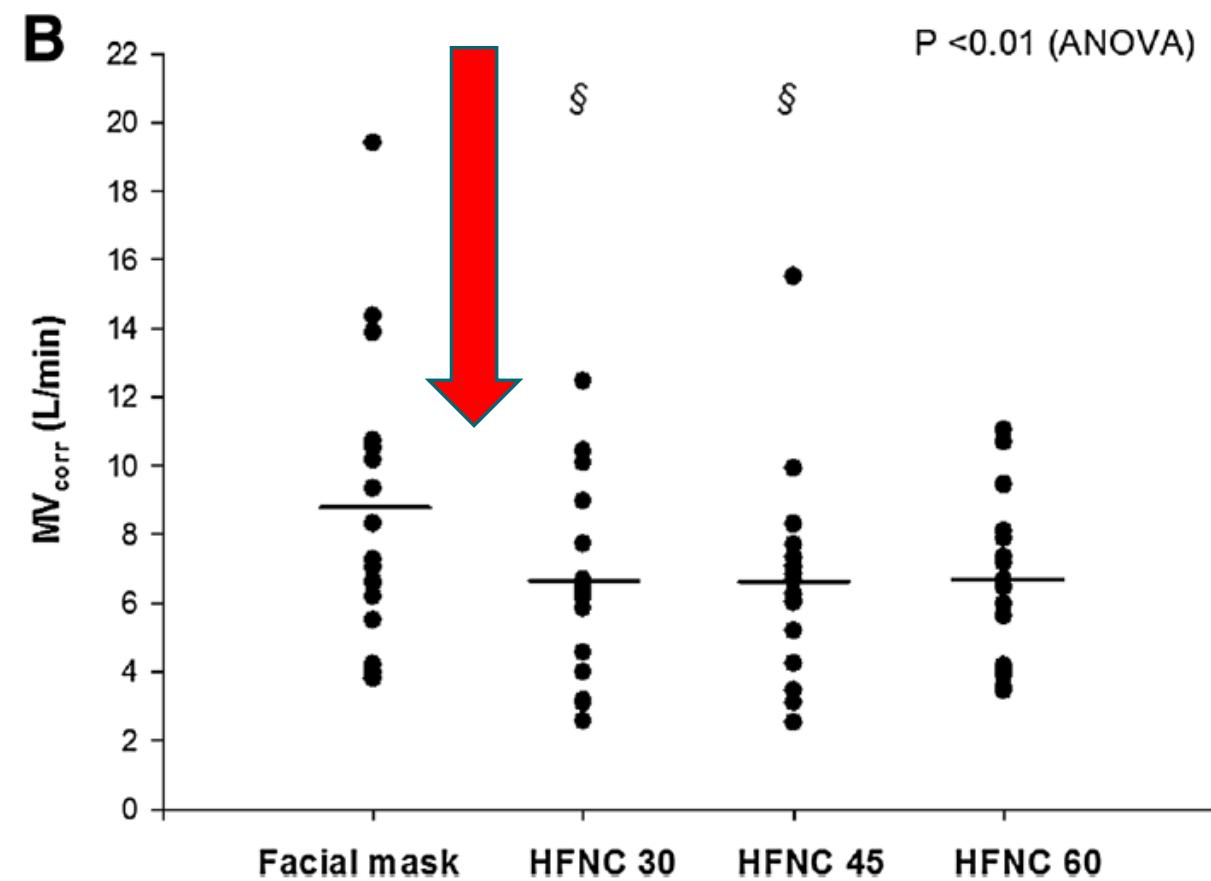
**Prospective randomized cross-over study, 17 AHRF, PAFl≤300 mmHg.**

**HFNC at flow rates of 30, 45 and 60 l/min randomly applied, constant FiO2**

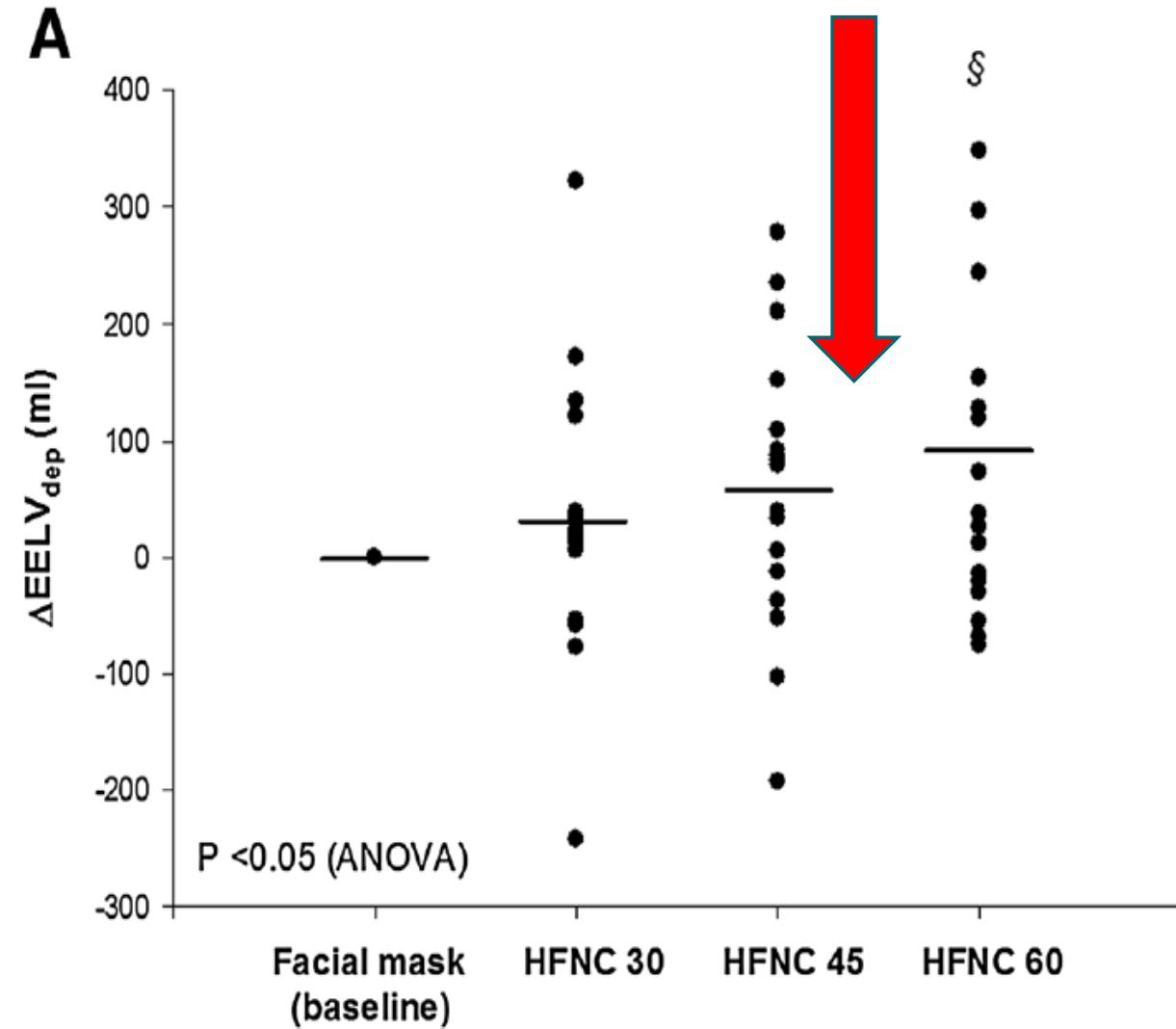
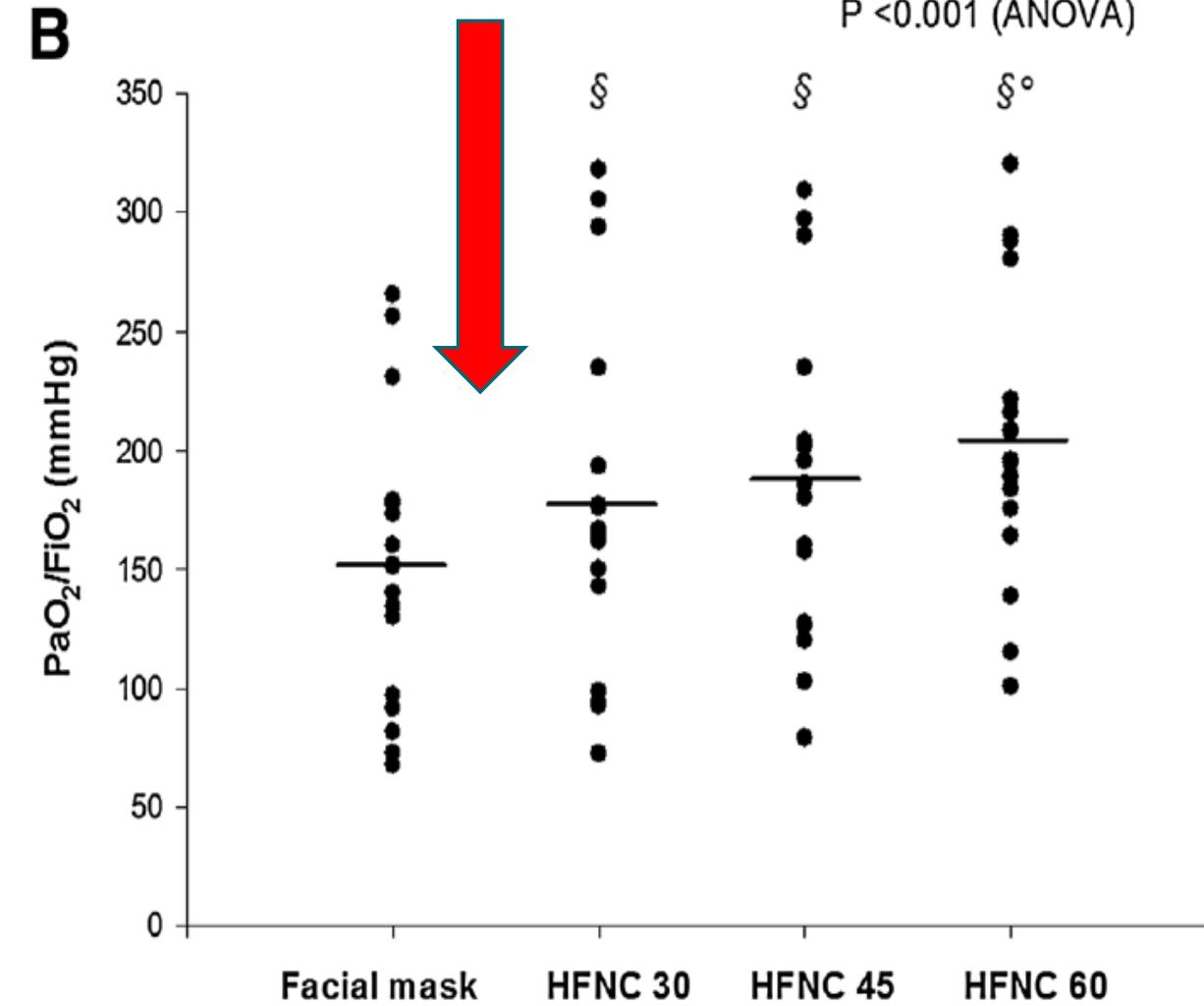
**At the end of each phase, ABG, we measured arterial blood gases, inspiratory effort ( $\Delta$ Pes) and on the esophageal pressure–time product (PTPPes), and lung volume, by electrical impedance tomography.**



inspiratory effort



CO2 « clearance »

**A****Changes in lung aeration****B****PaO<sub>2</sub> / FiO<sub>2</sub>**

Targeted physiologic variable

Optimum flow rate<sup>a</sup>

« OPTIMAL »

HFNC 30 l/min HFNC 45 l/min HFNC 60 l/min

$\Delta P_{\text{res}}$

✗

✗

✓

inspiratory effort

**Most of the effect on inspiratory workload and CO<sub>2</sub> clearance was already obtained at the lowest flow rate.**

**Increasing HFNC flow rate progressively decreased inspiratory effort and improved lung aeration, dynamic compliance and oxygenation.**

RR

✗

✗

✓

PaO<sub>2</sub>/FiO<sub>2</sub>

✗

✗

✓

RESEARCH

Open Access



# The effects of a 2-h trial of high-flow

	HFNO group (n = 52)	Venturi mask group (n = 48)	P value
Primary endpoint			
Number (%) of patients requiring mechanical ventilation	8 (15 %)	4 (8 %)	0.36
Noninvasive mechanical ventilation	6 <sup>a</sup>	3 <sup>a</sup>	
Invasive mechanical ventilation	4	2	
Secondary endpoints, median [25th–75th percentile]			
Discomfort VAS score <sup>b</sup> at 120 min	3 [1–5]	3 [0–5]	0.88
Dyspnea VAS score <sup>b</sup> at 120 min	3 [2 – 6]	3 [1–6]	0.87
Thirst VAS score <sup>b</sup> at 120 min	6 [3–8]	6 [5 – 9]	0.40
Respiratory rate at 120 min, breaths/min	25 [22–29]	25 [21–31]	
Heart rate at 120 min, beats/min	98 [90–110]	99 [83–112]	0.43

# Quel comparateur?

- Ventilation non invasive
  - Essai iVNIctus négatif: pas de bénéfice de la VNI
  - Essai FLORALI post hoc: plus d'intubation et surmortalité avec la VNI
  - Cohorte Efraim: pas de bénéfice de la VNI
  - Chez les patients les plus hypoxémiques, les VTe expliquent en partie l'échec de la VNI
  - **La VNI ça suffit? Et la c.CPAP?**
- Oxygène standard
  - Essai iVNIctus – analyse post hoc : pas de différence entre HFNO et O<sub>2</sub>
  - Essai FLORALI – analyse post hoc: pas de différence entre HFNO et O<sub>2</sub>
  - Cohorte Efraim: Réduction de l'intubation avec HFNO, mortalité idem
  - **Besoin d'essais randomisés**

# (encore) Une place pour la VNI ?

	iVNIctus JAMA 2015-CCM2017	Florali NEJM2015-LRM 2016	Efraim ICM 2017
	RCT N=374	RCT N=300 ( 82 ID)	Cohorte N=1611 (915 non intubés)
O2 Standard	NS par rapport HFNC	NS par rapport HFNC	NS par rapport à la VNI
VNI	Aucun bénéfice Aucun effet nocif Effet Vte?	Plus d'intubation Plus de mortalité Effet VTe	NS par rapport à O2 Aucun effet nocif Effet VTe
HFNC	NS par rapport O2	NS par rapport O2	Presque supériorité par rapport à l'O2 sur l'IOT mais NS sur la mortalité

VNI : nég ou nocive, surtout si PAFI bas, FR élevée et hauts Vte : **on arrête? Helmet?**

O2 vs. HFNO: peut-être un effet sur l'IOT, à confirmer. Rien sur la mortalité

Et toujours le prix à payer de l'intubation retardée

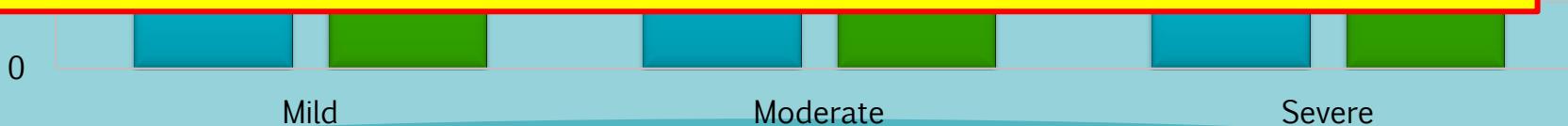
# NIV in patients with hypoxemic ARF: three RCTs

Study	Antonelli, NEJM 1998	Delclaux, JAMA 2000	Ferrer, AJRCCM
Patients	64 patients, 1 ICU hypoxemic acute resp. with criteria for MV Intubation vs. NPPV PAFI: 116 vs. 124	123 patients 6 ICUs hypoxemic acute resp.  Oxygen vs. CPAP PAFI 148 vs. 140	105 patients 1 ICU hypoxemic acute resp.  Oxygen vs. BiPAP PAFI 102 vs. 103
Endpoints	No S Improved oxygenation 31% intubation 53% vs. 72% (NIV) survival Complications: higher Lengths of ....: increased	Improved oxygenation 33% vs. 39% intubation 75% vs. 79% survival Complications NS Lengths: NS	/ 25% vs. 52% intubation 72% vs. 61% survival 10% vs. 24% NS
Comments	Pulmonary edema included	Pulmonary edema included	Pulmonary edema included

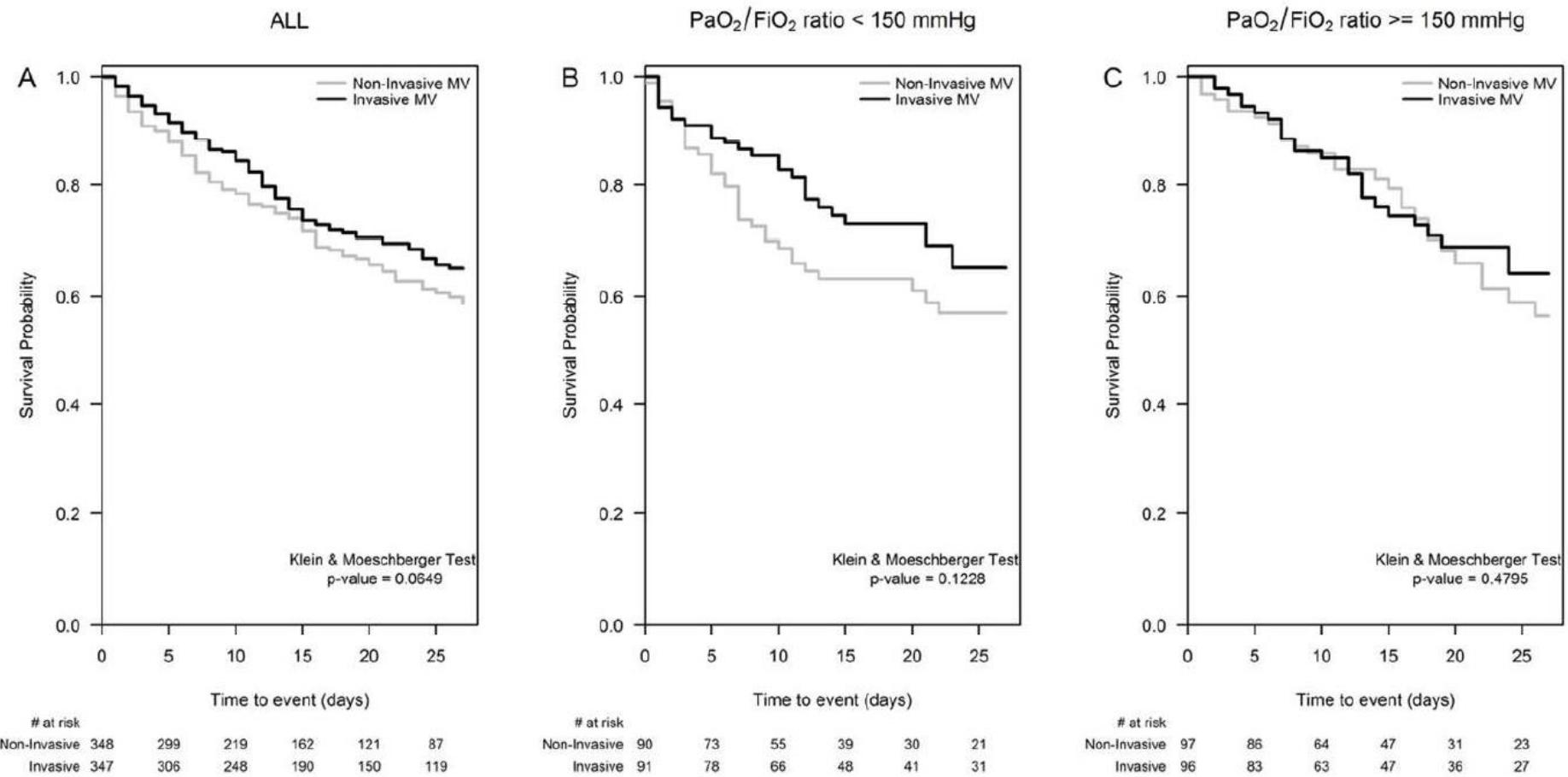
# NIV use and failure in ARDS patients (LUNG SAFE)

ARDS severity was associated with an increase in intensity of ventilatory support, NIV failure, and ICU mortality.

Overall, hospital mortality was 16.1 % in patients with NIV success and 45.4% in case of NIV failure.

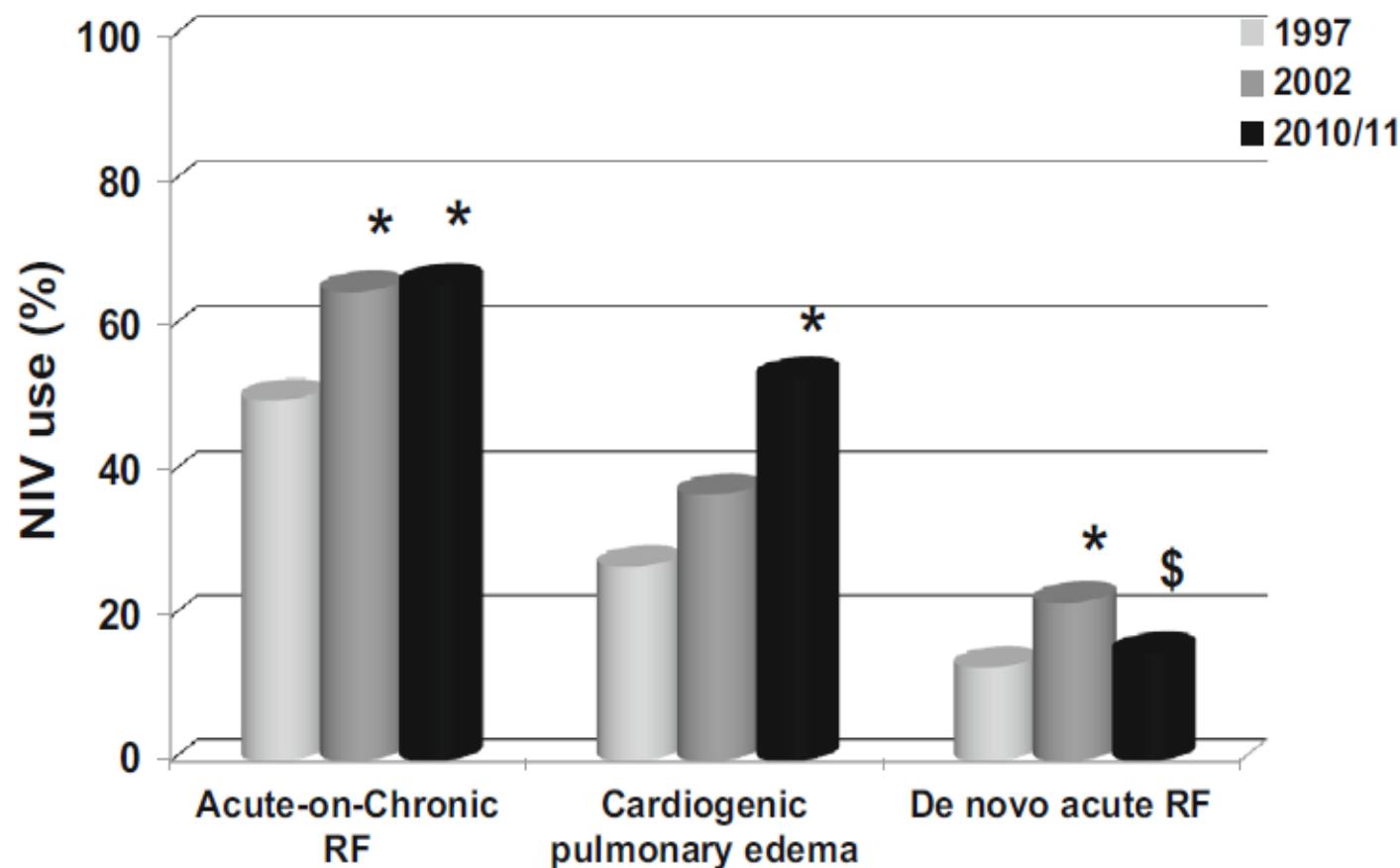


# Data from LUNG SAFE: 436/2813 (15%) NIV use

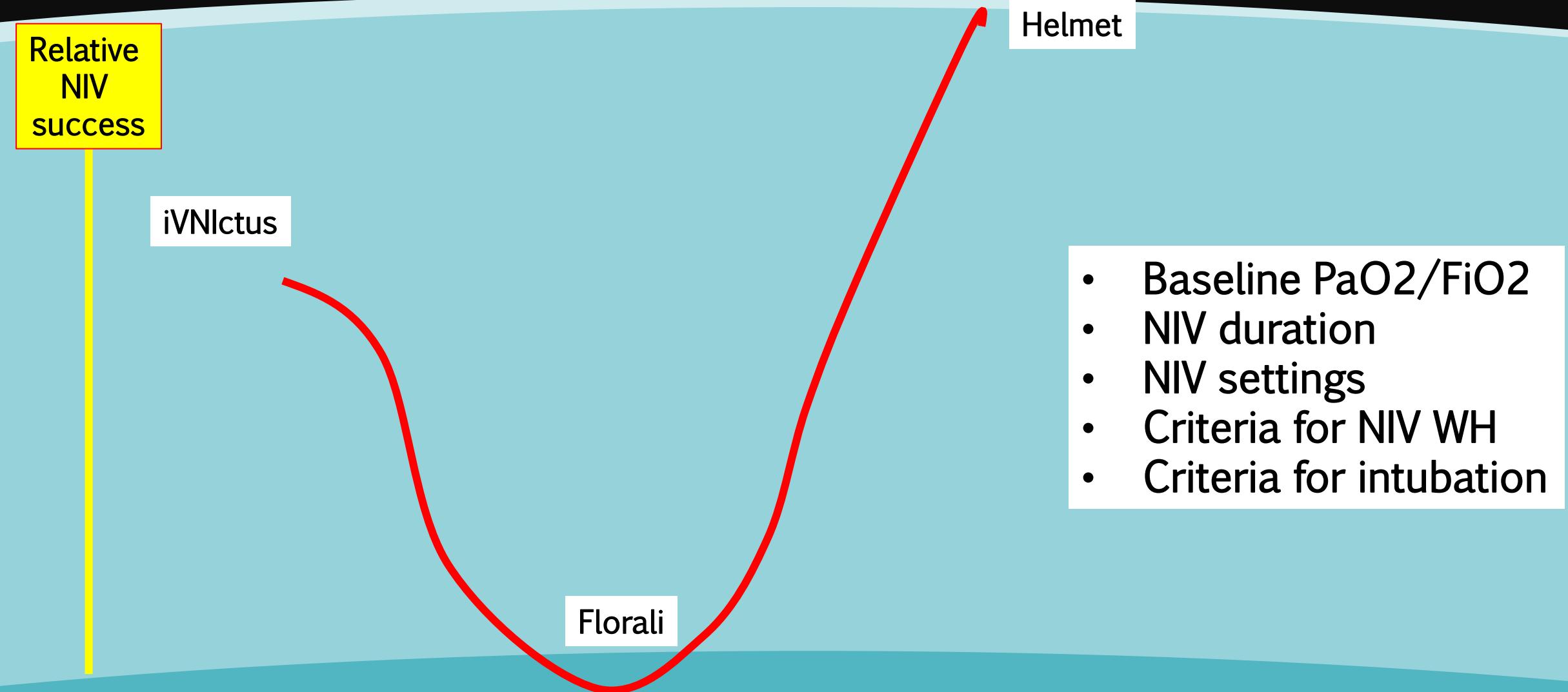


Alexandre Demoule  
Sylvie Chevret  
Annalisa Carlucci  
Achille Kouatchet  
Samir Jaber  
Ferhat Meziani  
Matthieu Schmidt  
David Schnell  
Céline Clergue  
Jérôme Aboab  
Antoine Rabbat  
Béatrice Eon  
Claude Guérin  
Hugues Georges  
Benjamin Zuber  
Jean Dellamonica  
Vincent Das  
Joël Cousson  
Didier Perez  
Laurent Brochard  
Elie Azoulay

# Changing use of noninvasive ventilation in critically ill patients: trends over 15 years in francophone countries



# A U shape paradoxical effect of NIV in hypoxemic ARF?



# 680 met all study inclusion criteria

306 were not included

81 met  $\geq 1$  exclusion criterion  
82 required immediate intubation  
55 had do-not-intubate orders  
33 declined participation  
19 were eligible but were not randomized  
10 were outside randomization window  
9 were previously included into the study  
17 had other reasons

374 were randomized

183 were assigned to oxygen therapy only

Three (1,5%) patients received rescue noninvasive ventilation

183 were included in the intent-to-treat analysis of the primary outcome

191 were assigned to Non-invasive ventilation

All patients received NIV  
14 (7,3%) received a single NIV session

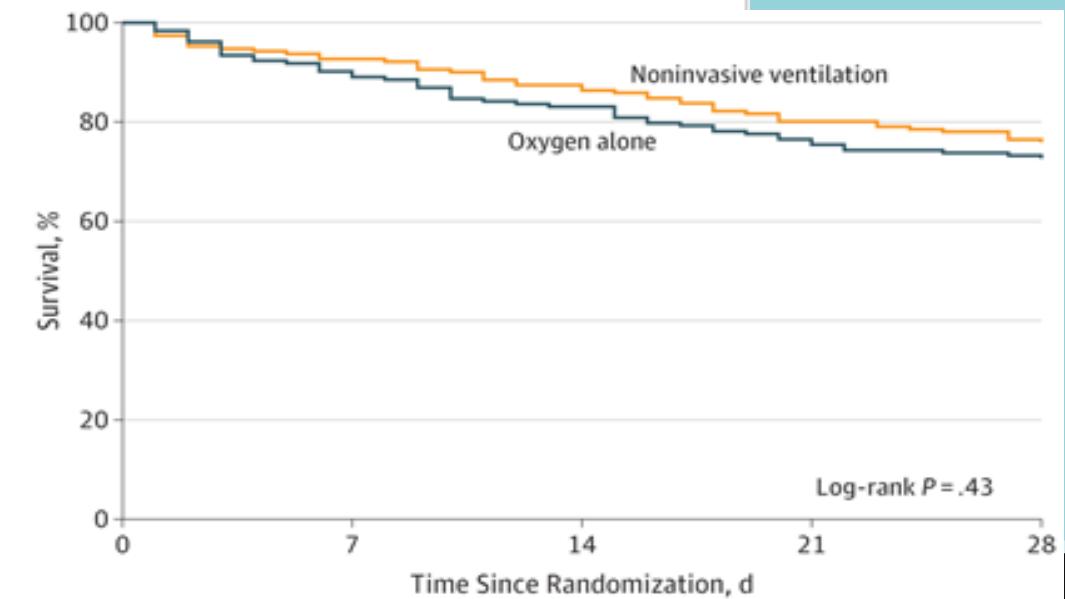
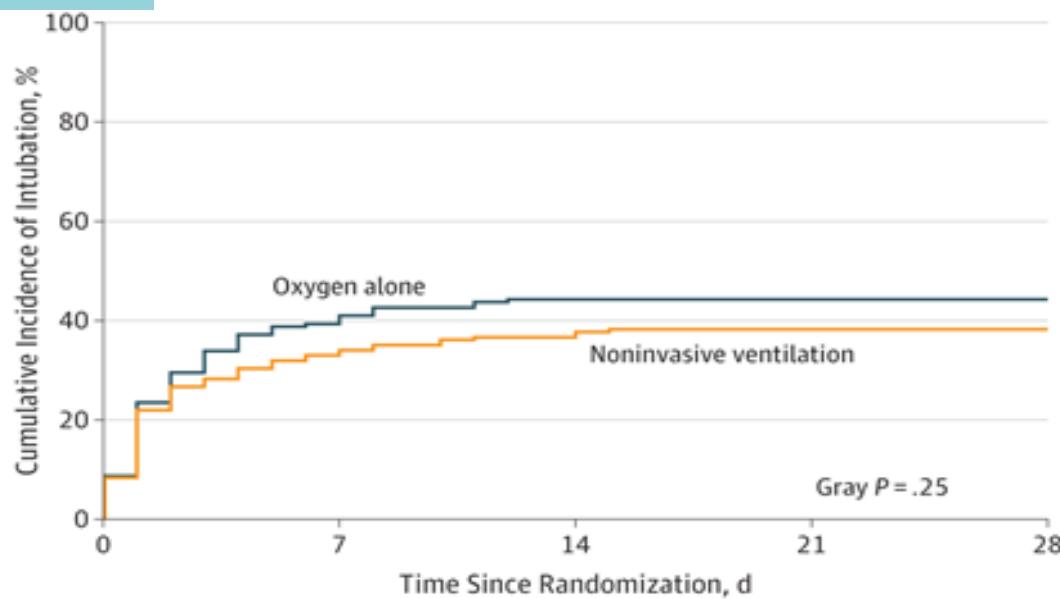
191 were included in the intent-to-treat analysis of the primary outcome



## **Effect of Noninvasive Ventilation vs Oxygen Therapy on Mortality Among Immunocompromised Patients With Acute Respiratory Failure**

A Randomized Clinical Trial **FREE** **ONLINE FIRST**

Virginie Lemiale, MD<sup>1</sup>; Djamel Mokart, MD<sup>2</sup>; Matthieu Resche-Rigon, MD, PhD<sup>3</sup>; Frédéric Pène, MD, PhD<sup>3</sup>; Julien Mayaux, MD<sup>4</sup>; Etienne Faucher, MD<sup>5</sup>; Martine Nyunga, MD<sup>6</sup>; Christophe Girault, MD, PhD<sup>7</sup>; Pierre Perez, MD<sup>8</sup>; Christophe Guitton, MD, PhD<sup>9</sup>; Kenneth Ekpe, MD<sup>10</sup>; Achille Kouatchet, MD<sup>11</sup>; Igor Théodore, MS<sup>3</sup>; Dominique Benoit, MD, PhD<sup>12</sup>; Emmanuel Canet, MD<sup>3</sup>; François Barbier, MD, PhD<sup>13</sup>; Antoine Rabbat, MD<sup>3</sup>; Fabrice Bruneel, MD<sup>14</sup>; Francois Vincent, MD<sup>15</sup>; Kada Klouche, MD, PhD<sup>16</sup>; Kontar Loay, MD<sup>17</sup>; Eric Mariotte, MD<sup>3</sup>; Lila Bouadma, MD, PhD<sup>3</sup>; Anne-Sophie Moreau, MD<sup>18</sup>; Amélie Seguin, MD<sup>19</sup>; Anne-Pascale Meert, MD, PhD<sup>20</sup>; Jean Reignier, MD, PhD<sup>21</sup>; Laurent Papazian, MD, PhD<sup>22</sup>; Ilham Mehzari, MD<sup>23</sup>; Yves Cohen, MD, PhD<sup>15</sup>; Maleka Schenck, MD<sup>24</sup>; Rebecca Hamidfar, MD<sup>25</sup>; Michael Darmon, MD, PhD<sup>26</sup>; Alexandre Demoule, MD, PhD<sup>3</sup>; Sylvie Chevret, MD, PhD<sup>1</sup>; Elie Azoulay, MD, PhD<sup>1</sup>; for the Groupe de Recherche en Réanimation Respiratoire du patient d'Onco-Hématologie (GRRR-OH)



No. at risk	Noninvasive ventilation				Oxygen alone				Risk			
Noninvasive ventilation	191	125	118	112	111	191	177	167	153	146		
Oxygen alone	183	107	95	92	91	183	165	152	140	134		

# High-Flow Nasal Cannula Oxygenation in Immunocompromised Patients With Acute Hypoxemic Respiratory Failure: A Groupe de Recherche Respiratoire en Réanimation Onco-Hématologique Study

Critical Care Medicine, online

**TABLE 2. Primary and Secondary Outcomes According to Oxygenation Strategy in the Matched Population**

	High-Flow Nasal Oxygen Group (n = 90)	O <sub>2</sub> Group (n = 90)	Hazard Ratio 95% CI	p
Primary endpoint				
All cause 28-d mortality	21 (23.3%)	23 (25.5%)	0.80 (0.45–1.43)	0.45
Secondary endpoints				
Need for invasive mechanical ventilation	40 (44.4%)	48 (53.3%)	0.42 (0.11–1.61)	0.20
Duration of mechanical ventilation	13 [4–46]	16 [8–33]		0.32
ICU-acquired infection	21 (23.3%)	28 (31.1%)	0.80 (0.39–1.66)	0.55
Length of ICU stay	8 [5–16]	8 [3–29]		0.59
Length of hospital stay	24 [14–51]	32 [19–52]		0.25



# Effect of non-invasive oxygenation strategies in immunocompromised patients with severe acute respiratory failure: a post-hoc analysis of a randomised trial

Jean-Pierre Frat, Stéphanie Ragot, Christophe Girault, Sébastien Perbet, Gwénael Prat, Thierry Boulain, Alexandre Demoule, Jean-Damien Ricard, Rémi Coudroy, René Robert, Alain Mercat, Laurent Brochard, Arnaud W Thille, for the REVA network

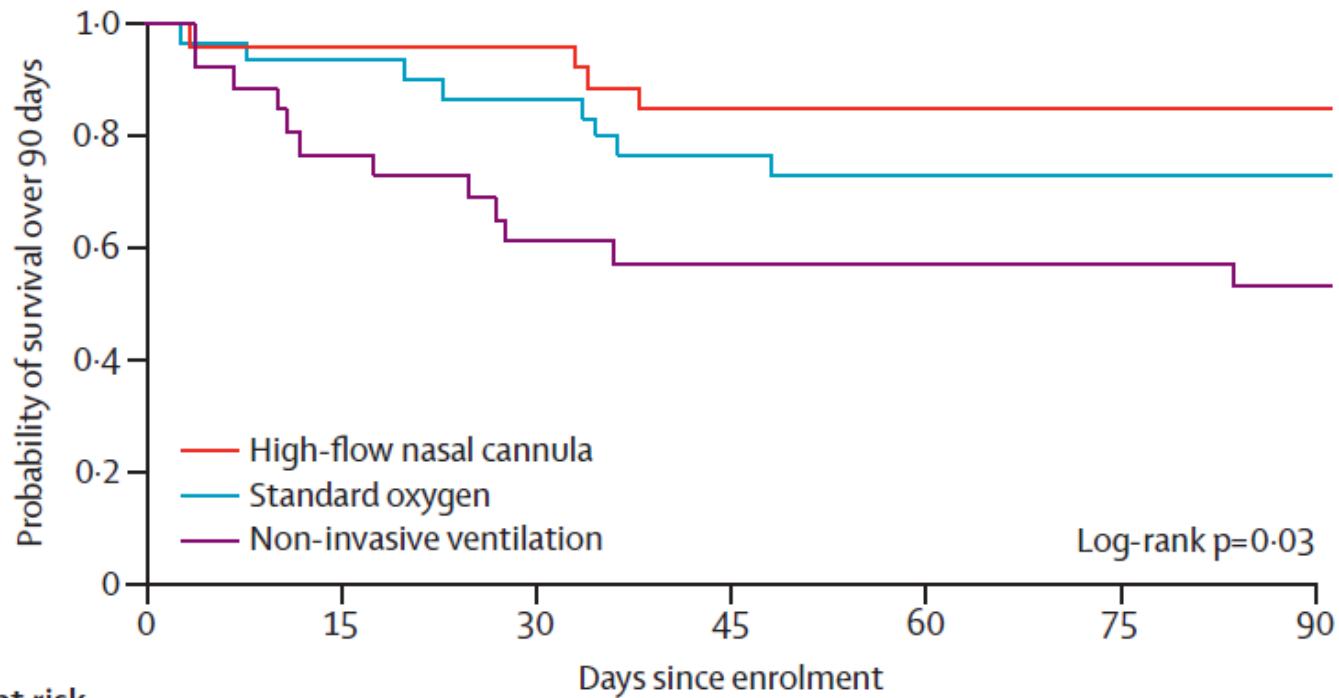
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	Number at risk						
High-flow nasal cannula group	26	25	25	22	22	22	22
Standard oxygen group	30	28	26	23	22	22	22
Non-invasive ventilation group	26	20	16	15	14	14	13

h endotracheal  
ne therapy.

## High-flow oxygen therapy in cancer patients with acute respiratory failure

Djamel Mokart  
Cyrille Geay  
Laurent Chow-Chine  
Jean-Paul Brun  
Marion Faucher  
Jean-Louis Blache  
Magali Bisbal  
Antoine Sannini

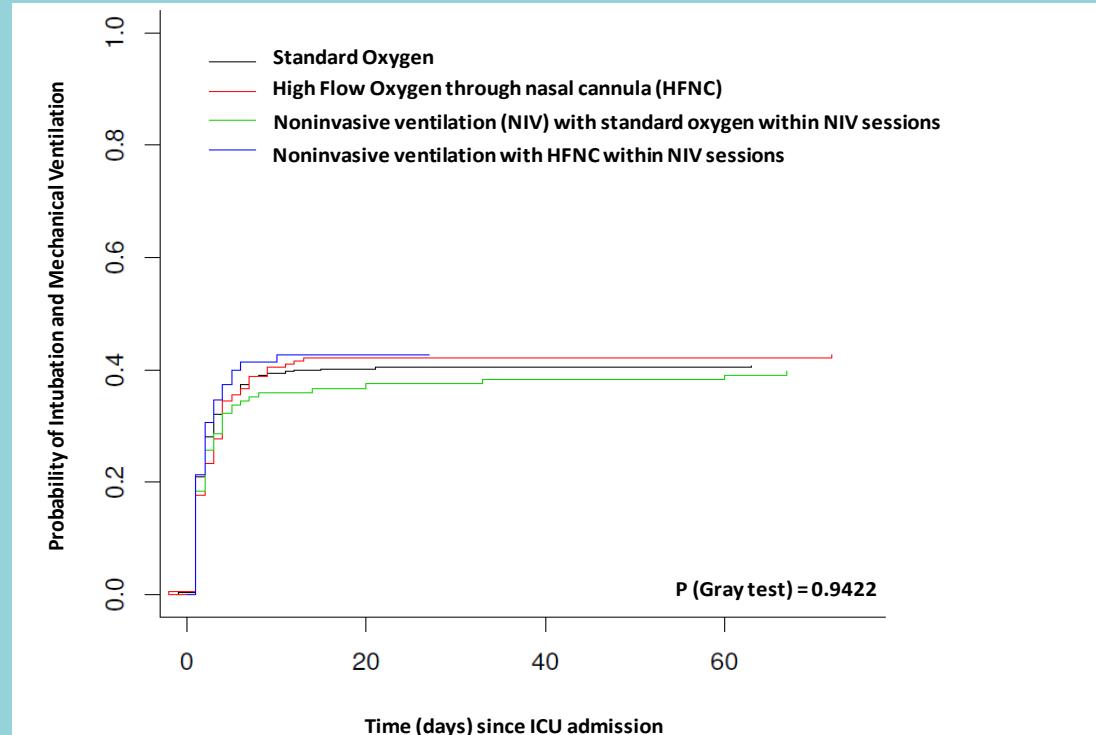
	HFNC + NIV N=69	OTHER N=69	P VALUE
SOFA	6 (4-9)	6 (4-8)	0.73
PaO <sub>2</sub> /FiO <sub>2</sub> <200	46%	51%	0.39
D-28 MORTALITY	36%	54%	0.01
MORTALITY IF VENTILATED	52%	72%	0.07



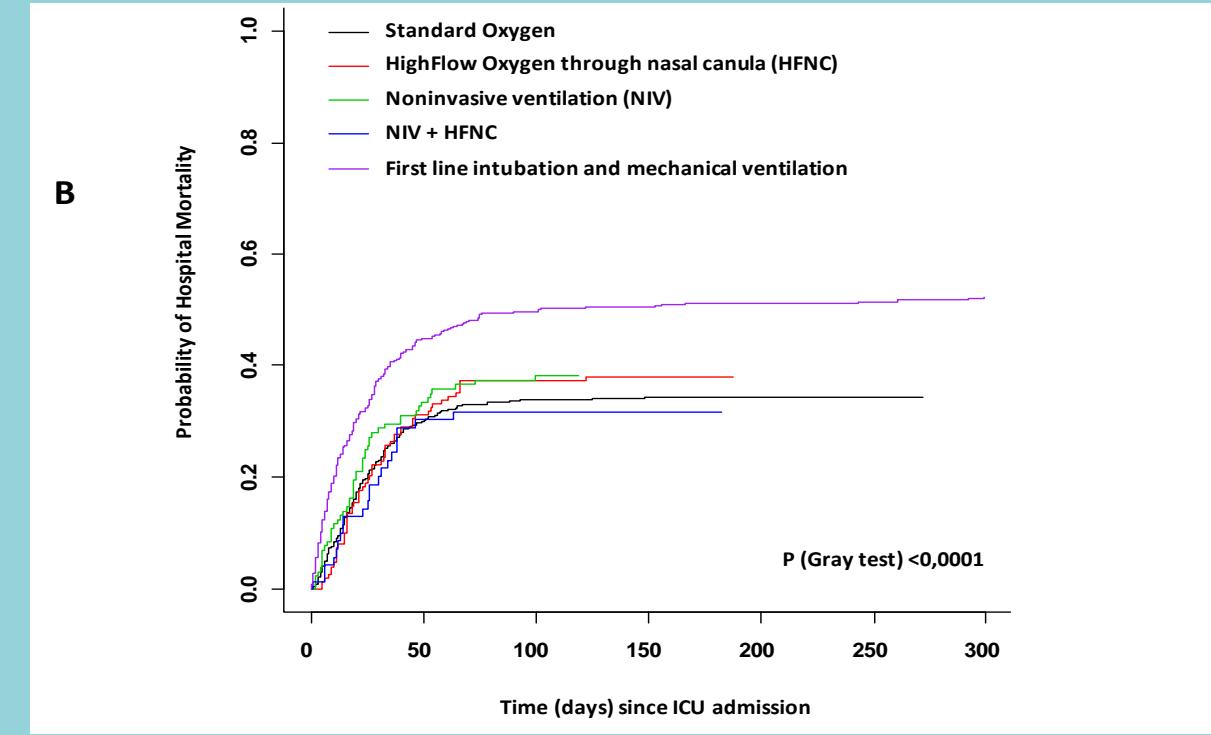
# Acute hypoxemic respiratory failure in immunocompromised patients: the Efraim multinational prospective cohort study

Elie Azoulay<sup>1\*</sup> , Peter Pickkers<sup>2</sup>, Marcio Soares<sup>3</sup>, Anders Perner<sup>4</sup>, Jordi Rello<sup>5</sup>, Philippe R. Bauer<sup>6</sup>, Andry van de Louw<sup>7</sup>, Pleun Hemelaar<sup>2</sup>, Virginie Lemiale<sup>1</sup>, Fabio Silvio Taccone<sup>8</sup>, Ignacio Martin Loeches<sup>9,10</sup>, Tine Sylvest Meyhoff<sup>4</sup>, Jorge Salluh<sup>3</sup>, Peter Schellongowski<sup>11</sup>, Katerina Rusinova<sup>12</sup>, Nicolas Terzi<sup>13</sup>, Sangeeta Mehta<sup>14</sup>, Massimo Antonelli<sup>15</sup>, Achille Kouatchet<sup>16</sup>, Andreas Barratt-Due<sup>17</sup>, Miia Valkonen<sup>18</sup>, Precious Pearl Landburg<sup>19</sup>, Fabrice Bruneel<sup>20</sup>, Ramin Brandt Bukan<sup>21</sup>, Frédéric Pène<sup>22</sup>, Victoria Metaxa<sup>23</sup>, Anne Sophie Moreau<sup>24</sup>, Virginie Soupart<sup>1</sup>, Gaston Burghi<sup>25</sup>, Christophe Girault<sup>26</sup>, Ulysses V. A. Silva<sup>27</sup>, Luca Montini<sup>15</sup>, François Barbier<sup>28</sup>, Lene B. Nielsen<sup>29,30</sup>, Benjamin Gaborit<sup>31</sup>, Djamel Mokart<sup>32</sup> and Sylvie Chevret<sup>33</sup> for the Efraim investigators and the Nine-I study group

# Probability of Intubation



# Probability of Mortality



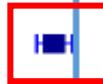
Azoulay et al. The Efraim Study. Intensive Care Medicine 2017

# INTUBATION

Age

Hazard Ratios  
(95% Confidence Intervals)

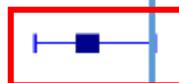
0.92 (0.86-0.99)



Initial ventilation strategy (with standard Oxygen as reference)

High Flow  
Oxygen

0.77 (0.59-1.01)



Noninvasive ventilation (NIV)

0.94 (0.69-1.28)



NIV + HFNC

0.74 (0.51-1.09)



Chronic Respiratory Insufficiency

0.76 (0.54-1.08)



SOFA at ICU admission

1.09 (1.06-1.13)



P/F ratio

1.47 (1.05-2.07)



Etiology of the Acute Respiratory Failure (ARF)

Pneumocystis jirovecii Pneumonia

2.11 (1.42-3.14)



IFI

Invasive Pulmonary Aspergillosis

1.85 (1.21-2.85)



Undetermined ARF etiology

1.46 (1.09-1.98)



Increased risk of intubation and mechanical ventilation

# MORTALITY

Odd Ratios  
(95% Confidence Intervals)

Intercept

0.06 (0.03-0.11)

Age

1.18 (1.09-1.27)

Direct ICU admission

0.69 (0.54-0.87)

SOFA score at Day 1

1.12 (1.08-1.16)

respiratory items

P/F ratio

0 (as the reference)

<100

1.60 (1.03-2.48)

100-199

1.46 (0.98-2.18)

200-299

1.30 (0.83-2.05)

Intubation

n and mechanical ventilation (IMV, with no intubation as the reference)

IMV after standard oxygen failure

4.16 (2.91-5.93)

IMV after high flow oxygen (HFNC) failure

5.54 (3.27-9.38)

IMV after noninvasive ventilation (NIV) failure

3.65 (2.05-6.53)

IMV after failure of NIV+HFNC

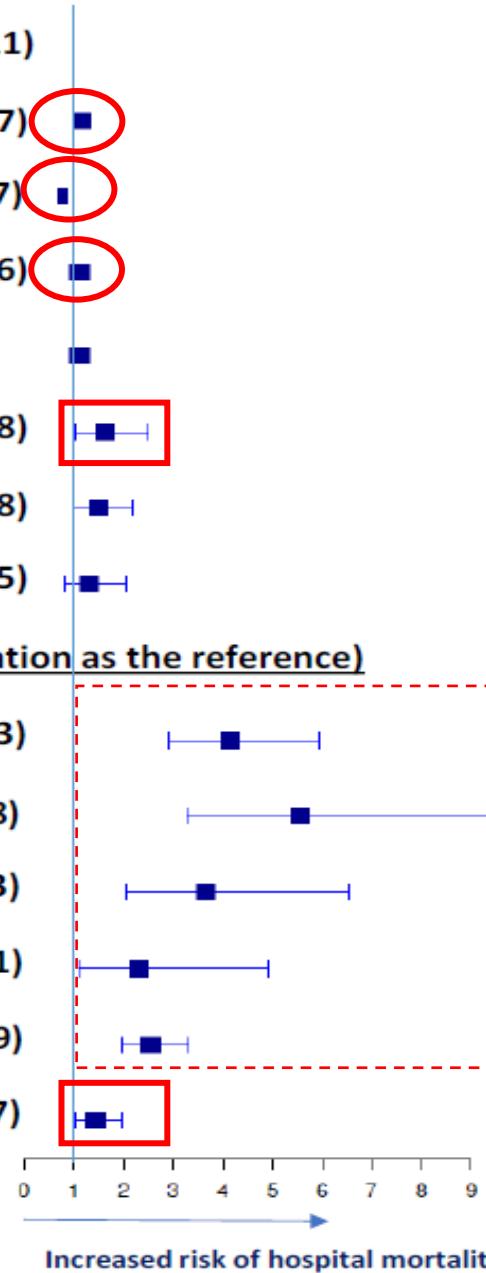
2.31 (1.09-4.91)

First line IMV

2.55 (1.94-3.29)

Undetermined ARF etiology

1.43 (1.04-1.97)



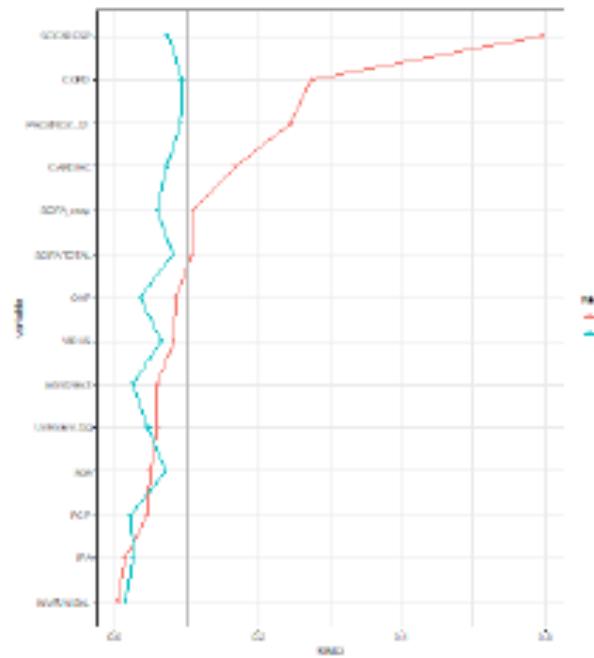
# Propensity Analysis

Initial oxygenation strategy has an impact on intubation

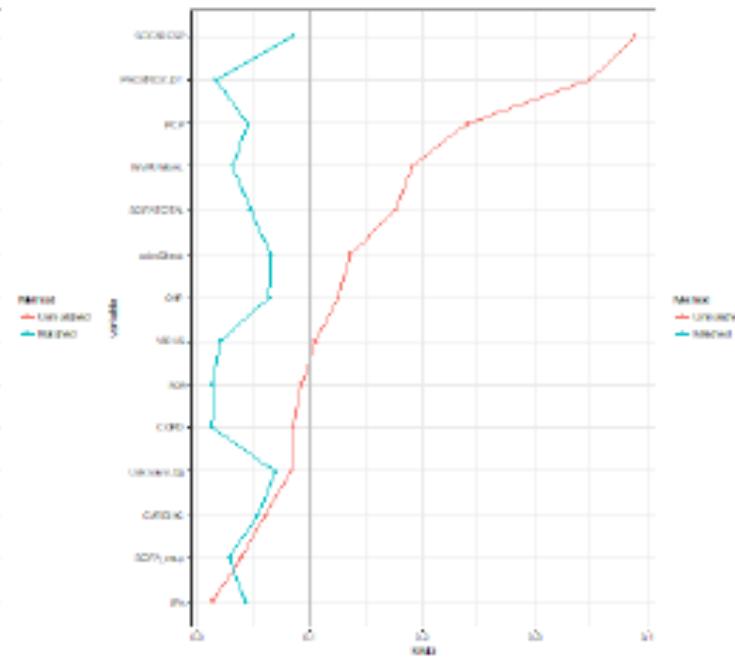
- *High flow oxygen ( $P=0.05$ ) , NIV not significant*

Initial oxygenation strategy has NO impact on mortality

(i) Propensity score for NIV exposure



(ii) Propensity score for HFNC exposure



# **Effect of high-flow nasal cannula oxygen therapy in adults with acute hypoxemic respiratory failure: a meta-analysis of randomized controlled trials**

**6 RCTs ( $n = 1892$ ).**

Xiaofeng Ou MD PhD, Yusi Hua MD MSc, Jin Liu MD PhD, Cansheng Gong MD PhD, Wenling Zhao MD MSc

- Compared with standard oxygen, HFNC oxygen therapy was associated with a lower intubation rate (risk ratio [RR] 0.60, 95% confidence interval [CI] 0.38 to 0.94;  $I^2 = 49\%$ ).
- No significant difference in the rate between HFNC oxygen therapy and noninvasive ventilation (RR 0.86, 95% CI 0.68 to 1.09;  $I^2 = 2\%$ ).
- No difference in mortality

Early HFNC failure group compared with the late HFNC failure group

Byung Ju Kang  
Younsuck Koh  
Chae-Man Lim  
Jin Won Huh  
Seunghee Baek  
Myongja Han  
Hyun-Suk Seo  
Hee Jung Suh  
Ga Jin Seo  
Eun Young Kim  
Sang-Bum Hong

**Failure of high-flow nasal cannula therapy may delay intubation and increase mortality**

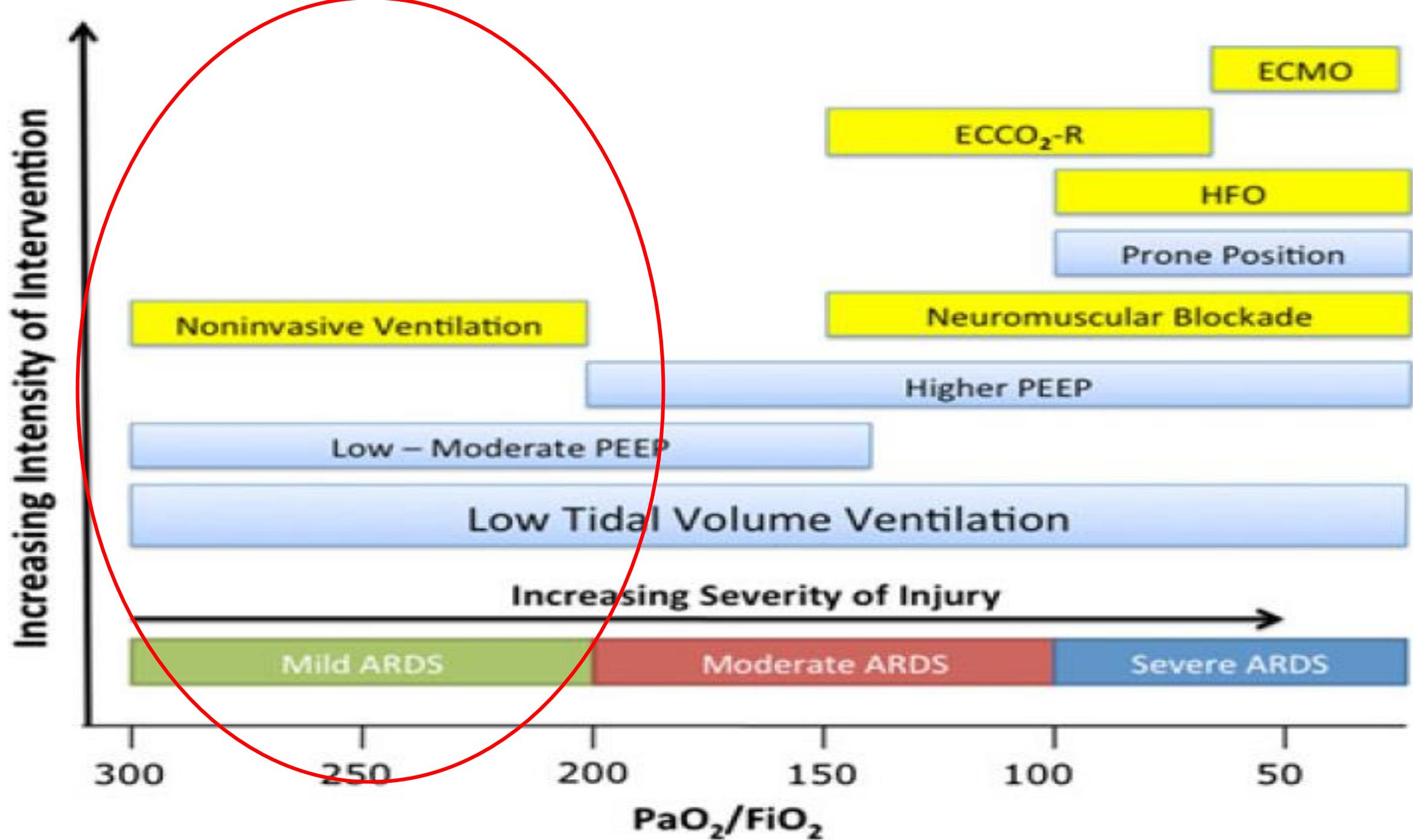
Variables	Crude		Propensity-adjusted <sup>a</sup>		Propensity-matched <sup>b</sup>	
	Odds ratio (95 % CI)	P value <sup>c</sup>	Odds ratio (95 % CI)	P value <sup>c</sup>	Odds ratio (95 % CI)	P value <sup>c</sup>
Primary outcome						
Overall ICU mortality	0.323 (0.158–0.658)	0.002	0.317 (0.143–0.700)	0.005	0.369 (0.139–0.984)	0.046
Secondary outcomes						
Extubation success	3.284 (1.361–7.923)	0.008	3.091 (1.193–8.013)	0.020	2.057 (0.746–5.672)	0.163
Ventilator-weaning	3.056 (1.470–6.351)	0.003	3.380 (1.492–7.656)	0.004	2.495 (1.039–5.991)	0.041
Ventilator-free days to day 28	0.542 (0.383–0.768) <sup>d</sup>	0.001 <sup>e</sup>	0.516 (0.349–0.763) <sup>d</sup>	0.001 <sup>e</sup>	0.639 (0.431–0.946) <sup>d</sup>	0.026 <sup>e</sup>
14-Day mortality from HFNC application	0.949 (0.455–1.977)	0.888	0.712 (0.312–1.622)	0.418	0.608 (0.231–1.606)	0.316
14-Day mortality from intubation	0.653 (0.325–1.311)	0.231	0.482 (0.218–1.067)	0.072	0.447 (0.168–1.184)	0.105
28-Day mortality from HFNC application	0.820 (0.416–1.616)	0.566	0.680 (0.318–1.457)	0.322	0.896 (0.440–1.824)	0.763
28-Day mortality from intubation	0.571 (0.287–1.138)	0.111	0.557 (0.258–1.198)	0.134	0.802 (0.380–1.692)	0.563
Length of ICU stay	0.827 (0.586–1.169) <sup>f</sup>	0.282 <sup>g</sup>	0.830 (0.552–0.800) <sup>f</sup>	0.372 <sup>g</sup>	1.329 (0.598–2.952) <sup>f</sup>	0.485 <sup>g</sup>



# ③ Éviter l'intubation ?

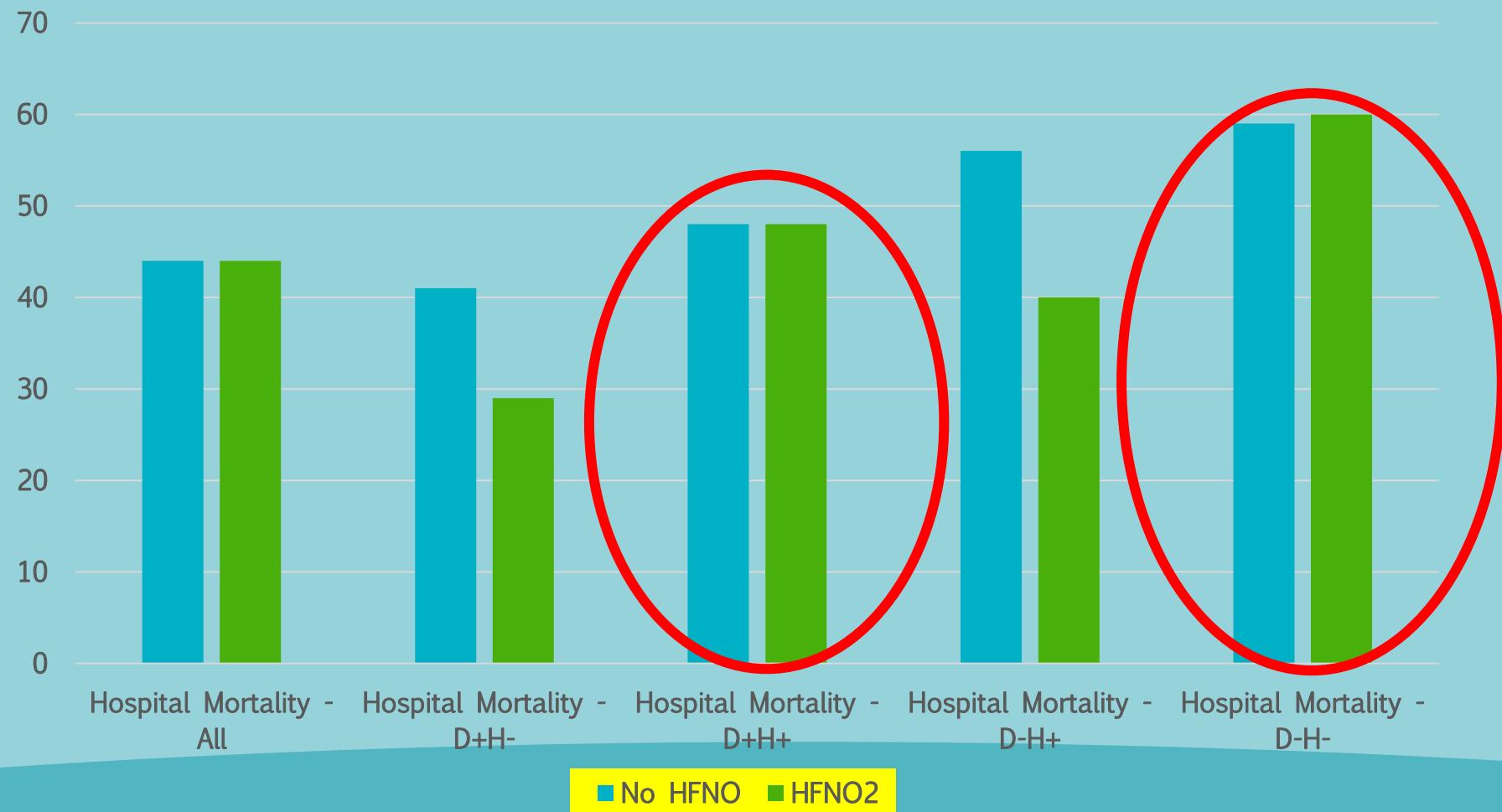
# Quelle sont les questions?

- Un *standard of care* pour les patients immunodéprimés en réanimation
- L'intubation chez les malades immunodéprimés?
- Prévenir l'intubation: bonne médecine vs. bon oxygène?
- Intubation précoce vs. Intubation tardive
- Ressusciter Helmet?



# High Flow Oxygen: a size for all?

Efraim



# Outcome for Cancer Patients Requiring Mechanical Ventilation

By Jeffrey S. Groeger, Peter White Jr, David M. Nierman, Jill Glassman, Weiji Shi, David Horak, and Kristen Price

*Journal of Clinical Oncology*, Vol 17, No 3 (March), 1999: pp 991-997

**Table 3. Results From Stepwise Logistic Regression Procedure**

Variable	Parameter Estimate	Odds Ratio	Confidence Interval	95% P
Intubated after 24 hours	0.7331	2.08	1.41-3.08	< .001
Tumor group: leukemia	0.6462	1.91	1.22-2.98	.005
Disease progression/recurrence	0.7042	2.02	1.40-2.91	< .001
Allogeneic BMT	0.8202	2.27	1.22-4.22	.01
Prior surgery with curative intent	-0.7093	0.49	0.33-0.74	< .001
Cardiac arrhythmias	0.7924	2.21	1.18-4.12	.01
DIC	1.3564	3.88	1.16-12.95	.03
Vasopressor therapy	0.6721	1.96	1.27-3.01	.002
Intercept	-0.1466			

# Outcome in Noninvasively and Invasively Ventilated Hematologic Patients With Acute Respiratory Failure\*

Pieter O. Depuydt, MD; Dominique D. Benoit, MD;

Variable	Parameter			
	Estimate	OR	95% CI	p Value
Female sex	-1.01	0.36	0.16–0.82	0.014
Intubation < 24 h	-1.25	0.29	0.11–0.78	0.015
Bacteremia < 48 h	-1.52	0.22	0.08–0.61	0.003
AML	1.004	2.73	1.05–7.11	0.04
SAPS II	0.08	1.07	1.04–1.11	< 0.001

# Patient-self inflicted lung injury (P-SILI)

- Spontaneously breathing patients with ARF have a high respiratory drive and breathe with large tidal volumes and potentially injurious transpulmonary pressures.
- In patients with existing lung injury, regional recruitment may lead to injurious effects on a remote part of the lung.
- The increase in transpulmonary pressure swings caused by inspiratory effort may contribute to the progression of lung injury.
- These patients may develop lung injury similar to VILI.
- Would the application of a lung protective ventilation, with sedation and intubation, might be considered a prophylactic therapy, rather than just a supportive therapy?

is MV able to Minimize Progression of Lung Injury in Acute Respiratory Failure.

EDITORIAL

Ventilation-induced lung injury exists in spontaneously breathing patients with acute respiratory failure: Yes

Laurent Brochard<sup>1,2\*</sup>

Intensive Care Med  
DOI 10.1007/s00134-016-4488-z



EDITORIAL

Ventilation-induced lung injury exists in spontaneously breathing patients with acute respiratory failure: No

Massimo Antonelli<sup>1\*</sup>

Intensive Care Med  
DOI 10.1007/s00134-016-4483-4



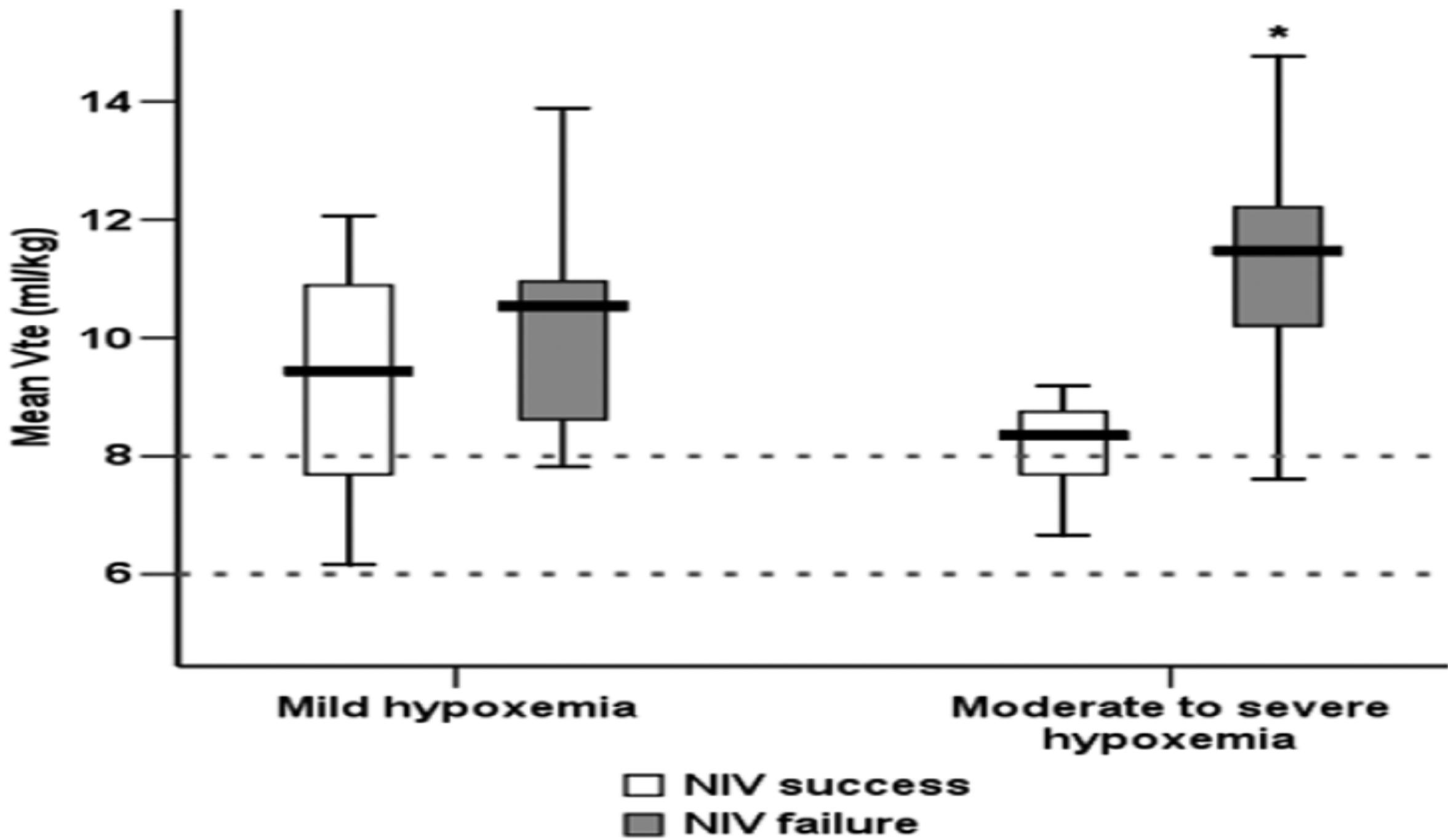
EDITORIAL

Ventilation-induced lung injury exists in spontaneously breathing patients with acute respiratory failure: We are not sure

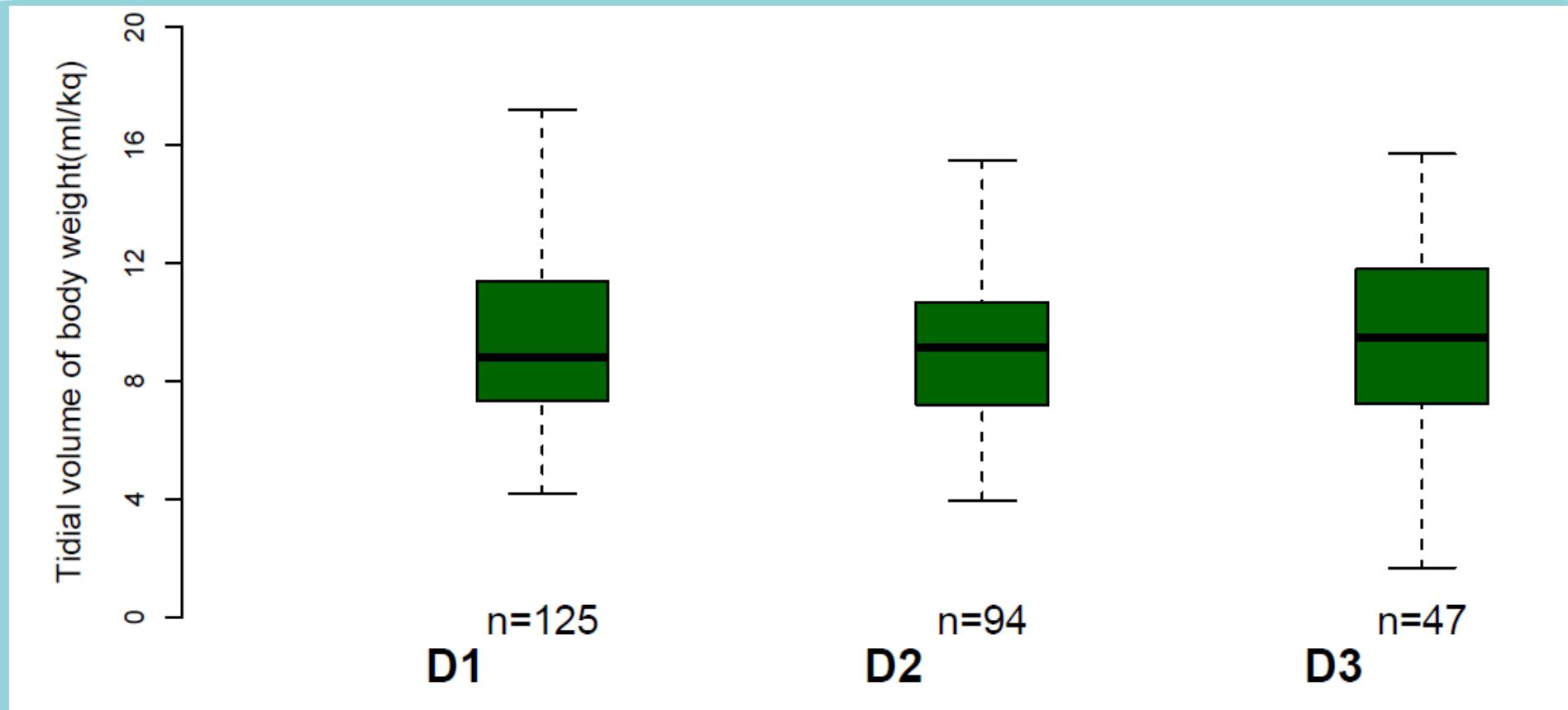


Luciano Gattinoni\*

**In the LUNG SAFE study, a propensity matched analysis, ICU mortality was higher in NIV than invasively ventilated patients with a  $\text{PaO}_2/\text{FiO}_2$  lower than 150 mmHg.**



# Reserve: Tidal volume in NIV group

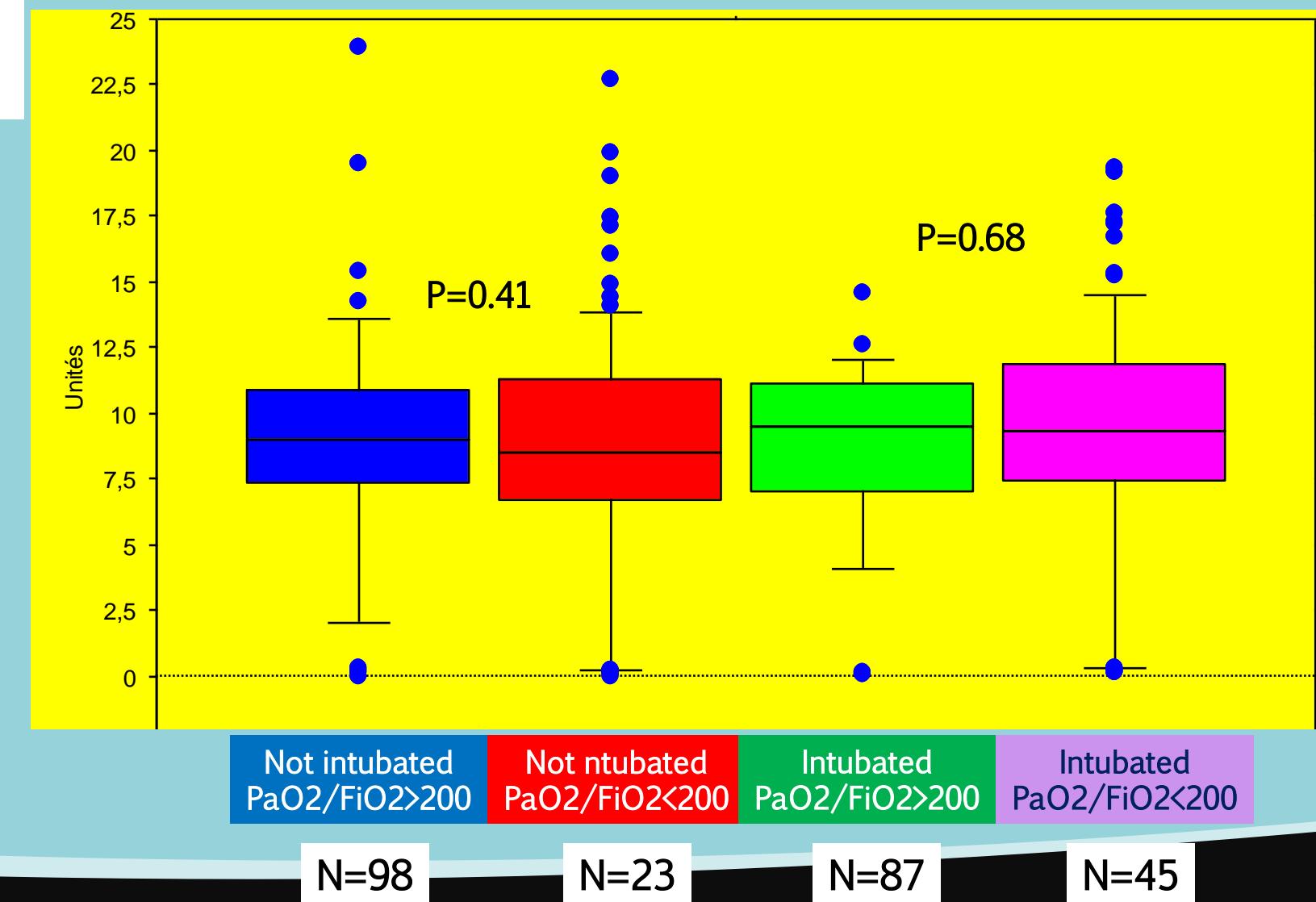


# Acute hypoxemic respiratory failure in immunocompromised patients: the Efraim multinational prospective cohort study



Elie Azoulay<sup>1\*</sup>, Peter Pickkers<sup>2</sup>, Marcio Soares<sup>3</sup>, Anders Perner<sup>4</sup>, Jordi Rello<sup>5</sup>, Philippe R. Bauer<sup>6</sup>, Andry van de Louw<sup>7</sup>, Pleun Hemelaar<sup>2</sup>, Virginie Lemiale<sup>1</sup>, Fabio Silvio Taccone<sup>8</sup>, Ignacio Martin Loeches<sup>9,10</sup>, Tine Sylvest Meyhoff<sup>4</sup>, Jorge Salluh<sup>3</sup>, Peter Schellongowski<sup>11</sup>, Katerina Rusinova<sup>12</sup>, Nicolas Terzi<sup>13</sup>, Sangeeta Mehta<sup>14</sup>, Massimo Antonelli<sup>15</sup>, Achille Kouatchet<sup>16</sup>, Andreas Barratt-Due<sup>17</sup>, Miia Valkonen<sup>18</sup>, Precious Pearl Landburg<sup>19</sup>, Fabrice Bruneel<sup>20</sup>, Ramin Brandt Bukan<sup>21</sup>, Frédéric Pène<sup>22</sup>, Victoria Metaxa<sup>23</sup>, Anne Sophie Moreau<sup>24</sup>, Virginie Soupart<sup>1</sup>, Gaston Burghi<sup>25</sup>, Christophe Girault<sup>26</sup>, Ulysses V. A. Silva<sup>27</sup>, Luca Montini<sup>15</sup>, François Barbier<sup>28</sup>, Lene B. Nielsen<sup>29,30</sup>, Benjamin Gaborit<sup>31</sup>, Djamel Mokart<sup>32</sup> and Sylvie Chevret<sup>33</sup> for the Efraim investigators and the Nine-l study group

# Efraim: ETV D1 (ml/kg ibw)

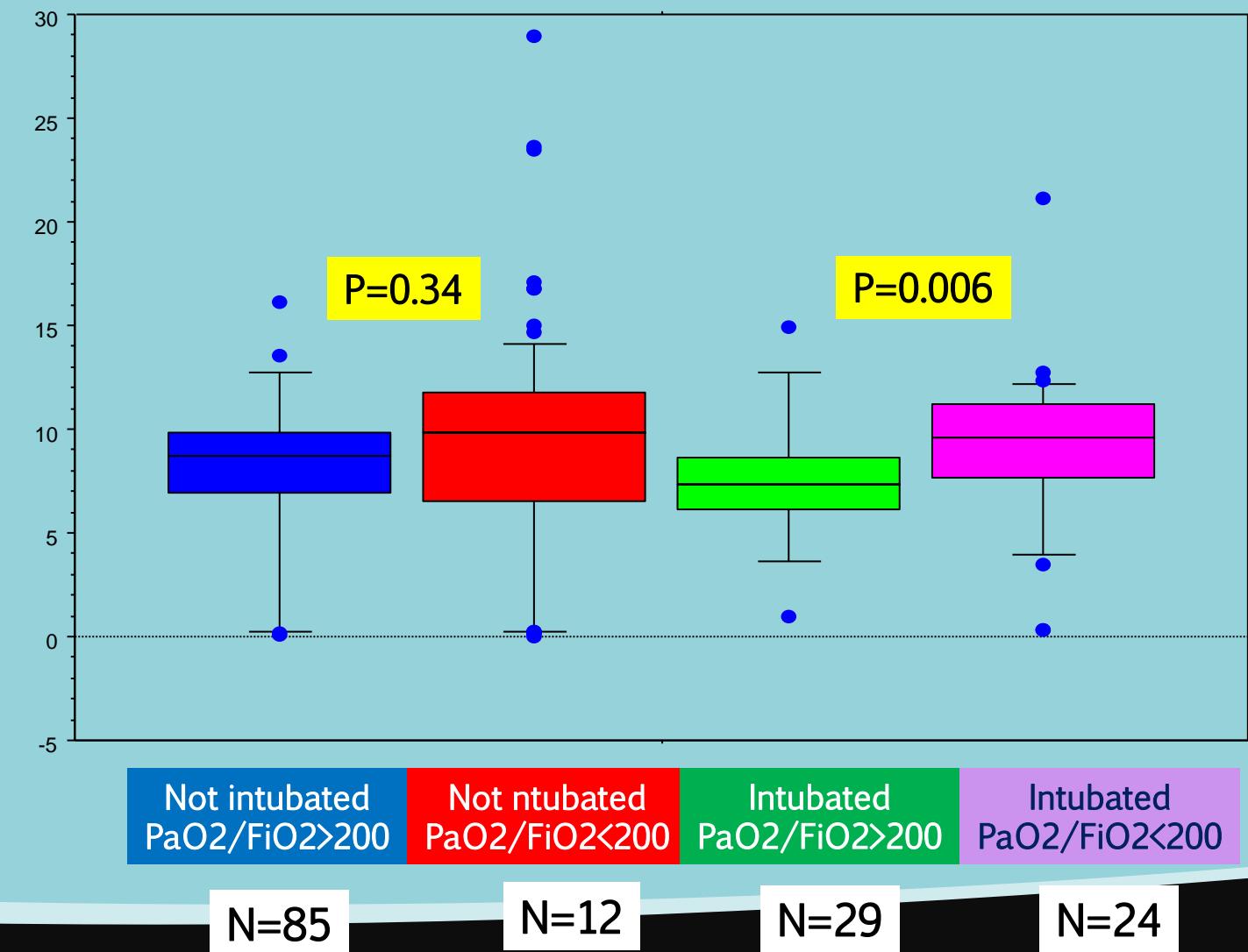




# Acute hypoxemic respiratory failure in immunocompromised patients: the Efraim multinational prospective cohort study

Elie Azoulay<sup>1\*</sup>, Peter Pickkers<sup>2</sup>, Marcio Soares<sup>3</sup>, Anders Perner<sup>4</sup>, Jordi Rello<sup>5</sup>, Philippe R. Bauer<sup>6</sup>,  
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Tine Sylvest Meyhoff<sup>4</sup>, Jorge Salluh<sup>3</sup>, Peter Schellongowski<sup>11</sup>, Katerina Rusinova<sup>12</sup>, Nicolas Terzi<sup>13</sup>,  
Sangeeta Mehta<sup>14</sup>, Massimo Antonelli<sup>15</sup>, Achille Kouatchet<sup>16</sup>, Andreas Barratt-Due<sup>17</sup>, Miia Valkonen<sup>18</sup>,  
Precious Pearl Landburg<sup>19</sup>, Fabrice Bruneel<sup>20</sup>, Ramin Brandt Bukan<sup>21</sup>, Frédéric Pène<sup>22</sup>, Victoria Metaxa<sup>23</sup>,  
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Sylvie Chevret<sup>33</sup> for the Efraim investigators and the Nine-l study group

# Efraim: ETV D2 (ml/kg ibw)



# Conclusion

Faut-il tout faire pour éviter  
l'intubation dans l'IRA hypoxémique  
du patient immunodéprimés?

Ou bien ... faut-il le protéger?  
identifier la cause de l'IRA?

# Thank you for your attention

