

Preventive ventilation In high risk patients for ARDS

Samir JABER

Department of Critical Care Medicine and Anesthesiology (DAR B)
Saint Eloi University Hospital and Montpellier School of Medicine – INSERM U1046
80 Avenue Augustin Fliche; 34295 Montpellier. **FRANCE**
Mail : s-jaber@chu-montpellier.fr ; Tel : +33 4 67 33 72 71

Hammamet – 28 mai 2016

Déclaration de liens

Déclare les liens suivants (consultants) :

- Drager France
- Maquet France
- Fisher Paykel
- Hamilton
- Baxter
- Medtronic

Objectifs

1. Qui sont les patients à risque?

2. Qu'est ce que la ventilation protectrice et préventive ?

3. Quelles sont les évidences ?

4. Take Home Messages

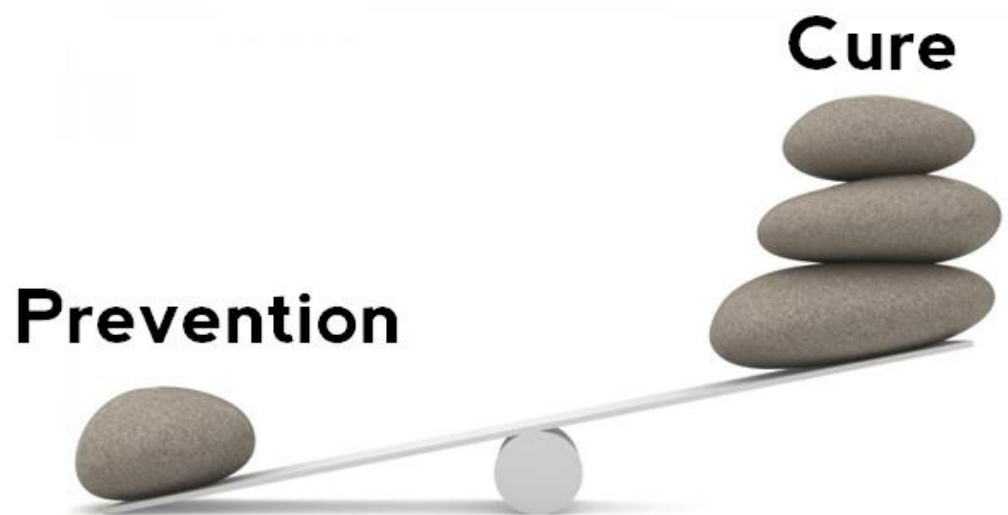
Benjamin Franklin



Born in Boston, Massachusetts,
The United States 1706-1790

Background

“An ounce of prevention is worth a pound of cure.”



Copyright © Pegasus Vertex, Inc.

“Prevention is better than cure”

The golden hours



“The most sophisticated
intensive care becomes
**unnecessarily expensive
terminal care...**”
when the pre ICU systems fail

Peter Safar

Quel "patient est à risque" ?

Medical history

(COPD, Cachexia, Cancer, immunosuppression, Muscle deconditioning...)

Supine Position

Transfusion

Mechanical Ventilation

Renal Failure

All ICU patients could be considered at patients at risk to develop lung injury (i.e, ARDS)

SEDATION

- Propofol, Morphine, NMBA

Antibiotic

Surgery

Pancreatitis

Abdo Cpt Synd

Inflammation

Shock

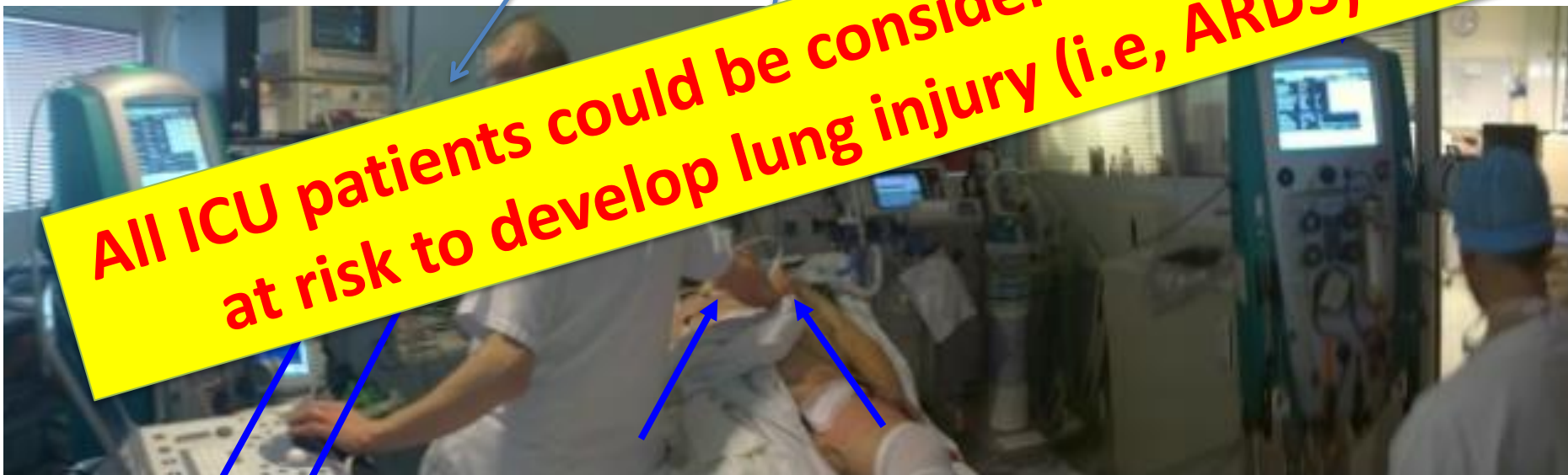
Sepsis

Electrolytes abnormalities

(Ca, Ph, Mg, K)

Hyperglycemia

Hypertriglyceridemia



Early Identification of Patients at Risk of Acute Lung Injury: Evaluation of Lung Injury Prediction Score in a Multicenter Cohort Study.

Gajic O, Dabbagh O, Park PK, Adesanya A, Chang SY, Hou P, Anderson Iii H, Hoth JJ, Mikkelsen ME, Gentile NT, Gong MN, Talmor D, Bajwa E, Watkins TR, Festic E, Yilmaz M, Iscimen R, Kaufman DA, Esper AM, Sadikot R, Douglas I, Sevransky J, Malinchoc M; on behalf of the US Critical Illness and Injury Trials Group: Lung Injury Prevention Study Investigators (USCIITG-LIPS).

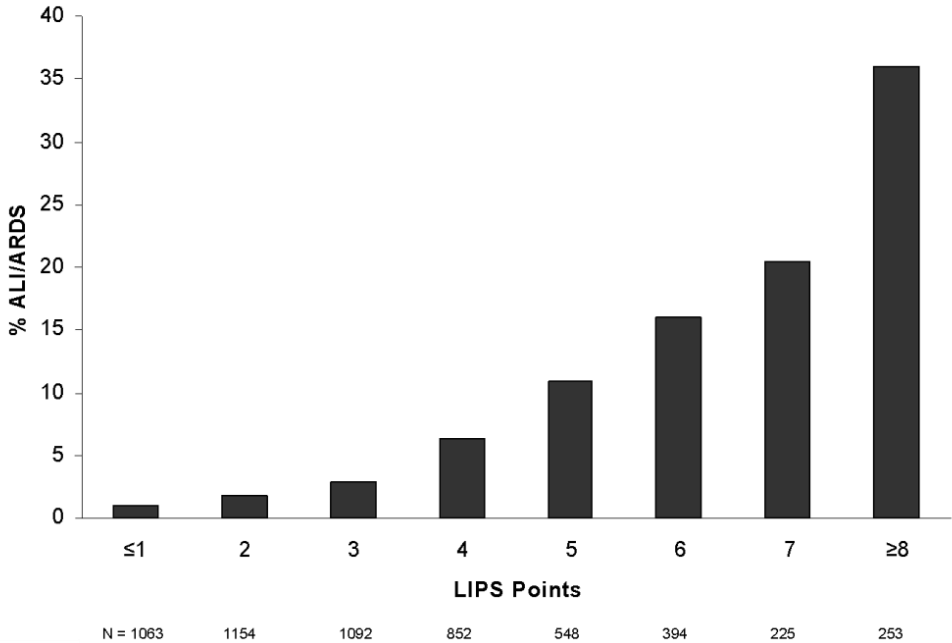
Department of Internal Medicine, Division of Pulmonary and Critical Care Medicine, Mayo Clinic College of Medicine, 200 First Street SW, Rochester, Minnesota, 55905, United States.

	Estimate	(95% CI)		p-value	LIPS points
Predisposing conditions					
Shock	0.77	0.19	1.32	0.008	2
Aspiration	0.79	0.07	1.45	0.024	2
Sepsis	0.37	-0.13	0.87	0.139	1
Pancreatitis	-1.07	-3.96	0.51	0.299	0
Pneumonia	0.83	0.33	1.34	0.001	1.5
High risk surgery*					
Thoracic (noncardiac)	-0.14	-2.00	1.15	0.896	-
Orthopedic spine	0.75	-0.11	1.53	0.071	1
Acute abdomen	0.93	0.06	1.72	0.028	2
Cardiac	1.32	0.67	1.96	<.001	2.5
Aortic vascular	1.78	0.93	2.56	<.001	3.5
High risk trauma					
Traumatic brain injury	1.29	0.67	1.91	<.001	2
Smoke inhalation	0.93	-0.21	1.41	0.438	2
Near drowning	1.68	-2.74	6.00	0.498	2
Lung contusion	0.40	-0.48	1.21	0.355	1.5
Multiple fractures	0.64	-0.21	1.41	0.117	1.5
Risk modifiers					
Male gender	0.02	-0.34	0.39	0.905	-
Alcohol abuse	0.51	-0.08	1.07	0.080	1
Obesity (BMI>30)	0.56	0.18	0.93	0.004	1
Hypoalbuminemia	0.46	0.04	0.87	0.029	1
Chemotherapy	0.46	-0.54	1.29	0.314	1
FiO ₂ >0.35 (>4L/min)	1.02	0.62	1.41	<.001	2
Tachypnea (RR>30)	0.69	0.11	1.25	0.017	1.5
SpO ₂ <95%	0.35	-0.04	0.73	0.078	1
Acidosis (pH<7.35)	0.55	0.09	1.00	0.017	1.5
Diabetes mellitus**	-0.59	-1.40	0.15	0.135	-1

*Add 1.5 point if emergency surgery

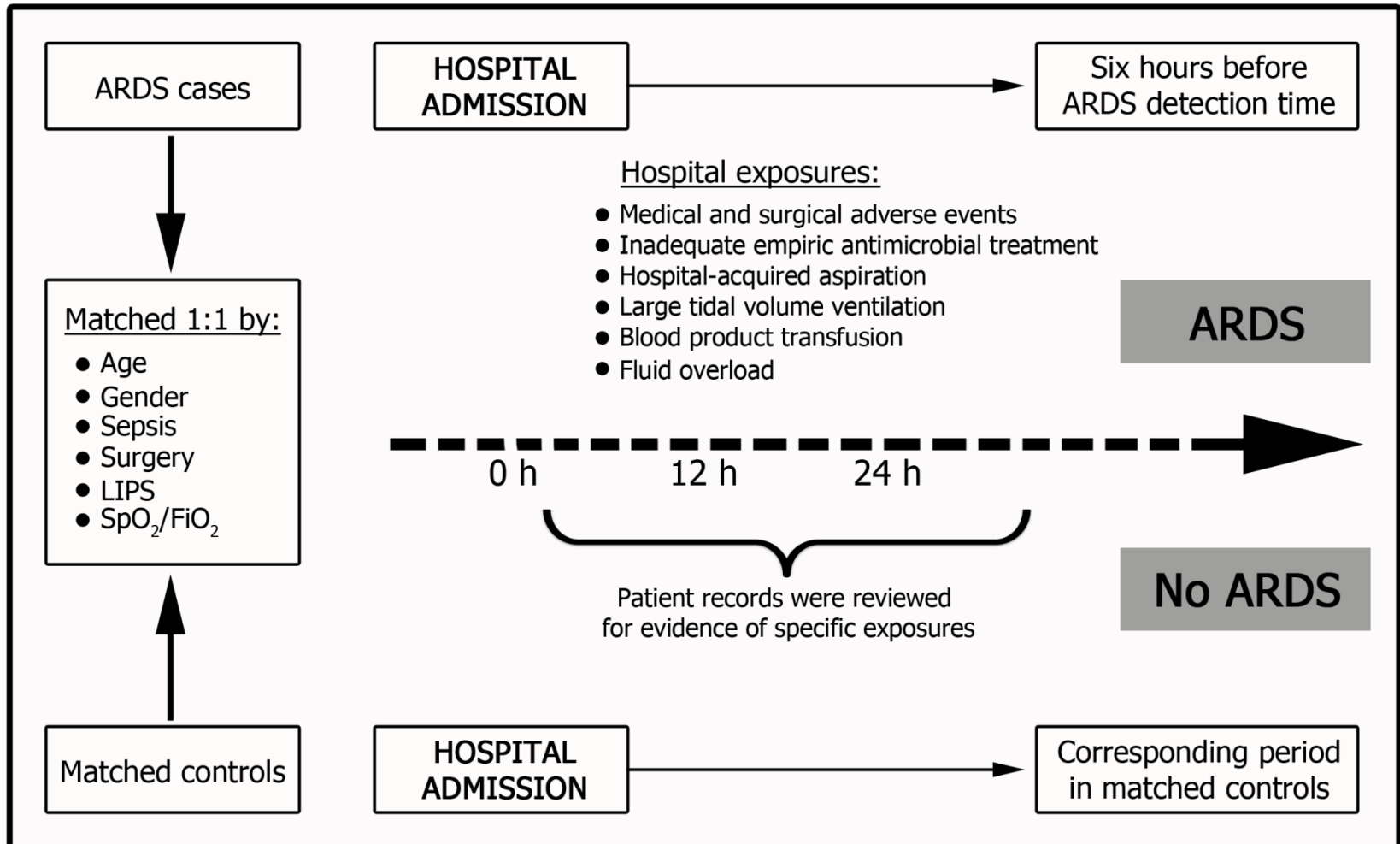
**Only if sepsis

Lung Injury Prediction Score (LIPS)

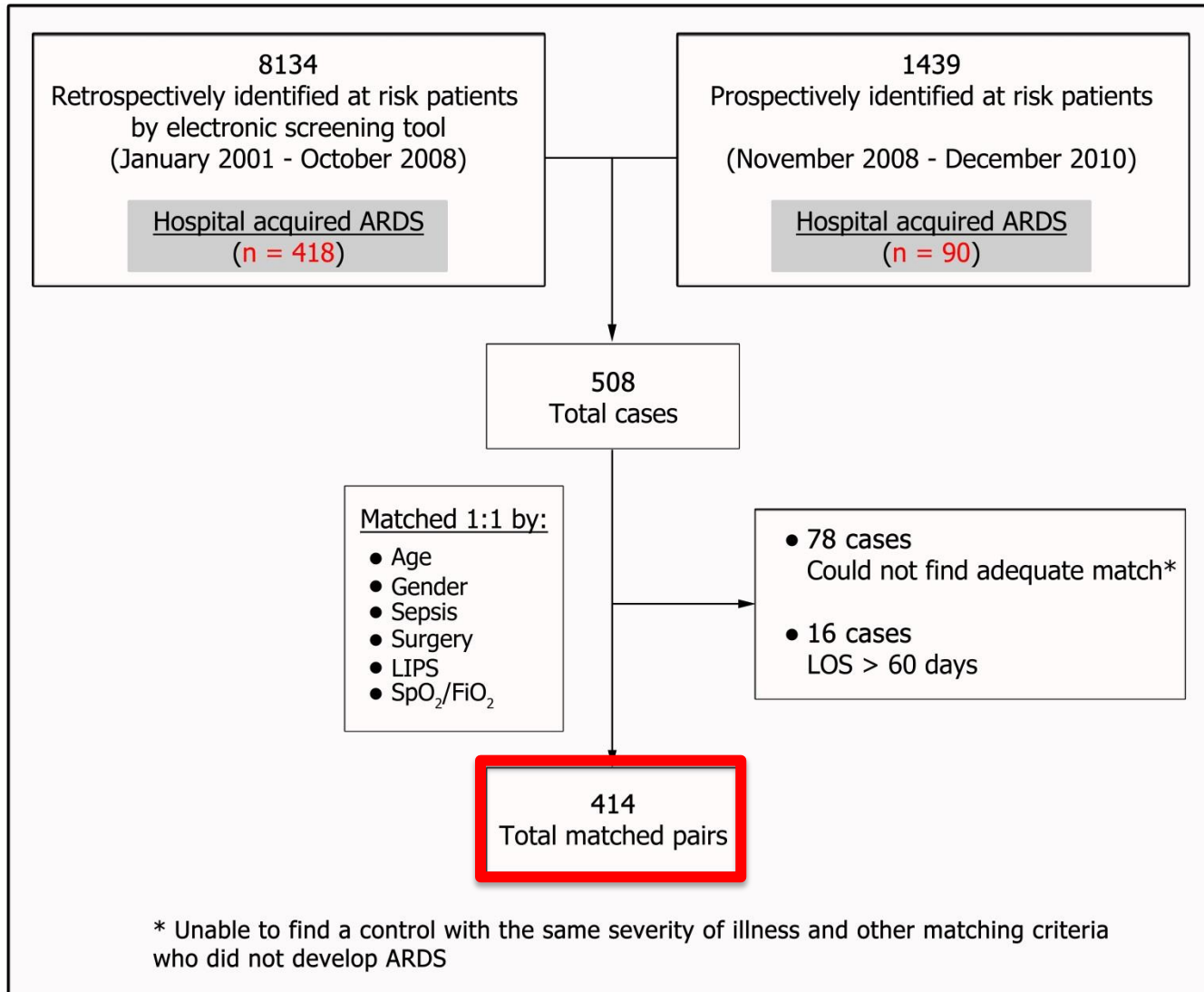


The Role of Potentially Preventable Hospital Exposures in the Development of Acute Respiratory Distress Syndrome: A Population-Based Study

Ahmed, Adil H. MBBS; Litell, John M. DO; Malinchoc, Michael MS; Kashyap, Rahul MBBS; Schiller, Henry J. MD; Pannu, Sonal R. MBBS; Singh, Balwinder MBBS; Li, Guangxi MD; Gajic, Ognjen MD, MSc



Study Flow chart



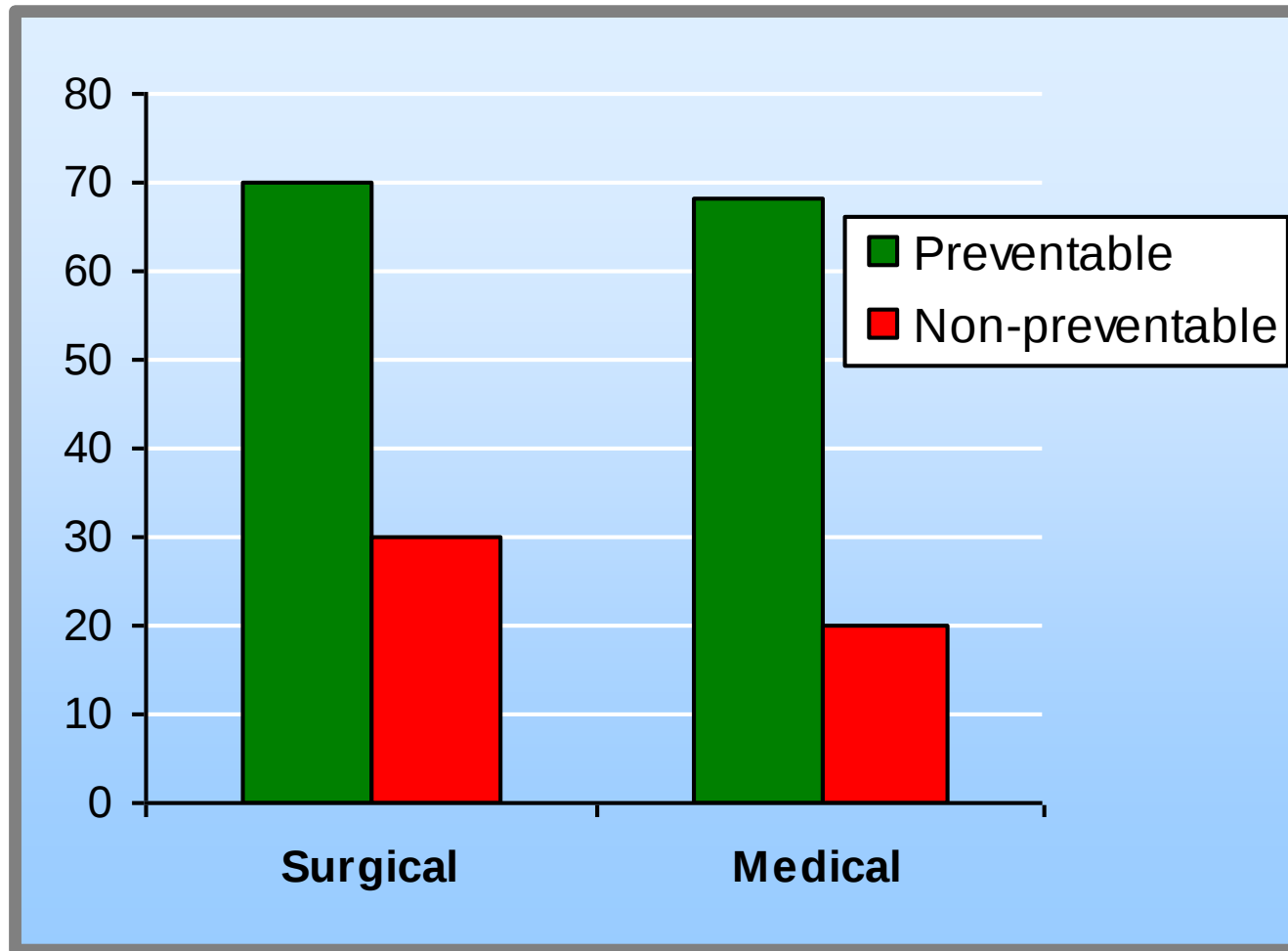
RESULTS

Exposures	ARDS Cases	Controls	OR(95% CI)	P value
Infection control (among those with suspected infection N=151 pairs)				
Inadequate antimicrobial, N (%)	61(40.4)	30 (19.9)	2.9(1.7-5.2)	<0.001
Time to adequate antimicrobial in hours M(IQR)	7.6(3.0-25.4)	3.9 (2.4-8.4)	1.3 (1.1-1.5)	<0.001
Hospital acquired aspiration and aspiration surrogates N (%)				
Hospital acquired aspiration	51 (12.3)	1 (0.2)	51 (7.1-369)	<0.001
Nasogastric tube	190 (45.9)	60 (14.5)	6.4(4.2-9.9)	<0.001
Head of bed elevation *($\geq 30^\circ$)	16 (61.5)	17 (65.4)	0.8 (0.4-2.4)	0.81
Documented intubation difficulty	32 (7.7)	10 (2.4)	3.2(1.5-6.5)	<0.001
IV Fluids -for one or more liter infused , median (IQR)				
Crystalloid first 24 hours- M (IQR)	1 (0.2,3.0)	1.4 (0.4,3.4)	0.9 (0.9-1.0)	0.26
Mechanical ventilation parameters (n=29 pairs)				
TV by PBW, median (IQR)	9 (7.3-10.8)	7.7 (6.9-8.5)	1.7 (1.1-2.6)	.025
FIO ₂ %, median (IQR)	54 (43-62)	55 (50-63)	0.9 (0.9-1.0)	.92
Blood product transfusion N (%)				
Red blood cells	163(39.4)	49 (11.8)	1.4 (1.3-1.6)	<0.001
Platelet	51 (12.3)	11 (2.7)	2.0 (1.4-2.9)	<0.001
Fresh frozen plasma	85 (20.5)	15 (3.6)	1.4 (1.2-1.6)	<0.001

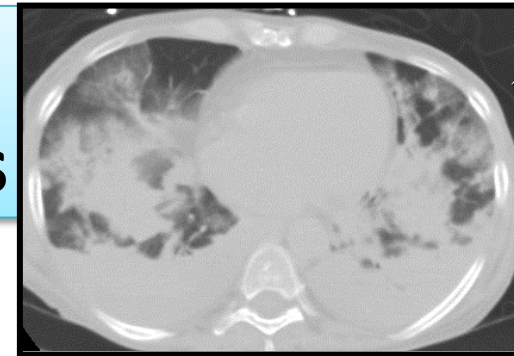
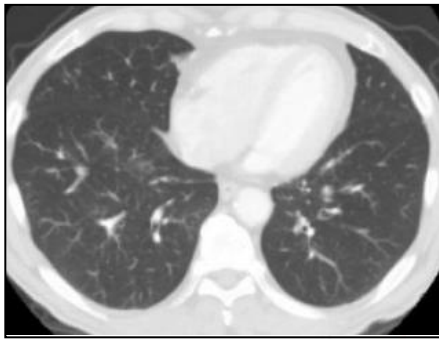
RESULTS

Exposures	ARDS	Controls	OR(95% CI)	P value
Adverse events (surgical and medical adverse events) N=828 **				
Accidental cut, puncture, perforation, or hemorrhage, N (%)	27 (6.5)	4(1.0)	12.5(3.0-52.8)	<0.001
Failure of sterile precautions, N (%)	4 (1.0)	0 (0.0)	-----	-----
Dosage failure, N (%)	113(27.3)	35 (8.5)	3.5(0.7-16.8)	0.118
Contaminated or infected blood, fluid, drug, or biological substance, N (%)	0 (0.0)	1 (0.2)	-----	-----
Other and unspecified N (%)	57 (13.8)	8 (1.9)	8 (3.6-17.6)	<0.001
Any adverse event N (%)	131 (31.6)	47 (11.4)	6.2(4.0-9.7)	<0.001

Medical and Surgical Adverse events



ARDS Pathogenesis: “Multiple hit” Hypothesis



Patient at risk (1st hit)

- Pneumonia
- Toxic inhalation
- Pancreatitis
- Aspiration
- Trauma
- Sepsis
- Shock
- Age
- SNPs
- Alcohol
- Tobacco
- Thoracic and vascular surgery
- Preexisting lung disease
- Vasculitis
- Radiation
- Chemotherapy



**HOSPITAL
ADMISSION**

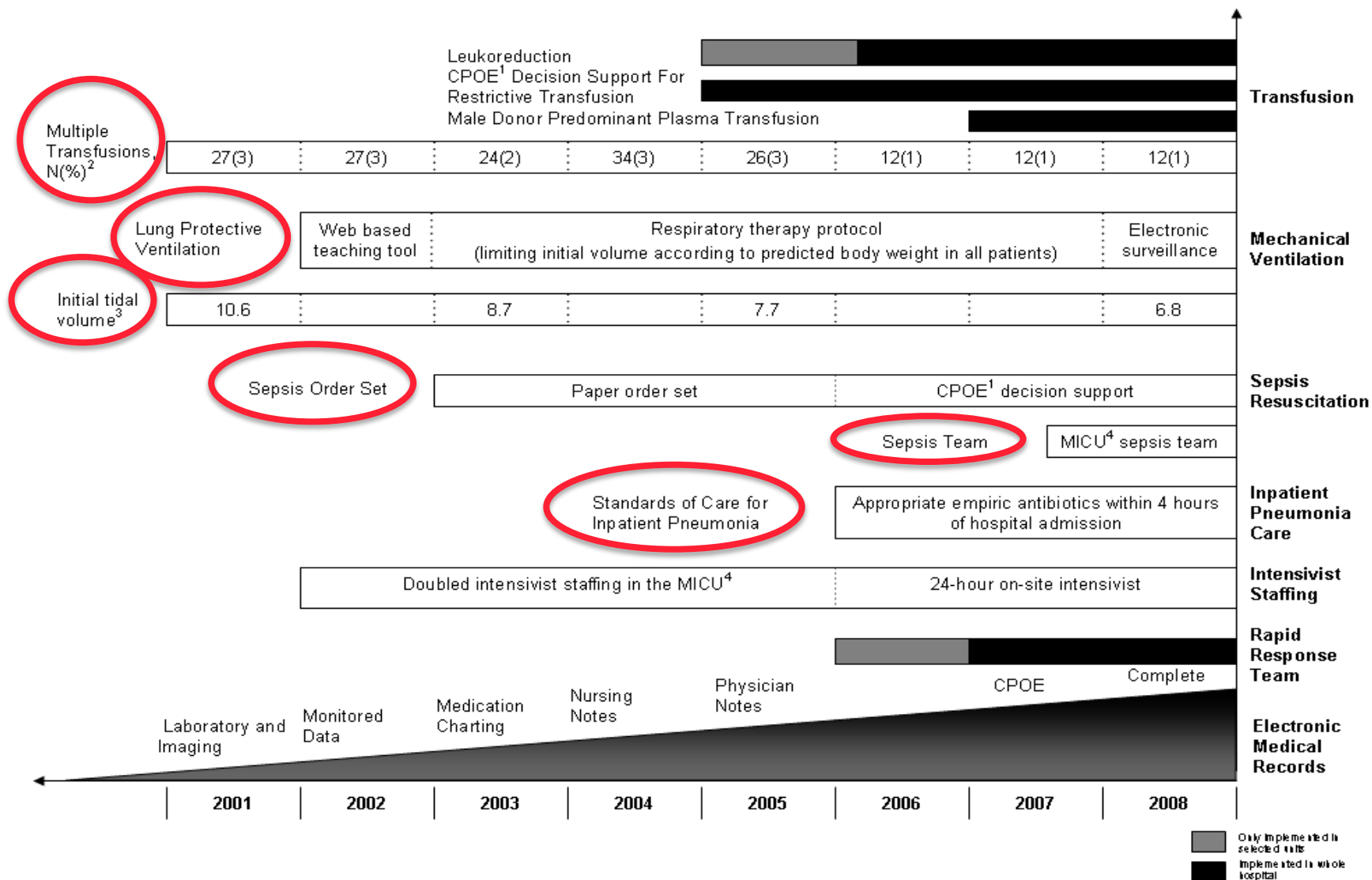
**ICU
ADMISSION**

Eight-Year Trend of Acute Respiratory Distress Syndrome - AJRCCM 2011

Systematic Changes in Acute Care Delivery

A Population-based Study in Olmsted County, Minnesota

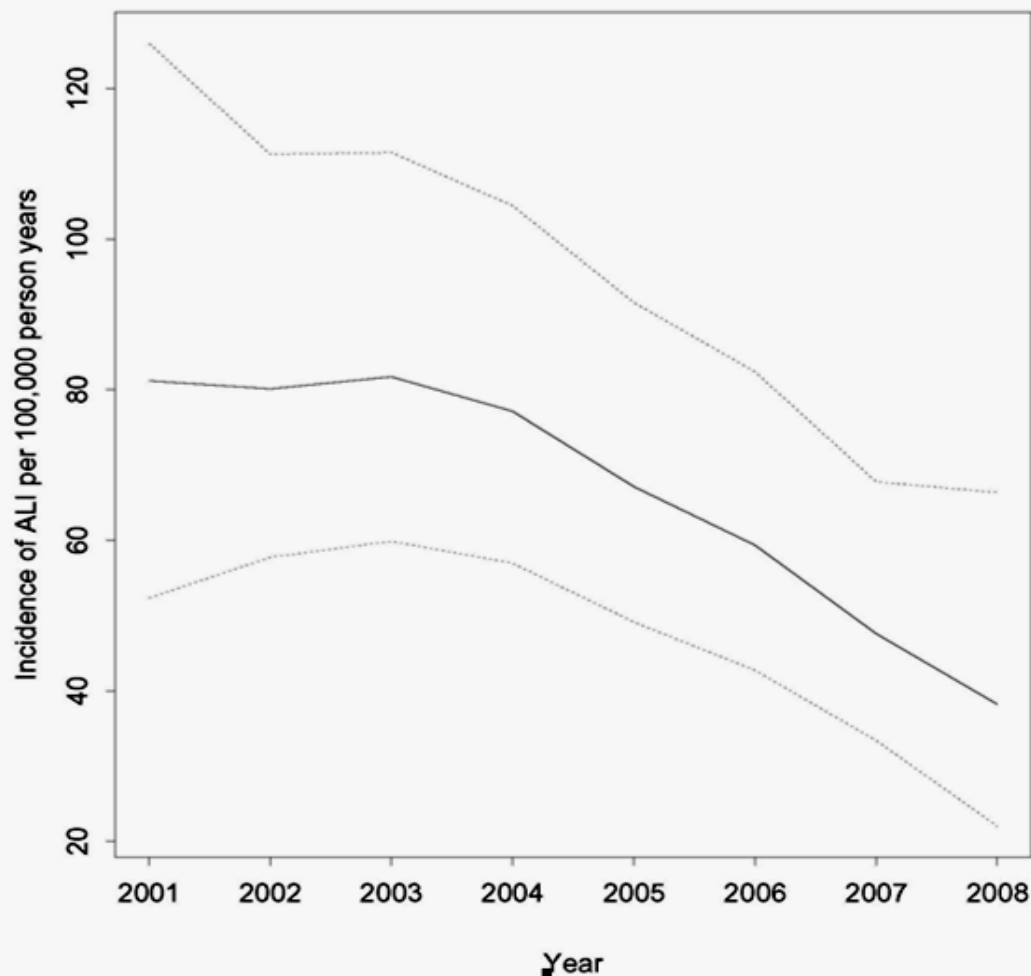
Guangxi Li^{1,2}, Michael Malinchoc^{1,3}, Rodrigo Cartin-Ceba¹, Chakradhar V. Venkata¹, Daryl J. Kor^{1,4}, Steve G. Peters¹, Rolf D. Hubmayr¹, and Ognjen Gajic¹



Eight-Year Trend of Acute Respiratory Distress Syndrome - AJRCCM 2011

A Population-based Study in Olmsted County, Minnesota

Guangxi Li^{1,2}, Michael Malinchoc^{1,3}, Rodrigo Cartin-Ceba¹, Chakradhar V. Venkata¹, Daryl J. Kor^{1,4}, Steve G. Peters¹, Rolf D. Hubmayr¹, and Ognjen Gajic¹



Incidence of ARDS in Olmsted County, Minnesota: Combined Effect?

Checklist for Lung Injury Prevention (CLIP)

Clip Elements	Definition
Lung protective mechanical ventilation	Tidal volume between 6-8 mL / kg predicted body weight and plateau pressure <30 cm H ₂ O; PEEP ≥ 5 cm H ₂ O, minimize FIO ₂ (target O ₂ sat 88-92% after early shock)

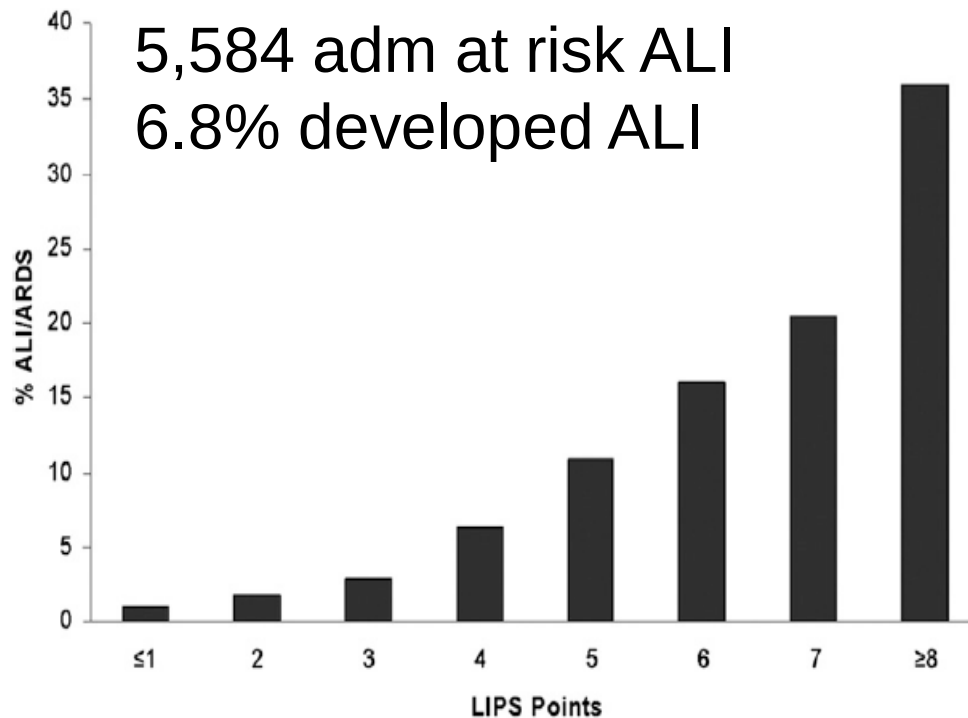


Early Identification of Patients at Risk of Acute Lung Injury

22 centres

5,584 adm at risk ALI

6.8% developed ALI



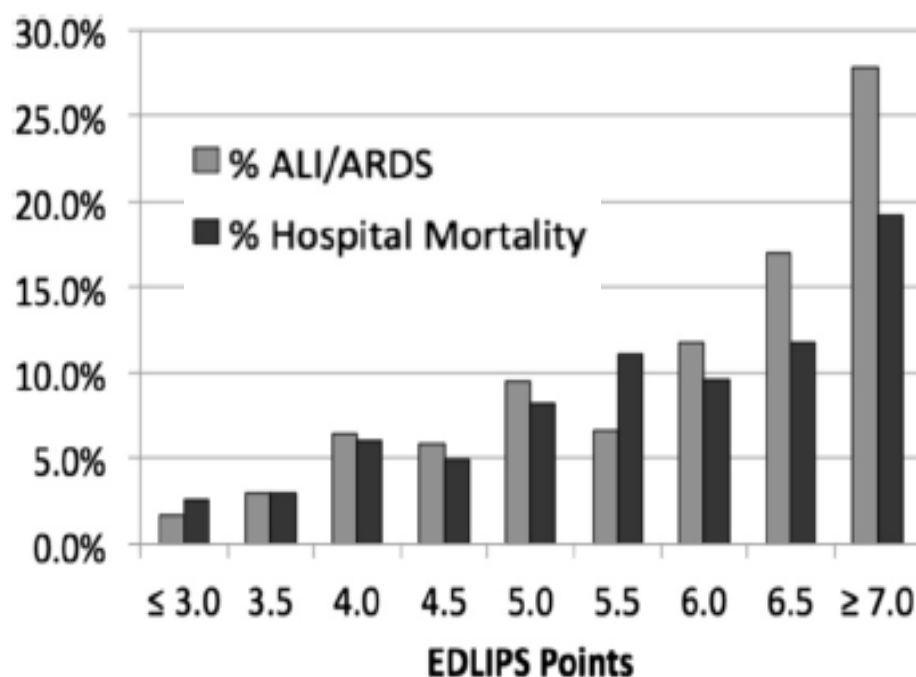
Predisposing Conditions

Shock	2
Aspiration	2
Sepsis	1
Pneumonia	1.5
High-risk surgery*	
Orthopedic spine	1
Acute abdomen	2
Cardiac	2.5
Aortic vascular	3.5
High-risk trauma	
Traumatic brain injury	2
Smoke inhalation	2
Near drowning	2
Lung contusion	1.5
Multiple fractures	1.5

Risk modifiers

Alcohol abuse	1
Obesity (BMI > 30)	1
Hypoalbuminemia	1
Chemotherapy	1
FiO ₂ > 0.35 (> 4 L/min)	2
Tachypnea (RR > 30)	1.5
SpO ₂ < 95%	1
Acidosis (pH < 7.35)	1.5
Diabetes mellitus [†]	-1

Lung injury prediction score for the emergency department: first step towards prevention in patients at risk



Pre-disposing conditions

Male 1, Aspiration 2, Pneumonia 1, Sepsis 1, Shock 2, Lung contusion 1, Smoke inhalation 1.5 Long bone fractures 2, Brain injury 2, Cardiac surgery 5, Aortic surgery 5, Spine surgery 5, Acute abdomen 2.5

Risk modifiers

Diabetes -0.5, Cirrhosis 1, Chemo 2, Obesity 1.5, Acidosis 2, FiO₂ > 0.35 2, Albumin 1.5, SpO₂ < 95% 1.5

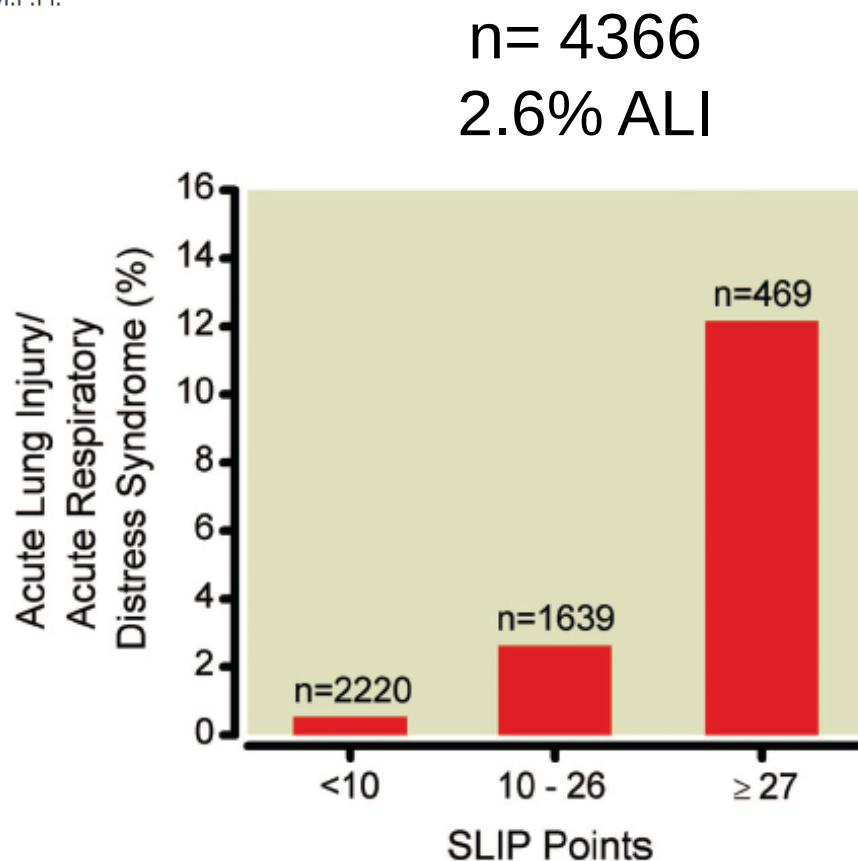
Predicting Risk of Postoperative Lung Injury in High-risk Surgical Patients

A Multicenter Cohort Study **ANESTHESIOLOGY 2014; 120:1168-81**

Daryl J. Kor, M.D., Ravi K. Lingineni, M.P.H., Ognjen Gajic, M.D., M.Sc., Pauline K. Park, M.D., James M. Blum, M.D., Peter C. Hou, M.D., J. Jason Hoth, M.D., Harry L. Anderson III, M.D., Ednan K. Bajwa, M.D., M.P.H., Raquel R. Bartz, M.D., Adebola Adesanya, M.D., Emir Festic, M.D., Michelle N. Gong, M.D., M.S., Rickey E. Carter, Ph.D., Daniel S. Talmor, M.D., M.P.H.

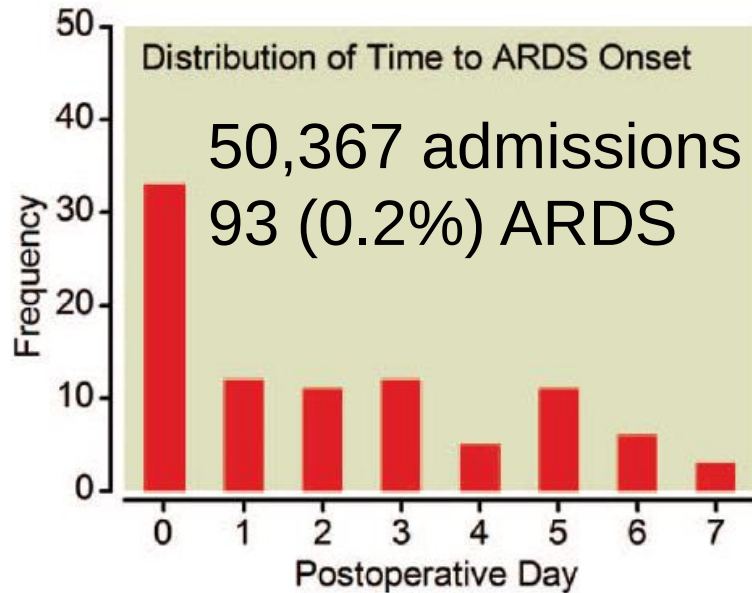
surgical lung injury prediction

Predictor Variables	SLIP Points
High-risk surgical procedure	
Cardiac	19
Vascular	32
Thoracic	16
Comorbidities	
Diabetes mellitus	6
COPD	10
GERD	7
Modifying conditions	
Alcohol abuse	11



Preoperative and Intraoperative Predictors of Postoperative Acute Respiratory Distress Syndrome in a General Surgical Population

Blum et al Anesthesiology 2013



Intra-operative predictors ARDS

RBC transfusion	OR 5.4 (1.4-11.1)
Median drive pressure	OR 1.2(1.1 – 1.3)
Median FiO ₂	OR 1.02(1-1.05)
Crystalloid	OR 1.4(1.2-1.9)

A RETENIR : Principaux facteurs associés à un risque de développer un SDRA

- 1. Traitement de la cause du SDRA insuffisant**
(ex: ATB inadapté, chirurgie non faite...)
- 2. Mauvais réglage du ventilateur :**
haut VT (>8-10 ml/kg); Pression motrice élevée...
- 3. Transfusion; surcharge volémique**
- 4. Post-opératoire de chirurgie « lourde »**
- 5. Co-morbidité(s)** (immunodépression, cirrhose, cancer...)

Objectifs

1. Qui sont les patients à risque?

2. Qu'est ce que la ventilation protectrice et préventive ?

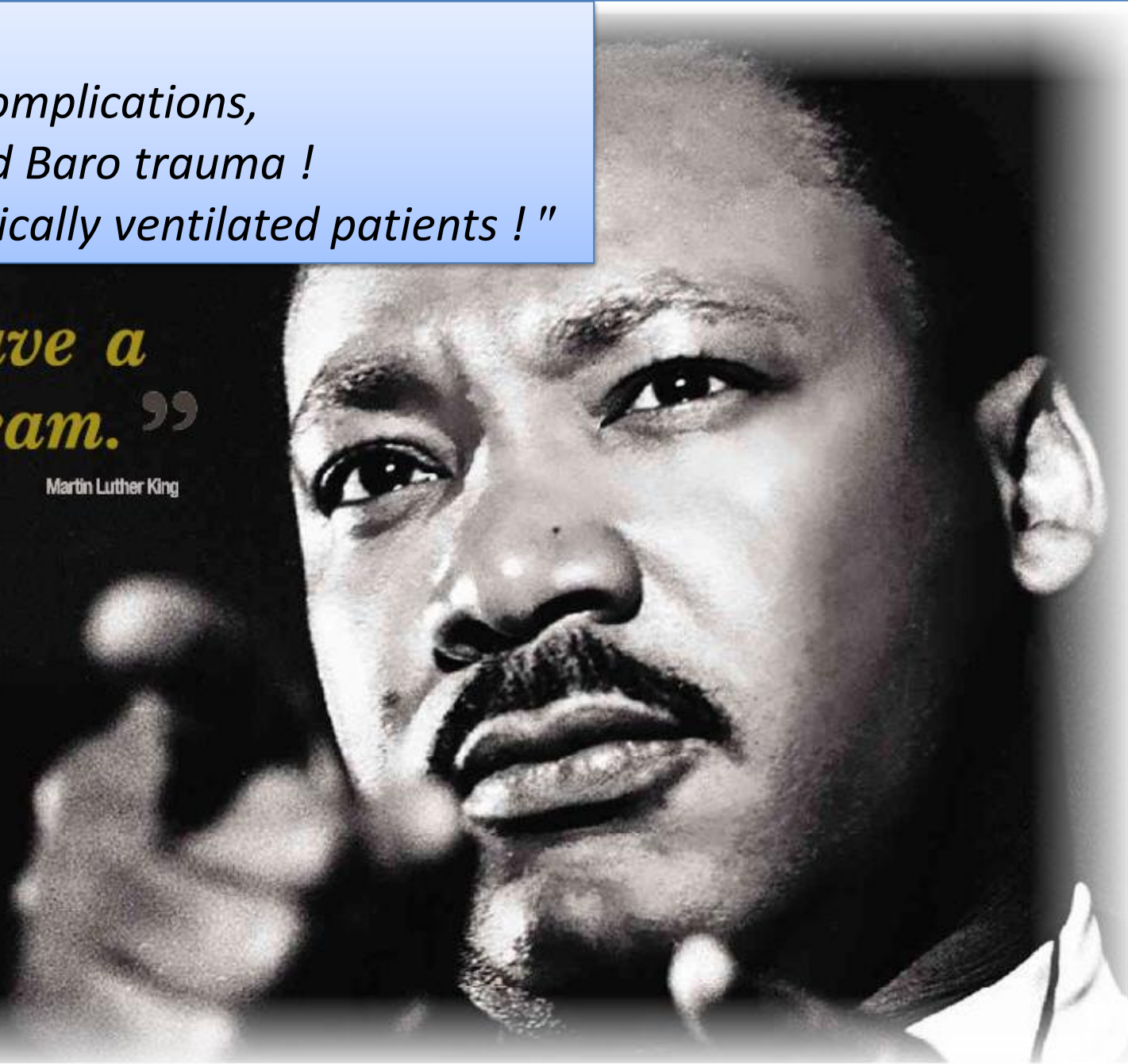
3. Quelles sont les évidences ?

4. Take Home Messages

*" No atelectasis,
No pulmonary complications,
No more volo and Baro trauma !
In « all » mechanically ventilated patients ! "*

*“ I have a
dream. ”*

Martin Luther King

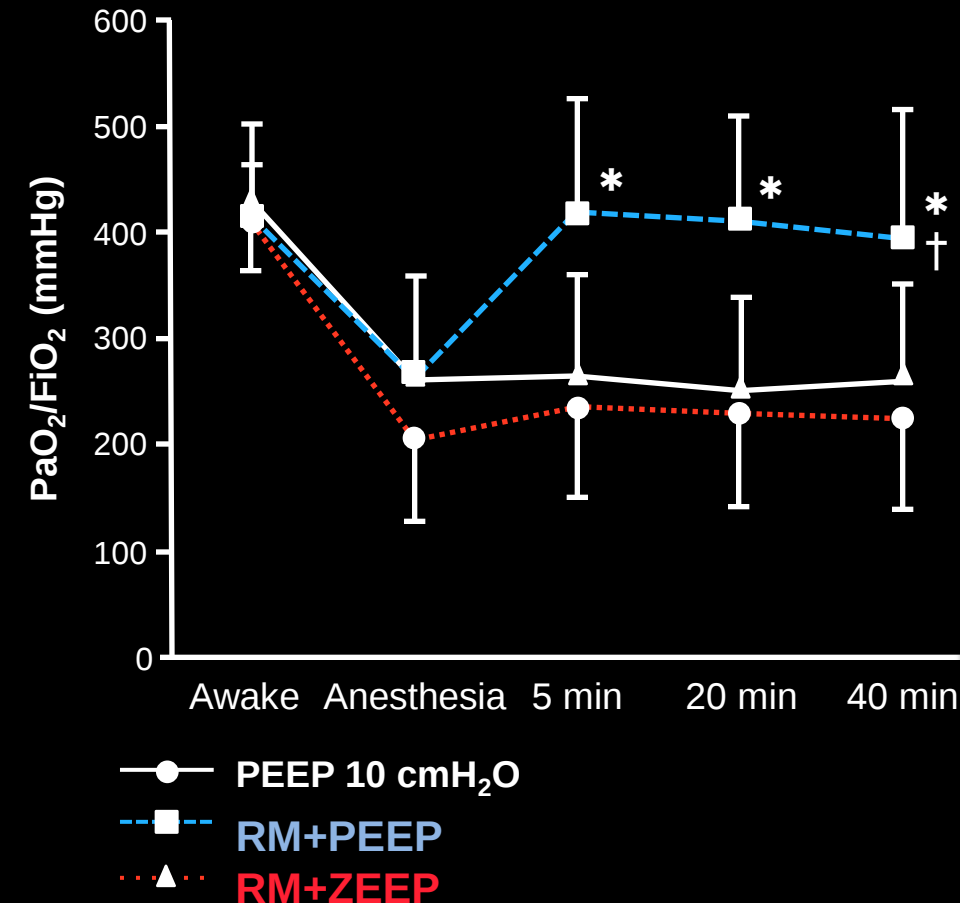


Prevention of Atelectasis in Morbidly Obese Patients during General Anesthesia and Paralysis (= 3 ventilatory strategies)

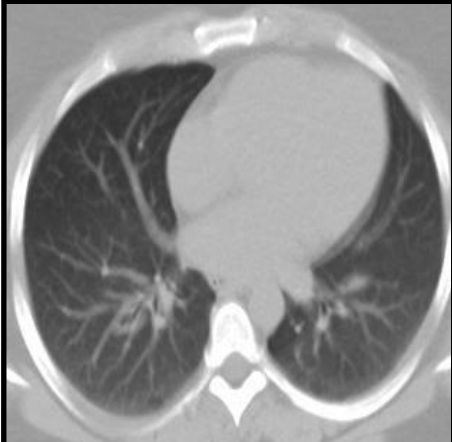
A Computerized Tomography Study

Henrik Reinius, M.D., Lennart Jonsson, M.D., Sven Gustafsson, M.D., Ph.D., Magnus Sundbom, M.D., Ph.D., Olov Duvernoy, M.D., Ph.D., Paolo Pelosi, M.D., Ph.D., Goran Hedenstierna, M.D., Ph.D., Filip Freden, M.D., Ph.D. **Anesthesiology** 2009

RM = CPAP 55 cmH₂O for 10 sec and/or PEEP=+10 cmH₂O



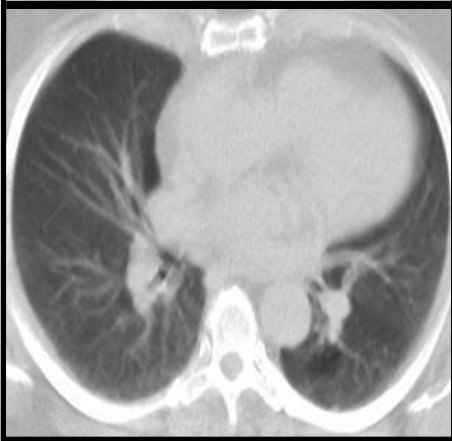
**P
E
E
P**



**R
M
+
P
E
E
P**



**R
M
+
Z
E
E
P**



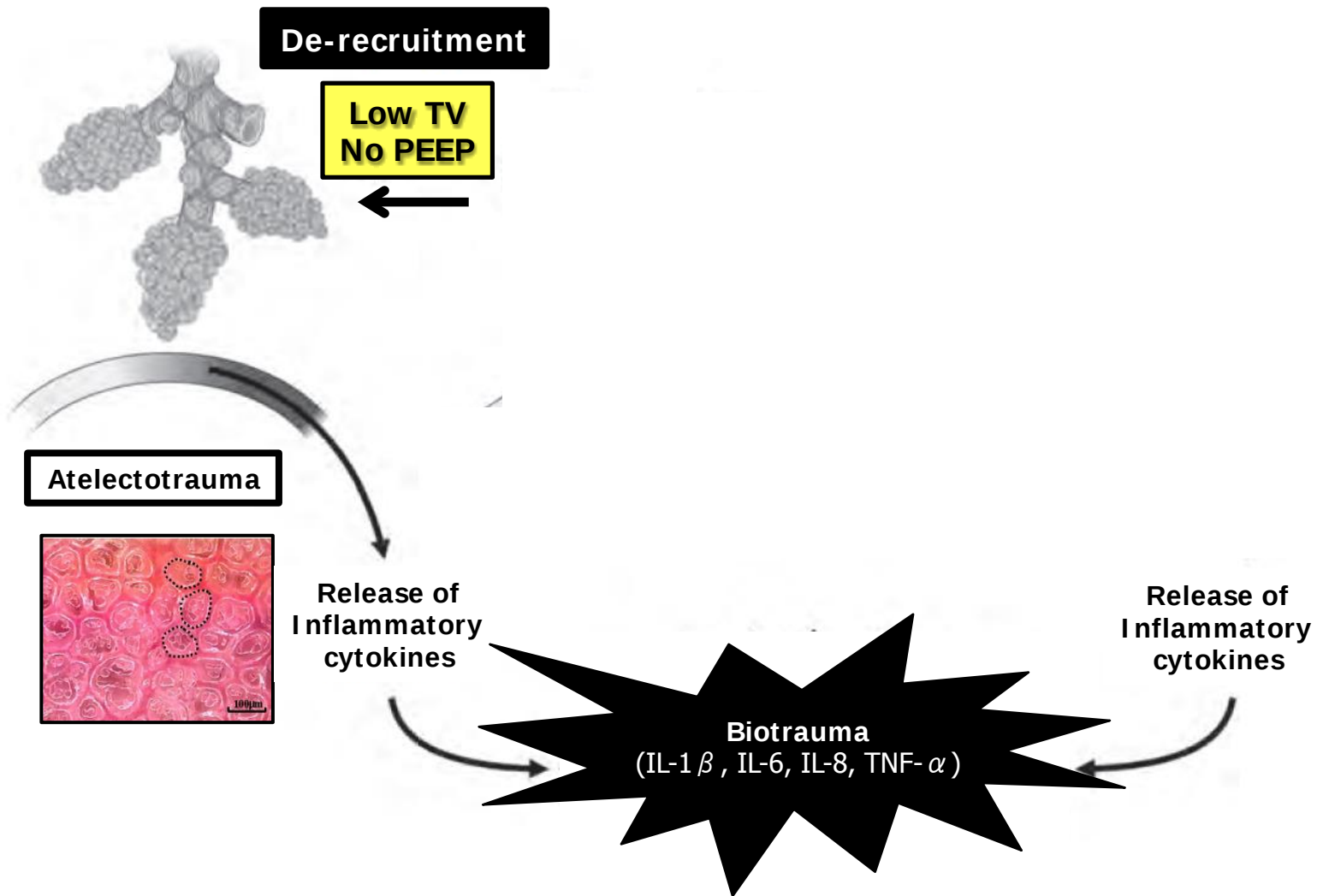
Awake

After induction

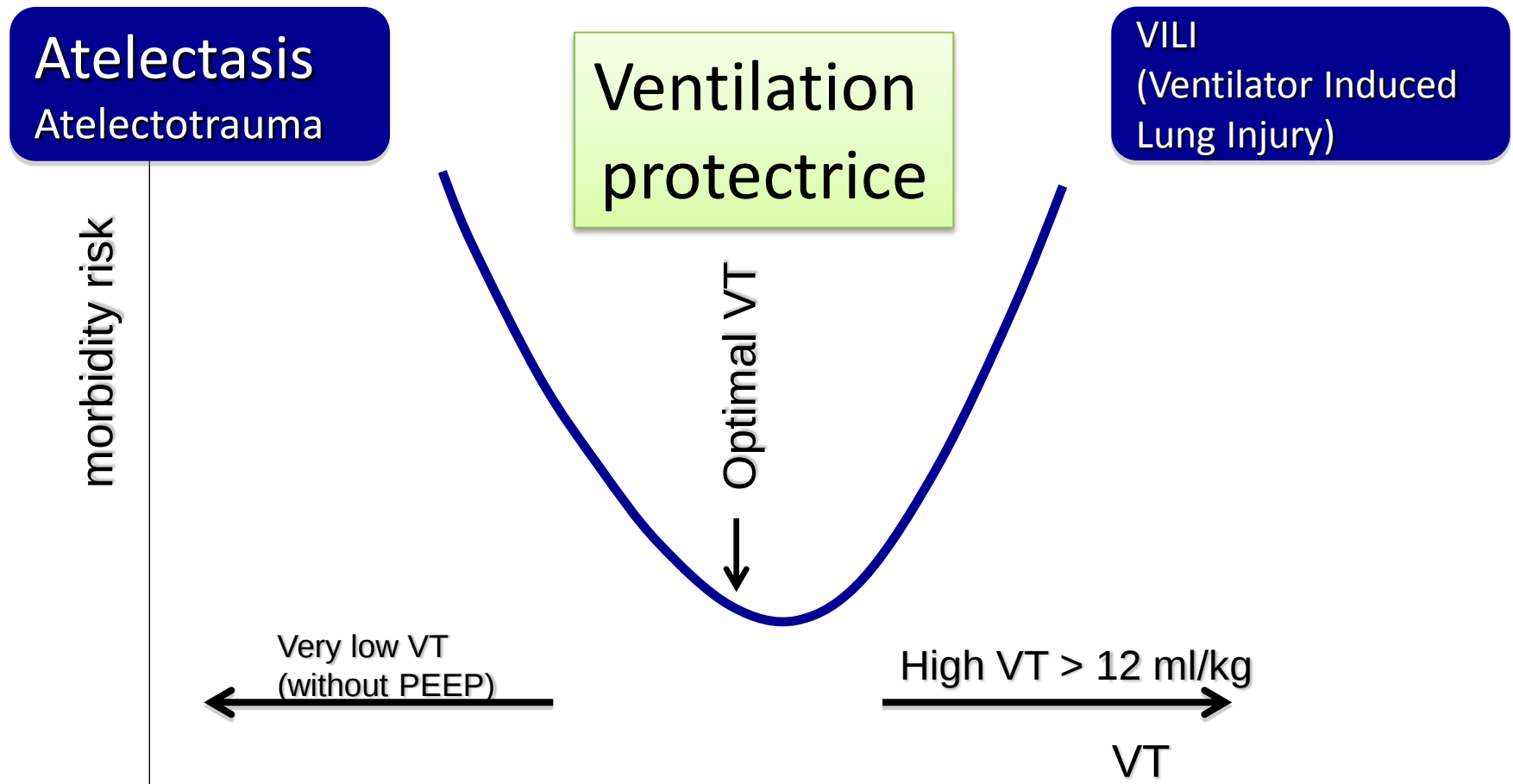
5 min

20 min

Ventilator-associated lung injury (VALI)



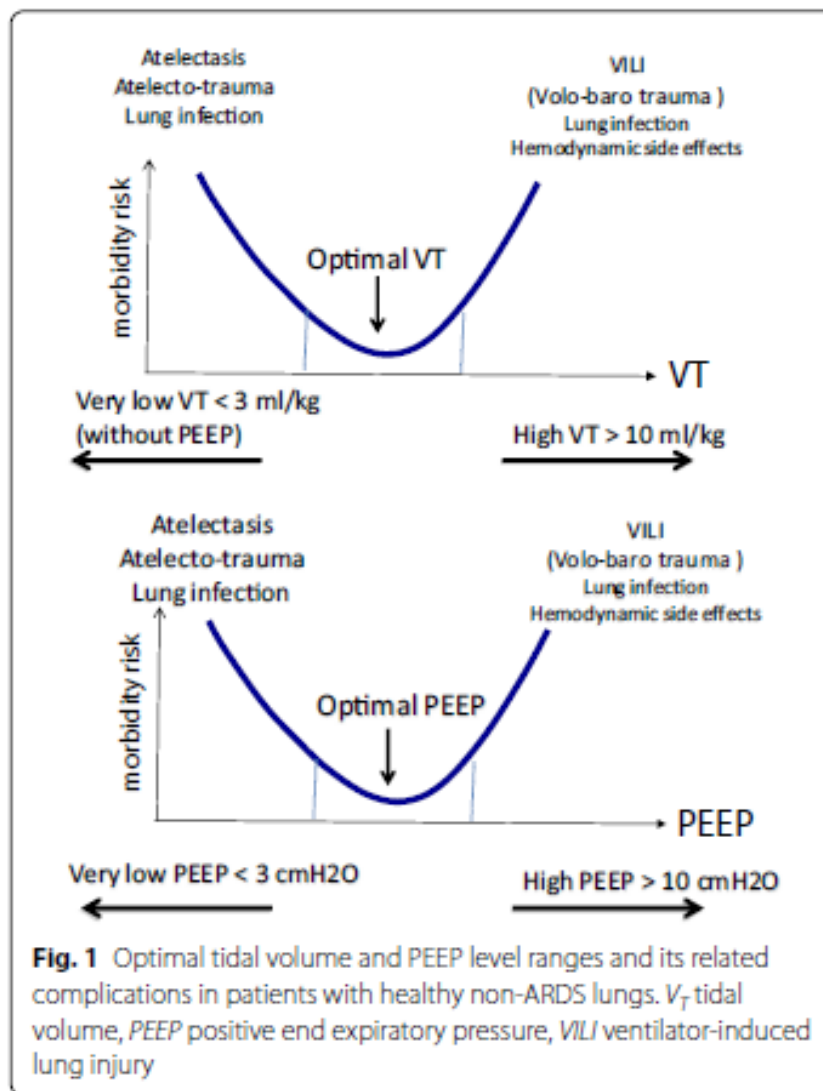
Tidal volume (VT) : how much is too much ?



What's new in mechanical ventilation in patients without ARDS: lessons from the ARDS literature

Ary Serpa Neto^{1,2,3} and Samir Jaber^{4*}

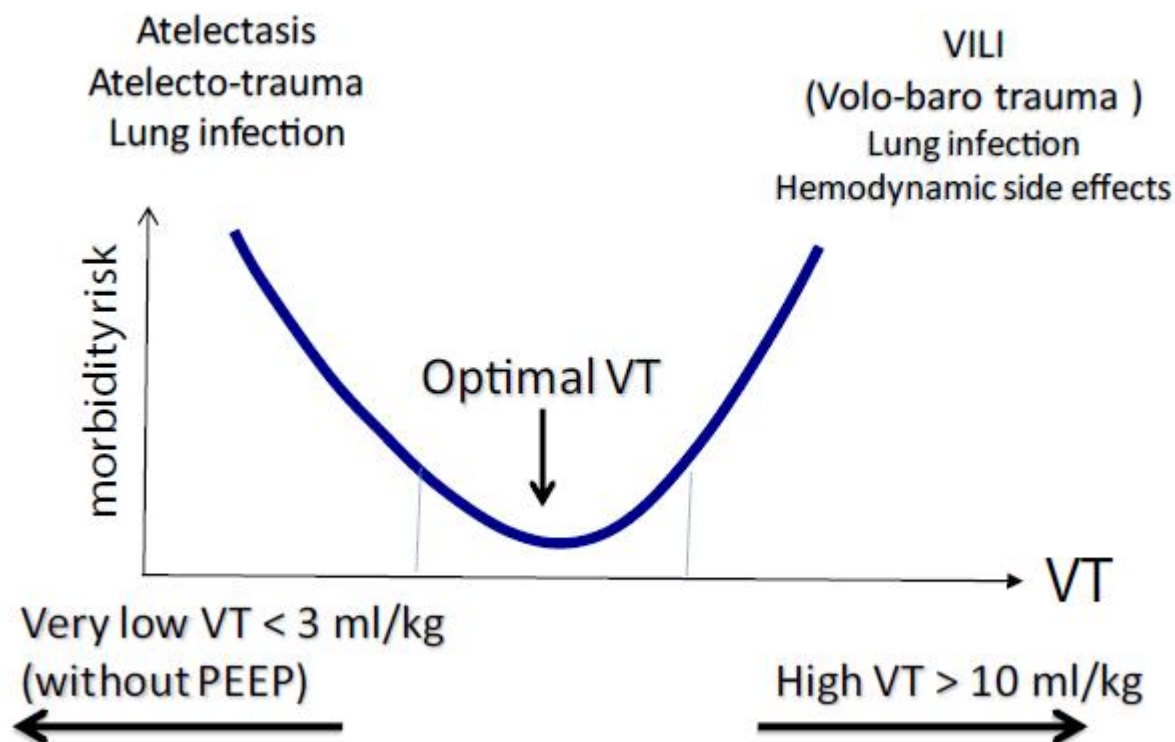
Intensive Care Med 2016 May



What's new in mechanical ventilation in patients without ARDS: lessons from the ARDS literature

Ary Serpa Neto^{1,2,3} and Samir Jaber^{4*}

$$4 < VT < 6 \text{ ml/kg}$$



Prevention and care of respiratory failure in obese patients

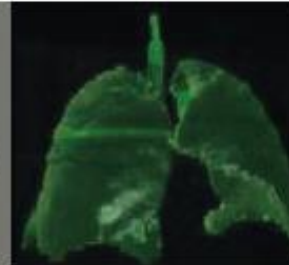
Jean Louis Pépin, Jean François Timsit, Renaud Tamisier, Jean Christian Borel, Patrick Lévy, Samir Jaber

A

Obese
Sex: female
Age: 53 years
Height: 155 cm
Weight: 125 kg
BMI=52 kg/m²



B



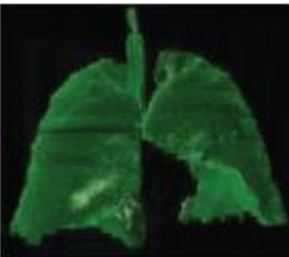
CT lung volume=3138 mL

C

Non-obese
Sex: female
Age: 65 years
Height: 164 cm
Weight: 57 kg
BMI=21 kg/m²



D



CT lung volume=3196 mL

Réglage du volume courant (VT)

- **Poids Idéal Théorique (P.I.T) :**

$$PIT = X + 0,91 (\text{taille en cm} - 152,4)$$

$$\text{Femmes : } X = 45,5$$

$$\text{Hommes : } X = 50$$

pour évaluer simplement

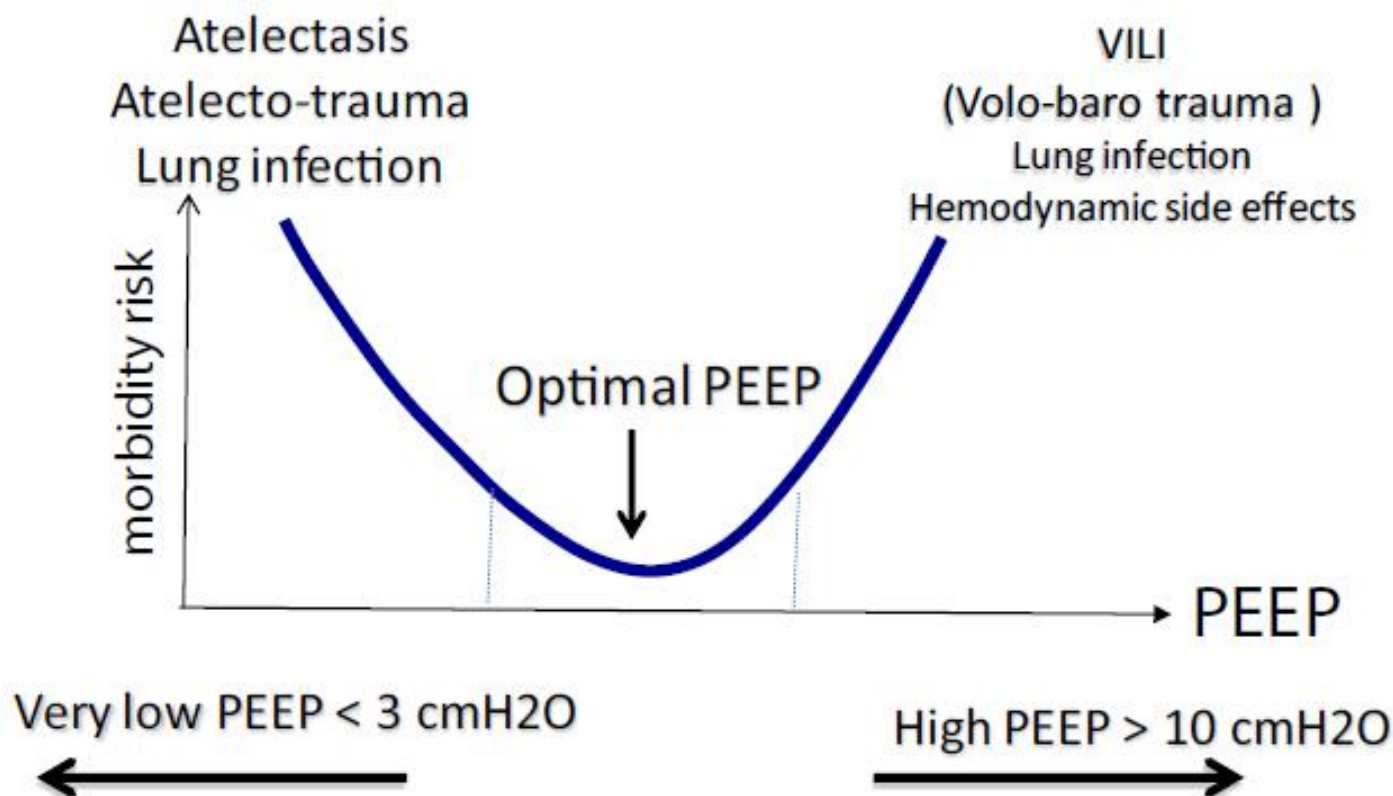
$$PIT \text{ (kg)} = \text{Taille (cm)} - 100 \quad \text{chez l'HOMME}$$

$$PIT \text{ (kg)} = \text{Taille (cm)} - 110 \quad \text{chez la FEMME}$$

What's new in mechanical ventilation in patients without ARDS: lessons from the ARDS literature

Ary Serpa Neto^{1,2,3} and Samir Jaber^{4*}

$5 < \text{PEEP} < 15 \text{ cmH}_2\text{O}$



Evolution of Mortality over Time in Patients Receiving Mechanical Ventilation

Andrés Esteban¹, Fernando Frutos-Vivar¹, Alfonso Muriel², Niall D. Ferguson³, Oscar Peñuelas¹, Victor Abaira², Konstantinos Raymondos⁴, Fernando Rios⁵, Nicolas Nin¹, Carlos Apezteguía⁵, Damian A. Violi⁶, Arnaud W. Thille⁷, Laurent Brochard⁸, Marco González⁹, Asisclo J. Villagomez¹⁰, Javier Hurtado¹¹, Andrew R. Davies¹², Bin Du¹³, Salvatore M. Maggiore¹⁴, Paolo Pelosi¹⁵, Luis Soto¹⁶, Vinko Tomicic¹⁷, Gabriel D'Empaire¹⁸, Dimitrios Matamis¹⁹, Fekri Abroug²⁰, Rui P. Moreno²¹, Marco Antonio Soares²², Yaseen Arabi²³, Freddy Sandi²⁴, Manuel Jibaja²⁵, Pravin Amin²⁶, Younsuck Koh²⁷, Michael A. Kuiper²⁸, Hans-Henrik Bülow²⁹, Amine Ali Zeggwagh³⁰, and Antonio **Anzueto**³¹



AJRCCM Vol. 188, No. 2 **(2013)** pp. 220-230

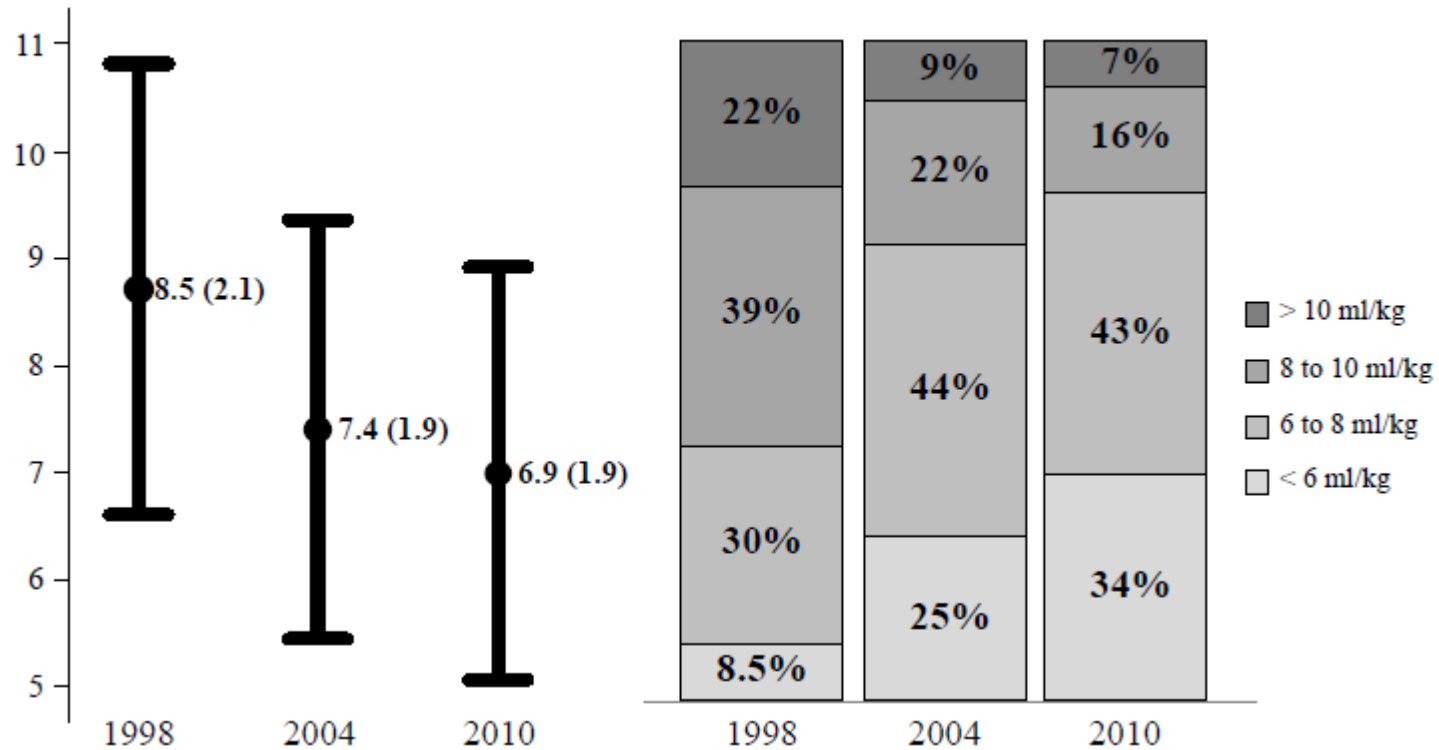


TABLE 2 - COMPARISON OF VARIABLES RELATED TO MANAGEMENT

	1998	2004	2010	P ^a
	N = 5183	N = 4968	N = 8151	
Non-invasive positive pressure ventilation prior to admission in the intensive care unit, n (%)	n.d.	n.d.	429 (5)	-
Non-invasive positive pressure ventilation at admission in the intensive care unit, n (%)	256(5)	479(10)	1169(14)	<0.001
Mode of ventilation, days of use per 1000 days of invasive mechanical ventilation (excluding days during weaning from mechanical ventilation process)				
Assist-control	627	412	330	<0.001
SIMV	66	21	34	<0.001
SIMV-PS	132	132	94	<0.001
Pressure support	65	125	237	<0.001
PCV	75	125	130	<0.001
APRV/BIPAP	n.d.	66	91	<0.001
PRVC	n.d.	98	61	<0.001
Other mode	35	21	23	<0.001
Tidal volume				
In ml/kg actual body weight, mean (SD)	8.8 (2.1)	7.6 (2.1)	6.9 (1.9)	<0.001
In ml/kg predicted body weight, mean (SD)	n.d.	9.3 (2.3)	8.2 (2.0)	<0.001
Total respiratory rate, mean (SD), breaths per minute	18 (11)	18 (6)	19 (6)	<0.001

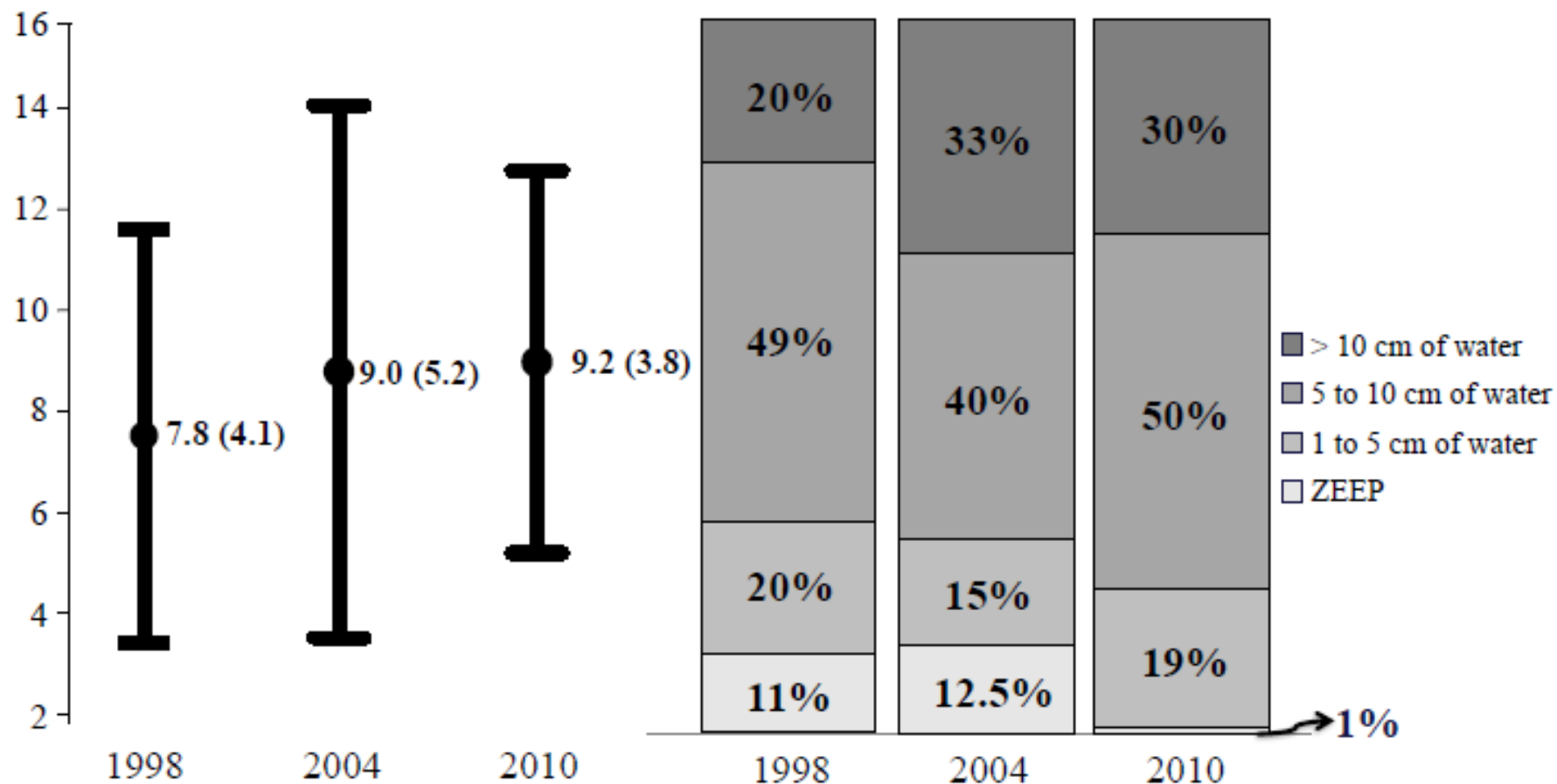
volume courant (VT)

Fig 1



PEEP

Fig 2



Objectifs

1. Qui sont les patients à risque?

2. Qu'est ce que la ventilation protectrice et préventive ?

3. Quelles sont les évidences ?

4. Take Home Messages

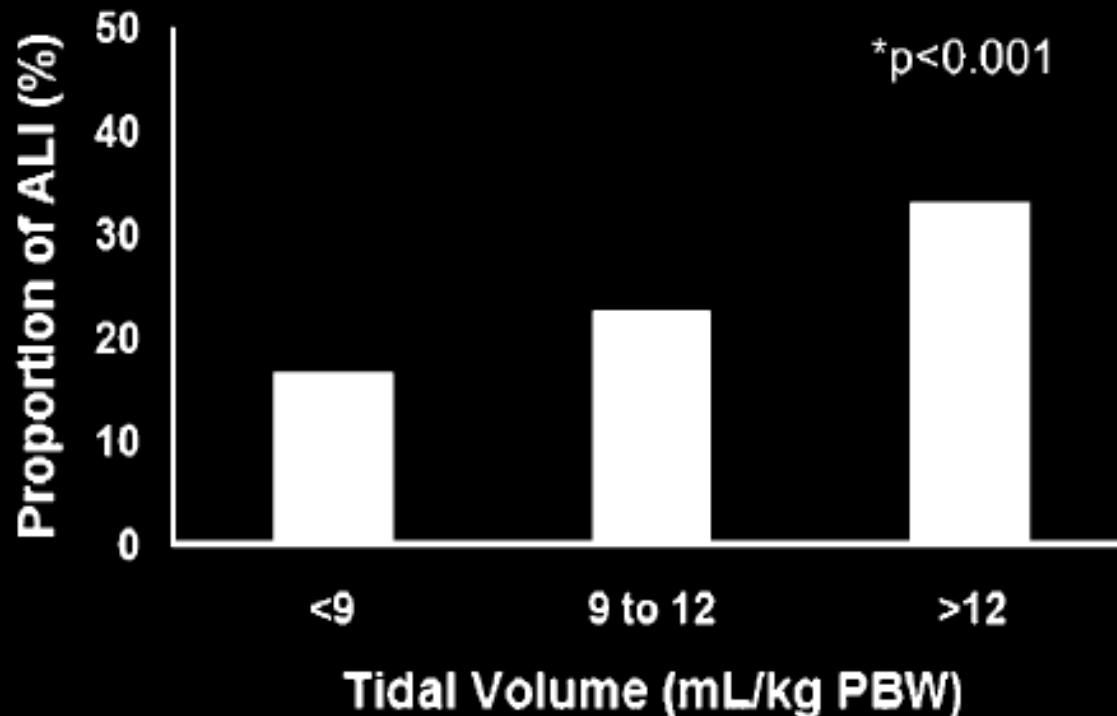
**Is there a rationale to
use lung protective
ventilation in patients
with normal lungs ?**

Ventilator-associated lung injury in patients without acute lung injury at the onset of mechanical ventilation

Ognjen Gajic, MD; Saqib I. Dara, MD; Jose L. Mendez, MD; Adebola O. Adesanya, MD; Emir Festic, MD; Sean M. Caples, MD; Rimki Rana, MD; Jennifer L. St. Sauver, PhD; James F. Lymp, PhD; Bekele Afessa, MD; Rolf D. Hubmayr, MD

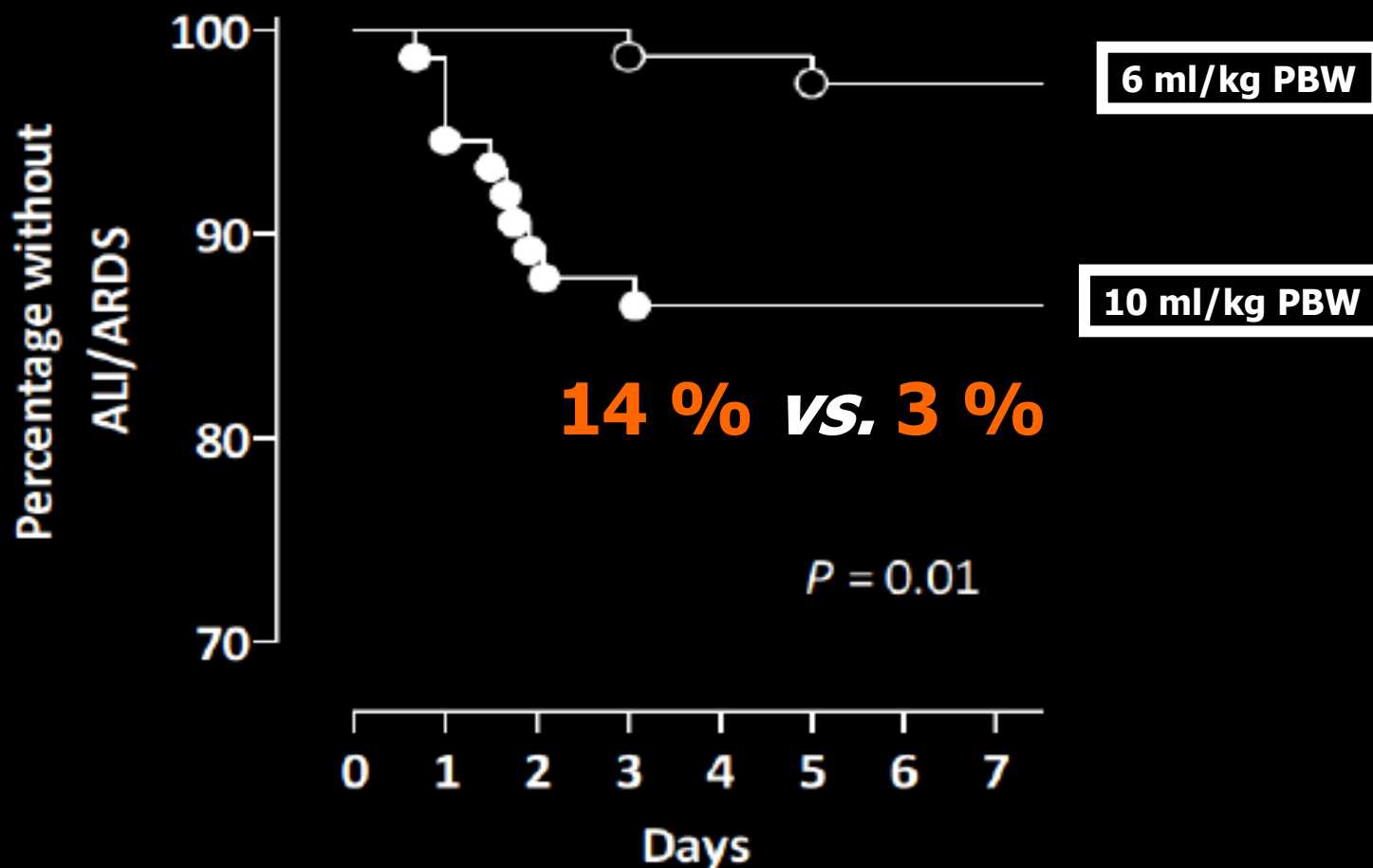
**A retrospective cohort study of 332 patients with « normal lungs » at the onset of mechanical ventilation and who received mechanical ventilation for ≥ 24 h
4 ICUs of a tertiary referral center**

Incidence ALI: N=80 patients (24%)



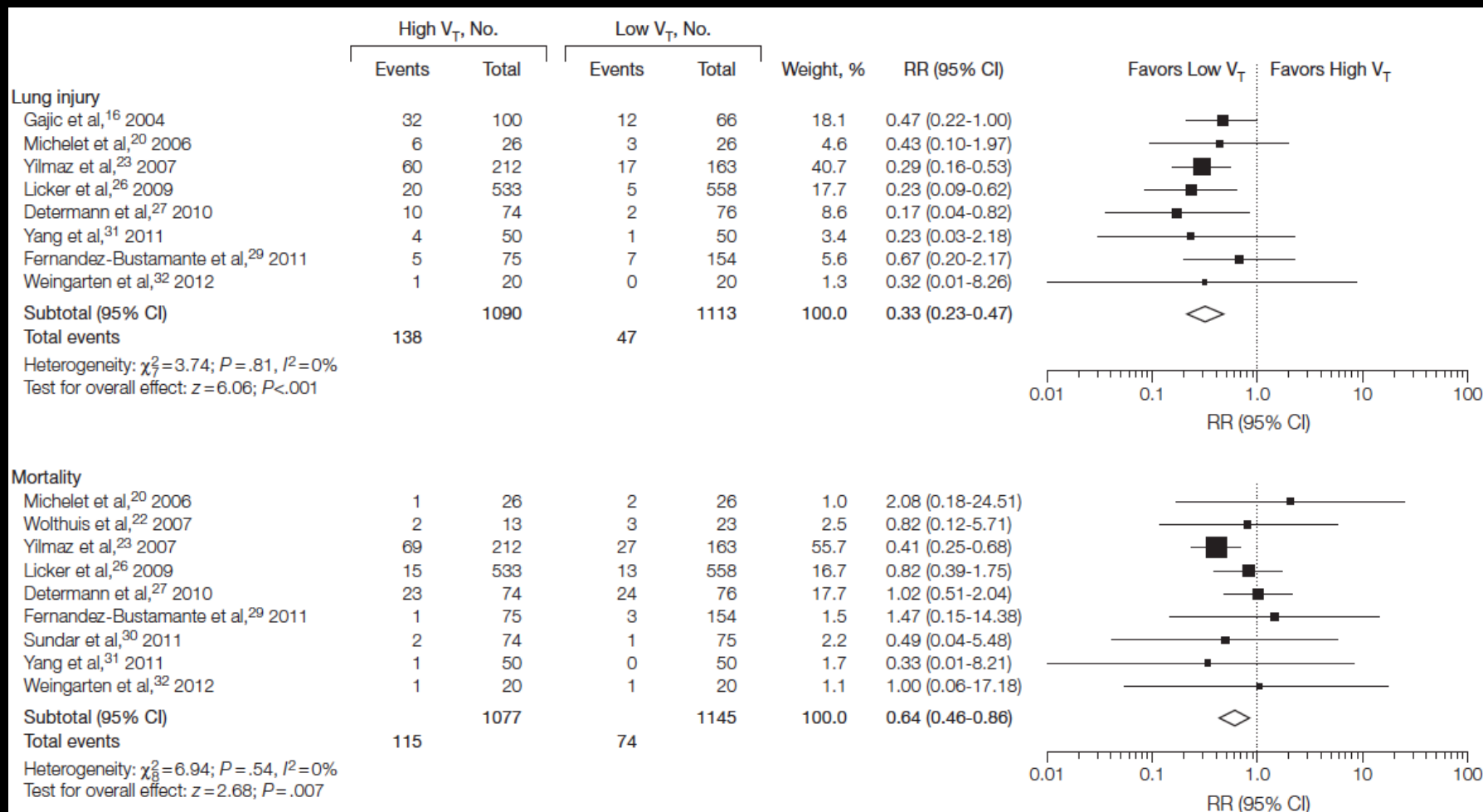
Ventilation with lower tidal volumes as compared to conventional tidal volumes for patients without acute lung injury - A preventive randomized controlled trial

Rogier M Determann, Annick Royakkers, Esther K Wolthuis, Alexander P Vlaar, Goda Choi, Frederique Paulus, Jorrit-Jan Hofstra, Mart J de Graaff, Johanna C Korevaar and Marcus J Schultz. . **Crit care 2010**



Lower Tidal Volumes in Patients without Preexisting Lung Injury

Serpa Neto A et al , JAMA. 2012 Oct 24;308(16):1651-9



Lower Tidal Volumes in Patients without Preexisting Lung Injury

Serpa Neto A et al , JAMA. 2012 Oct 24;308(16):1651-9

Table 3. Summary of Stratified Analyses of Pooled Relative Risks

Stratified Analysis	No. of Trials	No. of Patients	Risk Ratio (95% CI)	<i>P</i> Value	Heterogeneity, <i>Q</i>
Acute Lung Injury					
Setting					
Operation room	5	1512	0.34 (0.18-0.63)	<.001	0.73
ICU	3	691	0.33 (0.21-0.51)	<.001	0.43
Mortality					
Setting					
Operation room	6	1661	0.86 (0.46-1.60)	.63	0.94
ICU	3	561	0.57 (0.38-0.84)	.005	0.10

Lower Tidal Volumes in Patients without Preexisting Lung Injury in ICU: A Metaanalysis

Serpa Neto A et al , Curr Opin Crit Care 2014, 20:25–32

Author	Year	Effect of low tidal volume			
		Duration mechanical ventilation	Mortality	Inflammatory markers	Development of ARDS
Lee <i>et al.</i> [39]	1990	↓	↔	NR	↓
Gajic <i>et al.</i> [40]	2004	NR	↔	NR	↓
Wolthuis <i>et al.</i> [41]	2007	↔	↔	NR	NR
Yilmaz <i>et al.</i> [42]	2007	↓	↓	NR	↓
Determann <i>et al.</i> [8]	2010	↔	↔	↓	↓
Pinheiro de Oliveira <i>et al.</i> [43]	2010	↔	↔	↓	NR

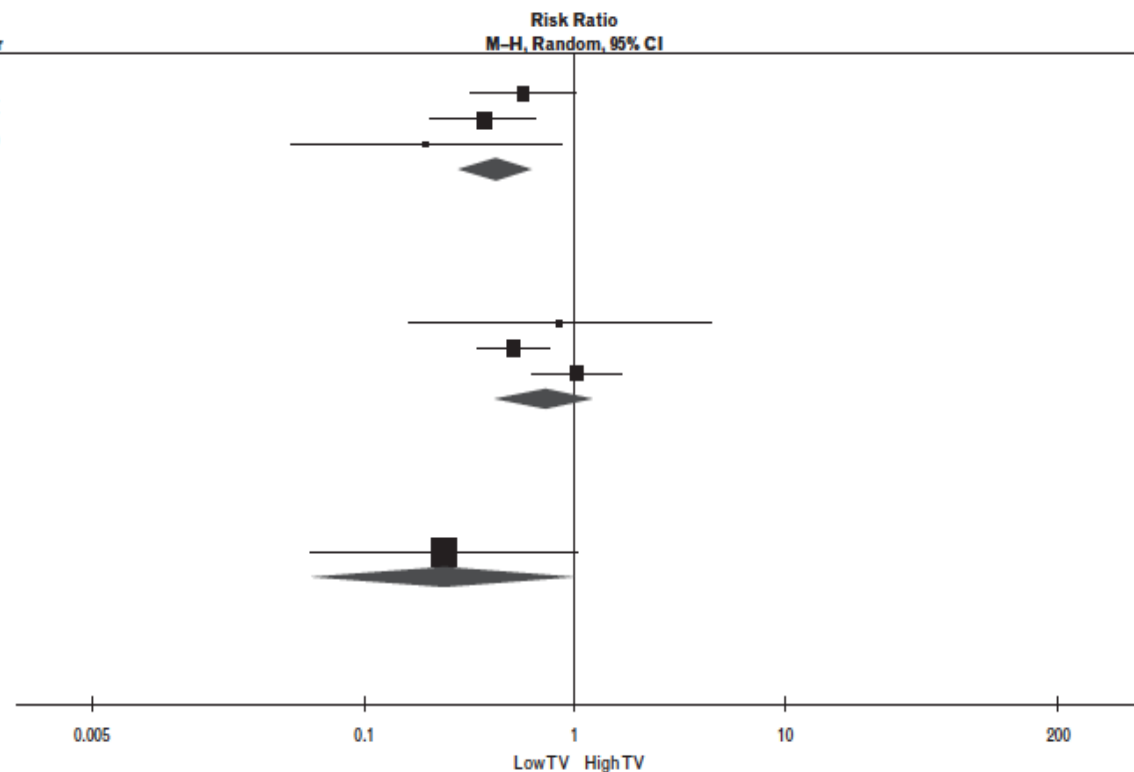
Lower Tidal Volumes in Patients without Preexisting Lung Injury in ICU: A Metaanalysis

Serpa Neto A et al , Curr Opin Crit Care 2014, 20:25–32

Study or Subgroup	Low TV		High TV		Weight	Risk Ratio M-H,Random,95%CI	Year
	Events	Total	Events	Total			
1.2.1 Lung Injury							
Gajic 2004 [40]	12	66	32	100	40.4%	0.57 [0.32, 1.02]	2004
Yilmaz 2007 [42]	17	163	60	212	52.0%	0.37 [0.22, 0.61]	2007
Determann 2010 [8]	2	76	10	74	7.6%	0.19 [0.04, 0.86]	2010
Subtotal (95%CI)		305		386	100.0%	0.42 [0.28, 0.63]	
Total events	31		102				
Heterogeneity: Tau ² = 0.02; chi ² = 2.35, df = 2 (P = 0.31); I ² = 15%							
Test for overall effect: Z = 4.10 (P < 0.0001)							

1.2.2 Mortality							
Wolthuis 2007 [41]	3	23	2	13	9.7%	0.85 [0.16, 4.44]	2007
Yilmaz 2007 [42]	27	163	69	212	47.4%	0.51 [0.34, 0.76]	2007
Determann 2010 [8]	24	76	23	74	42.9%	1.02 [0.63, 1.63]	2010
Subtotal (95%CI)	262	299	299	299	100.0%	0.72 [0.41, 1.26]	
Total events							
Heterogeneity: $\tau^2 = 0.13$; $\chi^2 = 4.95$, $df = 2$ ($P = 0.08$); $I^2 = 60\%$							
Test for overall effect: $Z = 1.15$ ($P < 0.25$)							

1.2.3 Pulmonary Infection							
Lee 1999 [39]	2	47	10	56	100.0%	0.24 [0.05, 1.03]	1999
Subtotal (95%CI)	47	56	56	56	100.0%	0.24 [0.05, 1.03]	
Total events	2		10				
Heterogeneity: Not applicable							
Test for overall effect: $Z = 1.92$ ($P < 0.06$)							



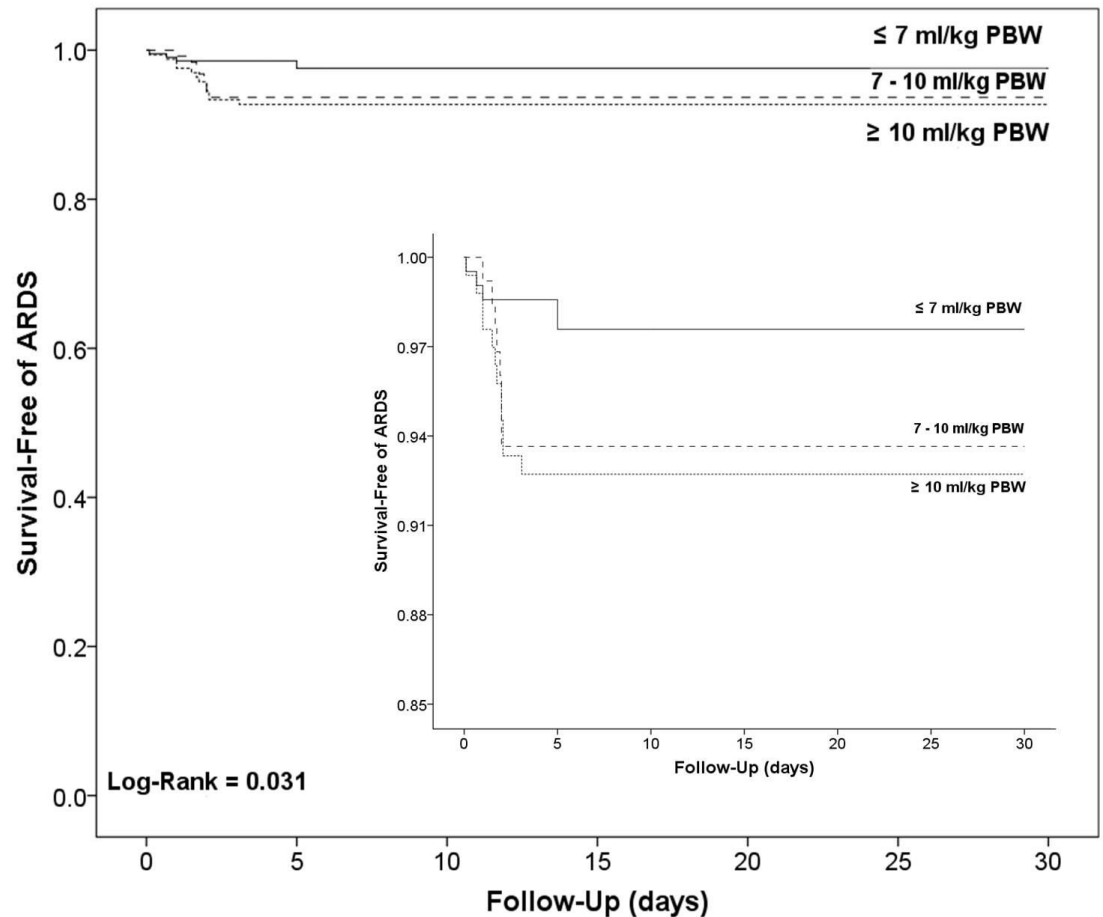
PROTECTIVE VENTILATION IN ICU



LUNG PROTECTIVE VENTILATION WITH LOWER TIDAL VOLUMES TO PREVENT ACUTE RESPIRATORY DISTRESS SYNDROME IN INTENSIVE CARE UNIT PATIENTS UNDER MECHANICAL VENTILATION: A systematic review and meta-analysis using individual data of 2,184 patients

Ary Serpa Neto MD MSc, Sabine NT Hemmes MD, Carmen SV Barbas MD PhD, Michelle Biehl MD, Rogier M Determann MD PhD, Jonathan Elmer MD PhD, Gilberto Friedman MD PhD, Ognjen Gajic MD PhD, Rita Linko MD PhD, Roselaine Pinheiro de Oliveira MD PhD, Esther K Wolthuis MD PhD, Marcelo Gama de Abreu MD PhD, Paolo Pelosi MD, Marcus J Schultz MD PhD

PROTECTIVE VENTILATION IN ICU

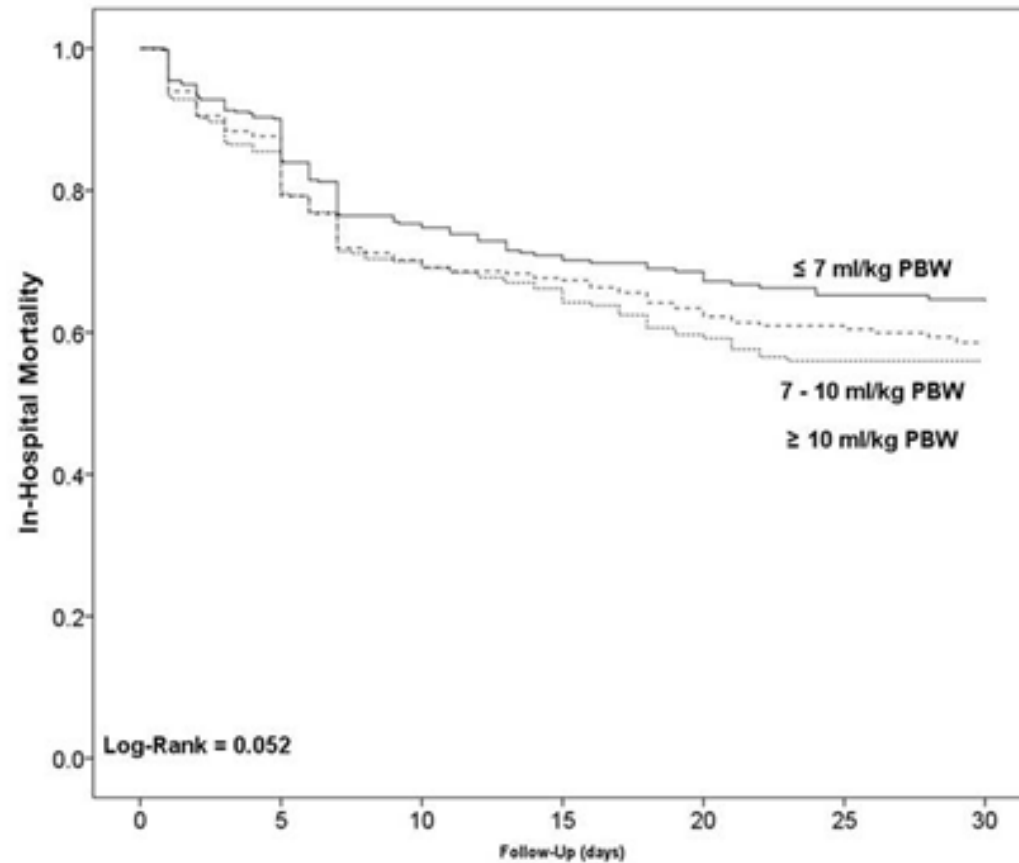


Number at Risk

	0	5	10	15	20	25	30
≤ 7 ml/kg PBW	506	387	278	192	101	85	34
7 - 10 ml/kg PBW	625	495	398	287	207	121	93
≥ 10 ml/kg PBW	464	293	198	117	78	54	29

PROTECTIVE VENTILATION IN ICU

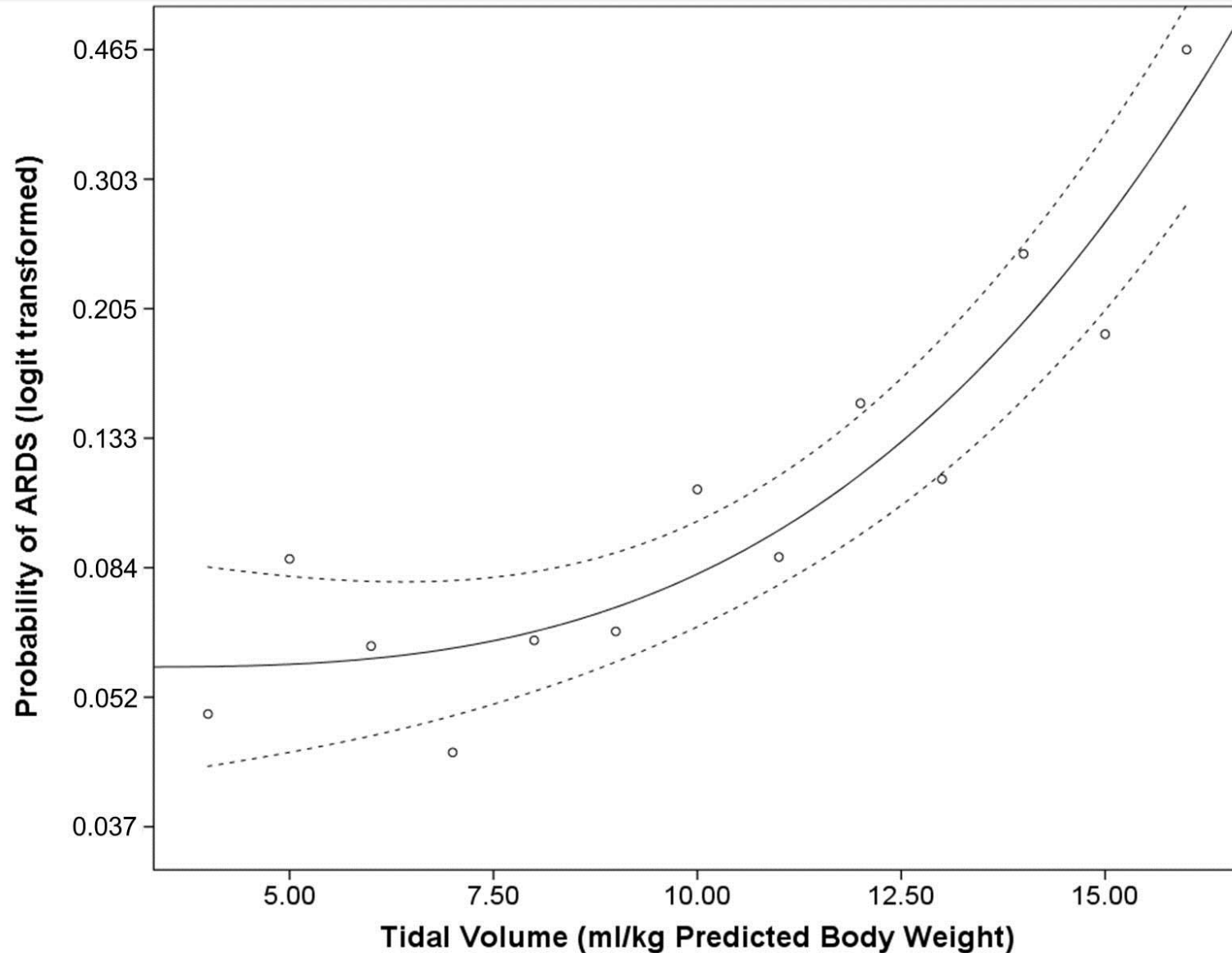
mortality



Number at Risk

≤ 7 ml/kg PBW	512	385	239	179	142	93	35
7 - 10 ml/kg PBW	598	436	243	189	145	101	45
≥ 10 ml/kg PBW	498	353	199	149	110	72	29

PROTECTIVE VENTILATION IN ICU



... and in the operating room,

What is the evidence ?

Lung Protective Mechanical Ventilation

Low TV and PEEP

In ICU patients

In the OR

Patients with
injured lungs

Patients with
healthy lungs

Patients with
healthy lungs

Improved morbidity
and mortality: **YES**

Improved morbidity: **YES**

Impact on
Postoperative outcome ?

Recommendations:
Limit VT < 10 ml/kg PBW
5 < PEEP < 15 cmH₂O

Recommendations:
VT ?
PEEP ?

1963

The New England Journal of Medicine

NOVEMBER 7, 1963

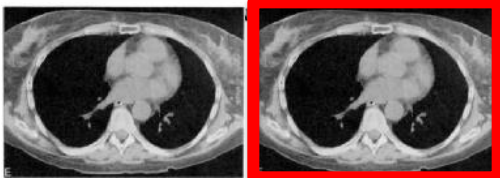
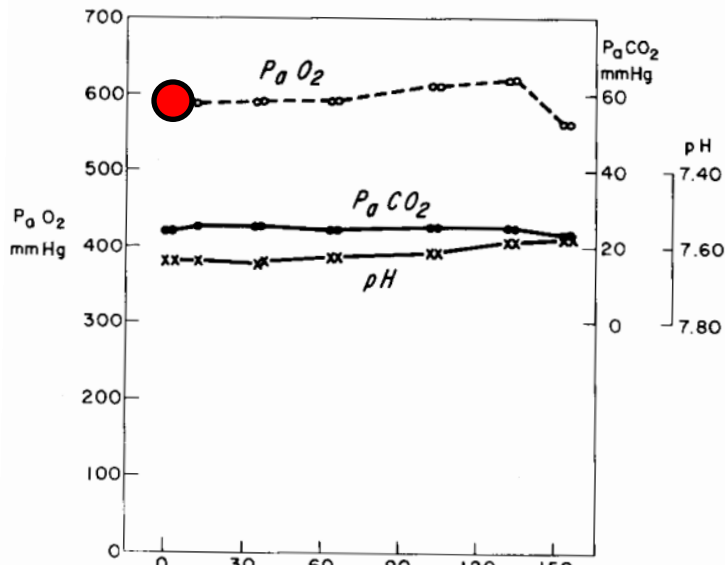
Number 19

IMPAIRED OXYGENATION IN SURGICAL PATIENTS DURING GENERAL ANESTHESIA WITH CONTROLLED VENTILATION*

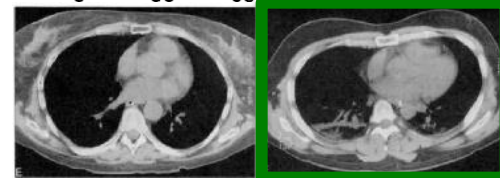
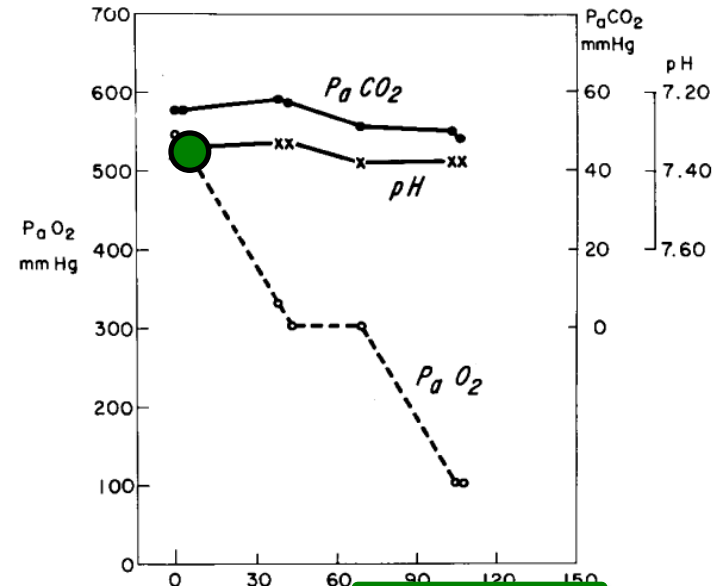
A Concept of Atelectasis

H. H. BENDIXEN, M.D.,† J. HEDLEY-WHYTE, M.B., B.CHIR.,‡ AND M. B. LAVER, M.D.§

Large TV and zero PEEP



Low TV and zero PEEP



Protective Mechanical Ventilation during General Anesthesia for Open Abdominal Surgery Improves Postoperative Pulmonary Function

Paolo Severgnini, Gabriele Selmo, Christian Lanza, Alessandro Chiesa, Alice Frigerio, Alessandro Bacuzzi, Gianlorenzo Dionigi, Raffaele Novario, Cesare Gregoretti, Marcelo Gama de Abreu, Marcus J. Schultz, Samir Jaber, Emmanuel Futier, Maurizio Chiaranda, **Paolo Pelosi** *Anesthesiology* 2013; 118:1307-21

N = 56 Patients undergoing elective open abdominal surgery

STANDARD VENTILATION

VT = 9 ml/kg PBW
zero PEEP

VS.

LUNG-PROTECTIVE VENTILATION

VT = 7 ml/kg PBW
10 cmH₂O PEEP + RM



Need for large RCT in Protective ventilation in anesthesia

● Protective Ventilation During General Anesthesia for Open Abdominal Surgery (PROVHILO study)



High versus low positive end-expiratory pressure during general anaesthesia for open abdominal surgery (PROVHILO trial): a multicentre randomised controlled trial

The PROVE Network Investigators* for the Clinical Trial Network of the European Society of Anaesthesiology *Lancet* 2014; 384: 495-503

The NEW ENGLAND JOURNAL of MEDICINE

N ENGL J MED 369;5 NEJM.ORG AUGUST 1, 2013

ORIGINAL ARTICLE

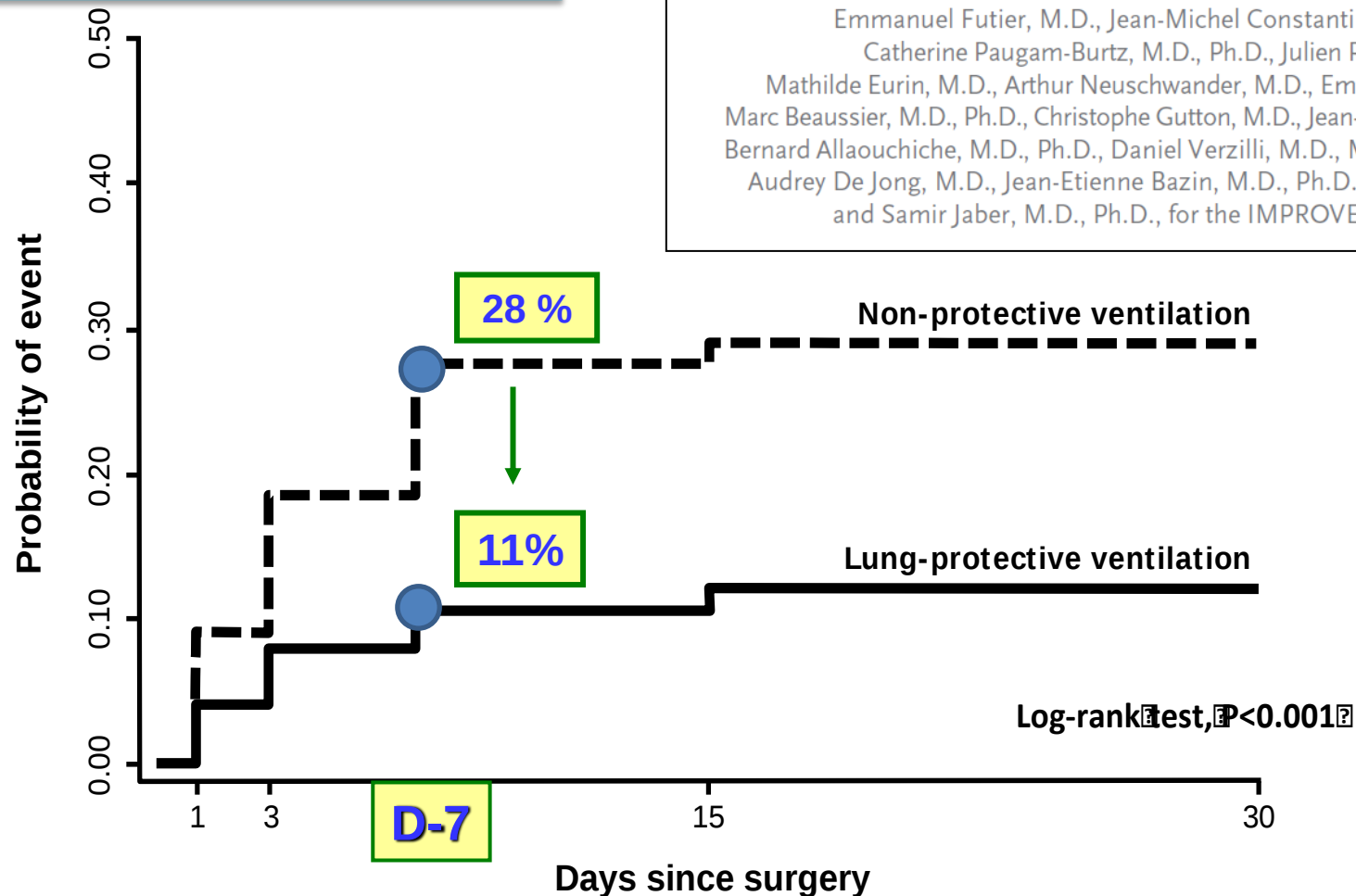
A Trial of Intraoperative Low-Tidal-Volume Ventilation in Abdominal Surgery

Emmanuel Futier, M.D., Jean-Michel Constantin, M.D., Ph.D.,
Catherine Paugam-Burtz, M.D., Ph.D., Julien Pascal, M.D.,
Mathilde Eurin, M.D., Arthur Neuschwander, M.D., Emmanuel Marret, M.D.,
Marc Beaussier, M.D., Ph.D., Christophe Gutton, M.D., Jean-Yves Lefrant, M.D., Ph.D.,
Bernard Allaouchiche, M.D., Ph.D., Daniel Verzilli, M.D., Marc Leone, M.D., Ph.D.,
Audrey De Jong, M.D., Jean-Etienne Bazin, M.D., Ph.D., Bruno Pereira, Ph.D.,
and Samir Jaber, M.D., Ph.D., for the IMPROVE Study Group*

Postoperative Pulmonary and Extra-pulmonary Complications

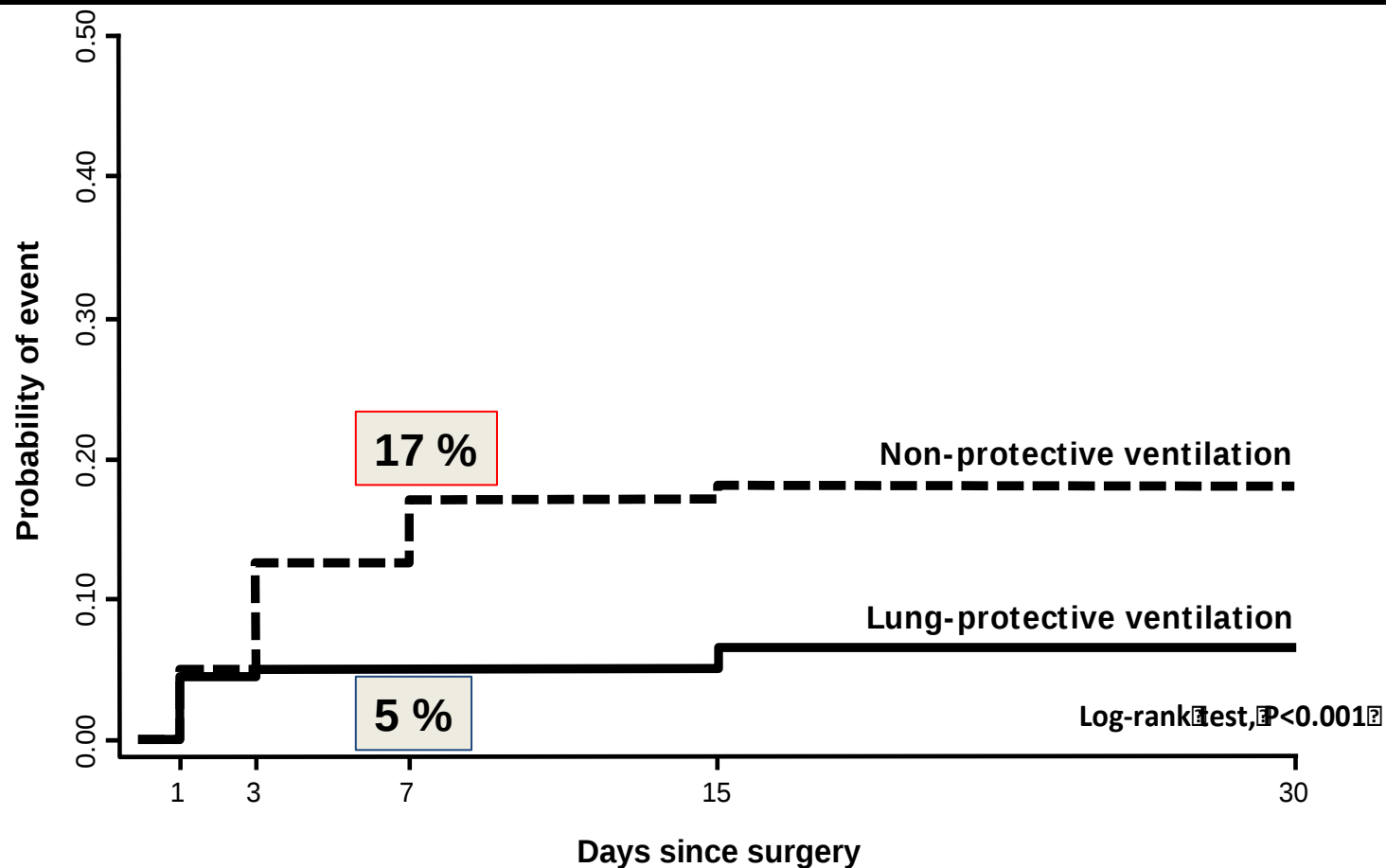
A Trial of Intraoperative Low-Tidal-Volume Ventilation in Abdominal Surgery

Emmanuel Futier, M.D., Jean-Michel Constantin, M.D., Ph.D.,
Catherine Paugam-Burtz, M.D., Ph.D., Julien Pascal, M.D.,
Mathilde Eurin, M.D., Arthur Neuschwander, M.D., Emmanuel Marret, M.D.,
Marc Beaussier, M.D., Ph.D., Christophe Gutton, M.D., Jean-Yves Lefrant, M.D., Ph.D.,
Bernard Allaouchiche, M.D., Ph.D., Daniel Verzilli, M.D., Marc Leone, M.D., Ph.D.,
Audrey De Jong, M.D., Jean-Etienne Bazin, M.D., Ph.D., Bruno Pereira, Ph.D.,
and Samir Jaber, M.D., Ph.D., for the IMPROVE Study Group*



Intraoperative Protective Ventilation in Abdominal Surgery : The IMPROVE Trial

Probability for requiring intubation or non-invasive ventilation for ARF to Postoperative Day 30



High versus low positive end-expiratory pressure during general anaesthesia for open abdominal surgery (PROVHILO trial): a multicentre randomised controlled trial

The PROVE Network Investigators for the Clinical Trial Network of the European Society of Anaesthesiology*

Vt = 8 ml/kg IBW

Patients scheduled for nonlaparoscopic abdominal surgery at high or intermediate or risk for postoperative pulmonary complications

Excluded

Informed consent

Excluded

Randomized
900 patients

Protective strategy
450 pt

Intra-operative
PEEP 12 cm H₂O
+ Recruitment maneuvers

Drop-out

Conventional strategy
450 pt

Intra-operative
PEEP 2 cm H₂O
- Recruitment maneuvers

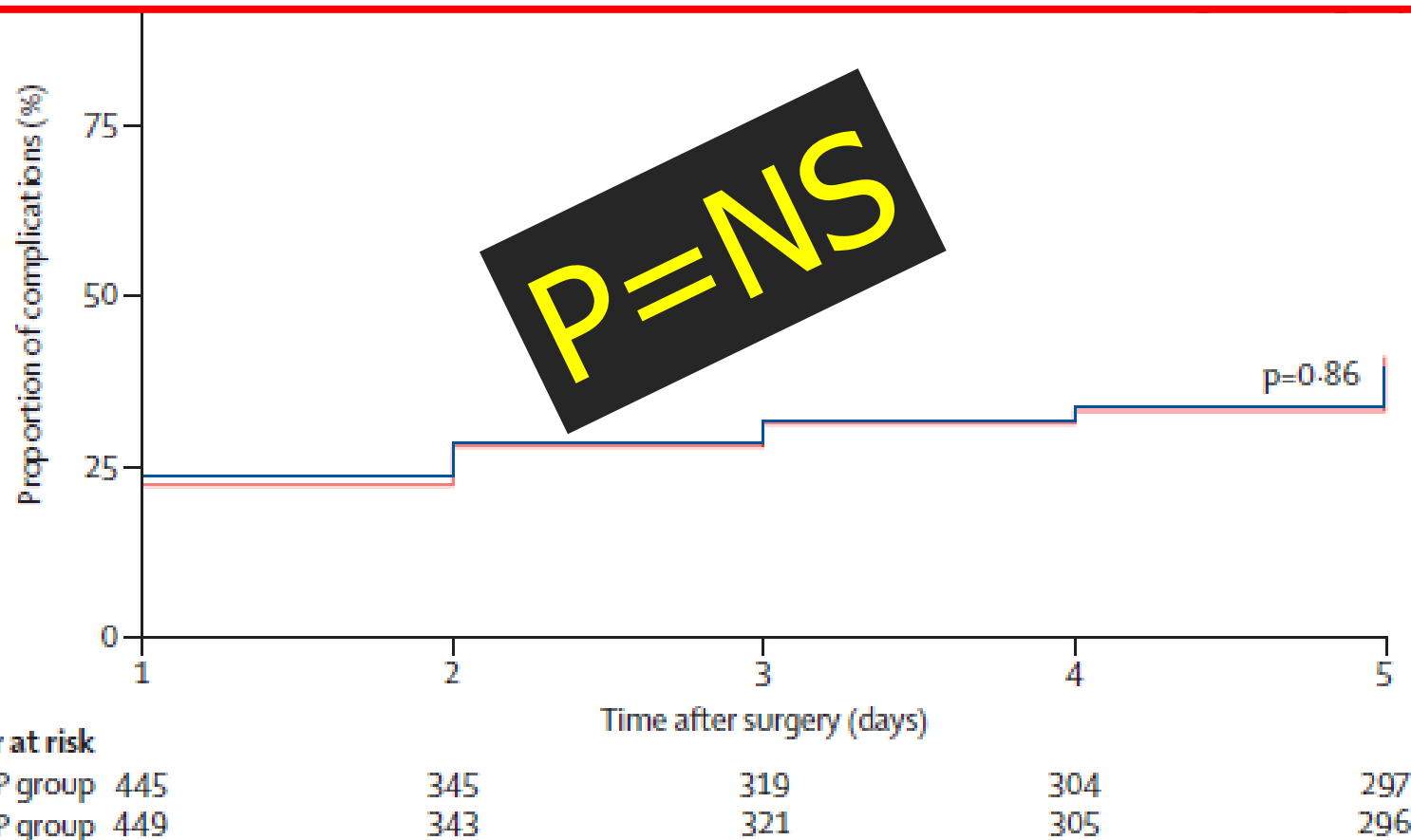
Drop-out

Post-operative Followup
Day 1,3,5 and
day before discharge or day 90 when still admitted

Lost to Follow-up

High versus low positive end-expiratory pressure during general anaesthesia for open abdominal surgery (PROVHILO trial): a multicentre randomised controlled trial

The PROVE Network Investigators* for the Clinical Trial Network of the European Society of Anaesthesiology



Controversy or complementary results ?

IMPROVE vs. PROVILO ?



IMPROVE trial

N=400

**PROVHILO trial**

N=900

**Non-protective group
(n=200)****Protective group
(n=200)****High PEEP group
(n=445)****Low PEEP group
(n=449)****VT, ml/kg IBW****11.1±1.1****6.4±0.8**

7.2±1.5

7.1±1.2

PEEP, cmH₂O**0****6 [6-8]**

12 [12-12]

2 [0-2]

RM**NO****CPAP 30-30
Every 30-45 min**Increase in VT (step of 4
cmH₂O) until a Pplat of
30-35 cmH₂O**NO****9 [6-12]**After intubation: 99%
Before extubation: 85%**FiO₂, %****47.2±7.6****46.4±7.3**

40 [40-49]

41 [40-50]

**Duration of
surgery****2-4 hr: 39.6%
4-6 hr: 39.1%
>6 hr: 21.4%****2-4 hr: 38.5%
4-6 hr: 39.0%
>6 hr: 22.6%**

200 [140-300] min

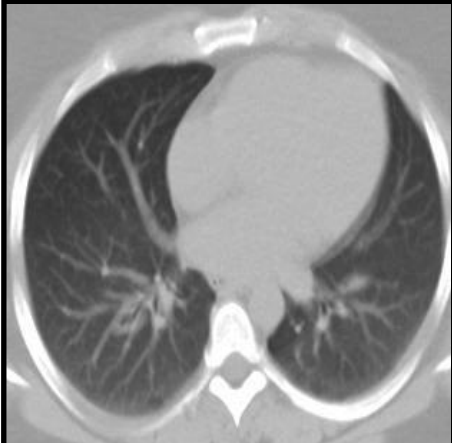
190 [140-262] min

**Laparoscopic
surgery****21 %**

Not included

***PEEP or RM
alone are not
enough !***

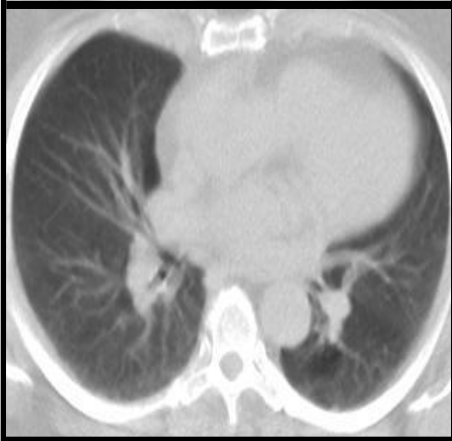
**P
E
E
P**



**R
M
+
P
E
E
P**



**R
M
+
Z
E
E
P**



Awake

After induction

5 min

20 min

QUESTION n°2

Quelle est la procédure qui marche le plus et/ou le mieux ?



« petit
volume courant »
(6 ml/kg)

ou



PEP

ou



Manœuvre de
recrutement

Réponse (probable) n°2

Quelle est la procédure qui marche le plus et/ou le mieux ?



« petit
volume courant »
(6 ml/kg)



PEP



Manœuvre de
recrutement

Réponse (probable) n°2

Quelle est la procédure qui marche le plus et/ou le mieux ?



« petit
volume courant »
(6 ml/kg)

~~X~~
+

PEP

~~X~~
+

Manœuvre de
recrutement

= Association des 3 procédures ++
= VENTILATION PROTECTRICE

Intraoperative Mechanical Ventilation

INTRAOPERATIVE VENTILATOR SETTINGS AND POSTOPERATIVE ACUTE RESPIRATORY DISTRESS SYNDROME:

An individual data meta-analysis of 3,659 patients

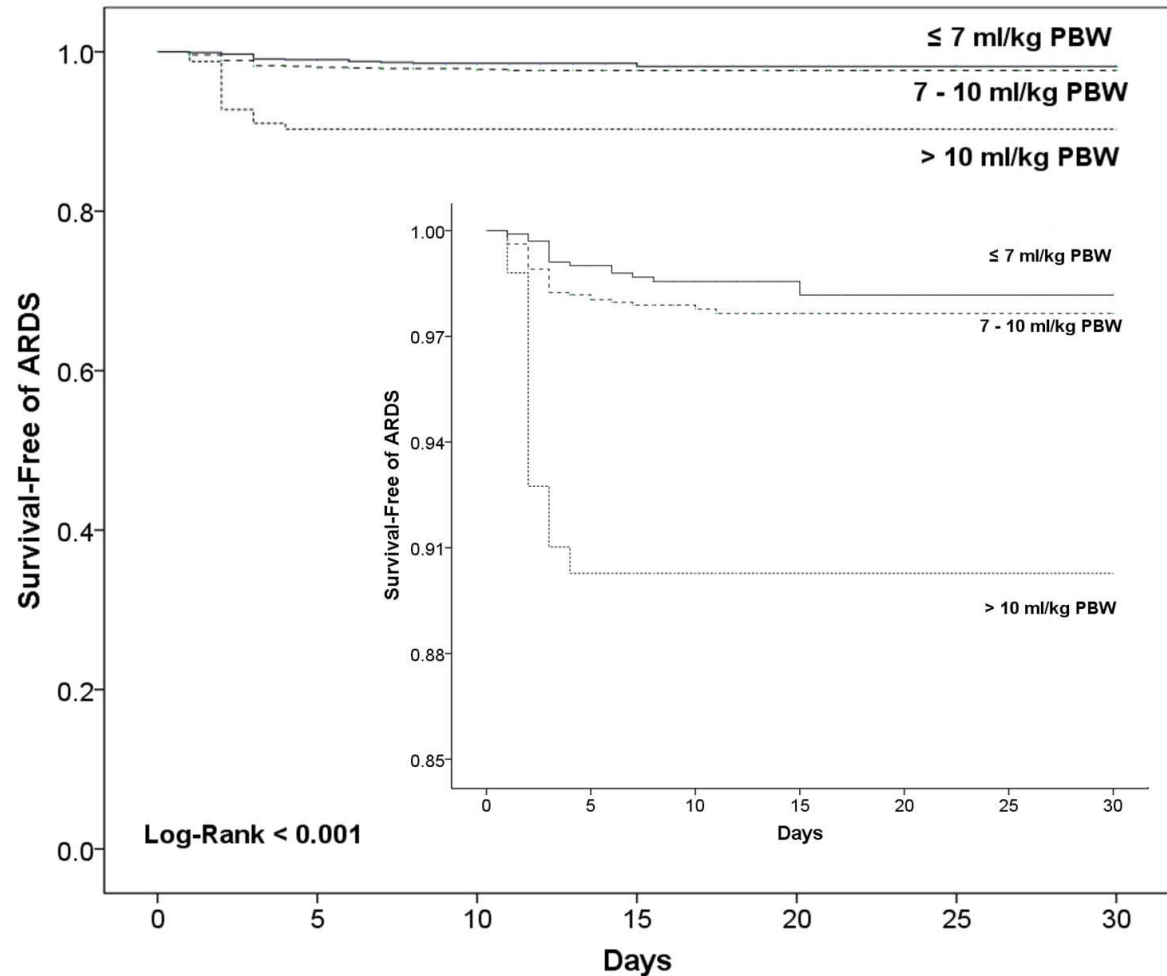
Incidence of mortality and morbidity related to postoperative lung injury in patients who have undergone abdominal or thoracic surgery: a systematic review and meta-analysis

Ary Serpa Neto, Sabine NT Hemmes, Carmen SV Barbas, Martin Beiderlinden, Ana Fernandez-Bustamante, Emmanuel Futier, Markus W Hollmann, Samir Jaber, Alf Kozian, Marc Licker, Wen-Qian Lin, Pierre Moine, Federica Scavonetto, Thomas Schilling, Gabriele Selmo, Paolo Severgnini, Juraj Sprung, Tanja Treschan, Carmen Unzueta, Toby N Weingarten, Esther K Wolthuis, Hermann Wrigge, Marcelo Gama de Abreu, Paolo Pelosi, Marcus J Schultz, for the the PROVE Network investigators

Serpa Neto et al. LANCET, respiratory 2014.

Intraoperative Mechanical Ventilation

**PROTECTIVE
VENTILATION
NETWORK**



Number at Risk

	0	5	10	15	20	25	30
≤ 7 ml/kg PBW	1,004	785	429	202	128	92	37
7 - 10 ml/kg PBW	1,512	1,144	674	356	201	139	59
> 10 ml/kg PBW	403	285	179	109	71	48	18

Original Investigation | CARING FOR THE CRITICALLY ILL PATIENT

Effect of Noninvasive Ventilation on Tracheal Reintubation Among Patients With Hypoxemic Respiratory Failure Following Abdominal Surgery A Randomized Clinical Trial

Samir Jaber, MD, PhD; Thomas Lescot, MD, PhD; Emmanuel Futier, MD, PhD; Catherine Paugam-Burtz, MD, PhD; Philippe Seguin, MD, PhD; Martine Ferrandiere, MD; Sigismund Lasocki, MD, PhD; Olivier Mimoz, MD, PhD; Baptiste Hengy, MD; Antoine Sannini, MD; Julien Pottecher, MD; Paër-Sélim Abback, MD; Beatrice Riou, MD; Fouad Belafia, MD; Jean-Michel Constantin, MD, PhD; Elodie Masseret, MD; Marc Beaussier, MD, PhD; Daniel Verzilli, MD; Audrey De Jong, MD; Gerald Chanques, MD, PhD; Laurent Brochard, MD, PhD; Nicolas Molinari, PhD; for the NIVAS Study Group

IMPORTANCE It has not been established whether noninvasive ventilation (NIV) reduces the need for invasive mechanical ventilation in patients who develop hypoxemic acute respiratory failure after abdominal surgery.

OBJECTIVE To evaluate whether noninvasive ventilation improves outcomes among patients developing hypoxemic acute respiratory failure after abdominal surgery.

DESIGN, SETTING, AND PARTICIPANTS Multicenter, randomized, parallel-group clinical trial conducted between May 2013 and September 2014 in 20 French intensive care units among 293 patients who had undergone abdominal surgery and developed hypoxemic respiratory failure (partial oxygen pressure <60 mm Hg or oxygen saturation [SpO₂] ≤90% when breathing room air or <80 mm Hg when breathing 15 L/min of oxygen, plus either [1] a respiratory rate above 30/min or [2] clinical signs suggestive of intense respiratory muscle work and/or labored breathing) if it occurred within 7 days after surgical procedure.

INTERVENTIONS Patients were randomly assigned to receive standard oxygen therapy (up to 15 L/min to maintain SpO₂ of 94% or higher) (n = 145) or NIV delivered via facial mask (inspiratory pressure support level, 5–15 cm H₂O; positive end-expiratory pressure, 5–10 cm H₂O; fraction of inspired oxygen titrated to maintain SpO₂ ≥94%) (n = 148).

MAIN OUTCOMES AND MEASURES The primary outcome was tracheal reintubation for any cause within 7 days of randomization. Secondary outcomes were gas exchange, invasive ventilation-free days at day 30, health care-associated infections, and 90-day mortality.

RESULTS Among the 293 patients (mean age, 63.4 [SD, 13.8] years; n=224 men) included in the intention-to-treat analysis, reintubation occurred in 49 of 148 (33.1%) in the NIV group and in 66 of 145 (45.5%) in the standard oxygen therapy group within 7 days after randomization (absolute difference, −12.4%; 95% CI, −23.5% to −1.3%; P = .03). Noninvasive ventilation was associated with significantly more invasive ventilation-free days compared with standard oxygen therapy (25.4 vs 23.2 days; absolute difference, −2.2 days; 95% CI, −0.1 to 4.6 days; P = .04), while fewer patients developed health care-associated infections (43/137 [31.4%] vs 63/128 [49.2%]; absolute difference, −17.8%; 95% CI, −30.2% to −5.4%; P = .003). At 90 days, 22 of 148 patients (14.9%) in the NIV group and 31 of 144 (21.5%) in the standard oxygen therapy group had died (absolute difference, −6.5%; 95% CI, −16.0% to 3.0%; P = .15). There were no significant differences in gas exchange.

CONCLUSIONS AND RELEVANCE Among patients with hypoxemic respiratory failure following abdominal surgery, use of NIV compared with standard oxygen therapy reduced the risk of tracheal reintubation within 7 days. These findings support use of NIV in this setting.

TRIAL REGISTRATION clinicaltrials.gov Identifier: NCT01971892

JAMA. doi:10.1001/jama.2016.2706
Published online March 15, 2016.

- Editorial
- Related article
- Supplemental content at jama.com

Author Affiliations: Author affiliations are listed at the end of this article.

Group Information: The Noninvasive Ventilation for Postextubation Respiratory Failure After Abdominal Surgery (NIVAS) study investigators are listed in eAppendix 1 in Supplement 1.

Corresponding Author: Samir Jaber, MD, PhD, Département d'Anesthésie-Réanimation B (DAR B), 80 Avenue Augustin Fliche, 34295 Montpellier, France (s-jaber@chu-montpellier.fr).
Section Editor: Derek C. Angus, MD, MPH, Associate Editor, JAMA (angusdc@upmc.edu).

JAMA[®]

The Journal of the American Medical Association

Jaber S and coauthors

Effect of Noninvasive Ventilation on Tracheal Reintubation Among Patients With Hypoxemic Respiratory Failure Following Abdominal Surgery: A Randomized Clinical Trial

Published online March 15, 2016

Available at jama.com and on The JAMA Network Reader at mobile.jamanetwork.com



The JAMA Network

PRIMARY OUTCOME = Re-intubation at D-7

Jaber et al. JAMA 2016

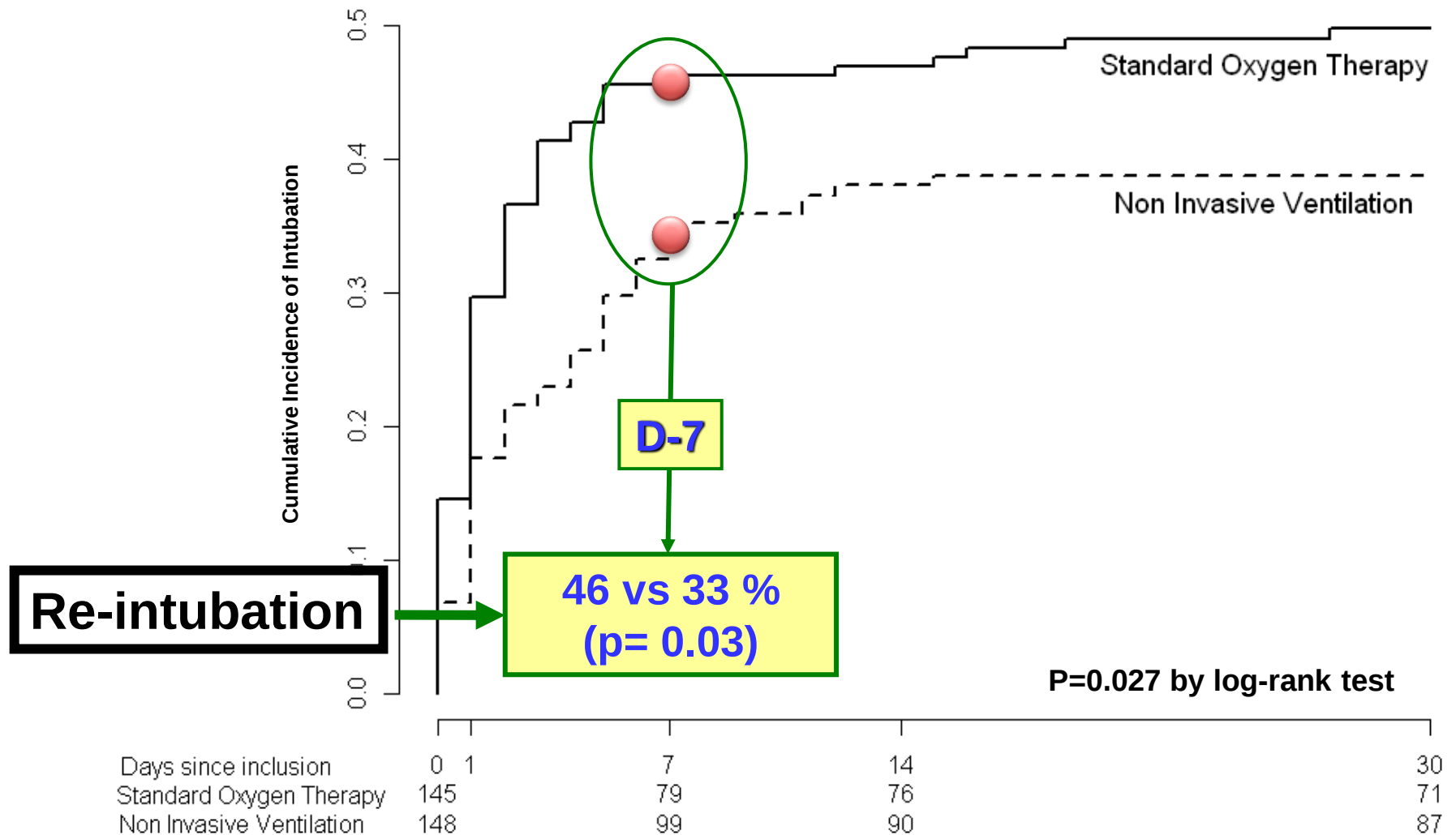
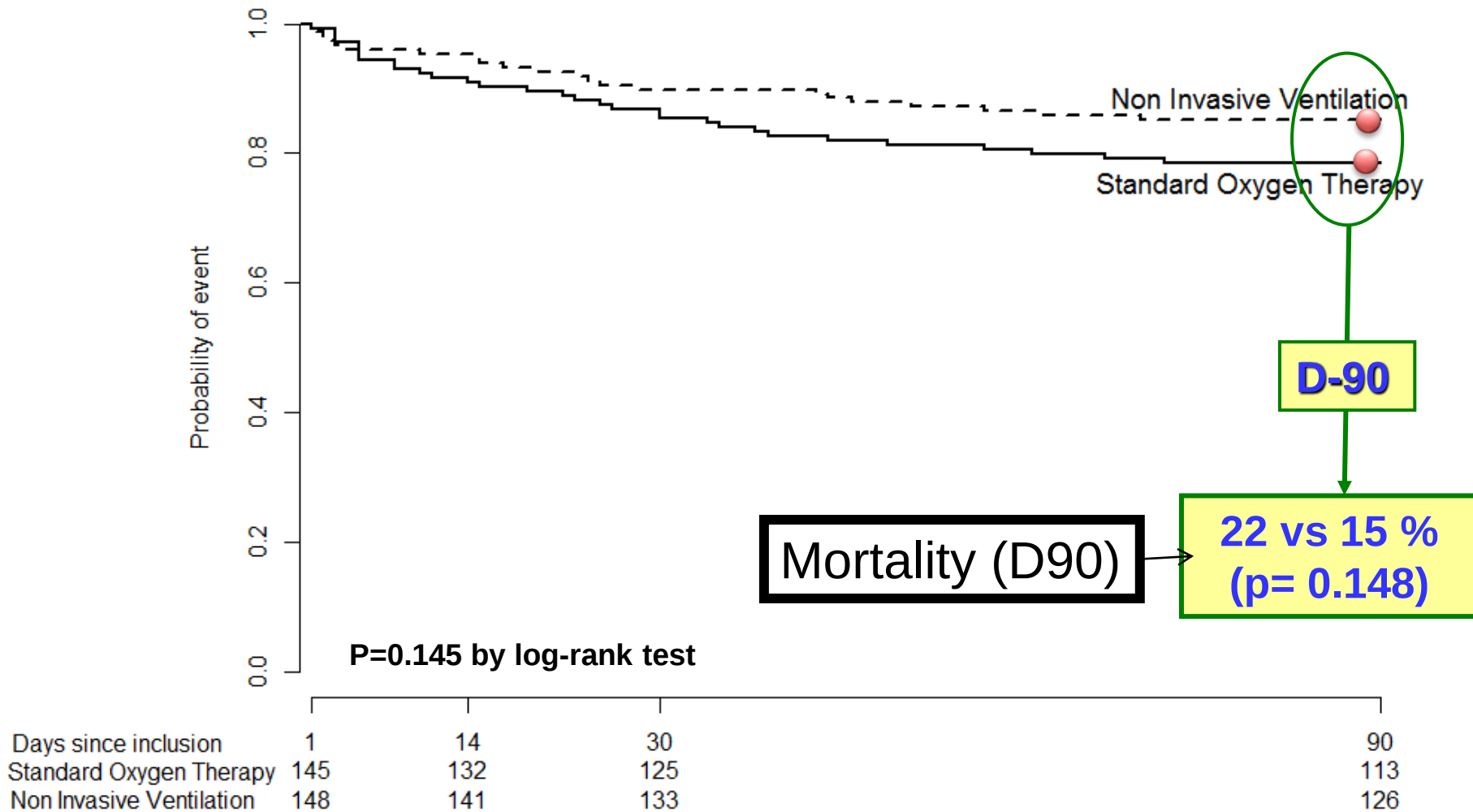


Figure = Kaplan-Meier Plots of the Cumulative Incidence of Intubation from Randomization to Day 30.

MORTALITY

Overall Survival



Objectifs

1. Qui sont les patients à risque?

2. Qu'est ce que la ventilation protectrice et préventive ?

3. Quelles sont les évidences ?

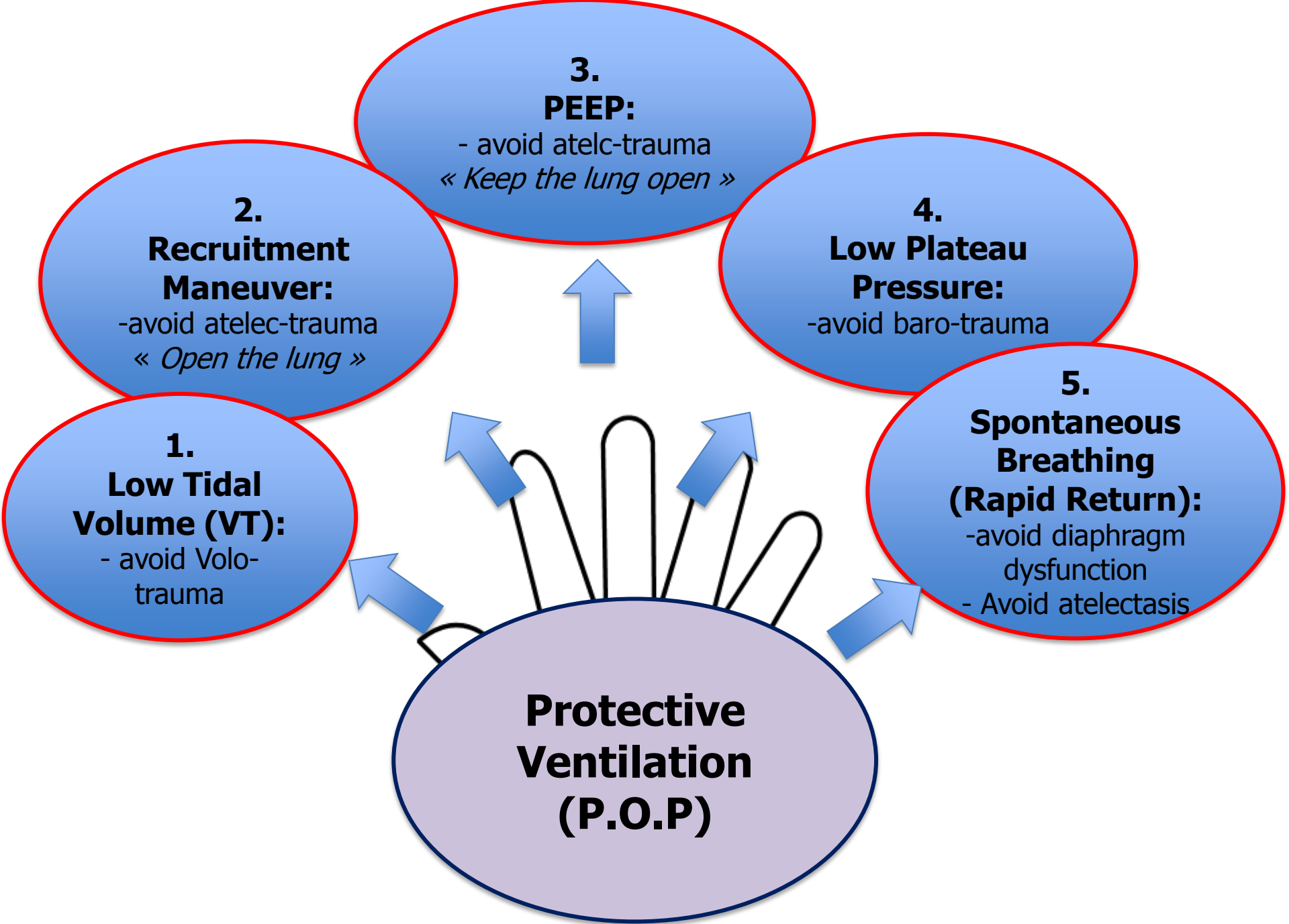
4. Take Home Messages

Take Home Message

5 messages



"Prevention is better than cure"





Montpellier *(France)*



Many thanks