

AER 2019



AER

ACTUALITÉS EN RÉANIMATION

25^{ème} AER : 19 & 20 novembre 2020



Que retenir de 2019 ? Insuffisance respiratoire aiguë



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Conflicts of interest

- Fees and/or travel expenses
 - Air Liquide Santé
 - Maquet
 - Drager
 - GE
 - Faron
 - GSK
 - Covidien
 - Janssen
 - Orion
 - Johnson & Johnson
 - Peninsula Pharma

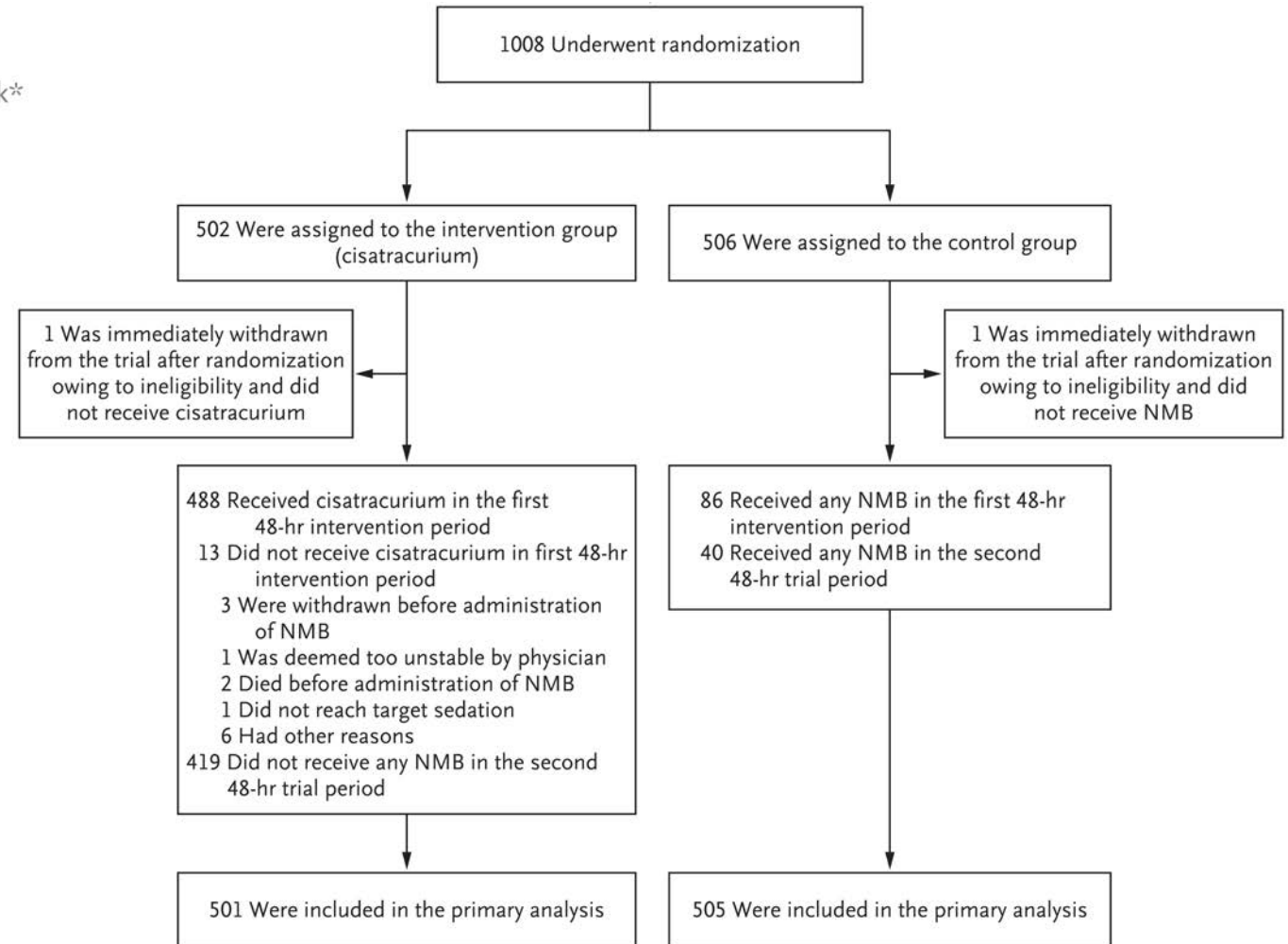
COI

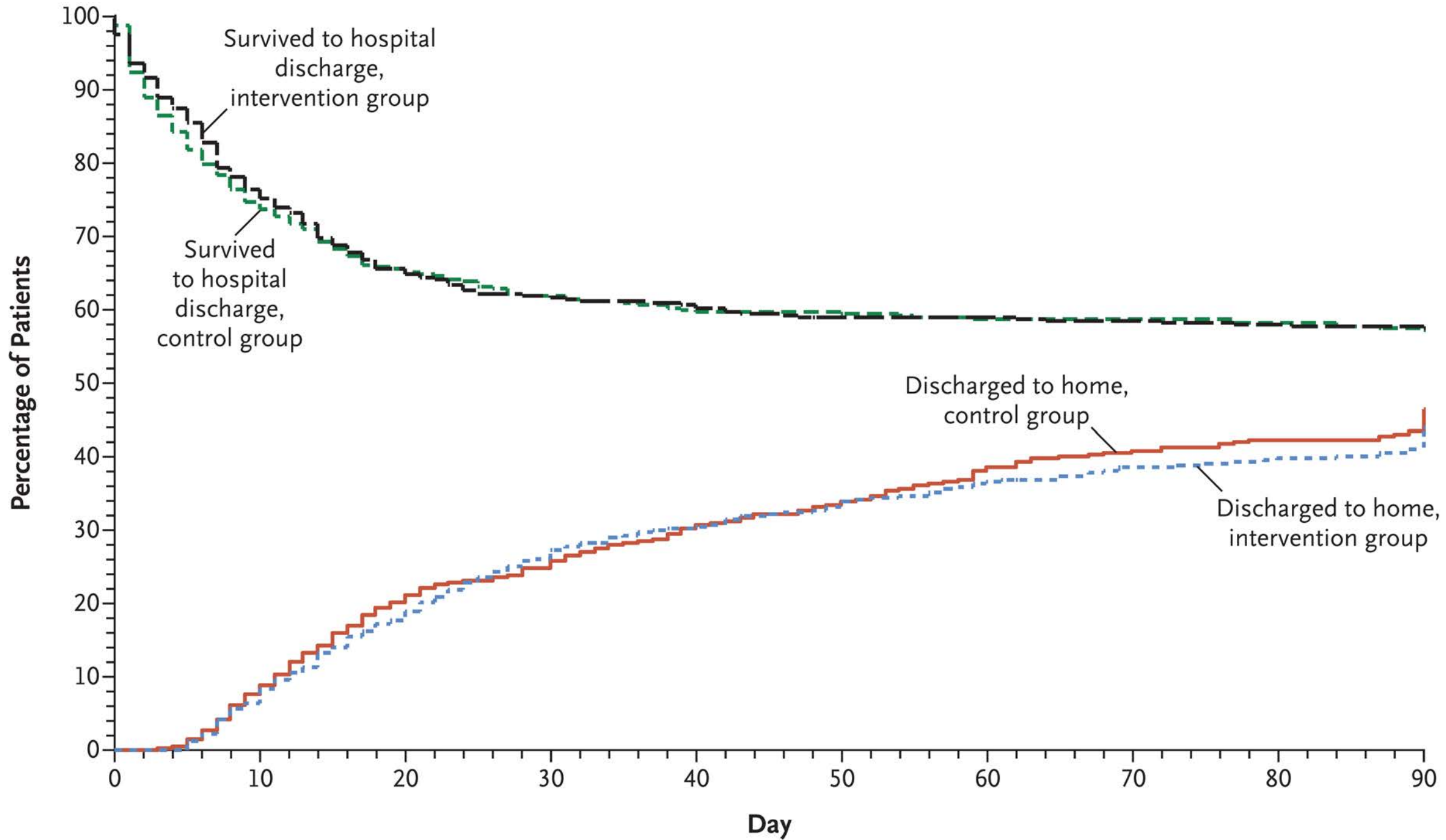
- GSK provided cisatracurium and placebo + 15k€ for ACURASYS

ORIGINAL ARTICLE

Early Neuromuscular Blockade in the Acute Respiratory Distress Syndrome

The National Heart, Lung, and Blood Institute PETAL Clinical Trials Network*





ROSE study

CLINICAL STUDY DESIGN

Design and Rationale of the Reevaluation of Systemic Early Neuromuscular Blockade Trial for Acute Respiratory Distress Syndrome

David T. Huang¹, Derek C. Angus¹, Marc Moss², B. Taylor Thompson³, Niall D. Ferguson⁴, Adit Ginde², Michelle Ng Gong⁵, Stephanie Gundel⁶, Douglas L. Hayden³, R. Duncan Hite⁷, Peter C. Hou³, Catherine L. Hough⁶, Theodore J. Iwashyna⁸, Kathleen D. Liu⁹, Daniel S. Talmor³, and Donald M. Yealy¹; for the Reevaluation of Systemic Early Neuromuscular Blockade Protocol Committee and the National Institutes of Health National Heart, Lung, and Blood Institute Prevention and Early Treatment of Acute Lung Injury Network Investigators*

Ann Am Thorac Soc Vol 14, No 1, pp 124–133, Jan 2017

Online Data Supplement

Design and Rationale of the Reevaluation of Systemic Early Neuromuscular Blockade (ROSE) Trial for Acute Respiratory Distress Syndrome

ACURASYS

- Median time from the diagnosis of ARDS to inclusion
 - 16 hrs (6-29)
- Median time from initiation of MV to inclusion
 - Cis 22 hrs (9-41)
 - Placebo 21 hrs (10-42)

ROSE

- Median of 7.6 hrs (3.7-15.6) after diagnosis of moderate-to-severe ARDS
- ?

Sedation/paralysis

ACURASYS

1. Sedation -> Ramsay 6
2. Ventilator settings
3. Cisatracurium/placebo
4. Stop at 48 hr

ROSE

Evolution of PaO₂/FiO₂ ratio

	Baseline		24h		72h	
	NMBA	Control	NMBA	Control	NMBA	Control
ACURASYS	106±36	115±41	164±72	168±72	166±70	157±68
ROSE	116±38	116±40	198±78	189±77	198±75	187±76

4848 Patients were assessed for eligibility

3840 Were excluded

658 Had $\text{PaO}_2:\text{FIO}_2 > 200$ mm Hg at time of randomization

655 Were receiving continuous NMB at enrollment

394 Declined to participate or had surrogate who declined

384 Were not expected to survive 24 hr

307 Were withdrawn by physician

270 Did not have surrogate available

245 Had been receiving mechanical ventilation for >120 hr

237 Had severe chronic liver disease

209 Had inclusion criteria for >48 hr

159 Decided to withhold life-sustaining treatment

124 Had body weight >1 kg/cm of height

113 Were receiving extracorporeal membrane oxygenation

109 Were expected to receive mechanical ventilation for <48 hr

561 Had other reason

1008 Underwent randomization

NMBA use

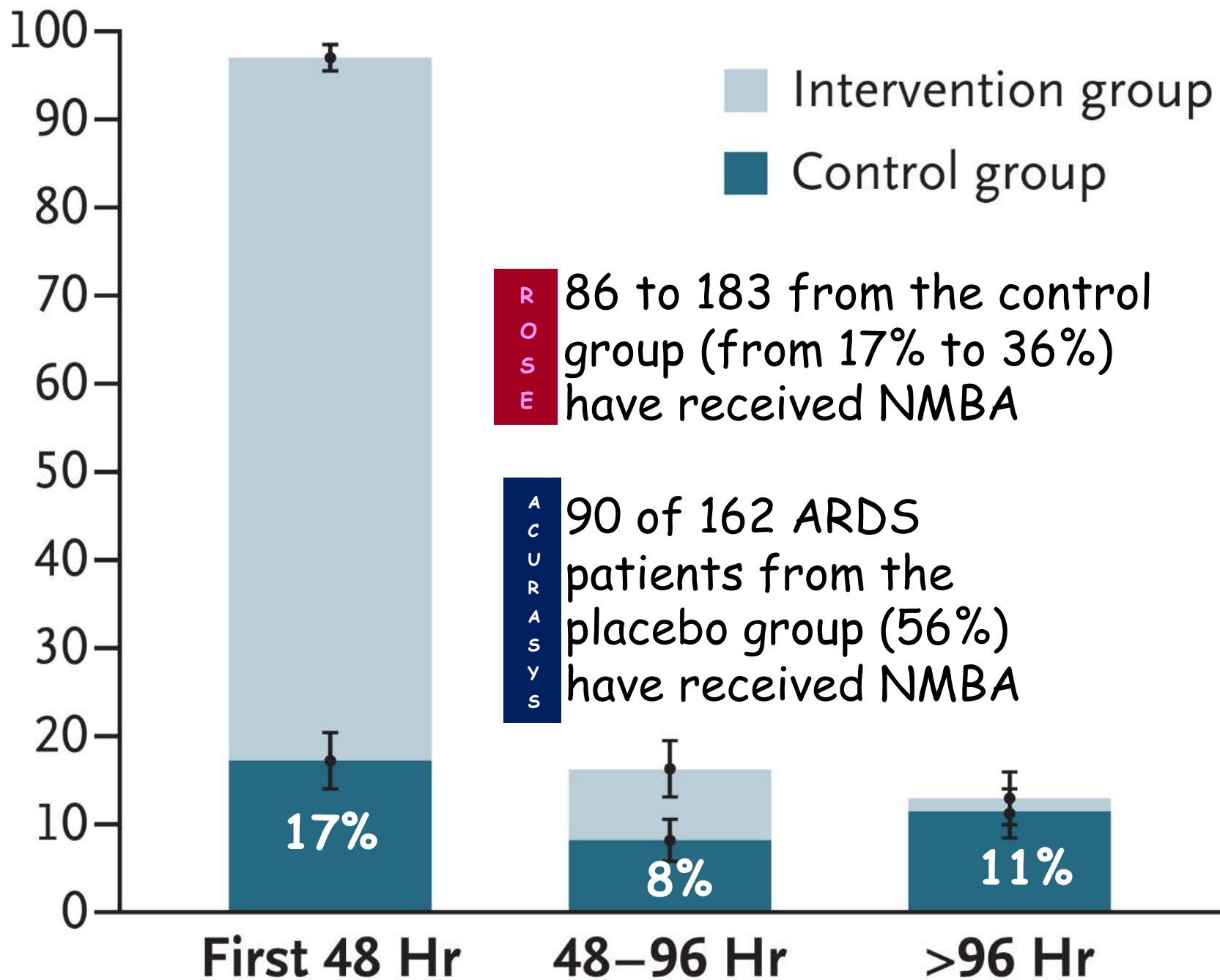
ACURASYS

- During the first 48 hrs
 - Open label NMBAs: according to the protocol
 - No stop
- After 48 hrs
 - protocol

ROSE

- During the first 48 hrs
 - Control group: protocol
 - Cis group: stop if $FiO_2 \leq 0.4$ and $PEEP \leq 8 / 12$ hrs
- After 48 hrs
 - Control group: protocol
 - Cis group: no protocol

Percentage of Patients Receiving NMB



4848 Patients were assessed for eligibility

3840 Were excluded

658 Had $P_{aO_2}:F_{iO_2} > 200$ mm Hg at time of randomization

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1008 Underwent randomization

Mechanical ventilation

ACURASYS

- 30-45°
- Volume assist-control mode
- PEEP/FiO₂ ARMA
- No RM
- Proning, iNO, almitrine

ROSE

- Volume or pressure-controlled
- PEEP/FiO₂ ALVEOLI high PEEP
- No proning for at least 12 hrs

Mechanical ventilation

ACURASYS

- 30-45°
- Volume assist-control mode
- PEEP/FiO2 ARMA
- No RM
- Proning, iNO, almitrine

ROSE

- Volume or pressure-controlled
- PEEP/FiO2 ALVEOLI high PEEP
- No proning for at least 12 hrs

ACURASYS protocol

Before increasing FiO2 from 0.50 to 0.60, you must have a PEEP set at

- 10 cmH2O in ACURASYS
- 20 cmH2O in ROSE

FiO2 %	PEEP cmH2O
30	5
40	5
40	8
50	8
50	10
60	10
70	10
70	12
70	14
80	14
90	14
90	16
90	18
100	18
100	20
100	22
100	24

TESTER SEVRABILITE
(APRES 72 h)



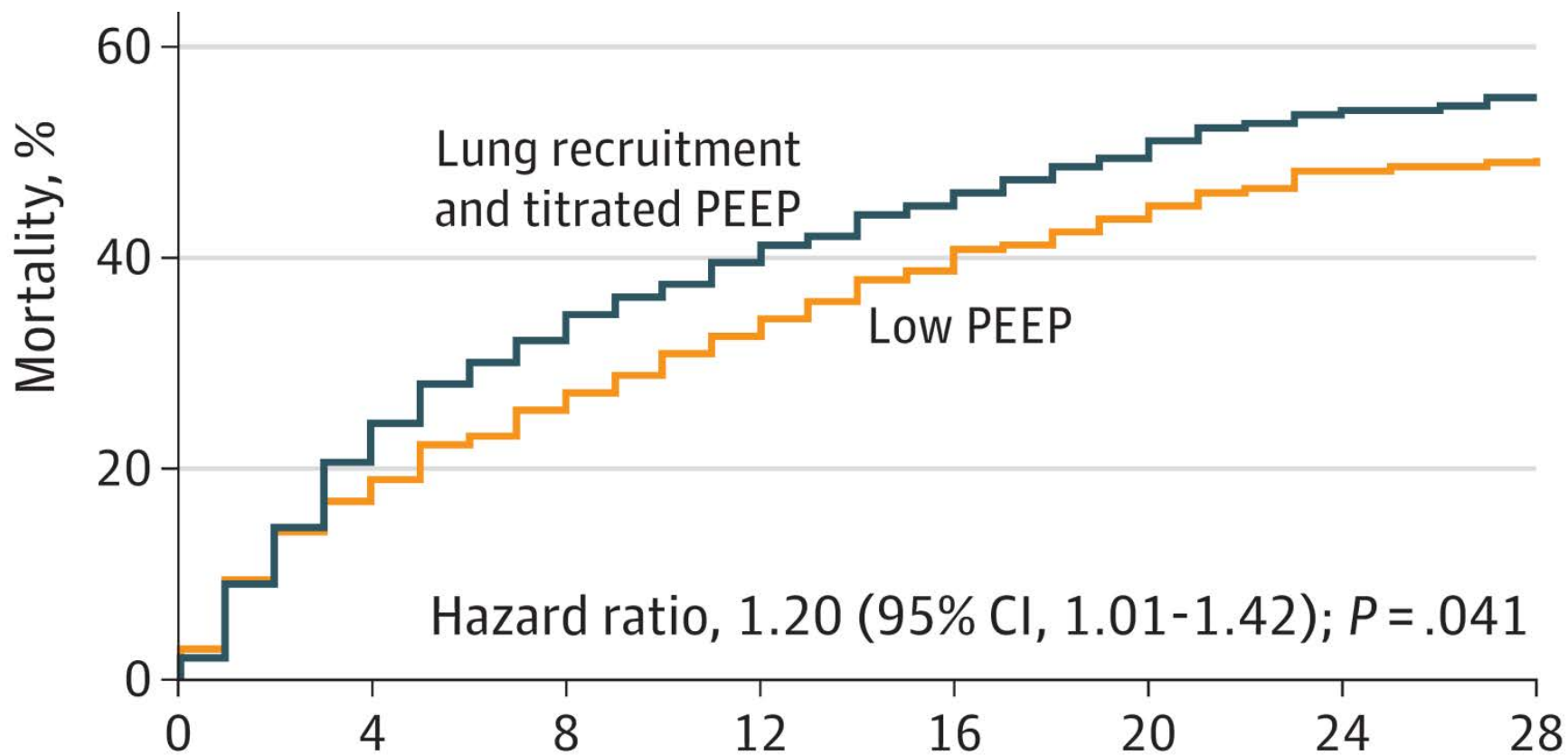
3. ROSE FiO2/PEEP table

FiO ₂	.30	.40	.40	.40	.40	.40	.40	.40	.50	.50	.50	.60	.70	.80	.80	.90	1.00	1.00
PEEP	5	5	6	8	10	12	14	16	16	18	20	20	20	20	22	22	22	24



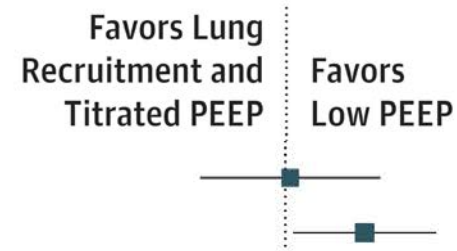
Effect of Lung Recruitment and Titrated Positive End-Expiratory Pressure (PEEP) vs Low PEEP on Mortality in Patients With Acute Respiratory Distress Syndrome: A Randomized Clinical Trial

Writing Group for the Alveolar Recruitment for Acute Respiratory Distress Syndrome Trial (ART) Investigators



Type of ARDS

Extrapulmonary	98/188 (52.1)	102/196 (52)	1.02 (0.74-1.40)
Pulmonary	179/313 (57.2)	149/313 (47.6)	1.32 (1.03-1.69)



On inclusion

	cisatracurium	placebo
Proning, iNO/alm	18.6%	14.2%
Corticosteroids for septic shock	39.5%	45.1%

	cisatracurium	control
Proning, iNO/alm		
Corticosteroids for septic shock		

During the study period

	cisatracurium	placebo
Proning	28%	29%
Proning, iNO/alm	42%	48%
Vasopressors	92%	89%

	cisatracurium	control
Proning		
Proning, iNO/alm		
Vasopressors		

Received NMBA=44%

	NMBA n = 177 no (%)	Placebo n= 162 no (%)	P value
Adjunctive therapies			
<i>Prone position</i>	50 (28%)	47 (29%)	0.88
<i>Inhaled nitric oxide</i>	50 (28%)	53 (33%)	0.37
<i>Almitrine bismesylate</i>	6 (3%)	10 (6%)	0.23
<i>Any of the three treatments above</i>	75 (42%)	77 (48%)	0.34

Characteristic	Day 0-2			Day 0-28		
	Intervention	Control	Difference (95% CI)	Intervention	Control	Difference (95% CI)
Any rescue therapy	93 (18.6)	90 (17.8)	0.7 (-4.0, 5.5)	130 (25.3)	125 (24.8)	1.2 (-4.2, 6.6)
Prone positioning	68 (13.6)	60 (11.9)	1.7 (-2.4, 5.8)	84 (16.8)	75 (14.9)	1.9 (-2.6, 6.4)
Inhaled epoprostenol	16 (3.2)	17 (3.4)	-0.2 (-2.4, 2.0)	26 (5.2)	27 (5.3)	-0.2 (-2.9, 2.6)
Recruitment maneuvers	14 (2.8)	16 (3.2)	-0.4 (-2.5, 1.7)	29 (5.8)	30 (5.9)	-0.2 (-3.1, 2.8)
Inhaled nitric oxide	4 (0.8)	12 (2.4)	-1.6 (-3.1, 0.0)	7 (1.4)	17 (3.4)	-2.0 (-3.8, -0.1)
ECMO	2 (0.4)	3 (0.6)	-0.2 (-1.1, 0.7)	3 (0.6)	10 (2.0)	-1.4 (-2.8, 0.0)

Weaning protocol

ACURASYS

- From **day 3** when $FiO_2 \leq 0.6$ (PEEP 10) => PS and \Downarrow PEEP to 5 => SBT
- Transition from ACV to SB = PS
 - PS 20-15-10-5 to maintain RR 26-35/min and $SpO_2 \geq 88\%$

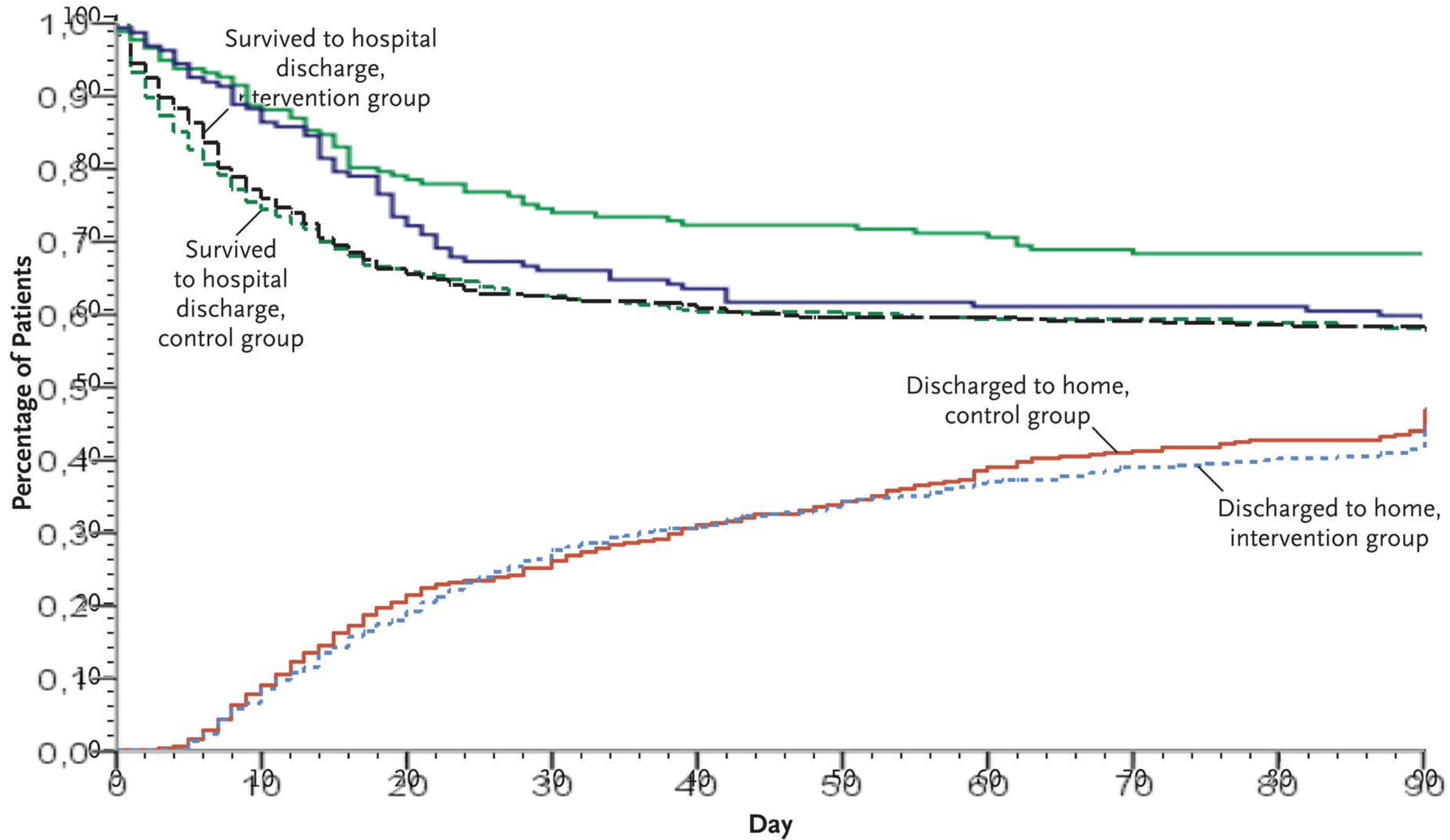
ROSE

• ACURASYS

Outcome	Cisatracurium (N=177)	Placebo (N=162)	Relative Risk with Cisatracurium (95% CI)	P Value
Death — no. (% [95% CI])				
At 28 days	42 (23.7 [18.1–30.5])	54 (33.3 [26.5–40.9])	0.71 (0.51–1.00)	0.05
In the ICU	52 (29.4 [23.2–36.5])	63 (38.9 [31.7–46.6])	0.76 (0.56–1.02)	0.06
In the hospital	57 (32.2 [25.8–39.4])	67 (41.4 [34.1–49.1])	0.78 (0.59–1.03)	0.08
No. of ventilator-free days†				
From day 1 to day 28	10.6±9.7	8.5±9.4		0.04

• ROSE

Variable	Intervention Group (N=501)	Control Group (N=505)	Between-Group Difference (95% CI)	P Value
<i>percentage points</i>				
Primary end point: in-hospital death by day 90 — no. (%)†	213 (42.5±2.2)	216 (42.8±2.2)	−0.3 (−6.4 to 5.9)	0.93
Secondary end points				
In-hospital death by day 28 — no. (%)	184 (36.7)	187 (37.0)	−0.3 (−6.3 to 5.7)	
Days free of ventilation at day 28‡	9.6±10.4	9.9±10.9	−0.3 (−1.7 to 1.0)	



Statistics

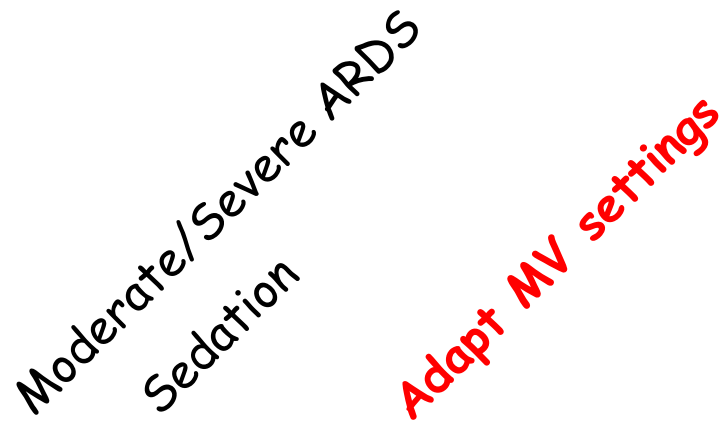
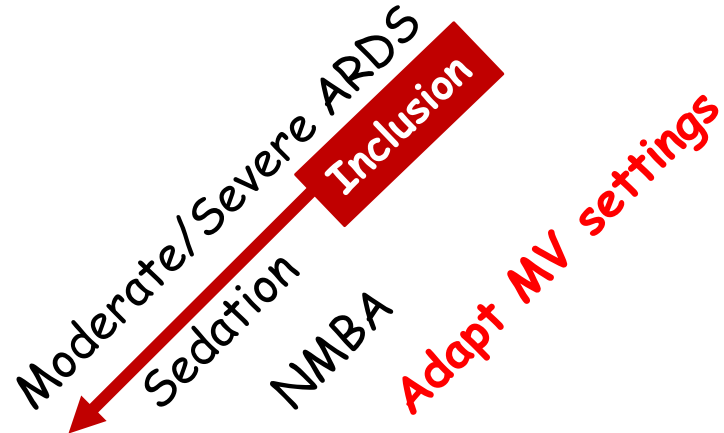
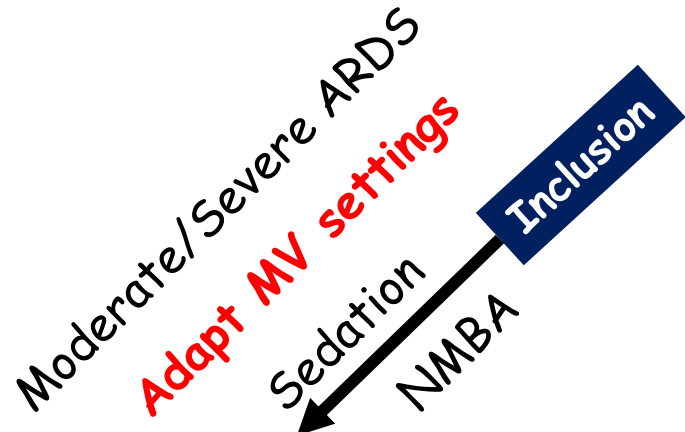
	Active Superior		Control Superior		Probability of Stopping	
Number of Subjects	Observed Mortality Difference Active-Control	P-value*	Observed Mortality Difference Active-Control	P-value*	Under the Null Hypothesis	Under the Alternative Hypothesis
470	-0.146	0.00031	0.146	0.99969	0.001	0.061
938	-0.078	0.00479	0.078	0.99521	0.010	0.528
1408	-0.049	0.02361	0.049	0.97639	0.050	0.900

* = These are one sided p values for the upper and lower boundaries.

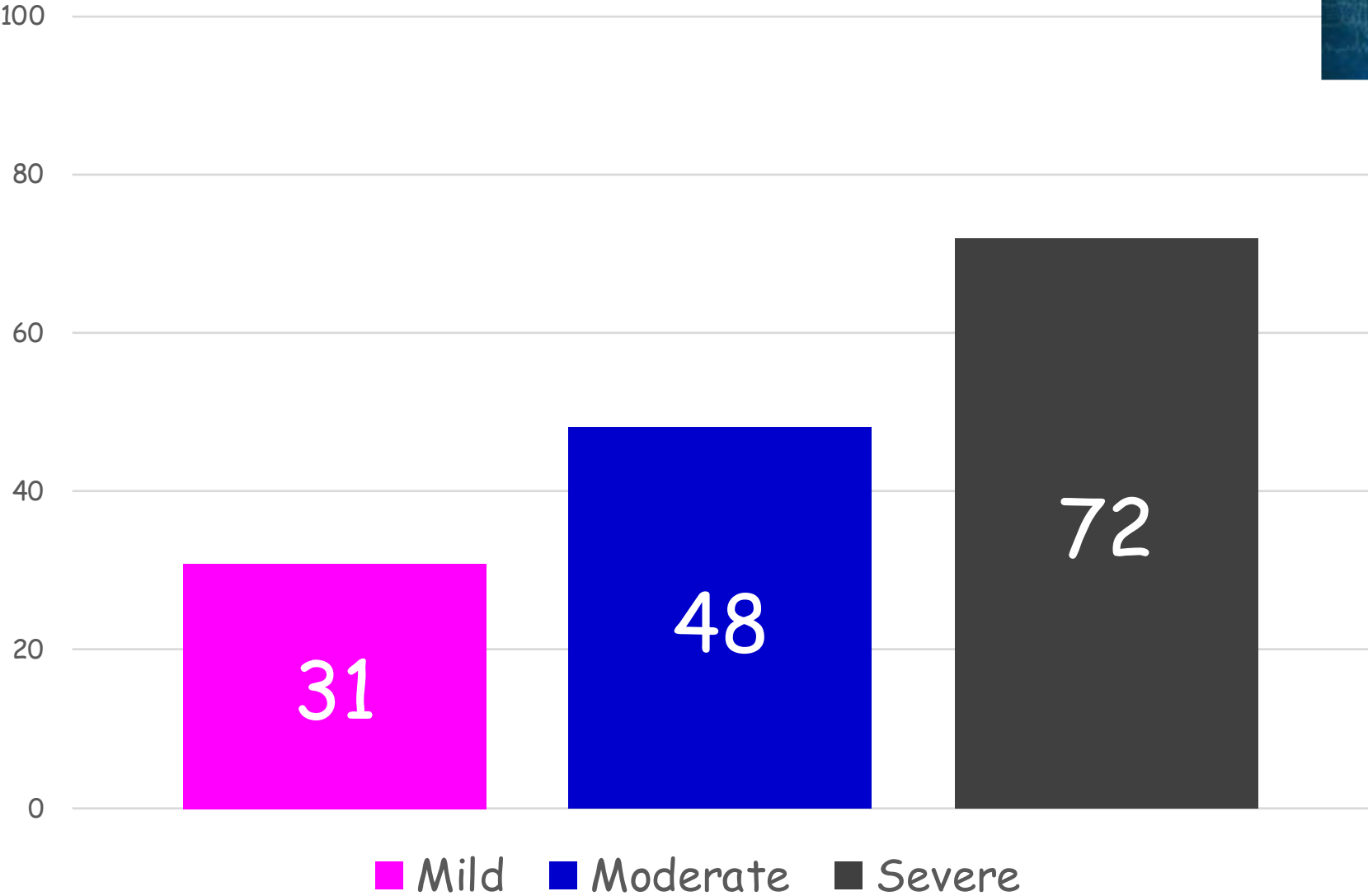
ACURASYS strategy

ROSE strategy

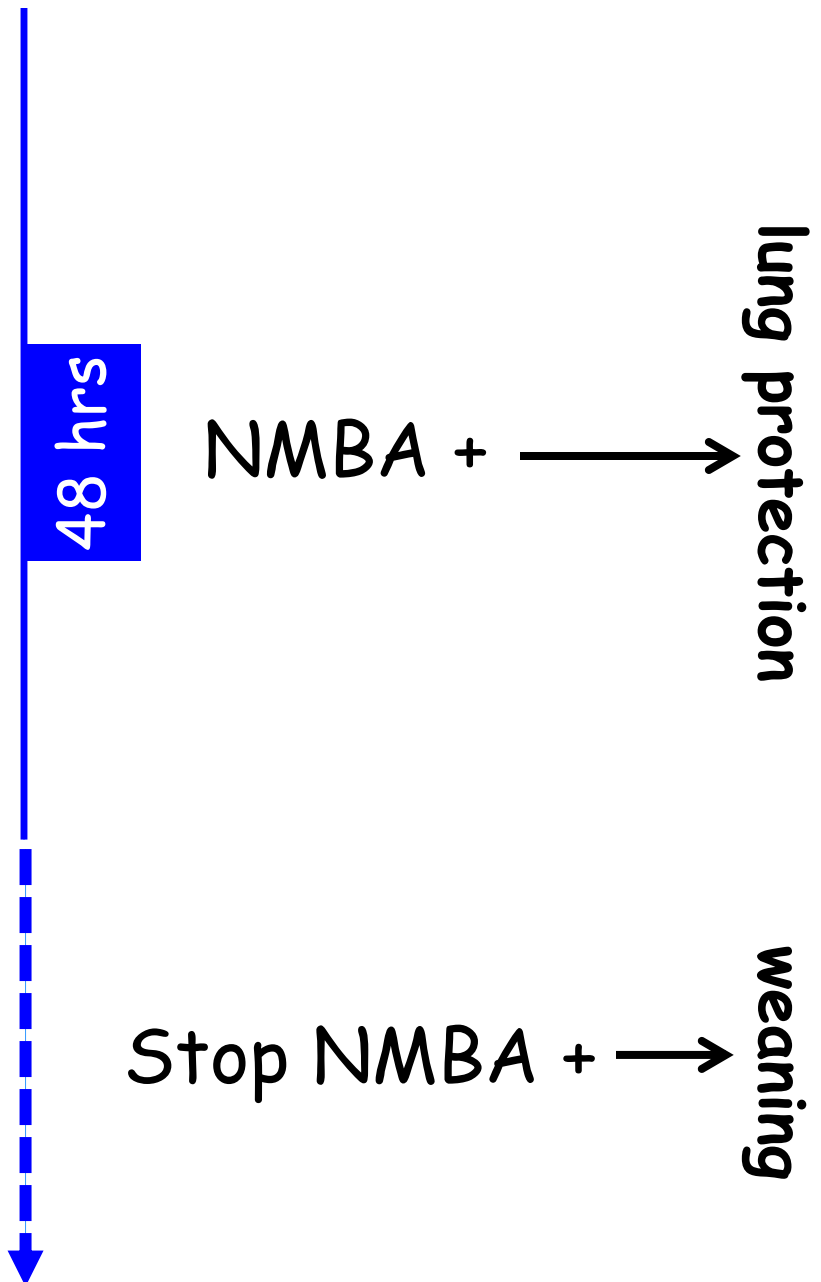
NMBA "rescue" strategy



LUNG SAFE - France - Curare



ACURASYS strategy



Variable
Ventilator mode: volume assist-control
Initial tidal volume: 6-8 ml/kg of predicted body weight
Plateau pressure: ≤ 32 cm of water
Plateau pressure limitation
ARMA PEEP/FiO2 Table «low PEEP»
iNO, almitrine, prone position
Control hypercapnia
jection of cisatracurium in a bolus of 20 mg (not to be given again if plateau pressure decreased by < 2 cm of water because further doses would probably be futile, but permitted if the drug had its intended effect)
Procedure to correct hypercapnia when pH is < 7.20 (in the following order, as needed): connect Y-piece directly to endotracheal tube, increase respiratory rate to a maximum of 35 cycles per min, and increase tidal volume to a maximum of 8 ml/kg
Weaning attempt: starting on day 3 if $FiO_2 < 0.6$
3rd day: ↘ PEEP
PS 20-15-10-5
SV



Feasibility and safety of ultra-low tidal volume ventilation without extracorporeal circulation in moderately severe and severe ARDS patients

J. C. Richard^{1,2,3*}, S. Marque⁴, A. Gros⁵, M. Muller⁶, G. Prat⁷, G. Beduneau^{8,9}, J. P. Quenot¹⁰, J. Dellamonica¹¹, R. Tapponnier¹², E. Soum¹³, L. Bitker^{1,2,3}, J. Richecoeur¹⁴ and the REVA research network

Ventilatory mode: volume-assist control

Instrumental dead space: minimize by using a heated humidifier and a low-volume endotracheal tube connector

Initial VT: stepwise reduction by 1 ml.kg⁻¹ PBW steps at intervals ≤ 2 h down to 4 ml.kg⁻¹ PBW

RR: increase up to 40 min⁻¹ to maintain VE constant (35 min⁻¹ if intrinsic PEEP > 2 cm H₂O)

Ratio of the duration of inspiration to the duration of expiration: adjust between 1:2 and 1:4 to maintain intrinsic PEEP ≤ 2 cm H₂O

Ventilatory goals: plateau pressure ≤ 30 cm H₂O; $55 \leq \text{PaO}_2 \leq 80$ mm Hg or $88\% \leq \text{SpO}_2 \leq 95\%$; $7.20 \leq \text{pH} \leq 7.45$

Allowable combinations of PEEP (cm of H₂O) and FiO₂: 5 and 30%, 8 and 30%, 10 and 30%, 12 and 30%, 14 and 30%, 14 and 40%, 16 and 40%, 16 and 50%, 18 and 50%, 20 and 50%, 20 and 60%, 20 and 70%, 20 and 80%, 22 and 80%, 22 and 90%, 22 and 100%, 24 and 100%

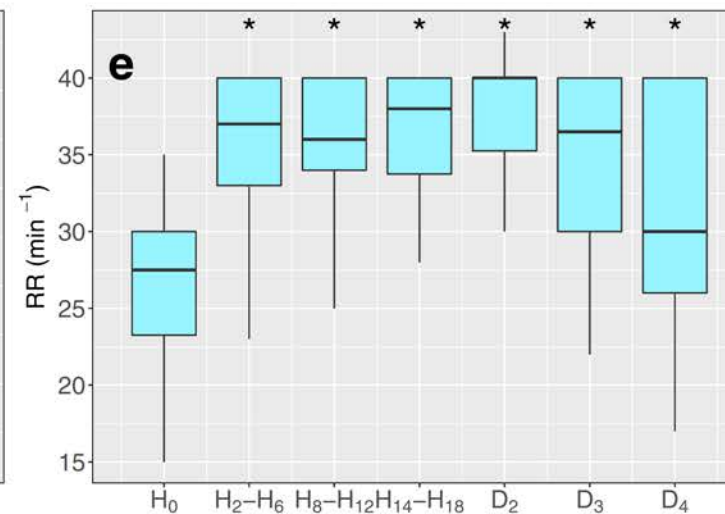
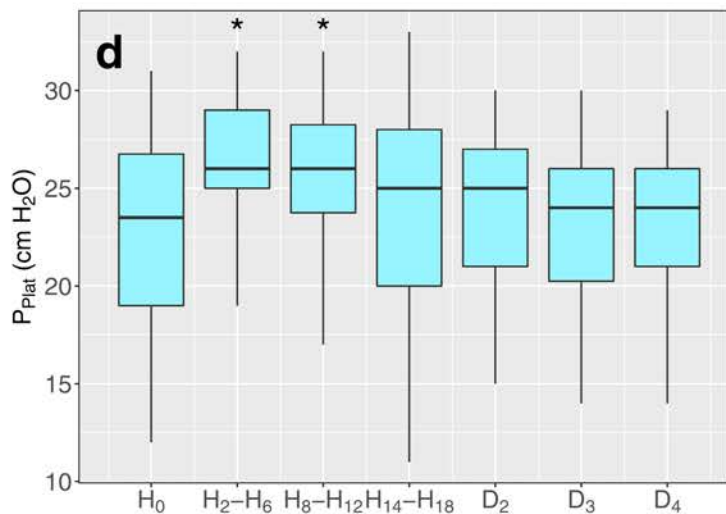
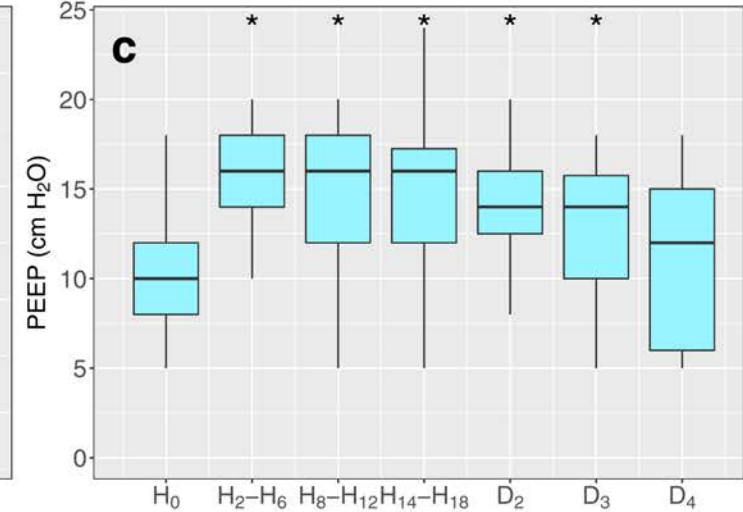
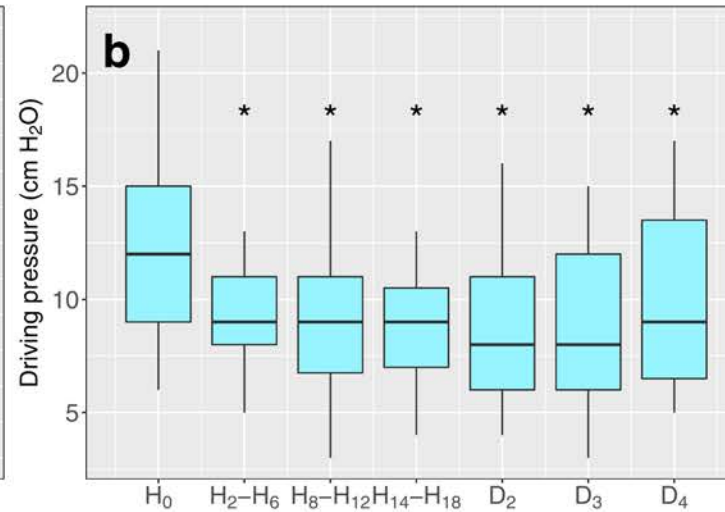
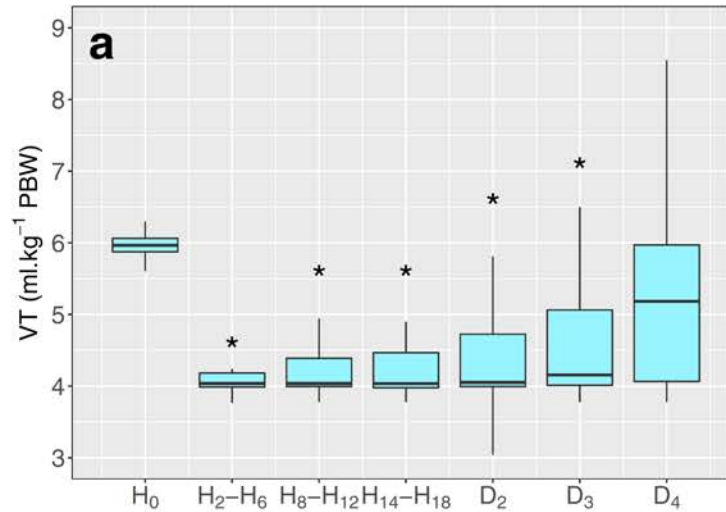
Procedure when PaO₂ < 55 mm Hg despite adjustments of FiO₂ and PEEP (in the following order as needed): (1) use PP if PaO₂/FiO₂ < 150 mm Hg with PEEP > 10 cm H₂O and FiO₂ > 60%; (2) add NMBA; (3) add iNO; (4) consider ECMO

Procedure when PaO₂ > 80 mm Hg (in the following order as needed): (1) stop iNO; (2) stop NMBA if administration > 48 h; (3) adjust FiO₂ and PEEP

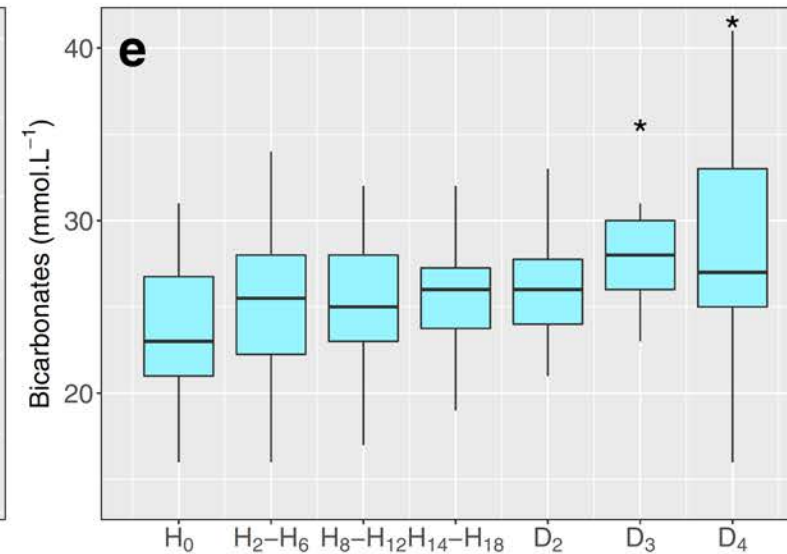
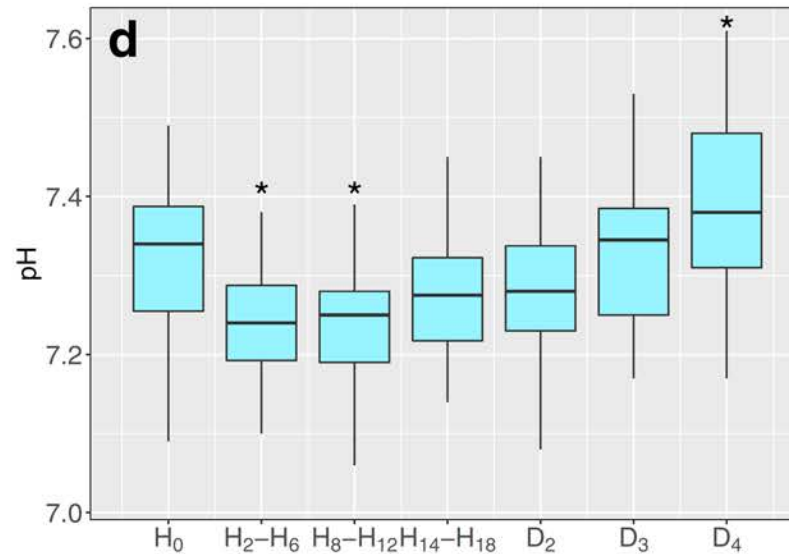
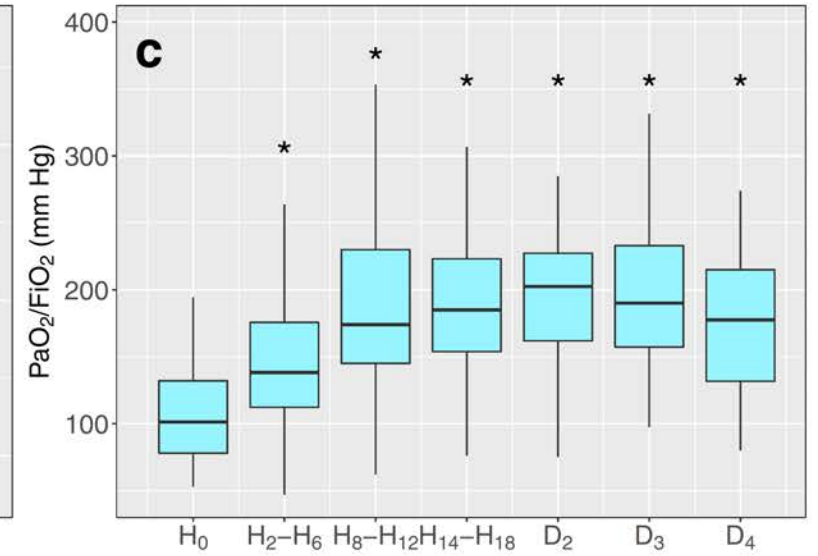
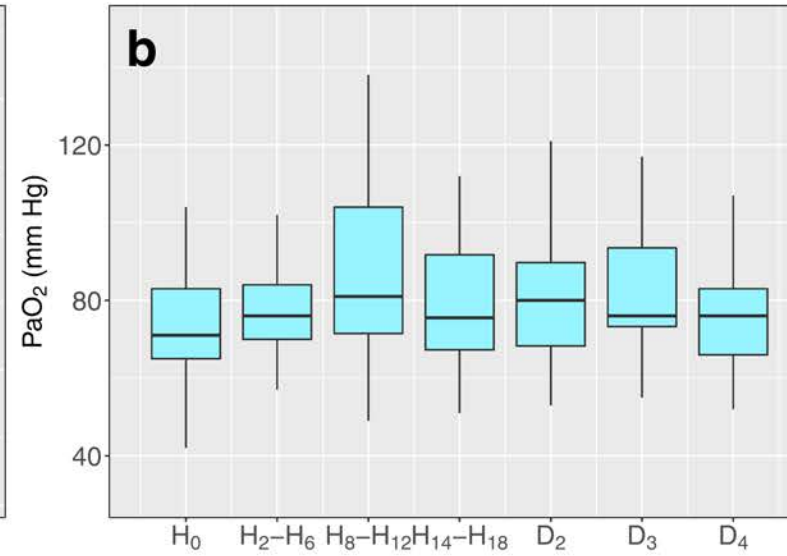
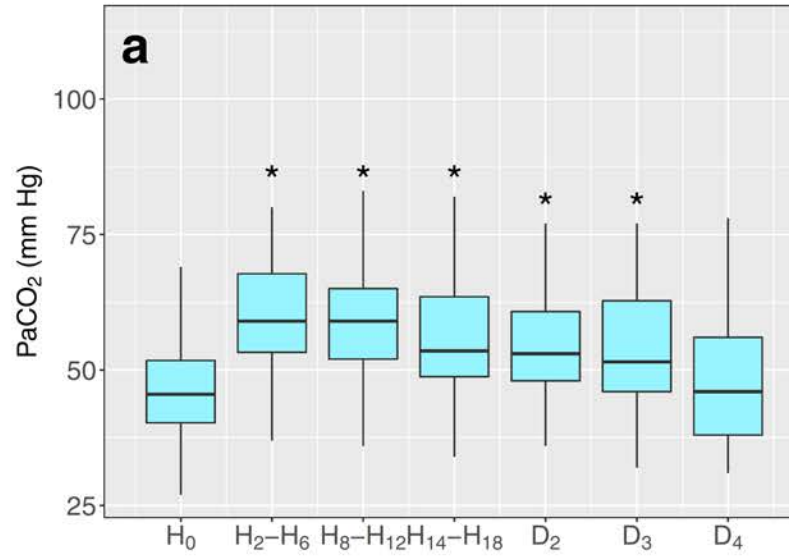
Procedure when plateau pressure is > 30 cm H₂O (in the following order as needed): (1) inject a bolus of NMBA; (2) reduce VT to 4 ml.kg⁻¹ PBW (if pH ≥ 7.2); (3) decrease PEEP down to a minimum of 5 cm H₂O

Procedure when pH < 7.20 (in the following order as needed): (1) increase sedation/NMBA dose to achieve good patient-ventilator synchrony; (2) increase RR up to 40 min⁻¹ (35 min⁻¹ if total PEEP > 2 cm H₂O); (3) may administer IV bicarbonate; (4) increase VT by 1 ml.kg⁻¹ PBW step up to 8 ml.kg⁻¹ PBW if pH < 7.15; (5) consider ECCO₂R or ECMO

Ventilatory parameters over the first 4 days following inclusion



Arterial blood gas over the first 4 days following inclusion



Online Resource 4. Physiological parameters over the first four days following inclusion.

Variables	H ₀	H ₂ -H ₆	H ₈ -H ₁₂	H ₁₄ -H ₁₈	D ₂	D ₃	D ₄
Patients alive under MV	34	34	34	34	34	31	31
ΔP (cm H ₂ O)	12 [9-15]	9 [8-11]†	9 [7-11]†	9 [7-11]†	8 [6-11]†	8 [6-12]†	9 [7-14]†
VT (mL.kg ⁻¹ PBW)	6.0 [5.9-6.1]	4.0 [4.0-4.2]†	4.0 [4.0-4.4]†	4.0 [4.0-4.5]†	4.1 [4.0-4.7]†	4.2 [4.0-5.1]†	5.3 [4.1-6.2]
VT < 4.2 mL.kg ⁻¹ PBW	0 (0%)	26 (76%)	23 (68%)	23 (68%)	22 (65%)	16 (52%)	9 (29%)
VT < 5.25 mL.kg ⁻¹ PBW	2 (6%)	32 (94%)	31 (91%)	28 (82%)	30 (88%)	24 (77%)	14 (45%)
RR (min ⁻¹)	28 [23-30]	37 [33-40]†	36 [34-40]†	38 [34-40]†	40 [35-40]†	37 [30-40]†	30 [26-40]†
Inspiratory time (s)	0.66 [0.60-0.86]	0.50 [0.45-0.58]†	0.50 [0.40-0.56]†	0.50 [0.36-0.56]†	0.50 [0.40-0.55]†	0.50 [0.45-0.62]	0.50 [0.44-0.66]
PEEP (cm H ₂ O)	10 [8-12]	16 [14-18]†	16 [12-18]†	16 [12-17]†	14 [13-16]†	14 [10-16]†	12 [6-15]
PEEPi (cm H ₂ O)	1 [0-1]	1 [0-1]	1 [0-1]	1 [0-1]	1 [0-2]	1 [0-1]	1 [0-1]
Plateau pressure (cm H ₂ O)	24 [19-27]	26 [25-29]†	26 [24-29]†	25 [20-28]	25 [21-27]	24 [20-26]	24 [21-26]
pH	7.34 [7.26-7.39]	7.24 [7.19-7.29]†	7.25 [7.19-7.28]†	7.28 [7.22-7.32]	7.28 [7.23-7.34]	7.35 [7.25-7.39]	7.38 [7.31-7.48]†
PaCO ₂ (mm Hg)	46 [40-52]	59 [53-68]†	59 [52-65]†	54 [49-64]†	53 [48-61]†	52 [46-63]†	46 [38-56]
PaO ₂ /FiO ₂ (mm Hg)	101 [78-132]	138 [112-176]†	174 [145-230]†	185 [154-223]†	202 [162-227]†	190 [157-233]†	178 [132-215]†
Bicarbonates (mmol/L ⁻¹)	23 [21-27]	26 [22-28]	25 [23-28]	26 [24-27]	26 [24-28]	28 [26-30]†	27 [25-33]†
Base excess (mmol/L ⁻¹)	-1.8 [-5.7-1.8]	-2.5 [-5.2-0.9]	-2.7 [-5.6-0.4]	-1.6 [-3-1.3]	-1.1 [-3-1.5]	0.5 [-1.9-4.9]†	2.0 [-1.2-8.2]†
Arterial lactate (mmol.L ⁻¹)	1.9 [1.3-2.8]	NR	NR	NR	1.6 [1.2-2.0]	1.4 [1.1-1.6]†	1.6 [1.3-2.0]

Values are median [IQR] or count (%); † p < 0.05 vs. H₀.

Adverse events	Number of episodes	Number of patients (%)
Metabolic events		
Severe mixed acidosis with pH < 7.15	16	11 (32%)
Other metabolic events	1	1 (3%)
Respiratory events		
Pneumothorax	2	2 (6%)
Refractory hypoxemia requiring ECMO	1	1 (3%)
Other	6	6 (18%)
Infectious events		
Nosocomial pneumonia	14	13 (38%)
Non-respiratory infection site	4	3 (9%)
Bacteremia	8	7 (21%)
Cardiovascular events		
Shock	5	5 (15%)
Cardiac arrest	3	3 (9%)
Acute cor pulmonale	2	2 (6%)
Supraventricular tachycardia	3	2 (6%)

Multivariate analysis of variables associated with severe mixed acidosis

Variables	OR [CI _{95%}]
Renal SOFA sub-score at inclusion (per 1 unit increase)	1.91 [1.08-3.71]
pH at inclusion (per 0.01 unit increase)	0.91 [0.80-0.99]

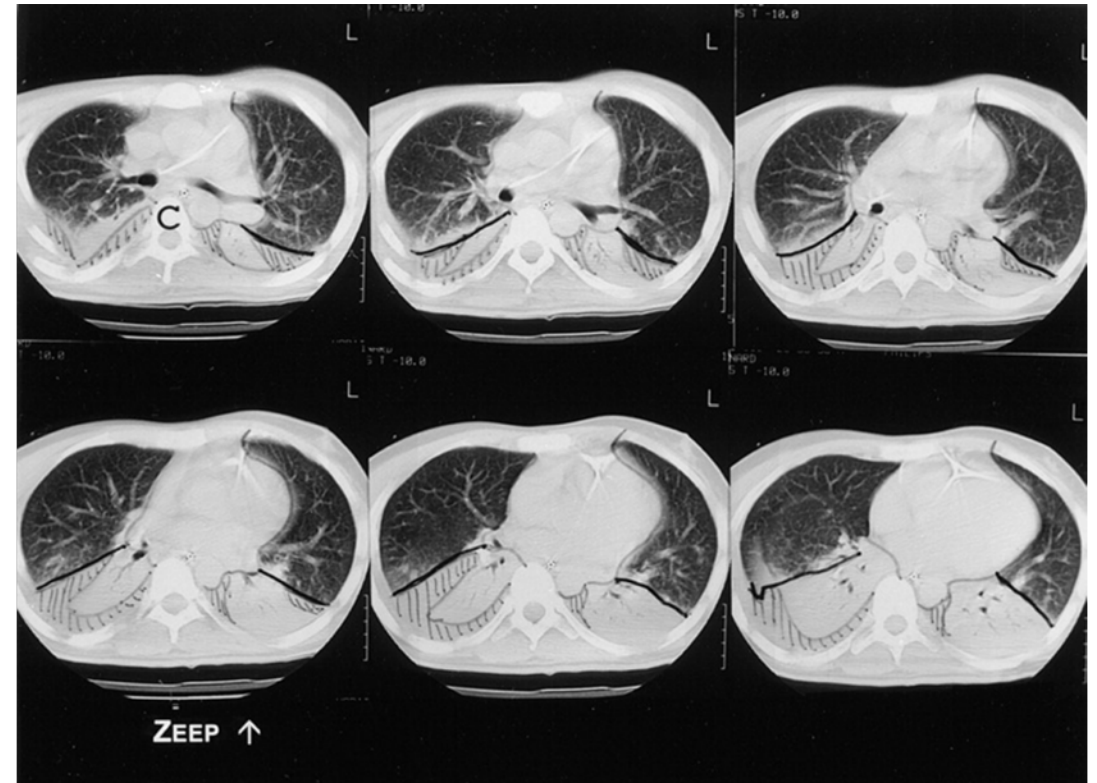
CI_{95%} = 95% confidence interval; OR = odd ratio.

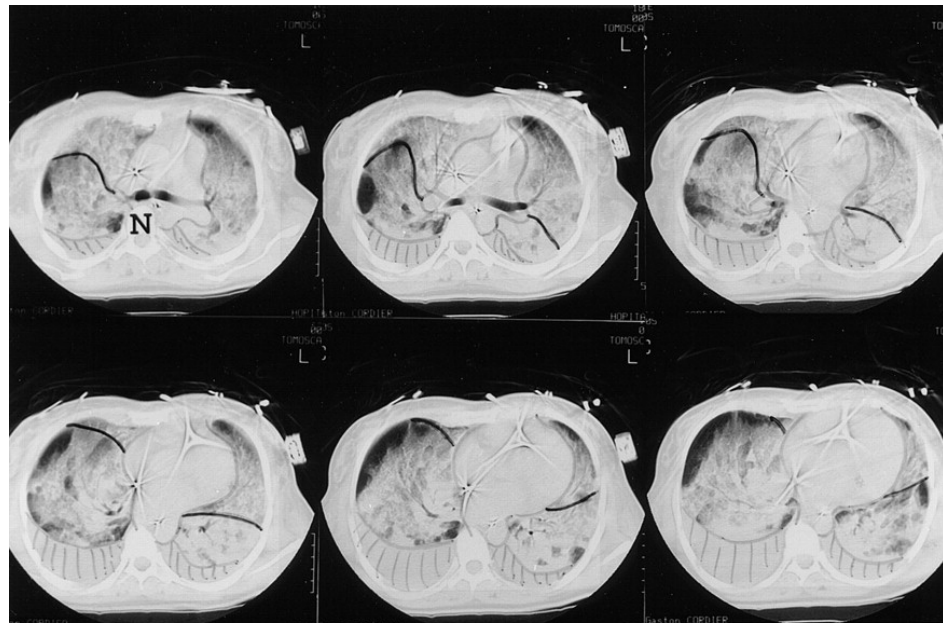
The following variables were entered into the multivariate model: tidal volume at inclusion, renal and cardiovascular SOFA sub-scores at inclusion, pH at inclusion, bicarbonates at inclusion, lactate at inclusion. Base excess was not entered into the multivariate model because of multicollinearity with pH. No significant interaction was identified between renal SOFA sub-score and pH at inclusion.

Personalised mechanical ventilation tailored to lung morphology versus low positive end-expiratory pressure for patients with acute respiratory distress syndrome in France (the LIVE study): a multicentre, single-blind, randomised controlled trial



*Jean-Michel Constantin, Matthieu Jabaudon, Jean-Yves Lefrant, Samir Jaber, Jean-Pierre Quenot, Olivier Langeron, Martine Ferrandière, Fabien Grelon, Philippe Seguin, Carole Ichai, Benoit Veber, Bertrand Souweine, Thomas Uberti, Sigismond Lasocki, François Legay, Marc Leone, Nathanael Eisenmann, Claire Dahyot-Fizelier, Hervé Dupont, Karim Asehnoune, Achille Sossou, Gérald Chanques, Laurent Muller, Jean-Etienne Bazin, Antoine Monsel, Lucile Borao, Jean-Marc Garcier, Jean-Jacques Rouby, Bruno Pereira, Emmanuel Futier, for the AZUREA Network**

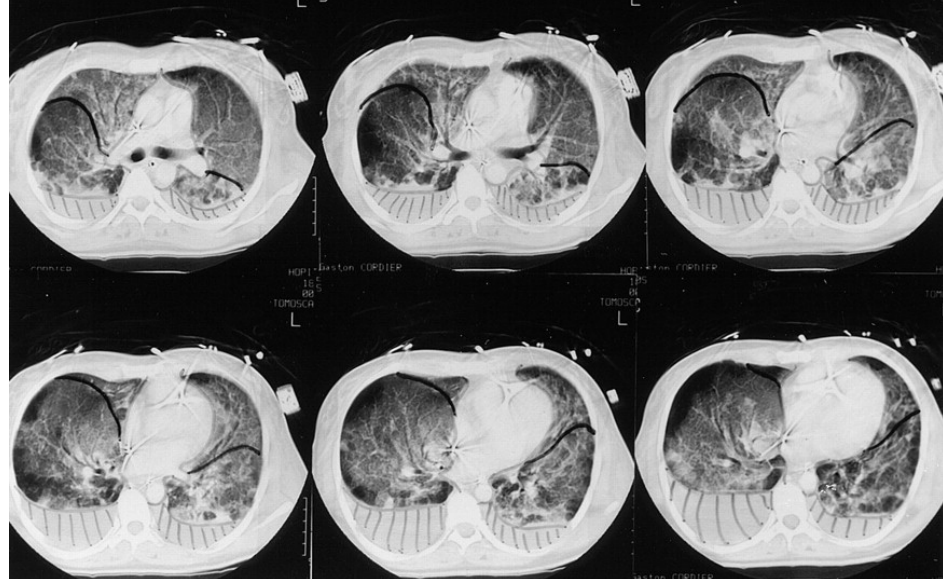




ZEEP ↑

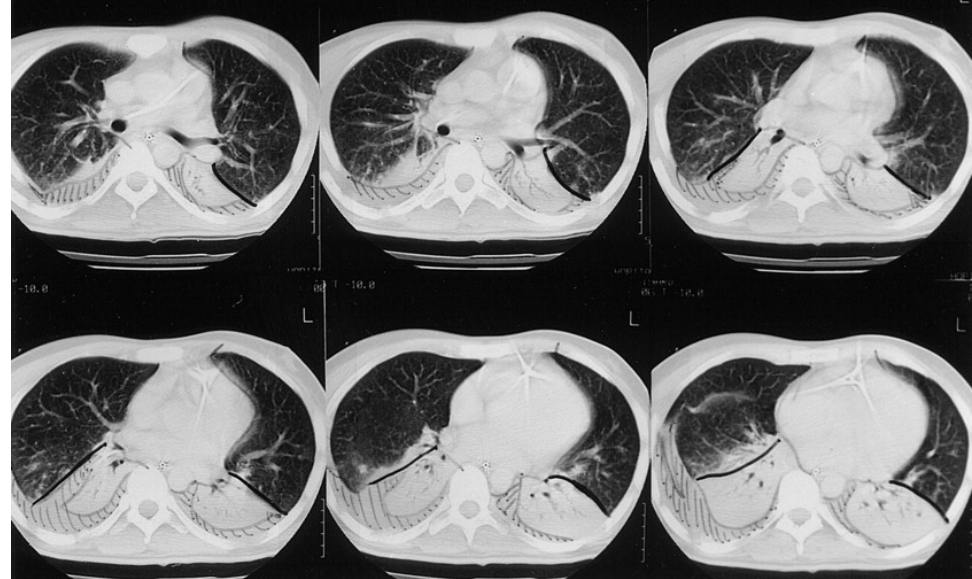
PEEP ↓

HOMOGENE: recrutement



ZEEP ↑

PEEP ↓ **HETEROGENE: pas/peu de recrutement**

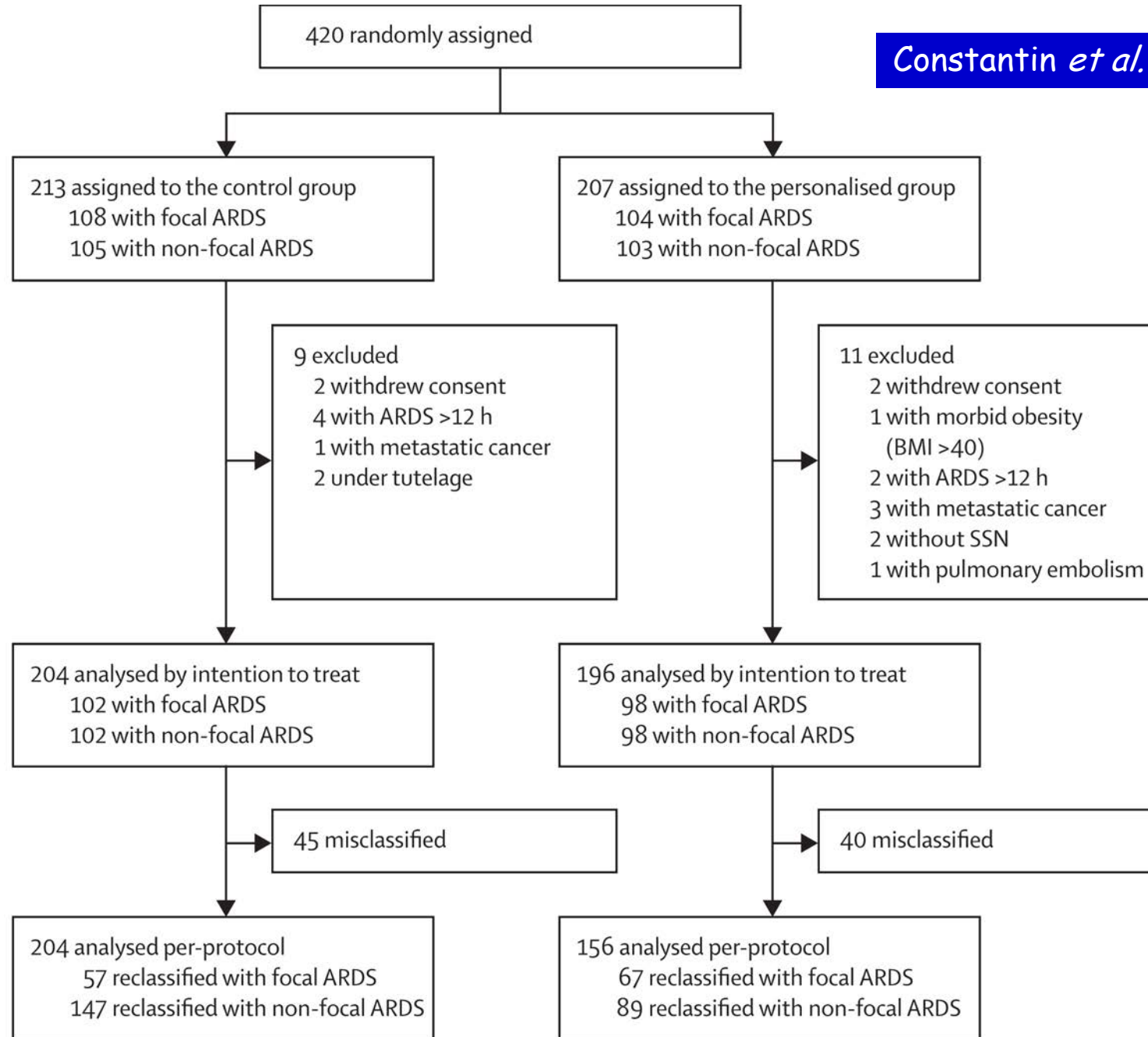


- CT scanning was used to assess lung morphology for 56 (29%) of 196 patients in the personalised group and 80 (39%) of 204 patients in the control group

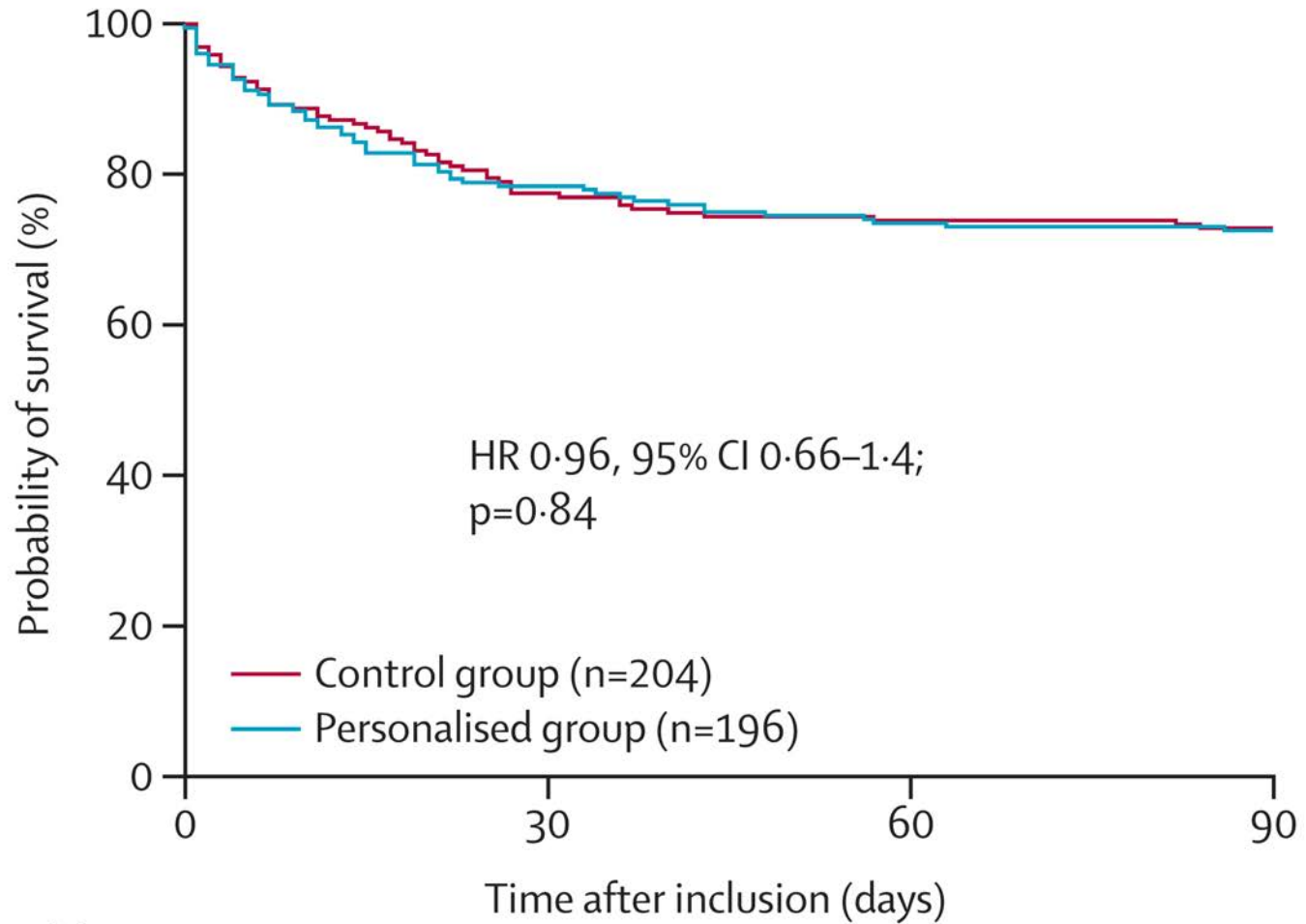
	Control group (n=204)	Personalised group (n=196)	
		Focal lung morphology	Non-focal lung morphology
Mode of ventilation	Volume control	Volume control	Volume control
Tidal volume	6 mL/kg IBW	8 mL/kg IBW	6 mL/kg IBW
PEEP	PEEP/FiO ₂	5–9 cm H ₂ O	To reach P _{plat} of 30 cm H ₂ O
PEEP-PSV	Free	5–9 cm H ₂ O	≥10 cm H ₂ O
Recruitment manoeuvre	Rescue	Rescue	Mandatory
Prone position	Encouraged	Mandatory	Rescue

IBW=ideal body weight. PEEP=positive-end expiratory pressure. FiO₂=fraction of inspired oxygen. P_{plat}=end-inspiratory plateau pressure. PEEP-PSV=positive-end expiratory pressure used during pressure support ventilation.

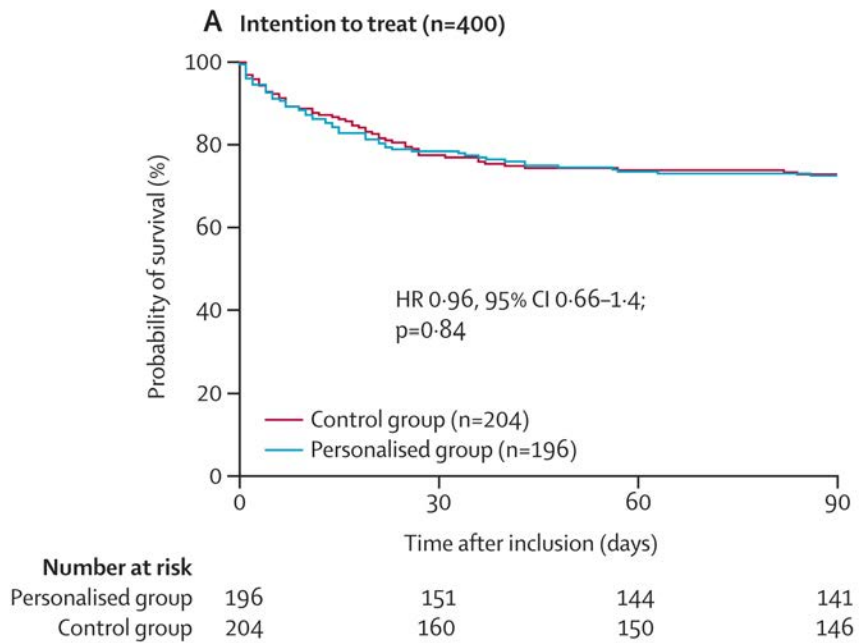
Table 1: Summary of ventilator settings according to lung morphology and randomisation group



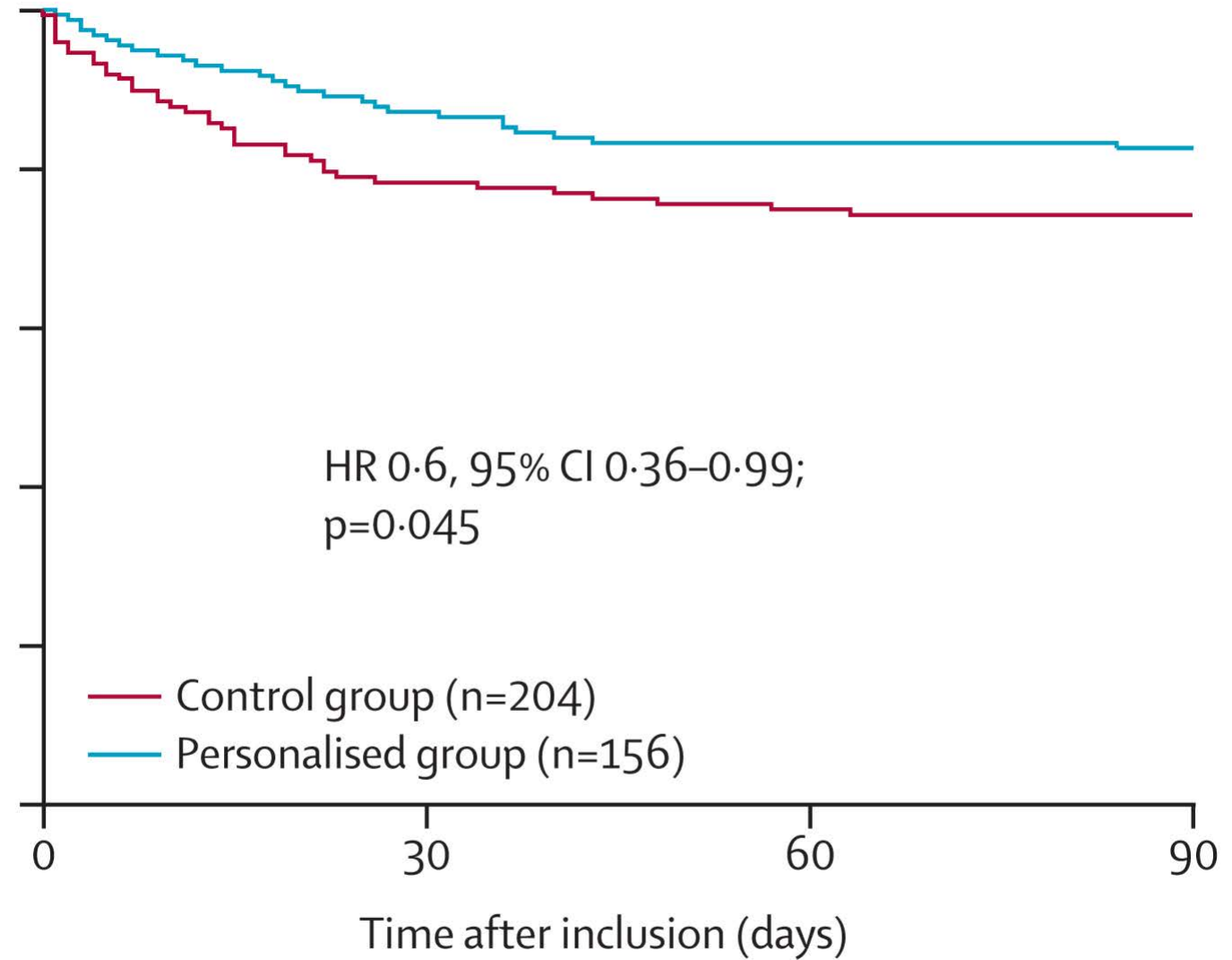
Intention to treat (n=400)



Number at risk					
Personalised group	196	151	144	141	
Control group	204	160	150	146	

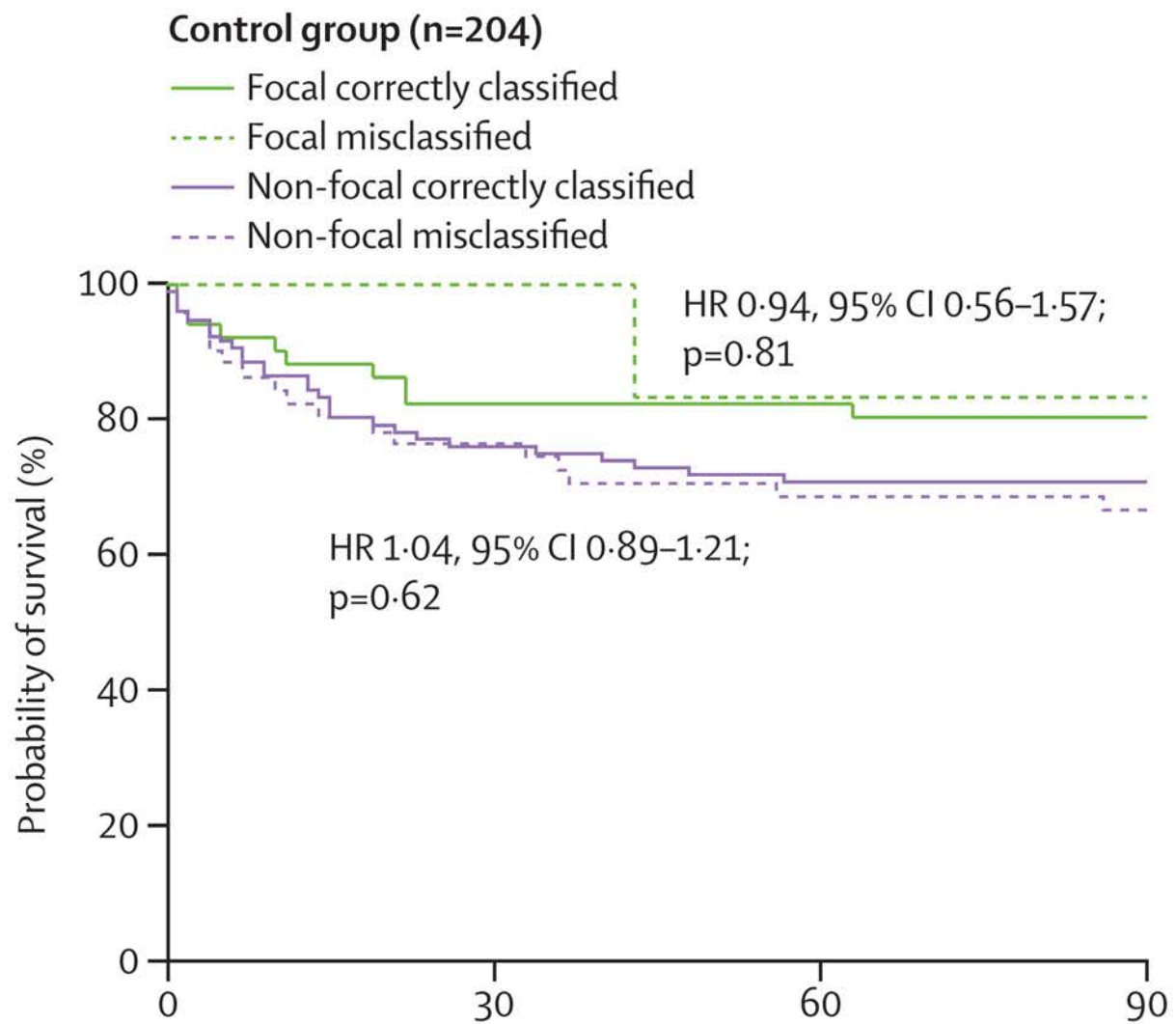


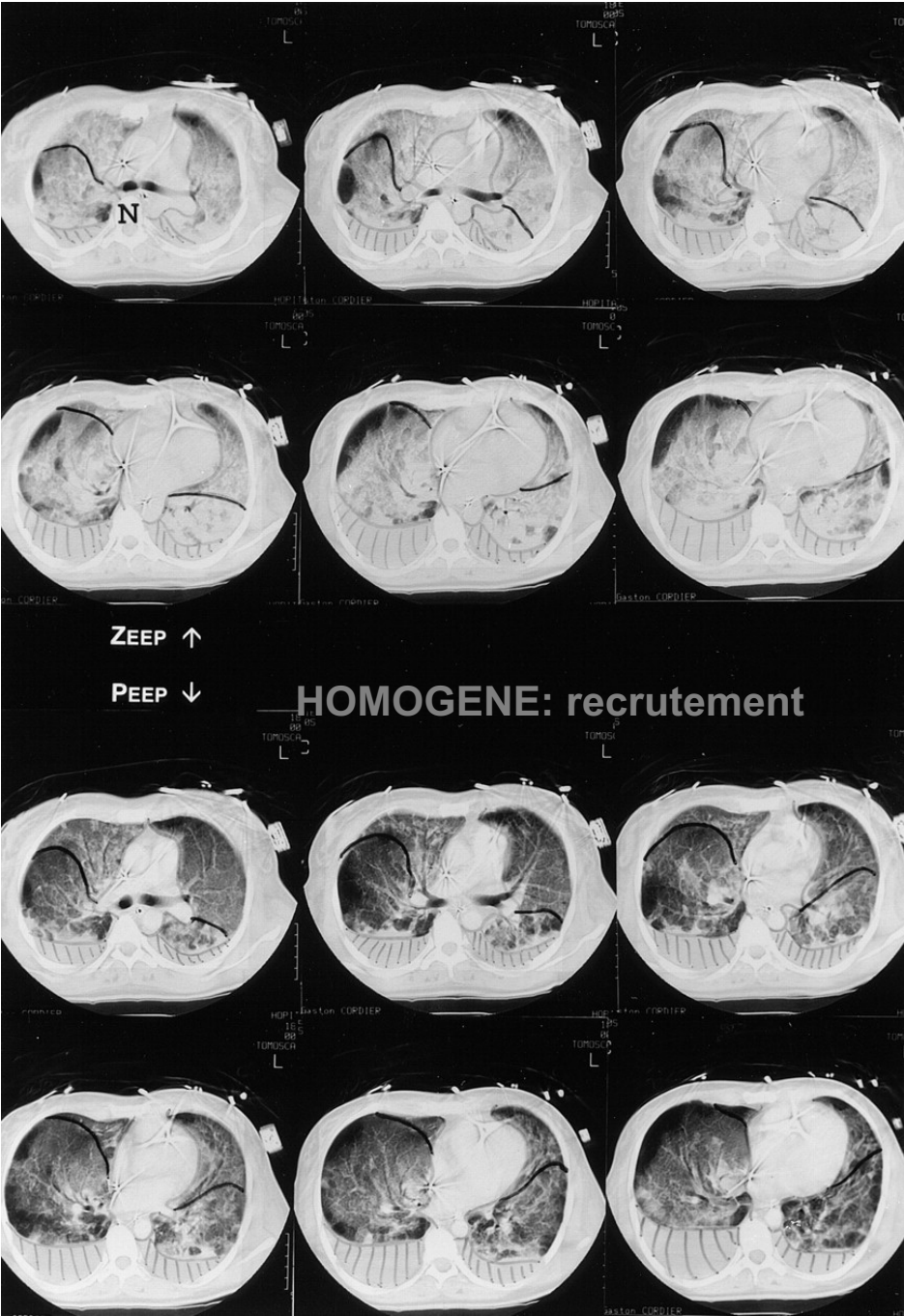
Per protocol (n=360)



Constantin *et al.* Lancet Resp Med 2019

	0	30	60	90
Personalised group	156	135	129	127
Control group	204	160	150	146





PEEP élevée
12-20...

PEEP modérée
8-10

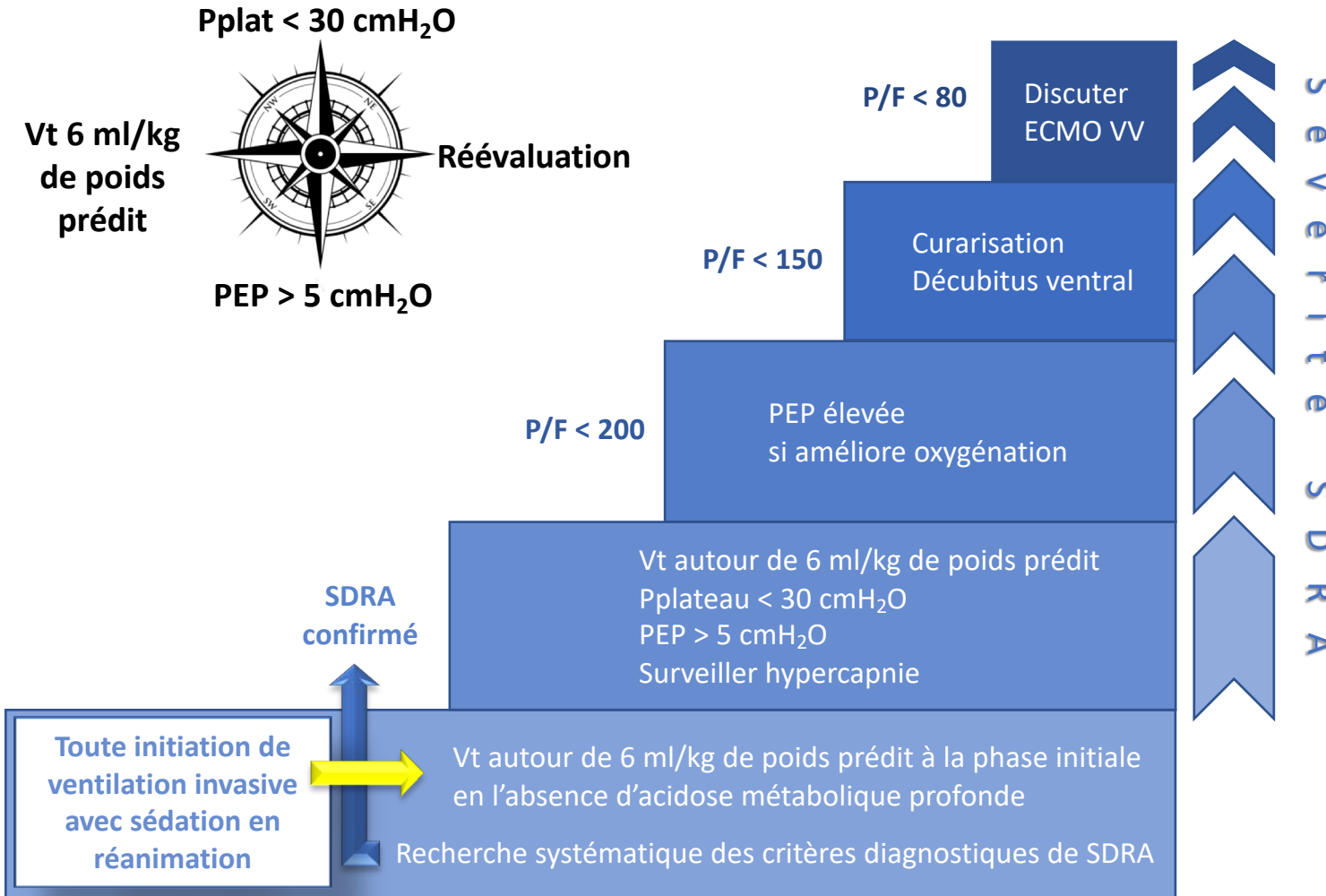
DV très précoce

Recommandations Formalisées d'Experts

Prise en charge du SDRA de l'adulte à la phase initiale

Cécile Aubron (Brest), Laurent Brochard (Toronto), Jean-Daniel Chiche (Paris), Alain Combes (Paris),
Didier Dreyfuss (Colombes), Jean-Marie Forel (Marseille), Claude Guérin (Lyon), Samir Jaber
(Montpellier), Armand Mekontso-Dessap (Créteil), Alain Mercat (Angers), Laurent Papazian (Marseille),
Jean-Christophe Richard (Annecy), Damien Roux (Colombes), Antoine Vieillard-Baron (Boulogne)

Prise en charge initiale du SDRA en 2019



ECMO veino-veineuse

- Si hypoxémie réfractaire ou ventilation protectrice non applicable
- A discuter avec un centre expert

Modalités de la curarisation : IVSE

- Précocement, dans les 48h du diagnostic

Modalités du décubitus ventral (DV)

- séance ≥ 16 heures, plusieurs séances

SDRA modéré ou sévère → Test PEP élevée (> 12 cmH₂O)

Utilisation PEP élevée si :

- Amélioration de l'oxygénation
- Sans dégradation significative de la compliance du système respiratoire et de l'hémodynamique
- Maintien Pplateau < 30 cmH₂O, monitoring continu

Critères du SDRA

- PaO₂/FiO₂ ≤ 300 mmHg
- PEP ≥ 5 cmH₂O
- Opacités bilatérales sur l'imagerie thoracique
- Non expliquées par défaillance ventriculaire gauche
- Évolution depuis moins de 7 jours

Traitement possible

- Monoxyde d'azote inhalé (iNO), si hypoxémie persistante en DV avant discussion de l'ECMO VV
- Ventilation spontanée après la phase aiguë avec Vt généré autour de 6 ml/kg sans dépasser 8 ml/kg

Pas de recommandation possible

- ECCO₂R
- Pression motrice
- Ventilation spontanée à la phase aiguë

Probablement ne pas faire

- Manœuvres de recrutement systématiques

Ne pas faire

- HFOV

Réévaluation des réglages et de la stratégie de prise en charge au moins toutes les 24h