



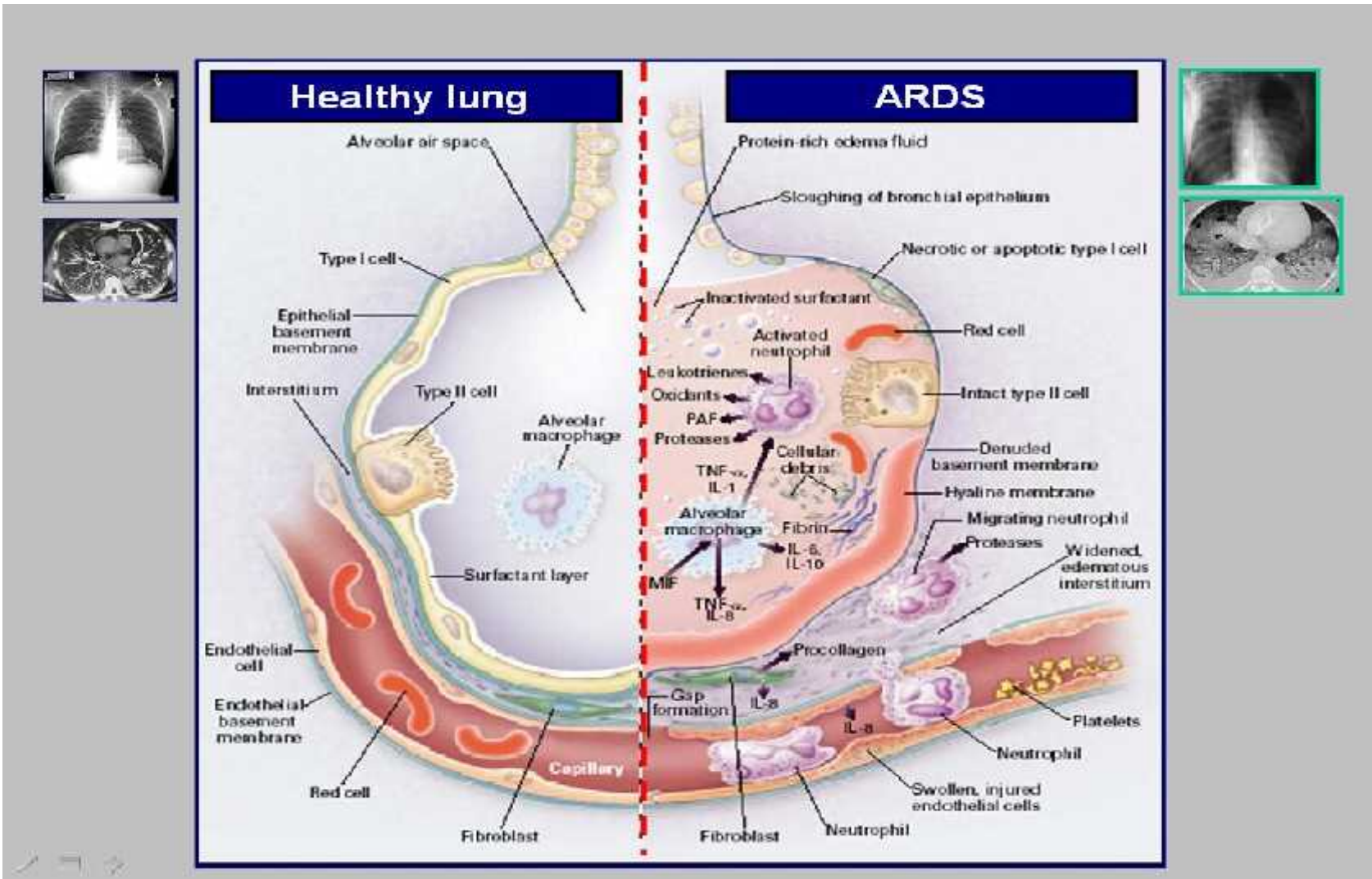
# UPV u ARDS Recruitment, PEEP

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# Fyziologický / patologický



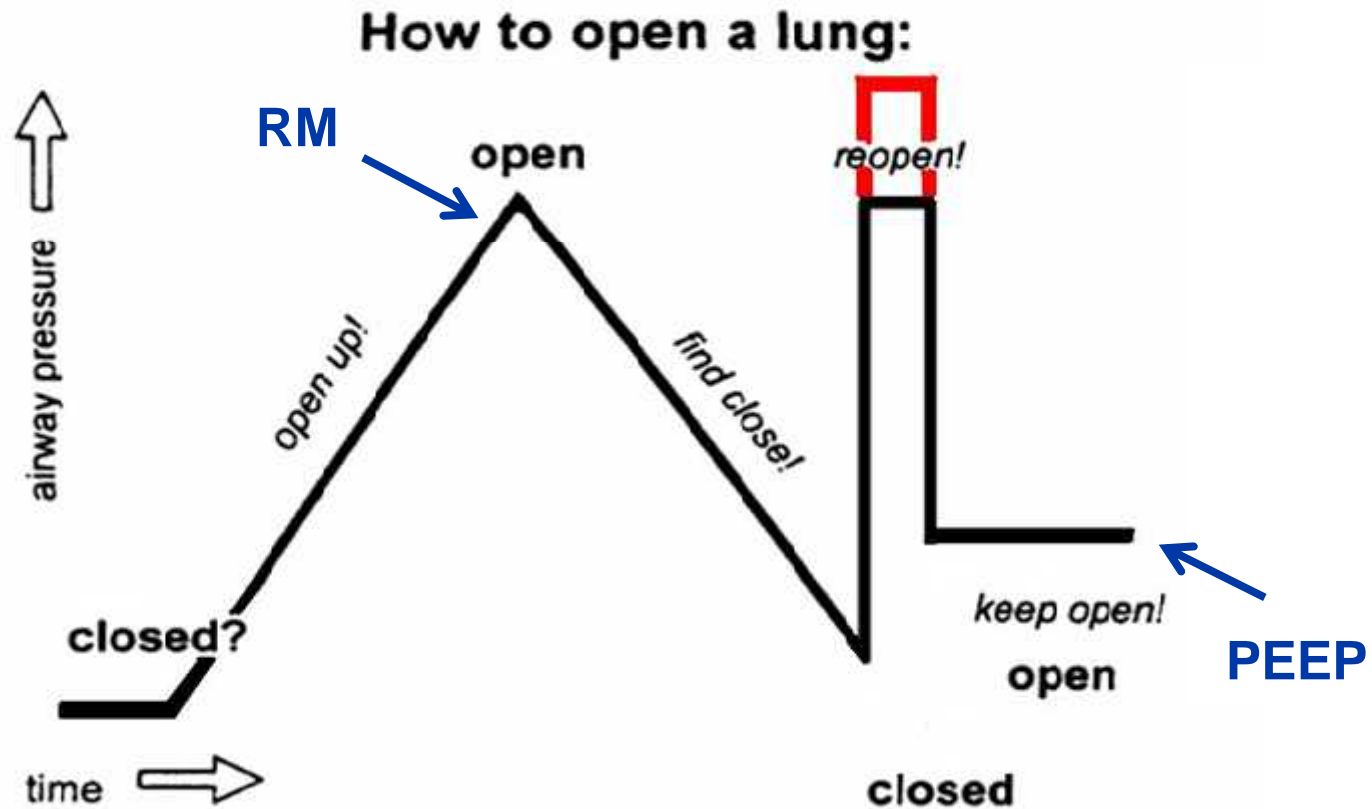
# ARDS nová definice

ARDS - Berlínská definice 2011			
stupň ARDS	mírné	střední	těžké
nástup	akutní začátek do 1 týdne od klinického insultu / prvních příznaků zhoršení		
hypoxémie	$\text{paO}_2/\text{FiO}_2$ 201-300 + PEEP 5	$\text{paO}_2/\text{FiO}_2$ 200 + PEEP 5	$\text{paO}_2/\text{FiO}_2$ 100 + PEEP 10
etiologie edému	respirační selhání není plně vysvětlitelné srdečním selháním nebo tekutinovým přetížením		
radiologický nález	bilaterální opacity	bilaterální opacity	bilaterální opacity minimálně v 3 kvadrantech
přítomnost přidružených patologických nálezů	x	x	$V_{\text{Ecorr}} > 10\text{L/min}$ nebo $C_{\text{RS}} < 40 \text{ ml/cmH}_2\text{O}$

$V_{\text{Ecorr}}$  korigovaná minutová ventilace  $V_{\text{E}} \times \text{pCO}_2/40$ ,  $C_{\text{RS}}$  compliance plic

# Open up the lung and keep the lung open

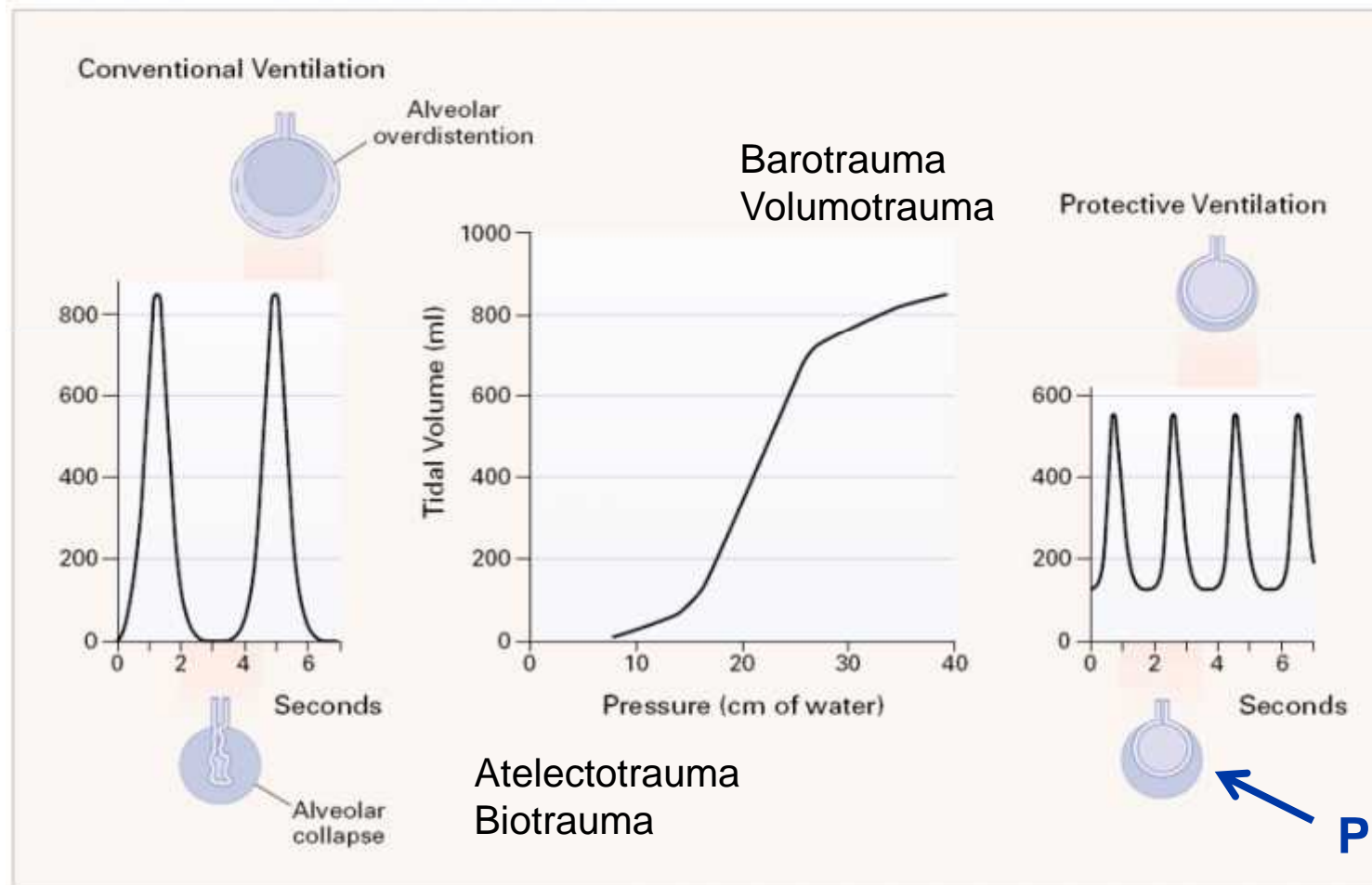
OPEN LUNG CONCEPT OF MECHANICAL VENTILATION



# Protektivní ventilace, PEEP

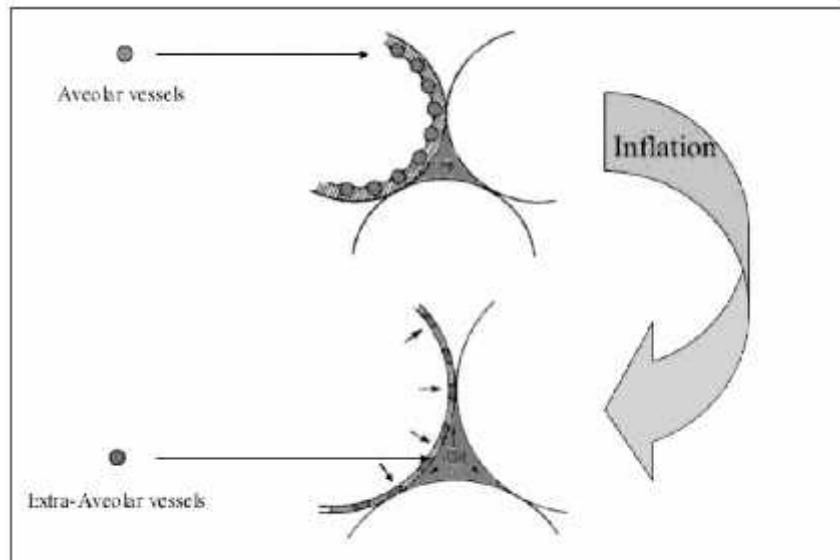
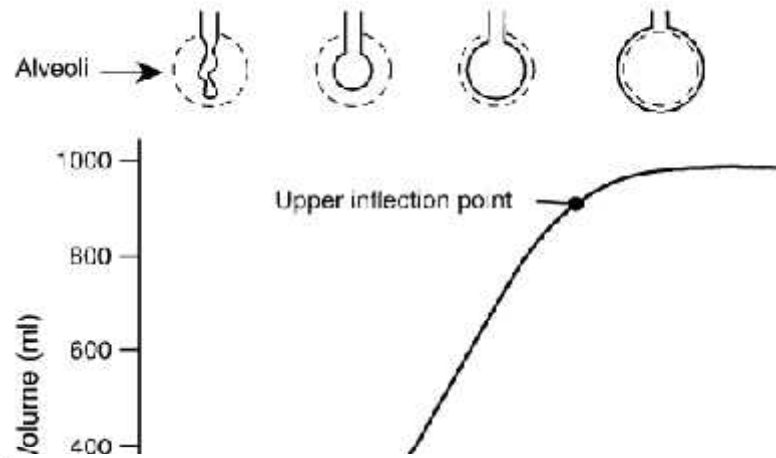
10 ml/kg, nízká DF

6 ml/kg, PEEP, vyšší DF



**PEEP**

# PEEP / RM



# Použití PEEPu



# Použití PEEP<sub>u</sub> / PCV-RM





# Efekt PEEPu na mortalitu ARDS?

## Higher vs Lower Positive End-Expiratory Pressure in Patients With Acute Lung Injury and Acute Respiratory Distress Syndrome Systematic Review and Meta-analysis

Characteristic	ALVEOLI, <sup>3</sup> 2004	LOVS, <sup>9</sup> 2008	EXPRESS, <sup>10</sup> 2008
Inclusion criteria	Acute lung injury with $P_{aO_2}:F_{iO_2} \leq 300^a$	Acute lung injury with $P_{aO_2}:F_{iO_2} \leq 250^a$	Acute lung injury with $P_{aO_2}:F_{iO_2} \leq 300^a$
Recruitment period	1999-2002	2000-2006	2002-2005
Recruiting hospitals (country)	23 (United States)	30 (Canada, Australia, Saudi Arabia)	37 (France)
Patients randomized to higher vs lower PEEP	276 vs 273	476 vs 500 <sup>b</sup>	385 vs 383 <sup>c</sup>
Validity			
Concealed allocation	Yes	Yes	Yes
Follow-up for primary outcome, %	100	100	100
Blinded data analysis	Yes	Yes	Yes
Stopped early	Stopped for perceived futility	No	Stopped for perceived futility
Experimental intervention	Higher PEEP according to $F_{iO_2}$ chart, recruitment maneuvers for first 80 patients	Higher PEEP according to $F_{iO_2}$ chart, required plateau pressures $\leq 40$ cm $H_2O$ , recruitment maneuvers	PEEP as high as possible without increasing the maximum inspiratory plateau pressure $>28-30$ cm $H_2O$
Control intervention	Conventional PEEP according to $F_{iO_2}$ chart, required plateau pressures $\leq 30$ cm $H_2O$ , no recruitment maneuvers	Conventional PEEP according to $F_{iO_2}$ chart, required plateau pressures $\leq 30$ cm $H_2O$ , no recruitment maneuvers	Conventional PEEP (5-9 cm $H_2O$ ) to meet oxygenation goals

# Efekt PEEPu na mortalitu ARDS? EXPRESS trial



Positive End-Expiratory Pressure Setting  
in Adults With Acute Lung Injury  
and Acute Respiratory Distress Syndrome  
A Randomized Controlled Trial

PEEP 5-9cmH<sub>2</sub>O **X** max. s cílem <Pplat. 28-30, SpO<sub>2</sub> 88-95%

Vt 6ml/kg, Ppl.>32 Vt 4ml/kg, df < 35/min pH 7.3-7.4

**RM povolen ale NEDOPORU EN**

**Table 3.** Respiratory Variables During the First 7 Days of Treatment<sup>a</sup>

Variable	Day 1			Day 3			Day 7		
	Minimal Distension	Increased Recruitment	P Value	Minimal Distension	Increased Recruitment	P Value	Minimal Distension	Increased Recruitment	P Value
Tidal volume, mL/kg of predicted body weight	6.1 (0.4)	6.1 (0.3)	.57	6.2 (0.6)	6.2 (0.5)	.83	6.4 (0.9)	6.8 (1.3)	.001
No. of patients	372	378		322	332		210	192	
Plateau pressure, cm H <sub>2</sub> O	21.1 (4.7)	27.5 (2.4)	<.001	20.7 (5.0)	26.5 (4.2)	<.001	21.1 (5.6)	24.3 (5.8)	<.001
No. of patients	365	378		314	329		173	163	
PEEP, cm H <sub>2</sub> O	7.1 (1.8)	14.6 (3.2)	<.001	6.7 (1.8)	13.4 (4.7)	<.001	6.2 (2.1)	8.9 (5.1)	<.001
No. of patients	372	380		333	351		264	252	
Total PEEP, cm H <sub>2</sub> O <sup>b</sup>	8.4 (1.9)	15.8 (2.9)	<.001	8.1 (2.0)	15.1 (4.3)	<.001	8.0 (2.5)	12.0 (5.4)	<.001
No. of patients	336	348		274	284		154	138	
PaO <sub>2</sub> /FiO <sub>2</sub>	150 (69)	218 (97)	<.001	175 (81)	245 (98)	<.001	184 (79)	206 (85)	.003
No. of patients	371	378		331	350		262	247	

# Efekt PEEPu na mortalitu ARDS? EXPRESS trial



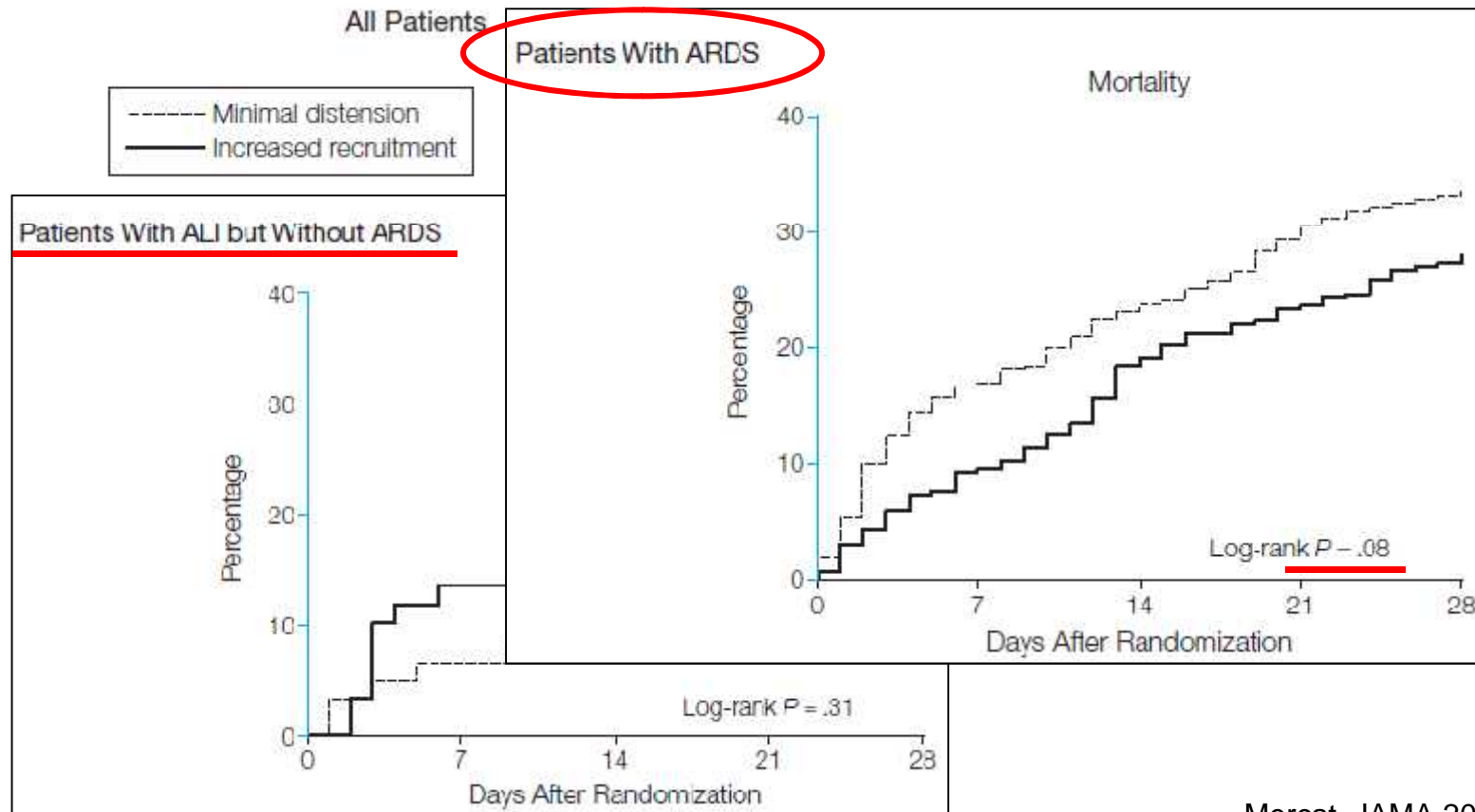
**Positive End-Expiratory Pressure Setting  
in Adults With Acute Lung Injury  
and Acute Respiratory Distress Syndrome**  
A Randomized Controlled Trial

**Table 5.** Cointerventions and Adjunctive Therapies

Intervention	No. (%) <sup>a</sup>		P Value
	Minimal Distension (n=382)	Increased Recruitment (n=385)	
During the first 72 h			
Fluid loading	255 (66.8)	290 (75.3)	.01
Volume of fluids, median (IQR), L <sup>b</sup>	0.5 (0-1.5)	1.0 (0.1-2.2)	<.001
During the first 7 d			
Epinephrine or norepinephrine	286 (74.9)	289 (75.1)	.95
Corticosteroids	198 (51.8)	199 (51.7)	.97
Neuromuscular blockade	209 (54.7)	204 (53)	.63
Recruitment maneuvers	49 (12.8)	27 (7.0)	.007
Adjunctive therapies during the first 7 d			
Prone position	72 (18.8)	34 (8.8)	<.001
Inhaled nitric oxide	98 (25.7)	57 (14.8)	<.001
Almitrine bismesylate	25 (6.5)	14 (3.6)	.07
Any therapy	132 (34.6)	72 (18.7)	<.001
Mortality in patients who received rescue therapy	62 (47.0)	37 (51.4)	.55

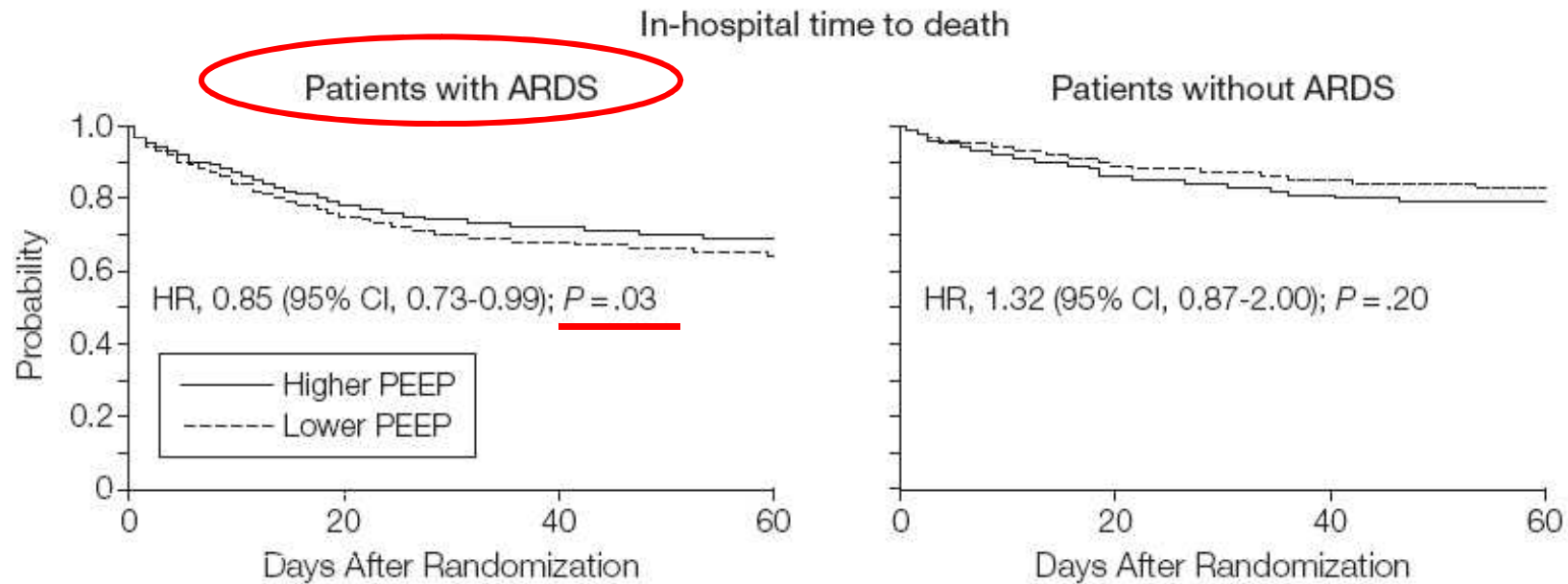
# Efekt PEEPu na mortalitu ARDS? EXPRESS trial

**Positive End-Expiratory Pressure Setting  
in Adults With Acute Lung Injury  
and Acute Respiratory Distress Syndrome**  
A Randomized Controlled Trial



# Efekt PEEP u na mortalitu ARDS?

**Higher vs Lower Positive End-Expiratory Pressure in Patients With Acute Lung Injury and Acute Respiratory Distress Syndrome**  
Systematic Review and Meta-analysis



## Recruitment maneuver (RM)

- r zné typy plicního poškození
  - u sekundárních / extra pulmonálních forem ARDS
- r zná tíže plicního postižení i r zná fáze
- transpulmonální tlak dosažený během RM
- typ recruitment manévru SI / PCV - RM
- nastavení PEEPu po provedeném RM
- difference v poloze pacienta (pronace)
- difference v intravazální náplni i vasoaktivních látkách, které ovlivní CO i plicní vask. rezistenci

## Primární / sekundární ARDS

**Primární** = direct (pulmonary) lung injury

- damaged alveolar epithelium, **alveolar filling by edema, fibrin, and neutrophilic aggregates**

**Sekundární** = indirect (extrapulmonary) lung injury

- inflammatory mediators released from extrapulmonary foci into the systemic circulation, microvessel congestion and interstitial edema with relative sparing of **intra-alveolar spaces**

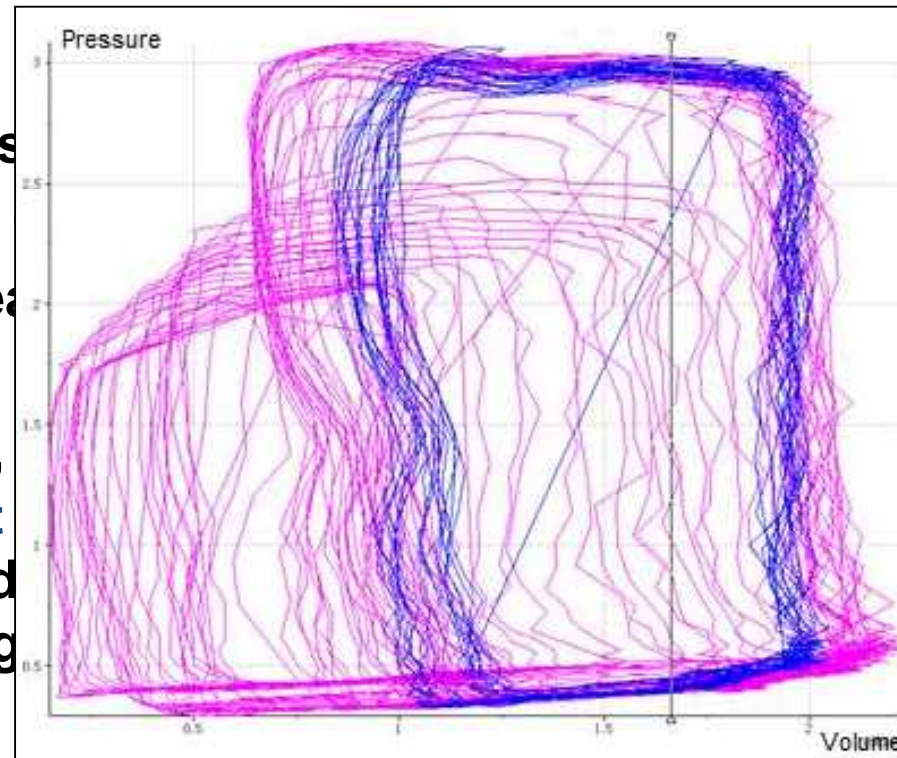
**„RM more effective to open atelectatic lung regions in indirect compared to direct lung injury“**

# Typy RM - SI

## Sustained Inflation technique

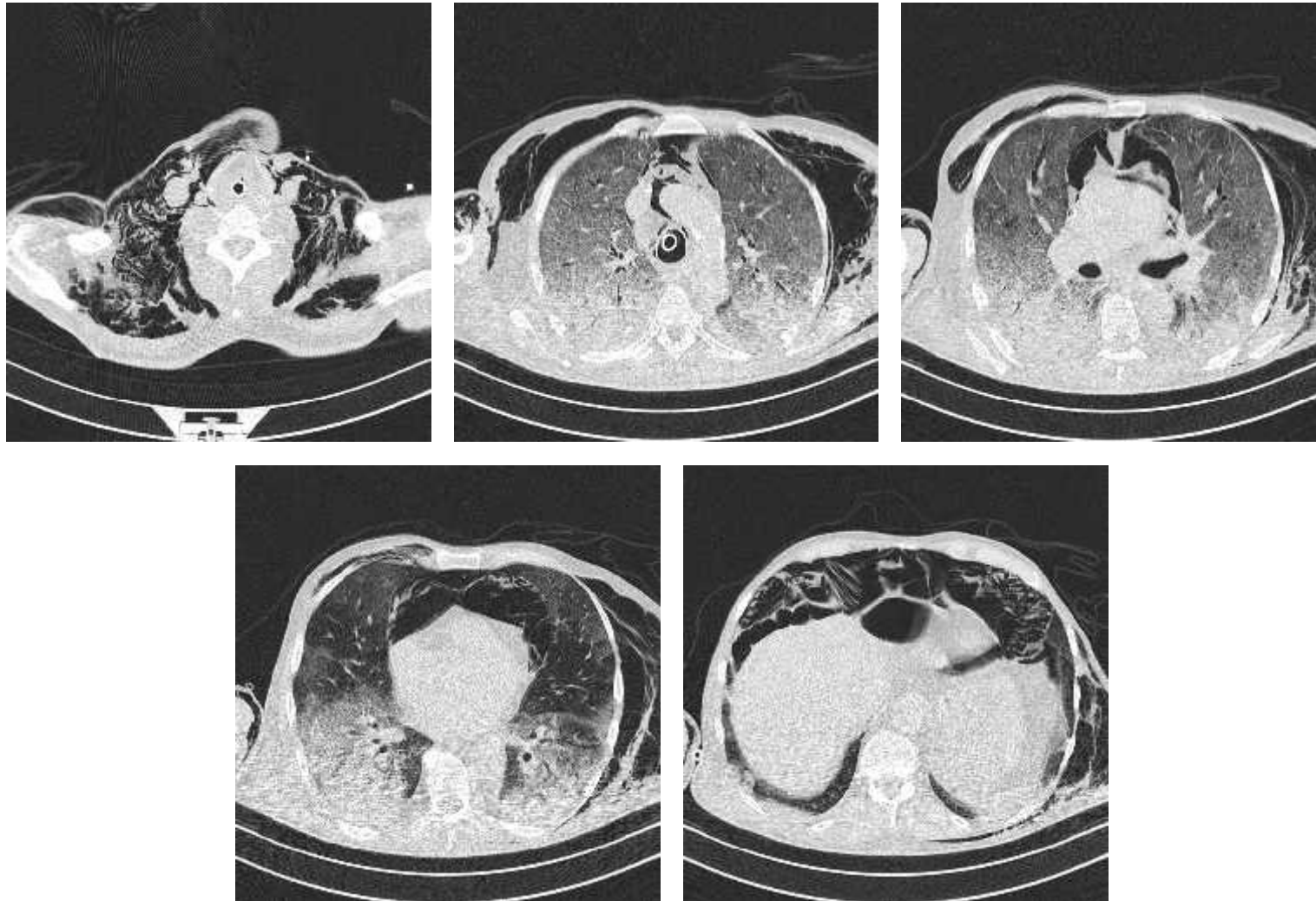
( 40 cmH<sub>2</sub>O to 40/60 sec, 45/45)

- reducing lung atelectasis and improving respiratory mechanics
- preventing endotracheal tube occlusion and derecruitment
- RM may be ineffective, circulatory impairment, baro/volutrauma, a reduction in oxygenation or even worsened oxygenation





# ARDS – barotrauma



## Typy RM - varianty

- 1) incrementally **increased PEEP** limiting the **maximum inspiratory pressure**

Rzezinski, *Respir Physiol Neurobiol* 2009

- 2) **PCV** applied **with escalating PEEP** and **constant driving pressure**

Villagr , *Am J Respir Crit Care Med* 2002

- 3) **SRM - staircase RM** progressive increase in PEEP (up to 40 cm H<sub>2</sub>O / higher pr. 55 cm H<sub>2</sub>O)

Hudgson, *CC* 2011

- 4) **intermittent sighs** to reach a **specific plateau pressure in volume**  
**or pressure control mode**

Steimback, *IMC*, 2009, Badet, *Respiratory Care*, 2009

- 5) (RAMP) **long slow increase in inspiratory pressure up to 40**  
**cmH<sub>2</sub>O = PV tool**

Riva, *Crit Care Med* 2009

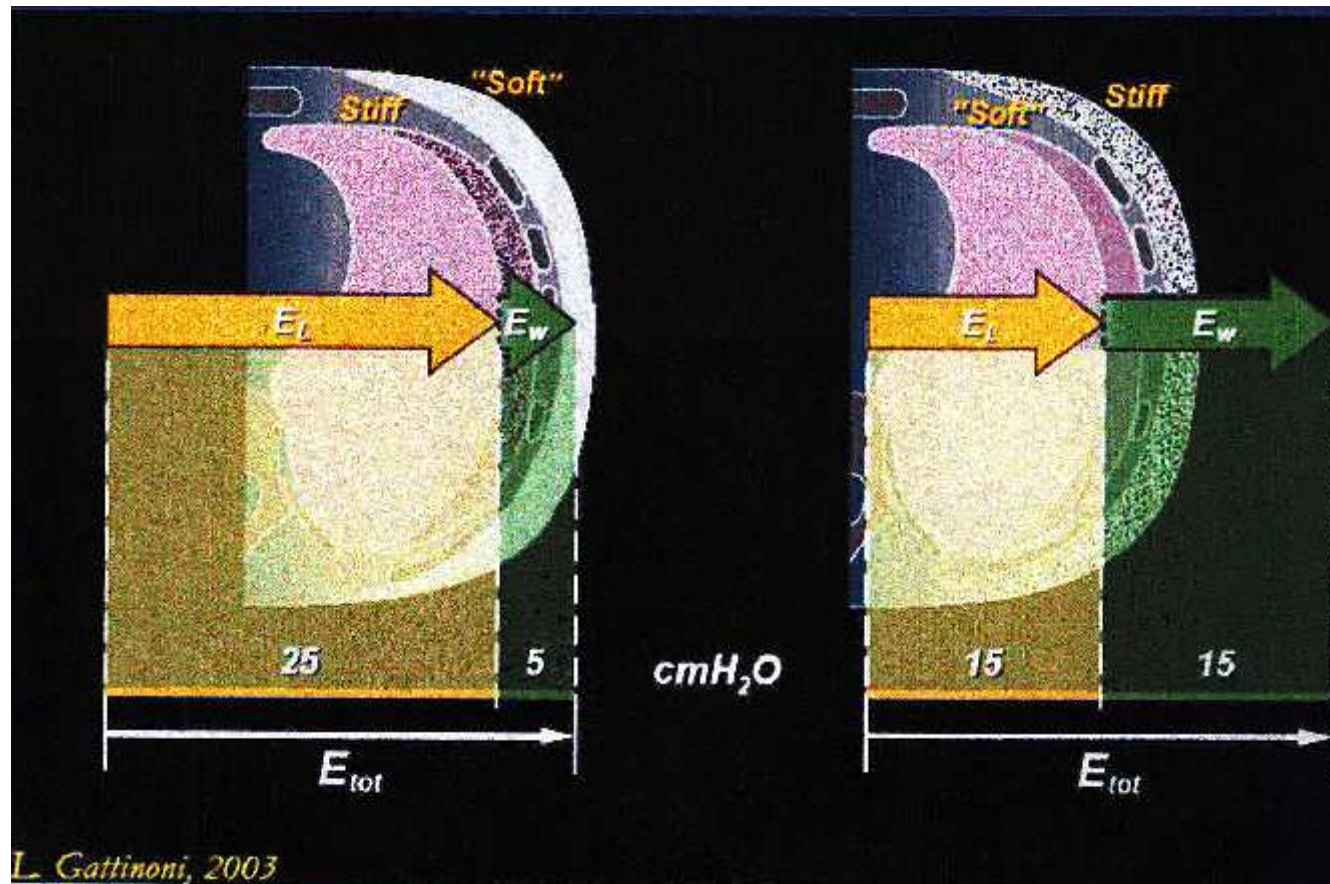
- 6) **PCV RM - eSigh: 10 cmH<sub>2</sub>O above LIP, 15 min**

Constantin, *CC* 2008

- 7) **PCV RM - PEEP 25 / P<sub>high</sub> 40, 30 min**

Borges, *AJRCCM*, 2006

# Inspira ní P / transpulmonální P



# Staircase RM

## SRM:

**PEEP 20 (2min)+15 PC  
30 (2min) before  
40 cmH2O (2min)**  
to check HD tolerance

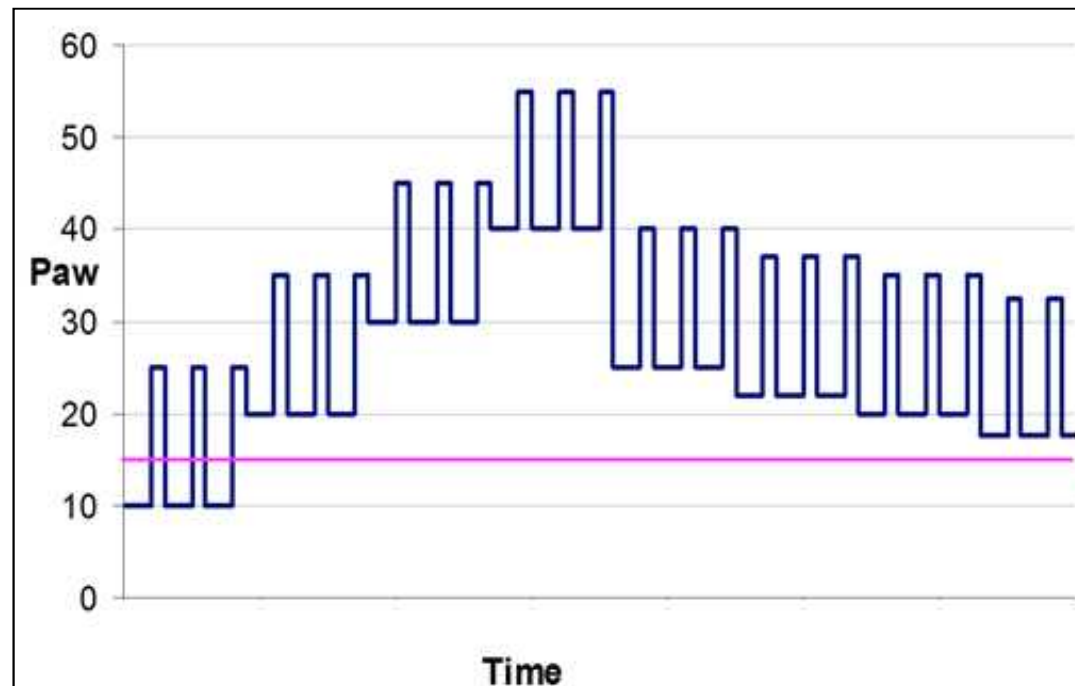
PEEP reductions to 25  
down in 2.5  
every 3 min  
derecruitment when  
SaO2 decreases by 1-2%

Incr. PEEP to 40 for 1min  
and return 2,5 above

20 pts:  
10 **PHARLAP SRM daily**  
10 ARDS protocol  
(Pplat<30, 6ml/kg)

A randomised controlled trial of an open lung strategy with staircase recruitment, titrated PEEP and targeted low airway pressures in patients with acute respiratory distress syndrome

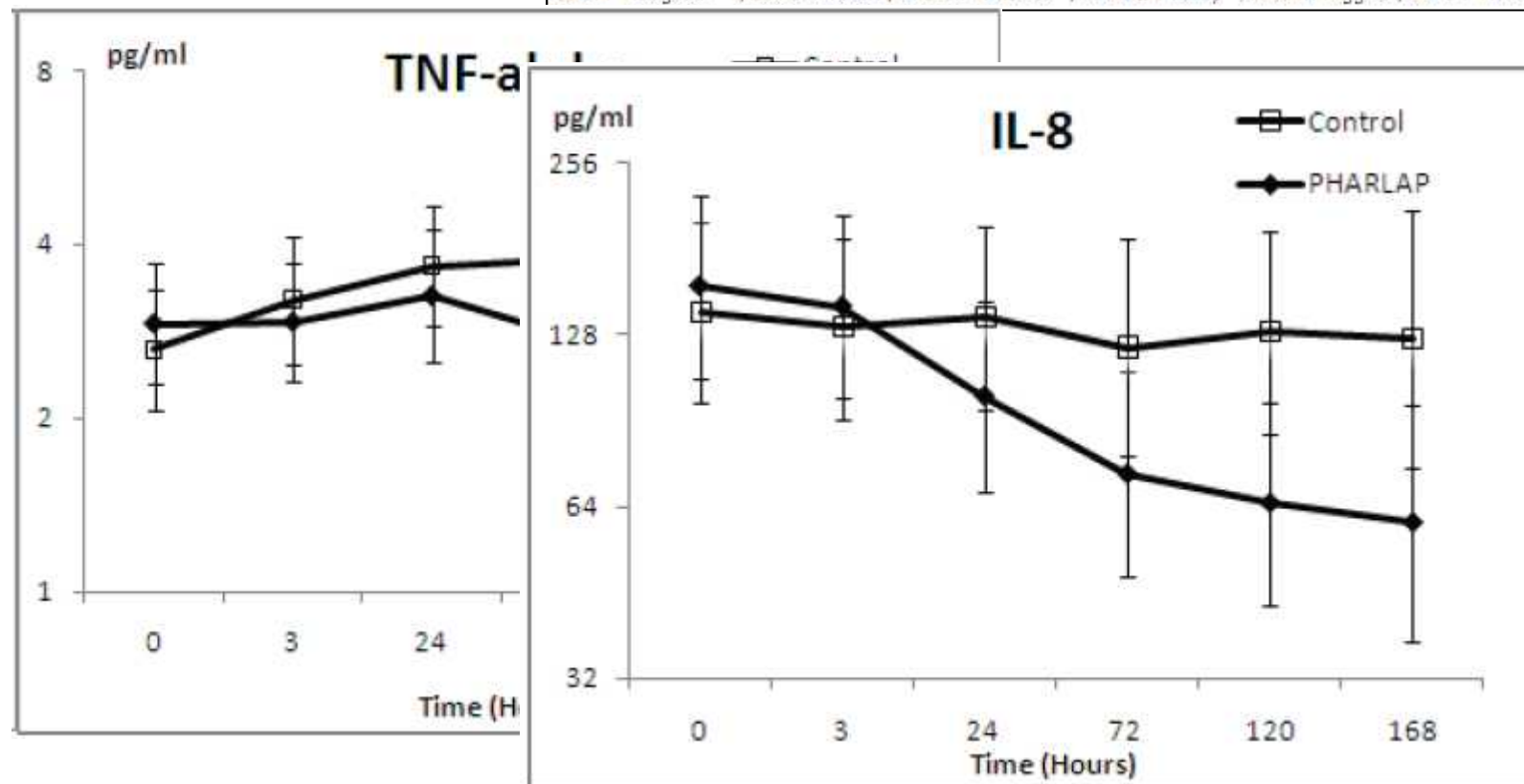
Carol J. Hodgson<sup>1,2,3\*</sup>, David V. Tuxen<sup>1</sup>, Andrew R. Davies<sup>1,2</sup>, Michael J. Bailey<sup>1,2</sup>, Alisa M. Higgins<sup>2</sup>, Anne F. Holland



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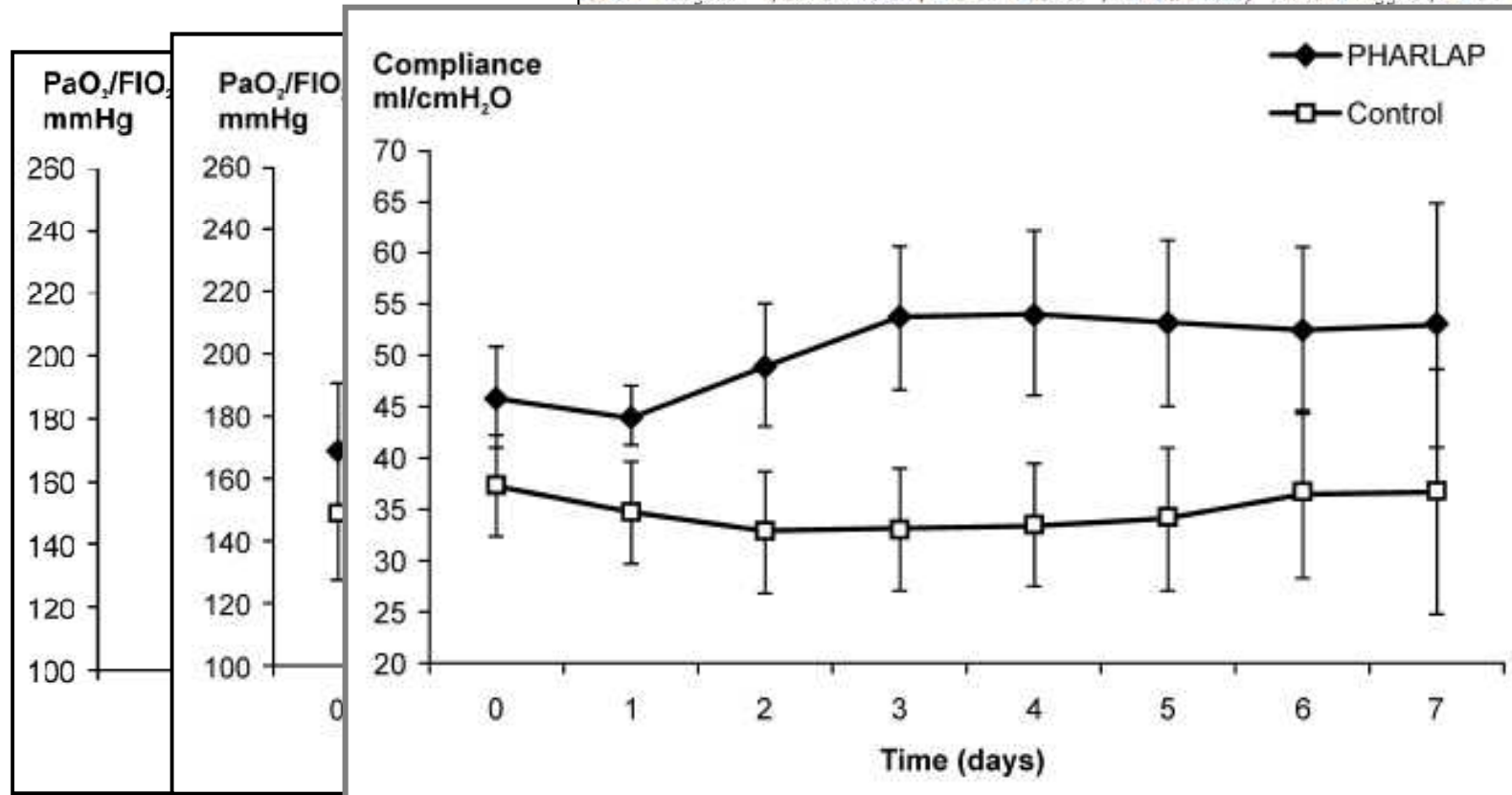
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**Table 4 Outcomes**

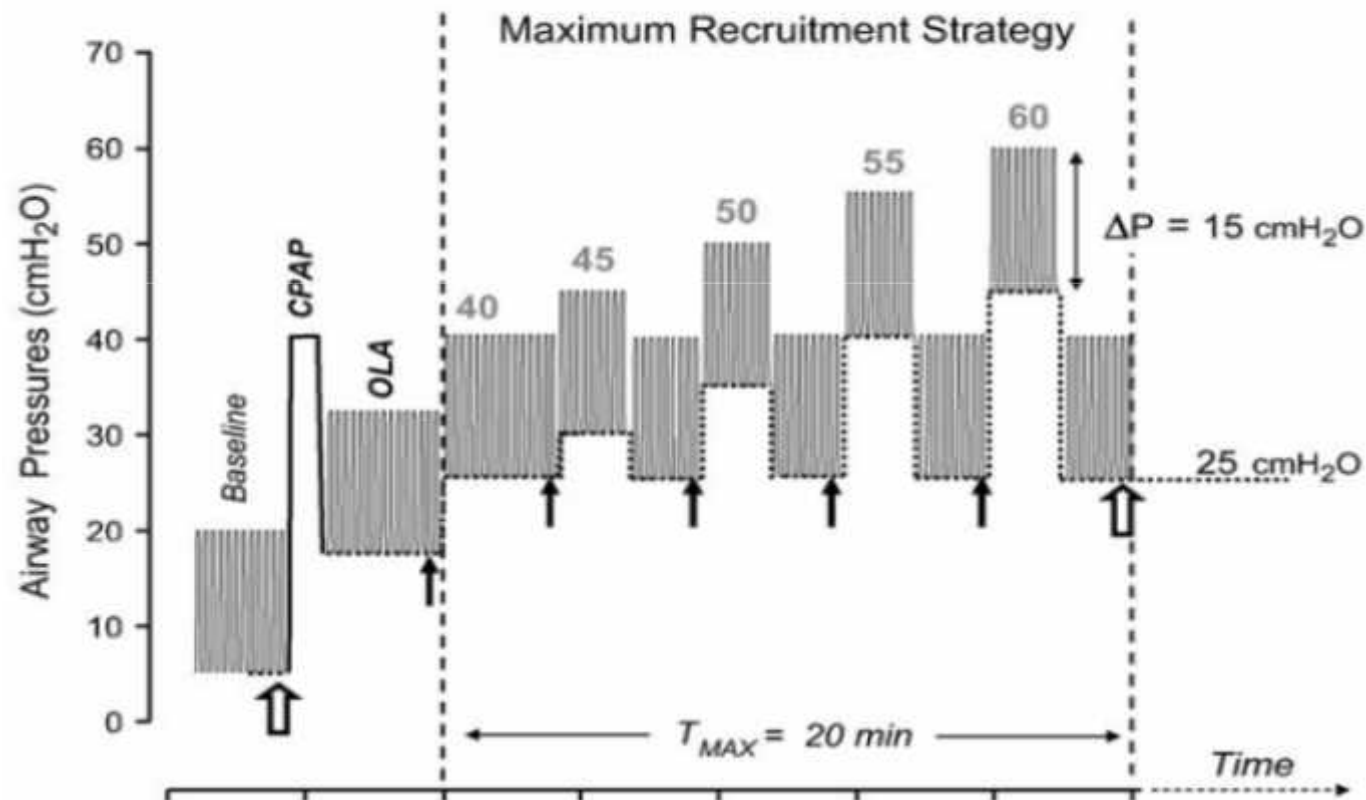
	PHARLAP	Control	<i>P</i>
Hospital mortality, number	3	2	0.61
LOV, hours	180 (87 to 298)	341 (131 to 351)	0.13
ICU LOS, days	9.9 (5.6 to 14.8)	16.0 (8.1 to 19.3)	0.19
Hospital LOS, days	17.9 (13.7 to 34.5)	24.7 (20.5 to 39.8)	0.16
Barotrauma, number	0	0	
Rescue therapies, number of patients	0	2	0.46
SOFA score (Day 7)	8.6 ± 0.3	8.4 ± 0.6	0.27

LOV, length of ventilation; ICU LOS, intensive care length of stay; LOS, length of stay;

# PCV RM

## Reversibility of Lung Collapse and Hypoxemia in Early Acute Respiratory Distress Syndrome

João B. Borges, Valdelis N. Okamoto, Gustavo F. J. Matos, Maria P. R. Caraméz, Paula R. Arantes,



SI RM 40/40, set PEEP LIP 19, Press + 5, paO<sub>2</sub> + pCO<sub>2</sub> > 400 mmHg, df 10/min

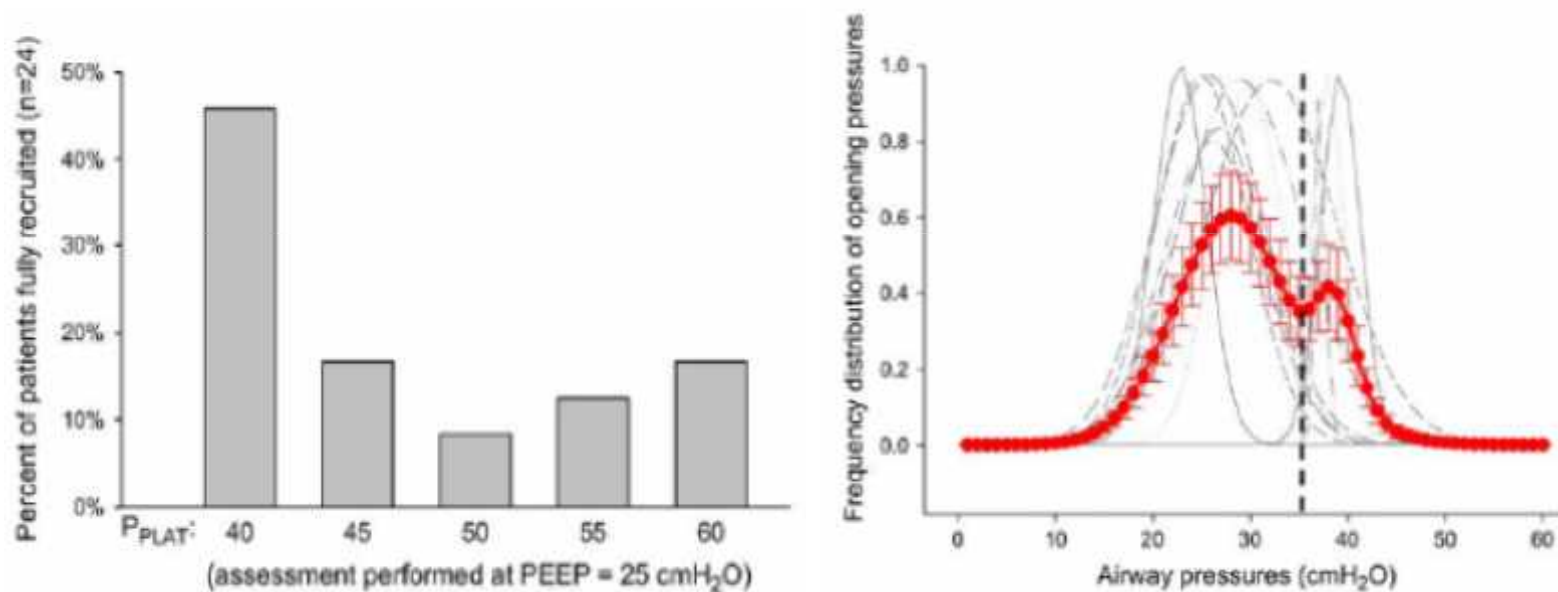
Borges, AJRCCM, 2006



# PCV RM

## Reversibility of Lung Collapse and Hypoxemia in Early Acute Respiratory Distress Syndrome

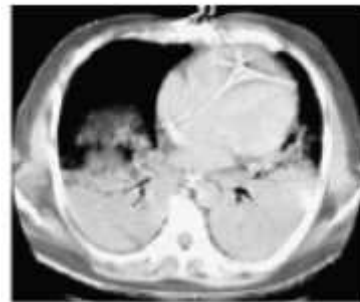
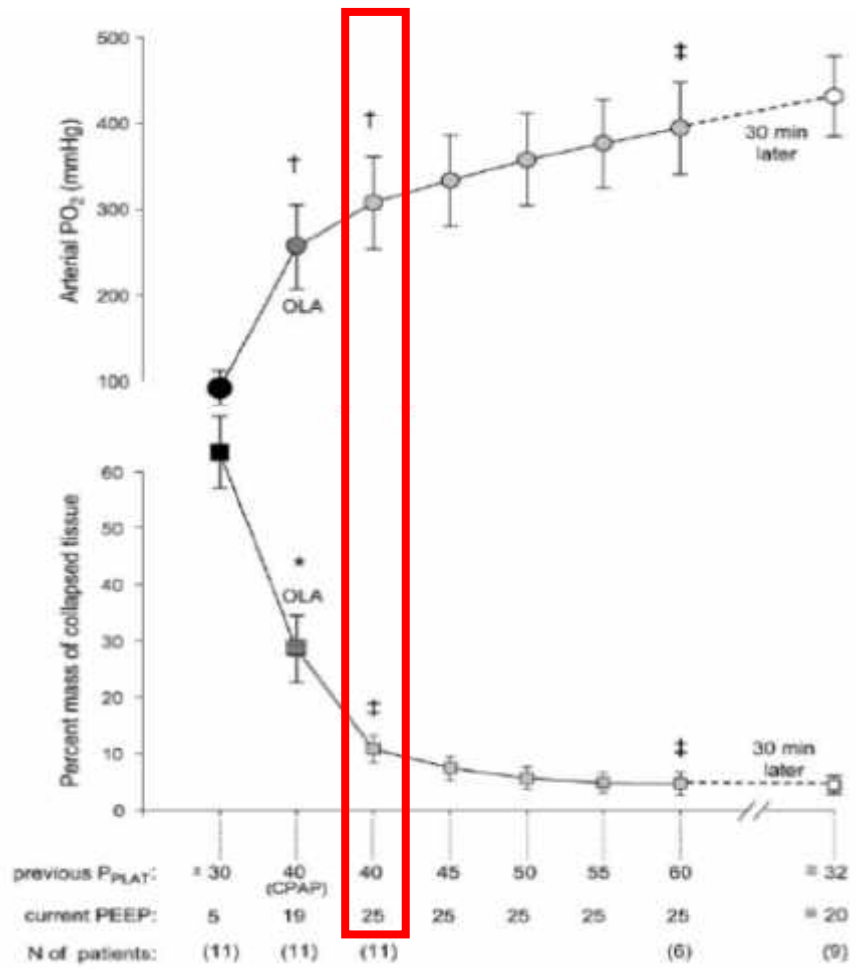
João B. Borges, Valdelis N. Okamoto, Gustavo F. J. Matos, Maria P. R. Caraméz, Paula R. Arantes,



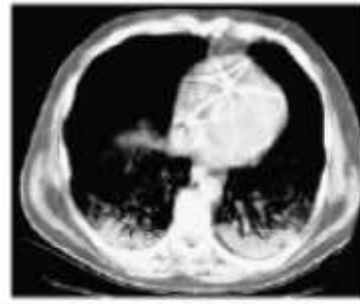
SI RM 40/40, set PEEP LIP 19, Press + 5, paO<sub>2</sub> + pCO<sub>2</sub> > 400 mmHg, df 10/min

RM nedosáhli 2/26, žádné barotrauma b hem

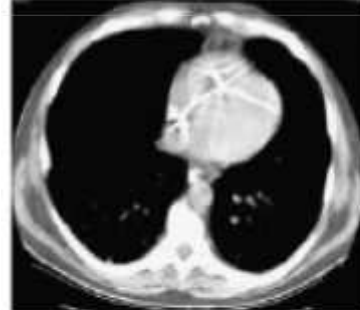
# PCV RM



PEEP = 5 cmH<sub>2</sub>O  
 collapsed Area = 54.3%  
 collapsed Mass = 69.2%



PEEP = 19 cmH<sub>2</sub>O (OLA)  
 collapsed Area = 21.9%  
 collapsed Mass = 36.8%

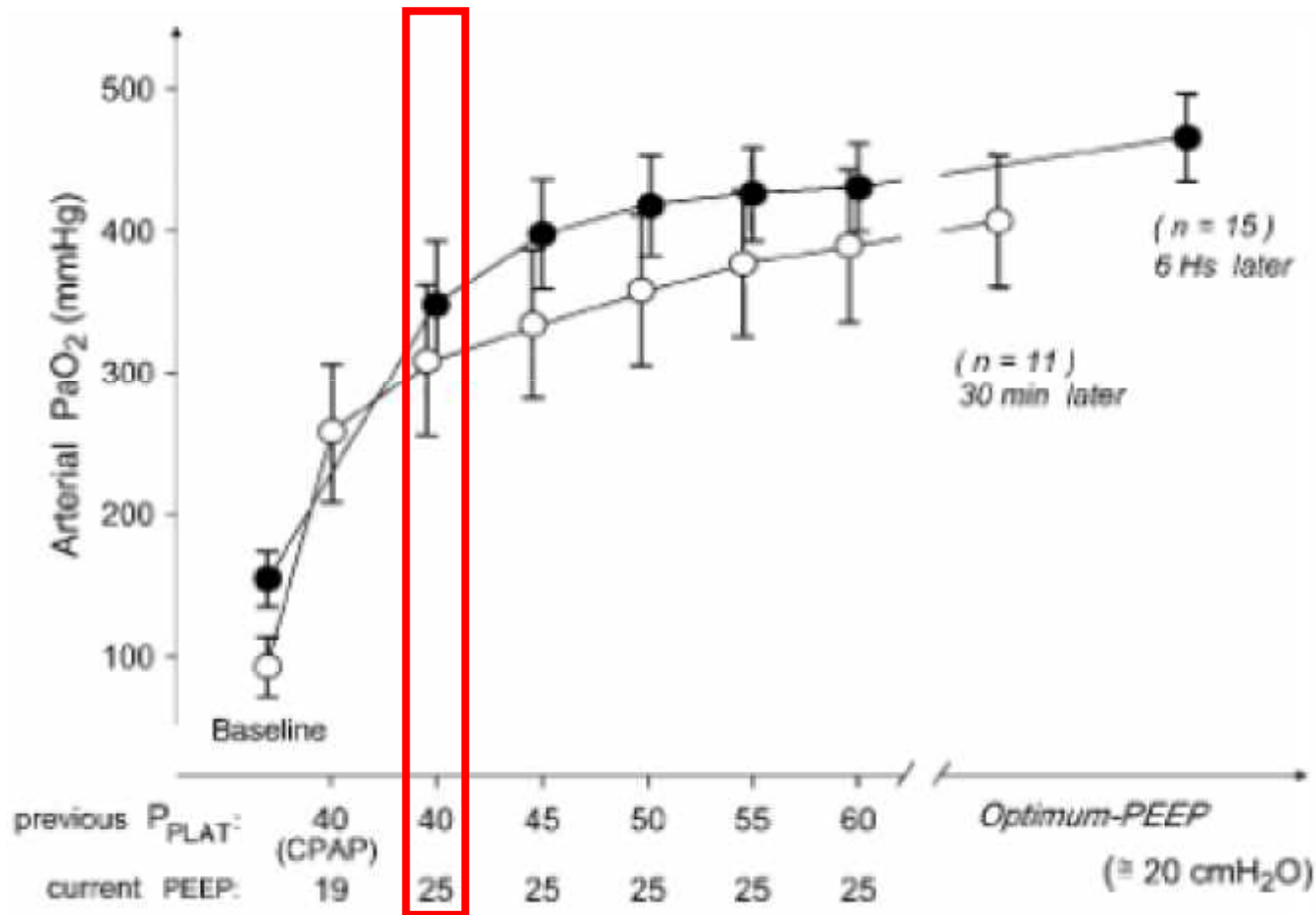


PEEP = 25 cmH<sub>2</sub>O (after P<sub>PLAT</sub> = 55)  
 collapsed Area = 0.4%  
 collapsed Mass = 0.9%



PEEP = 23 cmH<sub>2</sub>O (30 minutes later)  
 collapsed Area = 0.5%  
 collapsed Mass = 1.8%

# PCV RM 25/40 – 30min/6h?



# SI RM versus PCV RM

Different patterns of lung recruitment maneuvers in primary acute respiratory distress syndrome: effects on oxygenation and central hemodynamics

MIANNUZZI, A. DESIO, G. DE ROBERTIS, O. MAZZA, C. SBRVILLO, R. ILLIANO

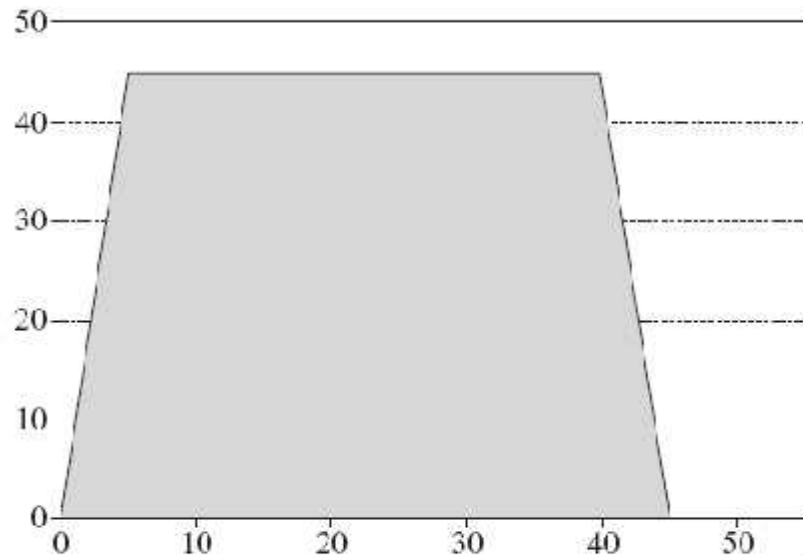


Figure 1. Pressure time product of sustained inflation (SI) recruitment maneuver. X axis: time (seconds); Y axis: airway

**SI RM : 45cm H<sub>2</sub>O na 40 sec.**  
**Pressure time product SI i PVC=1800**

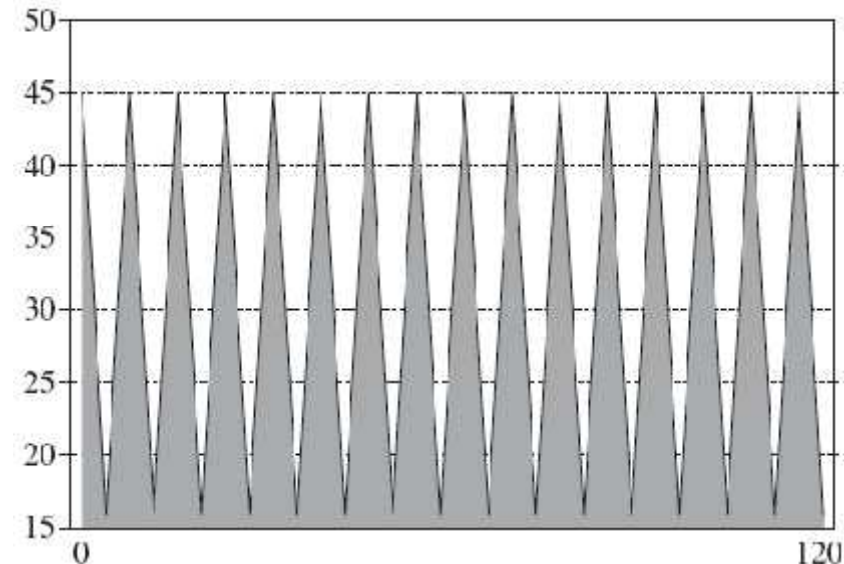


Figure 2.—Pressure-time product of pressure controlled ventilation (PCV) recruitment maneuver. X axis: time (seconds); Y axis: airway pressure (cmH<sub>2</sub>O).

**PCV RM: PEEP 16, PIP 45,**  
**I:E 1:2, df 8/min**

# SI RM versus PCV RM hemodynamika

TABLE II.—H

Author		Post-PCV RM
<u>PaO<sub>2</sub> (mmHg)</u>		157.6± 61.5*
<u>P/F (mmHg)</u>		238.8±86.5*
PaCO <sub>2</sub> (mmHg)		55.0±3.4
<u>C<sub>RS</sub></u>		62.0±12.5*
Q <sub>s</sub> /Q <sub>t</sub> (%)		3.4±0.8
C.I. (L/min)		19.7±2.0*
<u>PAOP (mmHg)</u>		30.2±3.5*
<u>MFAP (mmHg)</u>		247.0±15.2
PVRI (dyne · sec/cm <sup>5</sup> )		5.7±0.9*
RVSWI (g · m/100g)		19.7±2
CVP (mmHg)		106.5±13.1; 55.7±10.3
Sys/Dia (mmHg)		84.5±13.1
HR (bpm)		

TABLE III.—Echocardiographic evaluations before and after recruitment maneuvers (RM).

	Pre-SIRM	Post-SIRM	Pre-PCV	Post-PCV RM
LVEDA	19.0 (14.2 to 25.6)	7.4 (2.6-10.5)*	18.7 (13.2 to 24.2)	11.7 (7.5 to 19.2)
LVESA	9.2 (5.2 to 16.1)	5.4 (2.2 to 7.4)*	9.4 (5.5 to 13.3)	7.1 (3.2 to 15.4)
EI	1.15± 0.10	1.44±0.26*	1.18±0.20	1.21± 0.10

\*P<0.05. LVEDA: left ventricular end diastolic area; LVESA: left ventricular end systolic area; EI: eccentricity index.

# SIGH?



Comparison of Optimal Positive End-Expiratory Pressure  
and Recruitment Maneuvers During Lung-Protective  
Mechanical Ventilation in Patients With Acute Lung Injury/  
Acute Respiratory Distress Syndrome

Michel Badet MD, Frédérique Bayle MD, Jean-Christophe Richard MD PhD,  
and Claude Guérin MD PhD

- 12 pacient , asné ALI/ARDS
- SI RM 40 cm H<sub>2</sub>O for 30 s
- PEEP 24 cm H<sub>2</sub>O, regrese PEEP skokov 4 cm H<sub>2</sub>O každých 10 min, FiO<sub>2</sub> 0,8, paO<sub>2</sub>
- “optimal” PEEP - PEEP p ed poklesem PaO<sub>2</sub> > 20% (12)

Následn VCV 6mL/kg, optimal PEEP, Pplateau < 30 cm H<sub>2</sub>O,  
DF dle etCO<sub>2</sub> tak aby pH > 7.33 (350ml, df 25, Tinsp. 0,8)

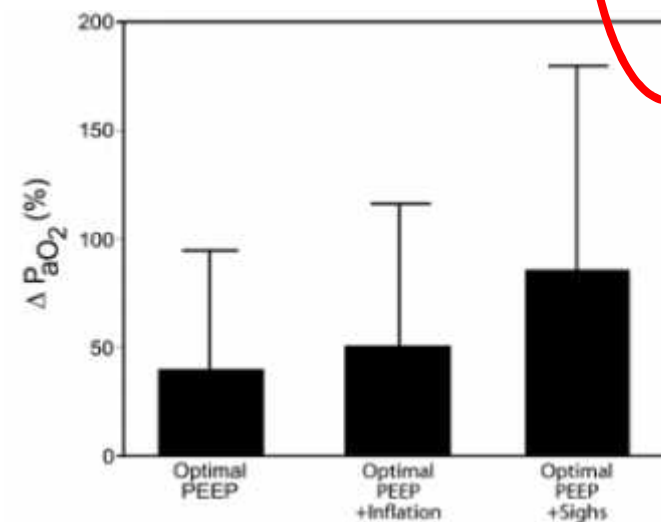
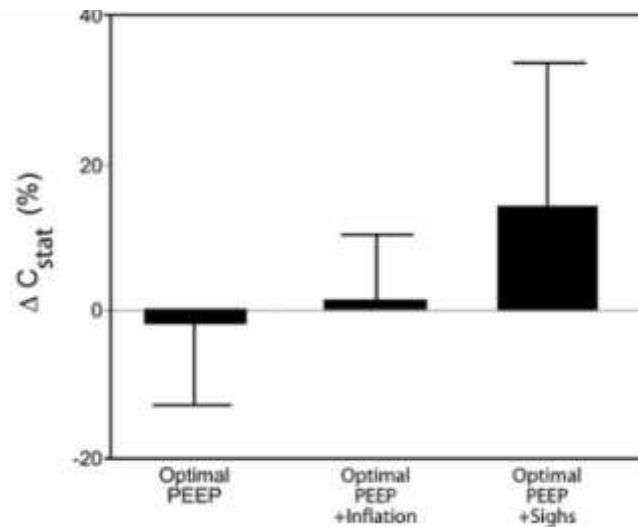
3 strategie: 1hod

- 1) optimal PEEP ( x maximal. O<sub>2</sub> transport )
- 2) optimal PEEP plus jednou SI RM (40 cm H<sub>2</sub>O/30 s)
- 3) optimal PEEP plus SIGH (Vt 12ml/, Pplateau < 40 cm H<sub>2</sub>O)  
každých 25 dech ( Vt 666ml )

# SIGH, intermitentní

Table 2. Respiratory Mechanics, Hemodynamics, and Arterial Blood Gas Values With 3 PEEP Strategies

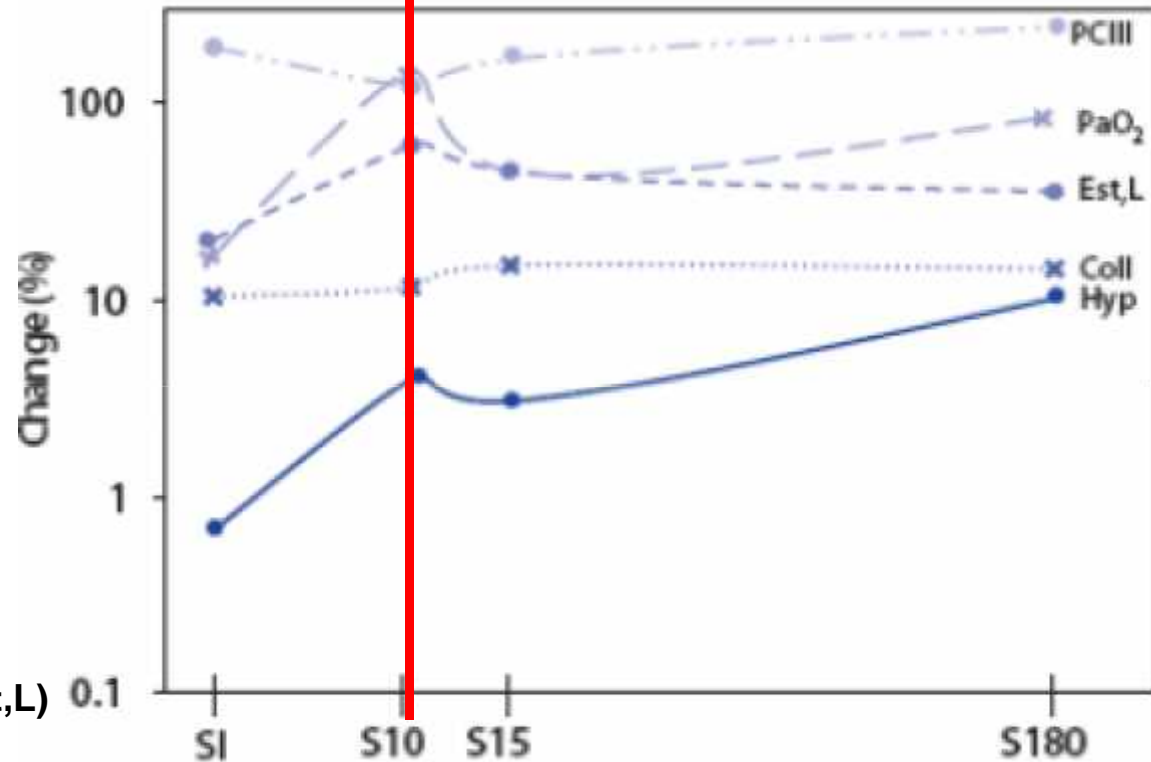
	Optimal PEEP Only (mean ± SD)		Optimal PEEP With a Sustained Inflation (mean ± SD)		Optimal PEEP With Sighs (mean ± SD)	
	Baseline	60 min	Baseline	60 min	Baseline	60 min
PEEP (cm H <sub>2</sub> O)	9 ± 5	12 ± 4	9 ± 5	12 ± 4	9 ± 5	12 ± 4
Total PEEP (cm H <sub>2</sub> O)	10 ± 4	13 ± 4	11 ± 4	13 ± 4	11 ± 4	14 ± 4
P <sub>plat</sub> (cm H <sub>2</sub> O)	20 ± 4	24 ± 5	20 ± 5	23 ± 5	21 ± 5	23 ± 5
C <sub>aw</sub> (mL/cm H <sub>2</sub> O)	37 ± 11	36 ± 10	38 ± 12	39 ± 13	37 ± 12	41 ± 13*
Mean systemic arterial pressure (mm Hg)	78 ± 8	78 ± 7	77 ± 8	75 ± 8	78 ± 9	81 ± 10
P <sub>aO<sub>2</sub></sub> (mm Hg)	111 ± 55	140 ± 53	114 ± 49	160 ± 63	110 ± 46	182 ± 66†
P <sub>aCO<sub>2</sub></sub> (mm Hg)	40 ± 8	41 ± 8	41 ± 8	40 ± 8	41 ± 8	40 ± 10
pH	7.39 ± 0.05	7.39 ± 0.05	7.39 ± 0.04	7.39 ± 0.04	7.39 ± 0.04	7.40 ± 0.04



# Sigh at different frequencies

(10, 15 and 180 per hour) to non-recruited acute lung injury rats

**Pplat = 40 cmH2O**



static lung lastance (Est,L)

oxygenation (PaO<sub>2</sub>)

fractional area of alveolar collapse (Coll)

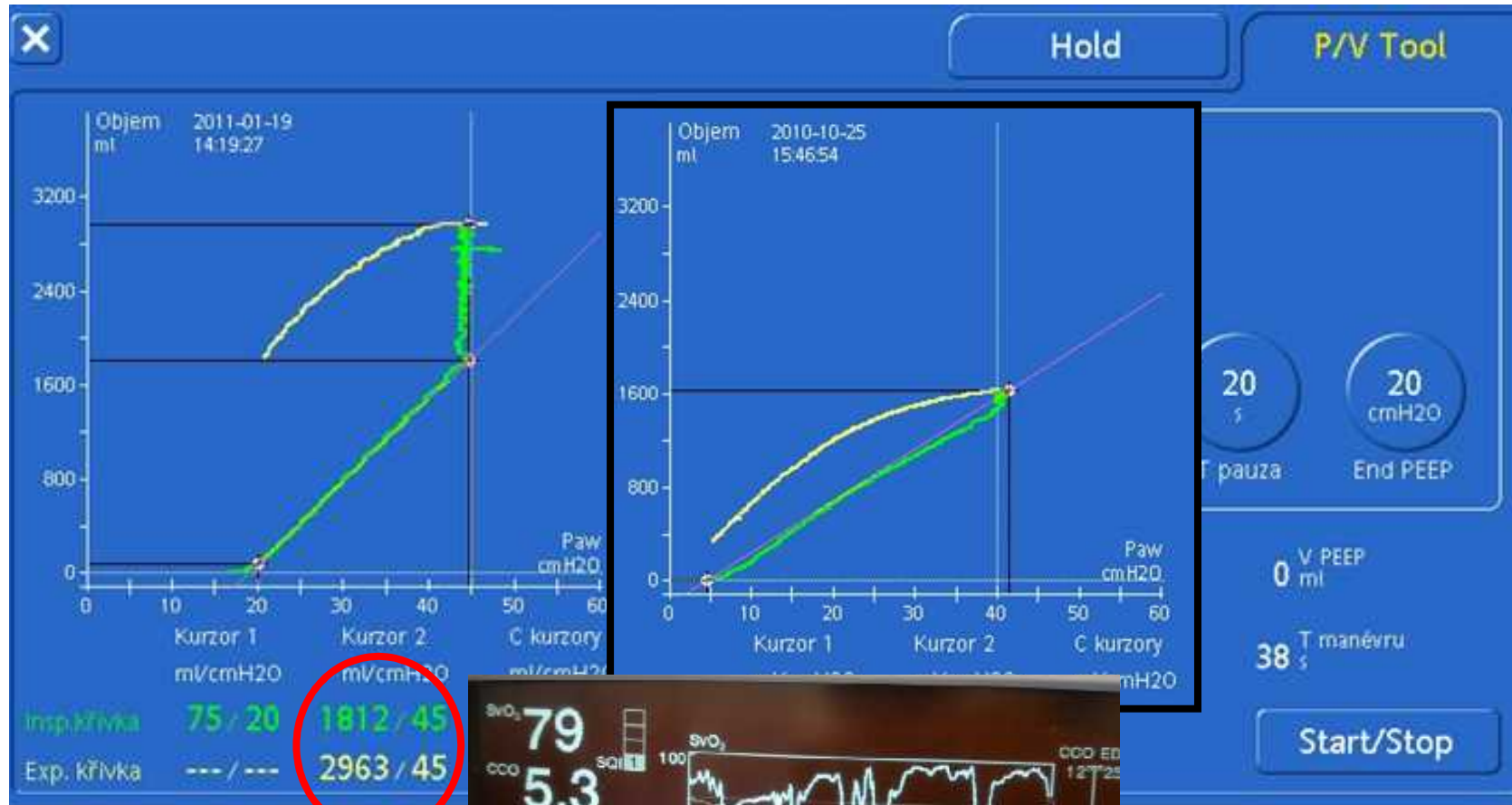
hyperinfl ation (Hyp),

mRNA expression of type III procollagen (PCIII)

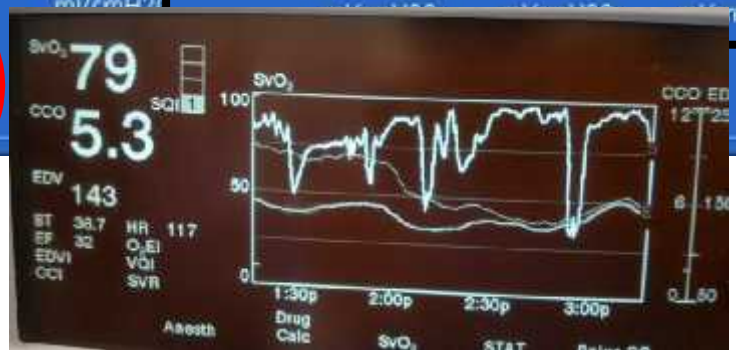
Steimback *et al.*: Eff ects of frequency and inspiratory plateau pressure during recruitment manoeuvres on lung and distal organs in acute lung injury. *Intensive Care Med* 2009



# PV tool



1812 / 45  
2963 / 45



# A co na to PK?

The Right Ventricle During the Acute Respiratory Distress Syndrome Revisited by Echocardiography

Antoine Vieillard-Baron

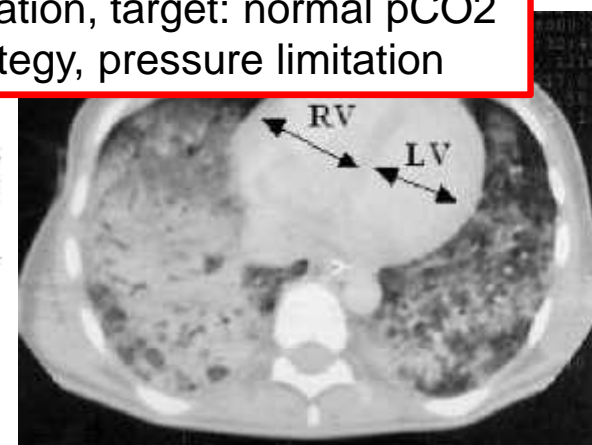
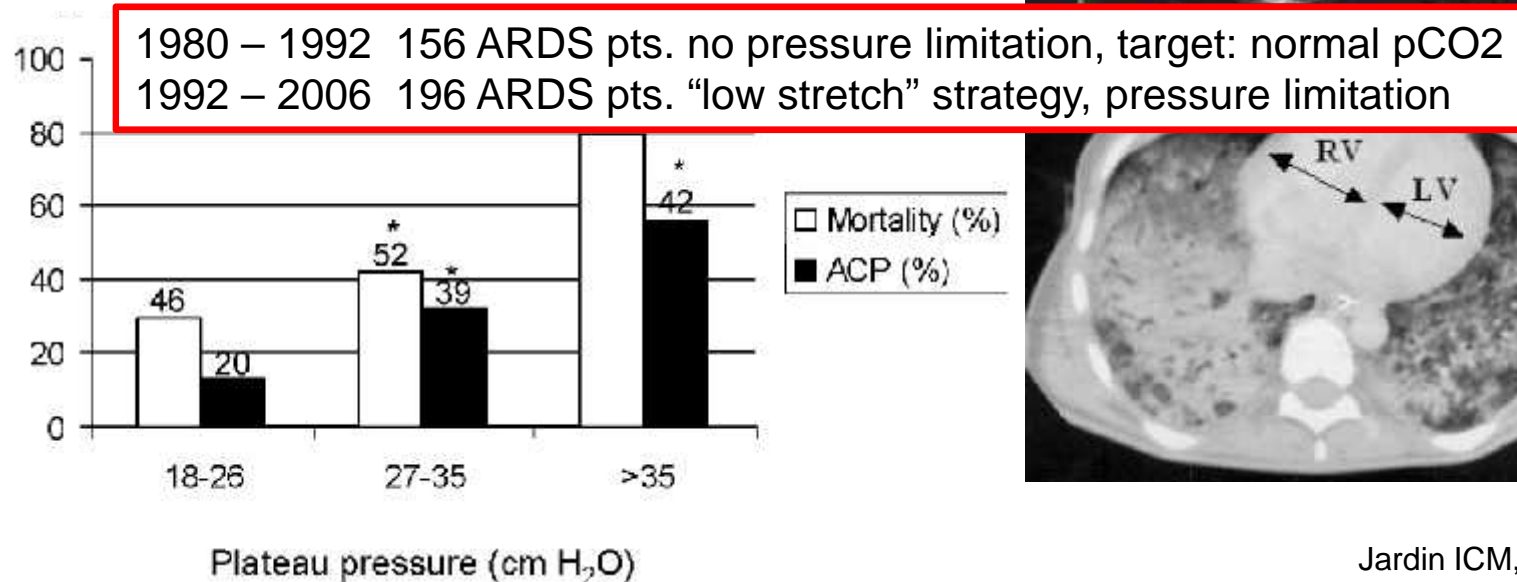
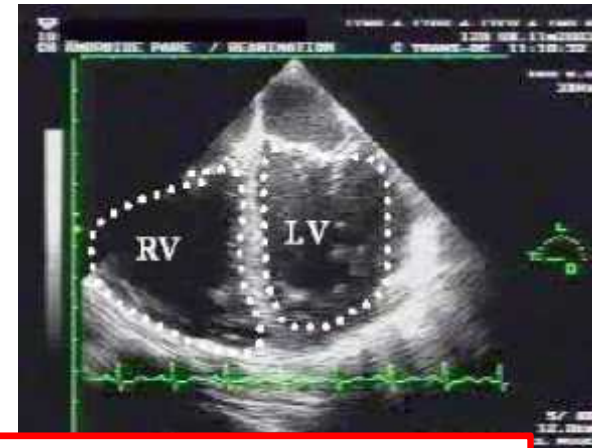
**Is there a safe plateau pressure in ARDS?**

**The right heart only knows**

François Jardin  
Antoine Vieillard-Baron

- Akutní cor pulmonale (ACP)  
PK/LK > 0.6 + septální dyskineza
- PK > LK 100% úmrtí
- **Signifikantní redukce mortality p i**

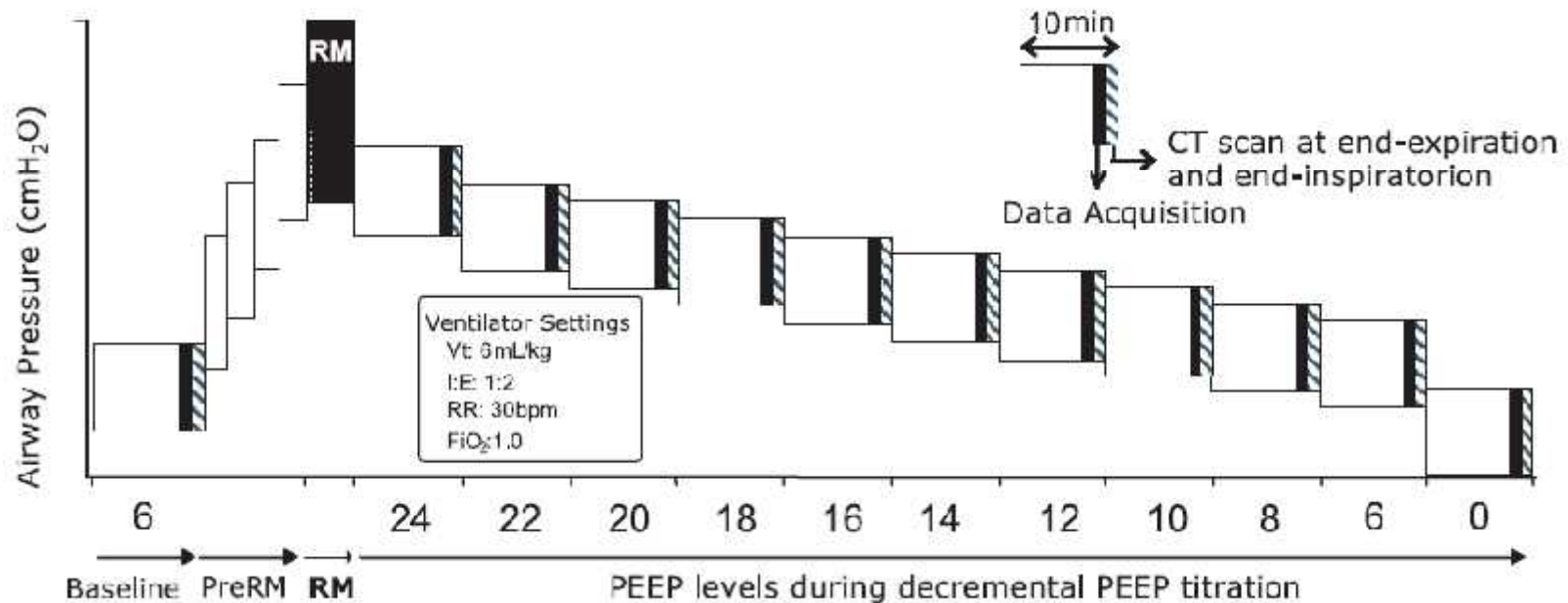
**Pplat < 27 cm H<sub>2</sub>O**



# Titrace PEEPu po RM pomocí Cdyn

Use of dynamic compliance for open lung positive end-expiratory pressure titration in an experimental study

Fernando Suarez-Sipmann, MD; Stephan H. Böhm, MD; Gerardo Tusman, MD; Tanja Pesch;

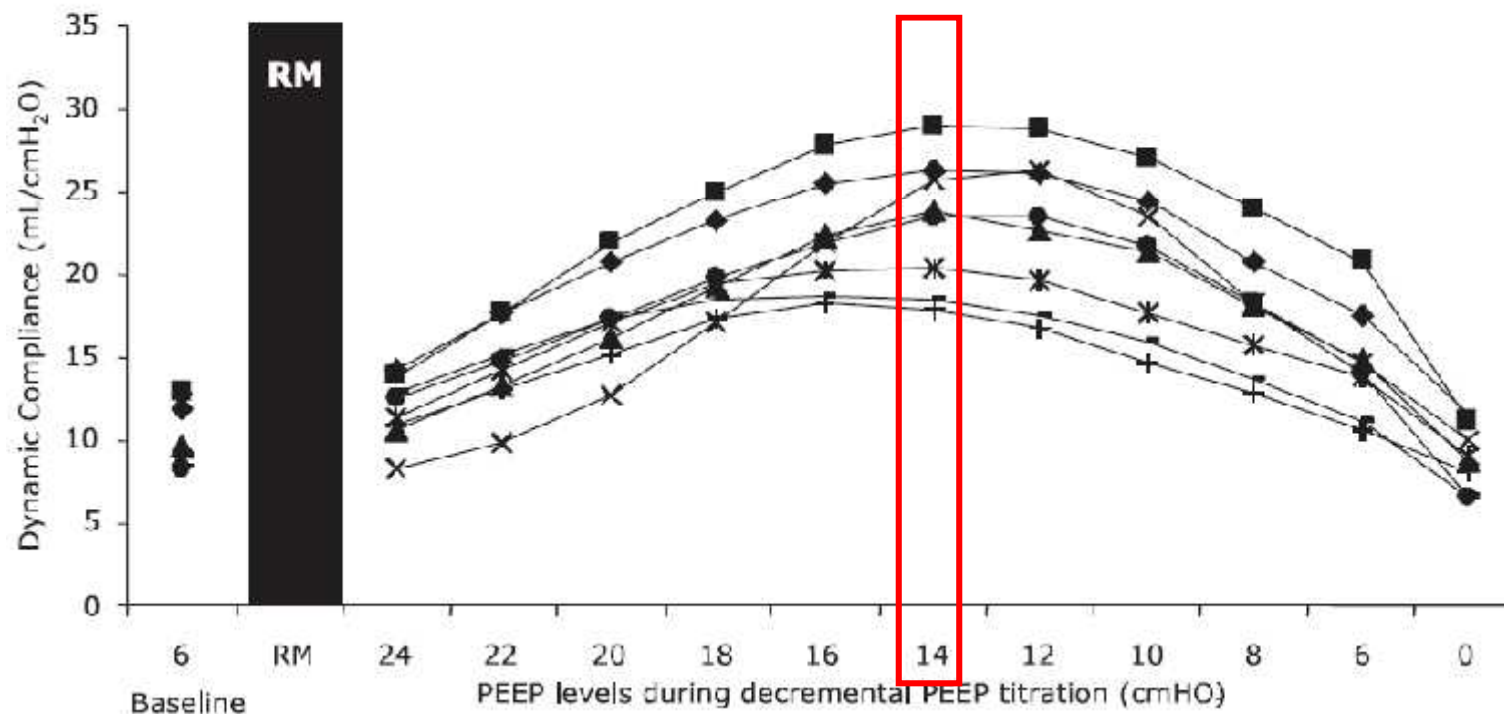


Prase í model, laváž FR á 5min až paO<sub>2</sub>/FiO<sub>2</sub><100, RM 30/60 1:1 2min  
 PEEP 24, regrese á 10min o -2cmH<sub>2</sub>O

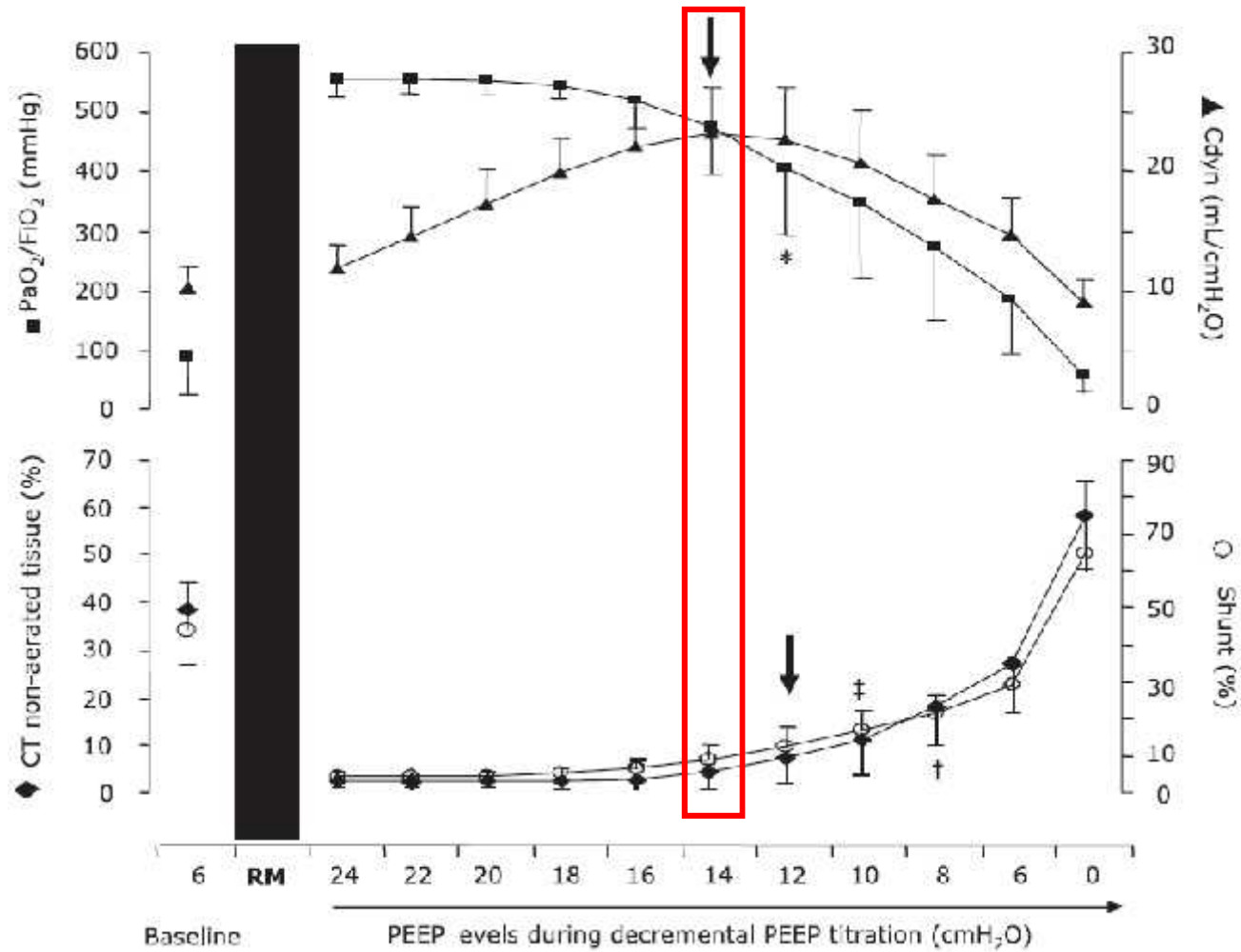
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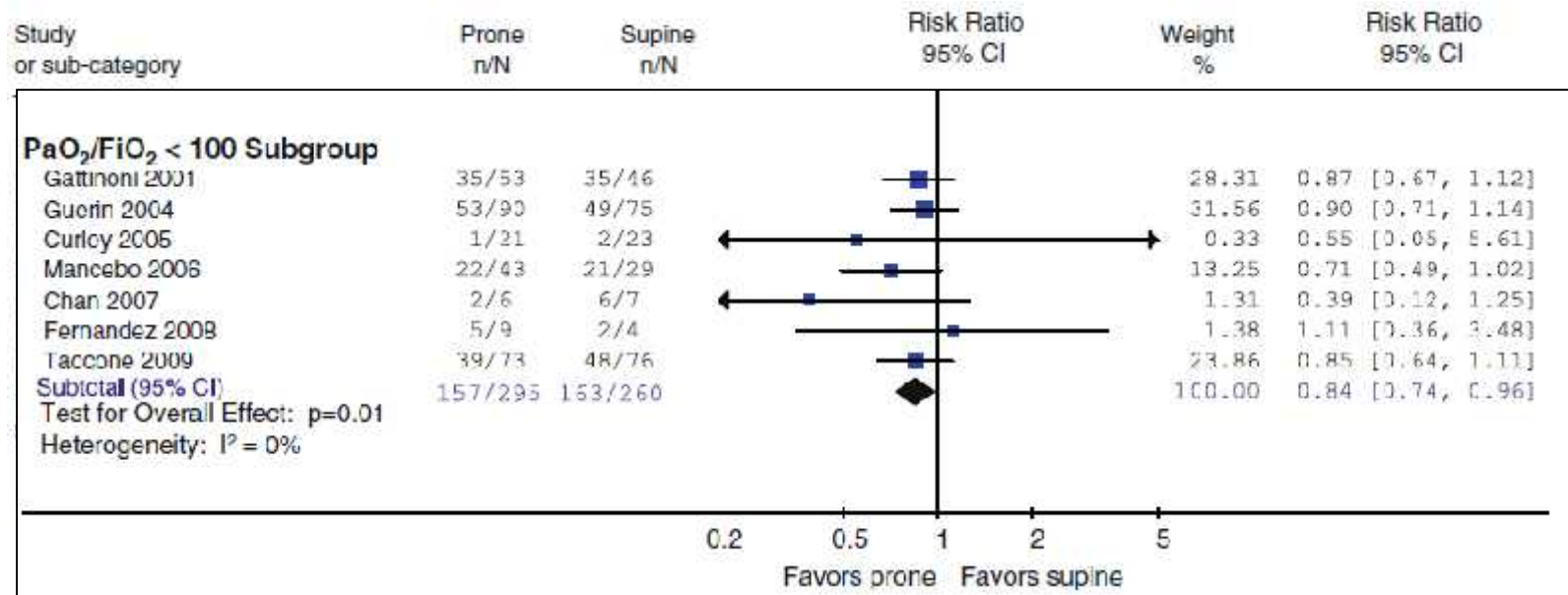


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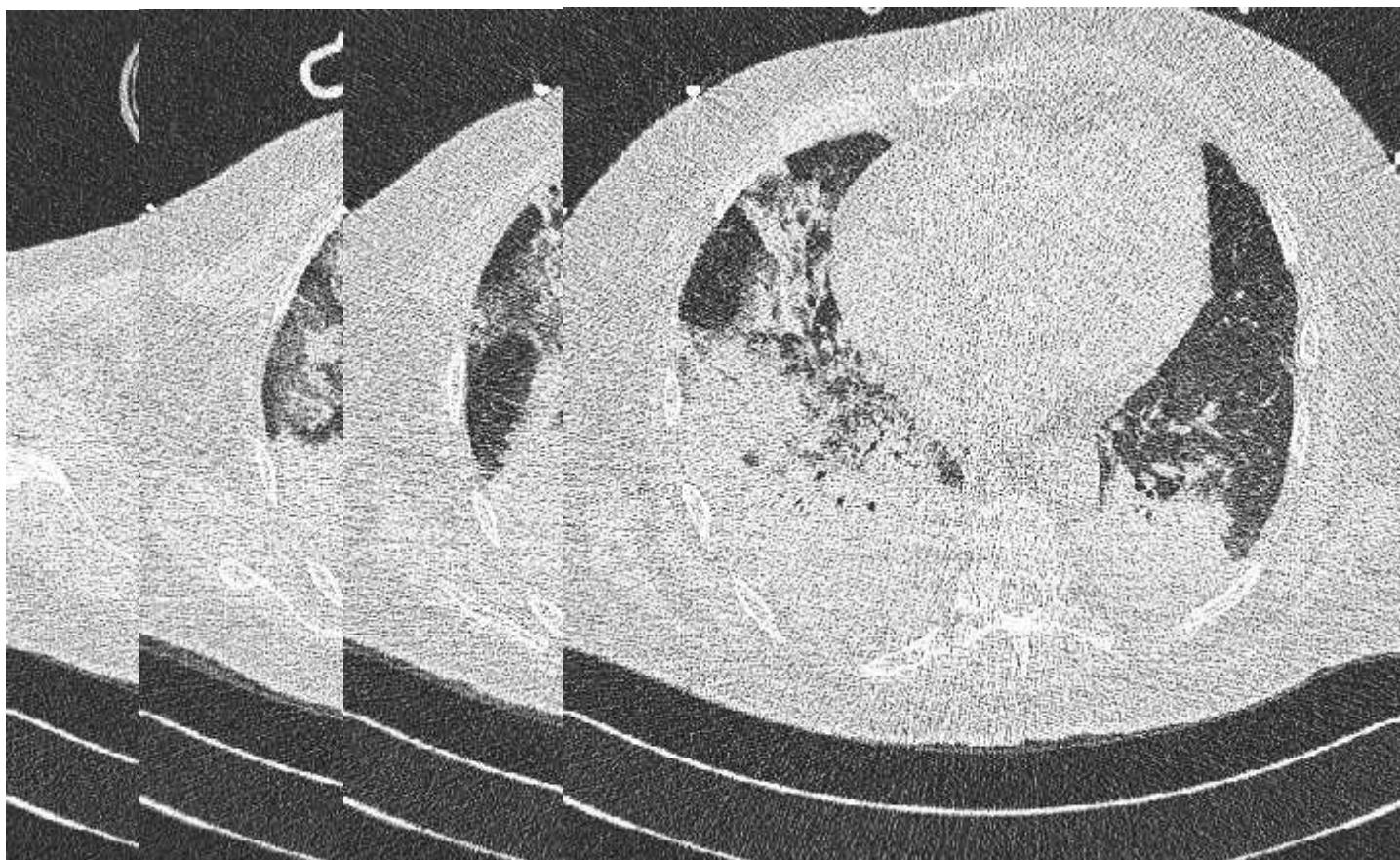


# Efekt PRONACE na mortalitu ARDS?

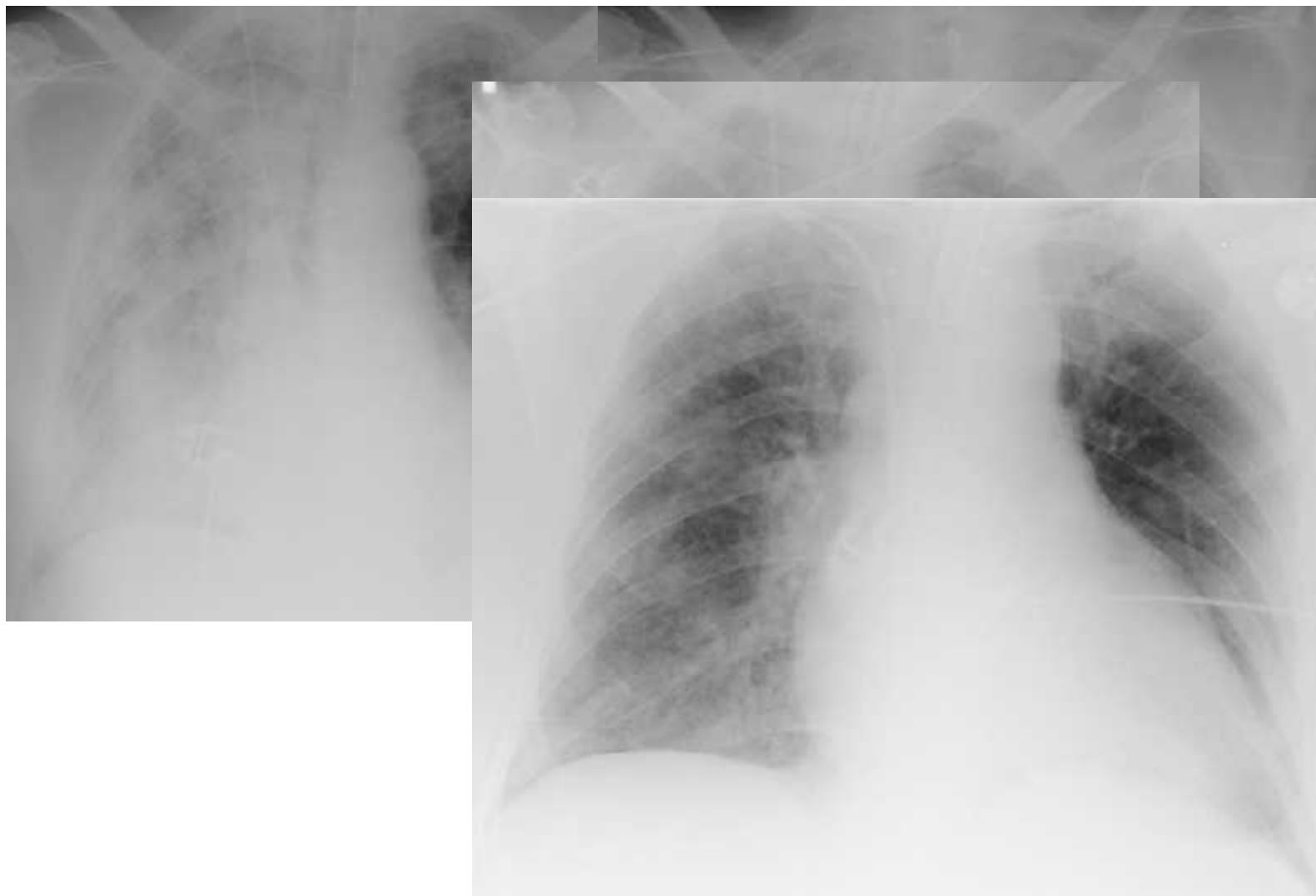
## Prone ventilation reduces mortality in patients with acute respiratory failure and severe hypoxemia: systematic review and meta-analysis



## Efekt SEMI - PRONACE



## Efekt SEMI - PRONACE



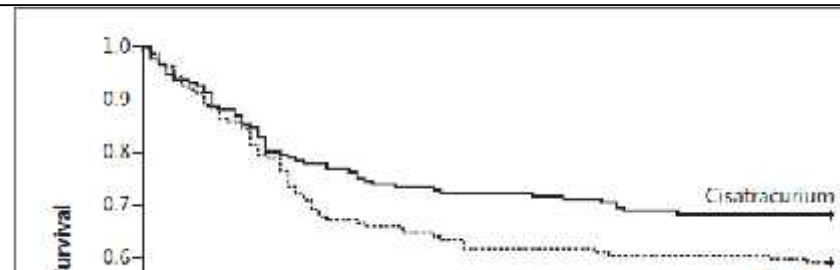


# Relaxace u ARDS

## Neuromuscular Blockers in Early Acute Respiratory Distress Syndrome

Laurent Papazian, M.D., Ph.D., Jean-Marie Forel, M.D.

**Multicentric, RCT, 20 ICUs**  
**340 pts. With ARDS for max. 48h.**  
**Cisatracurium vs. Placebo (178/162)**



Outcome	Cisatracurium (N=177)	Placebo (N=162)	Relative Risk with Cisatracurium (95% CI)	P Value
Barotrauma — no. (% [95% CI])†	9 (5.1 [2.7–9.4])	19 (11.7 [7.6–17.6])	0.43 (0.20–0.93)	0.03
Pneumothorax — no. (% [95% CI])	7 (4.0 [2.0–8.0])	19 (11.7 [7.6–17.6])	0.34 (0.15–0.78)	0.01
Patients without ICU-acquired paresis¶				
By day 28 — no./total no. (% [95% CI])	68/96 (70.8 [61.1–79.0])	52/77 (67.5 [56.5–77.0])		0.64
By ICU discharge — no./total no. (% [95% CI])	72/112 (64.3 [55.1–72.6])	61/89 (68.5 [58.3–77.3])		0.51

(P = 0.08)

Figure 2. Probability of Survival through Day 90, According to Study Group.

**rate of ICU-acquired paresis did not differ significantly between the two groups**

## Kontroverze

- nejsou „velké“ studie
- rozdílné kohorty, Berlínská definice?
- rozdílné provedení RM, nastavení PEEPu
- opakování RM?
- weaning .... mortalitní efekt?
  
- **ošetření derecruitmentu**  
(nebulizace, rozpojení, odsávání)  
„malý RM“ = inspirace ní hold PEEP>10, >14 RM?

## Závěr:

- PEEP není RM
- Recrutabilita? PV tool
- RM by mohl být součástí protektivní ventilace u ARDS - následně titrace optimálního PEEPu
- PK a její přetížení
- Pronašná poloha
- Relaxace v prvních 48h?
- PCV RM možná nejvýhodnější / PV tool



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**D kuji za pozornost**





# Workshop ventilace u ARDS

**VETRAM** - Ventilace ní TRénink na Animálním Modelu

**Praktický kurz ventilace, se speciálním zaměřením  
na model a management ARDS**

**30.11.2012**

**[www.karim-vfn.cz](http://www.karim-vfn.cz)**

