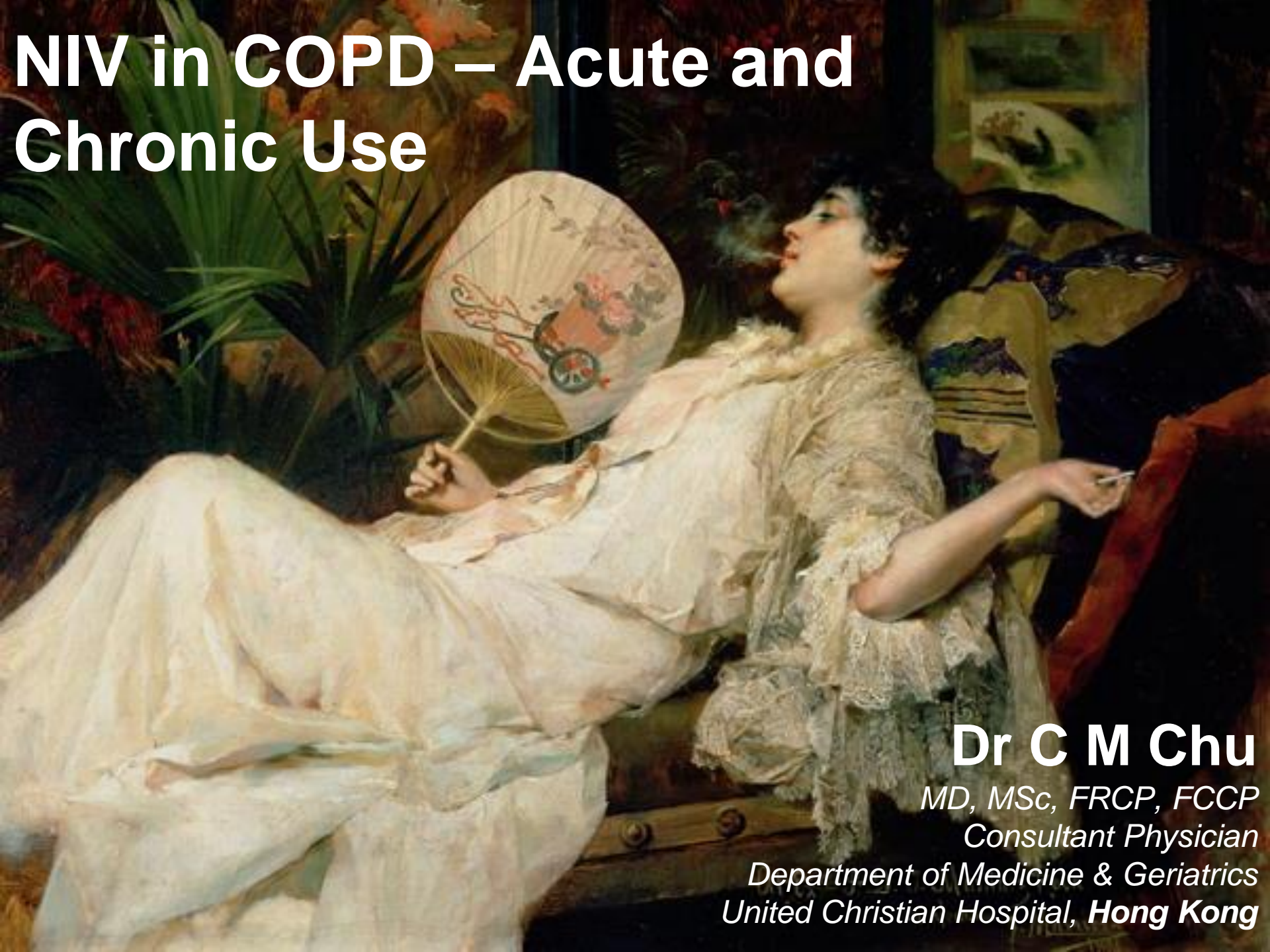


# NIV in COPD – Acute and Chronic Use



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*Consultant Physician*

*Department of Medicine & Geriatrics*

*United Christian Hospital, **Hong Kong***

# NIV in COPD

**I. AE-COPD/ARF**

**II. Weaning of  
intubated COPD**

**III. Home NIV**

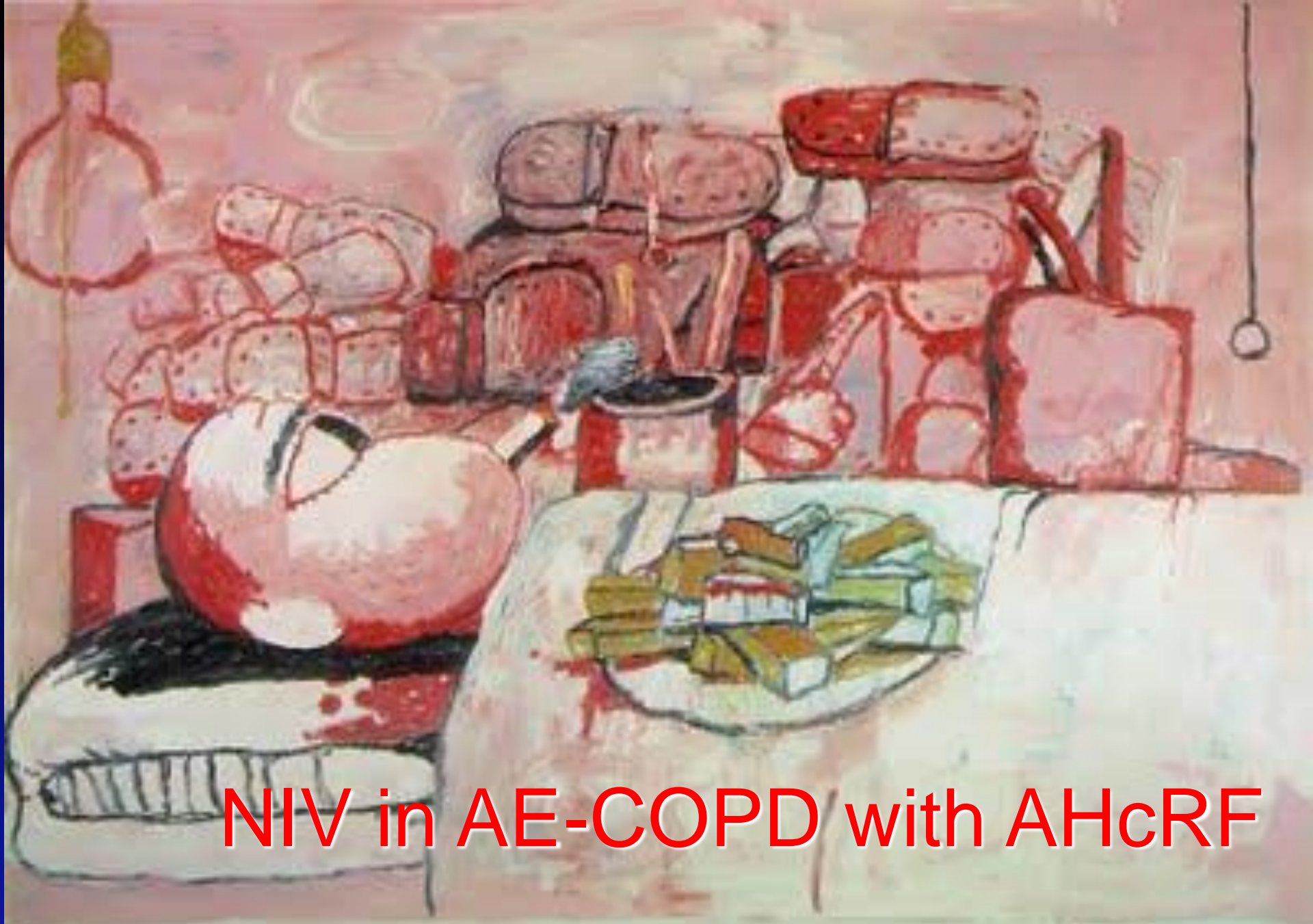
- 1. Stable pt.**
- 2. After ARF**

**1. COPD/OSA**

**2. COPD  
rehabilitation**

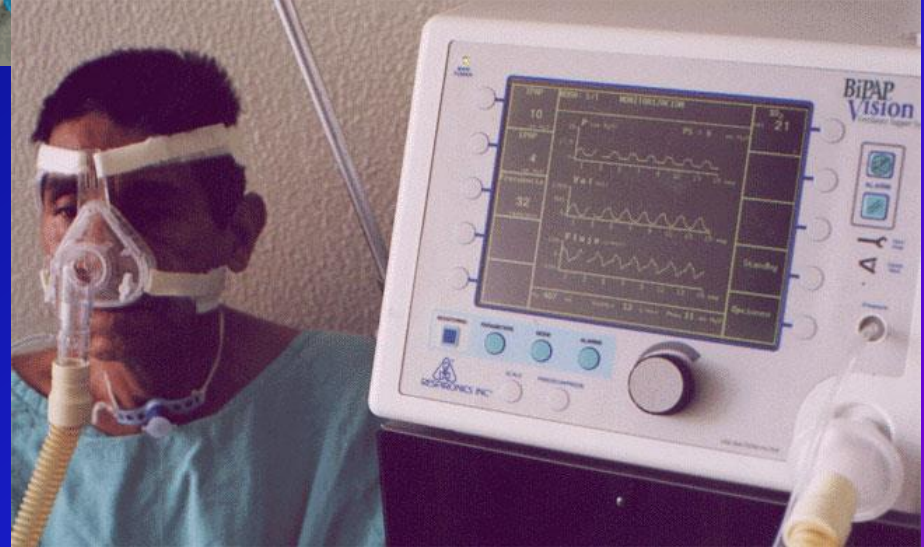
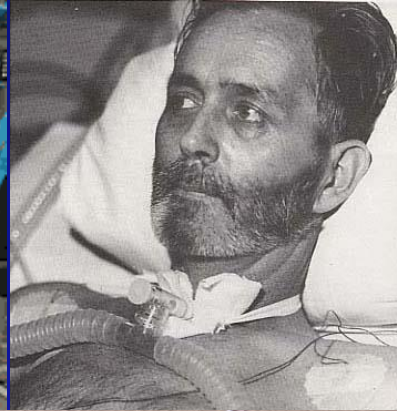
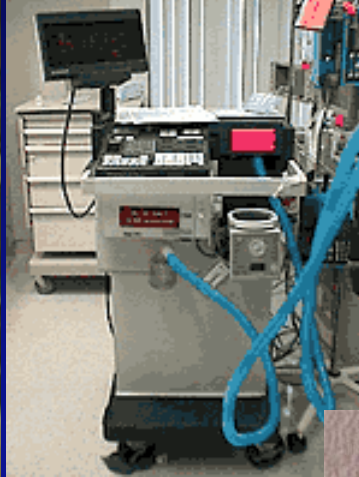
**3. Palliative care**





NIV in AE-COPD with AHcRF

# NIPPV in acute hypercapnic respiratory failure of COPD



# AE-COPD with AHcRF

- Persistent acidosis ( $\text{pH} < 7.35$ ) despite maximal medical Rx (NNT 10 to avoid 1 intubation)
  - $\text{pH} < 7.3$  strongly advised (50% intubation)
-



# Evidence base for acute NIV

Randomised controlled trials – reduced intubation, mortality

- Bott J, et al. Lancet 1993;341:1555.
  - Brochard L, et al. NEJM 1995;333:817.
  - Kramer N, et al. AJRCCM 1995;151:1799.
  - Angus RM, et al. Thorax 1996;51:1048.
  - Celikel T, et al. Chest 1998;114:1636.
  - Martin TJ, et al. AJRCCM 2000;161:807.
  - Plant PK, et al. Lancet 2000;355:1931.
-

# Meta-analyses

- COPD acute exacerbations with respiratory failure
  - NIV is associated with:
    - Lower mortality (RR = 0.41)
    - Lower intubation rate (RR = 0.42)
    - Lower treatment failure (RR 0.32)
    - Greater improvement in pH, PaCO<sub>2</sub>, RR at 1 hr
    - Fewer complications (RR = 0.51)
    - Shorter hospital stay  
(weight mean difference = - 3.24 days)
-

# Contraindications

- Cardiac/Respiratory arrest
- Unable to cooperate
- Unable to protect airway or clear secretions, high aspiration risk
- Facial surgery, injury or deformity
- Recent upper GI surgery/anastomosis
- Severely impaired mental state (relative)

**Hypercapnic encephalopathy NOT a contraindication, but higher failure rate (86% success with GCS < 8)**

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# NIV in AE-COPD with AHcRF

Trial of NIV should be given in  
most AE-COPD patients with  
AHcRF

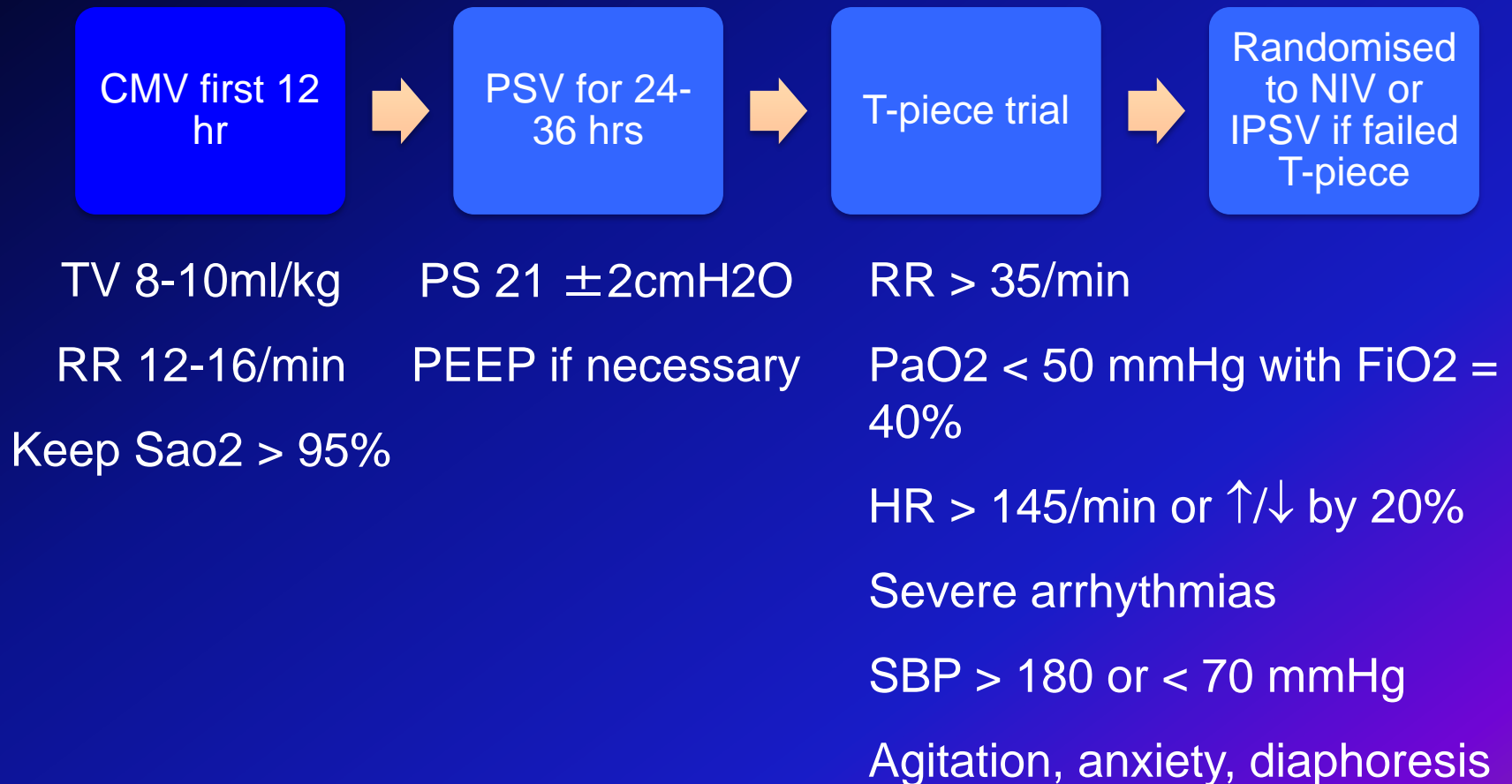


NIV in weaning of intubated  
COPD patients

# NIV in weaning of patients with Respiratory Failure due to COPD

- 2 group, parallel RCT, Multi-center
- COPD with RF, intubated
- Exclusion
  - Concomitant severe diseases
  - Cardiac arrest, AMI, APO
  - Shock
  - Sepsis
  - Trauma
  - GIB, obstruction, perforation
  - Metabolic coma, DKA
  - Drug overdose
  - Coagulopathy
  - Post-op

# Protocol





# Protocol

- NIV

- PS of  $19 \pm 2$  cm H<sub>2</sub>O to achieve RR < 25/min and acceptable ABG x 20 – 22 hr/day
- ↓ PS by 2 – 4 cm H<sub>2</sub>O per day
- Trial of spontaneous breathing (3 hours)
  - Sao<sub>2</sub> > 90% with FiO<sub>2</sub> ≤ 40%
  - pH ≥ 7.35
  - Hemodynamic stable; no resp distress, neurological stable

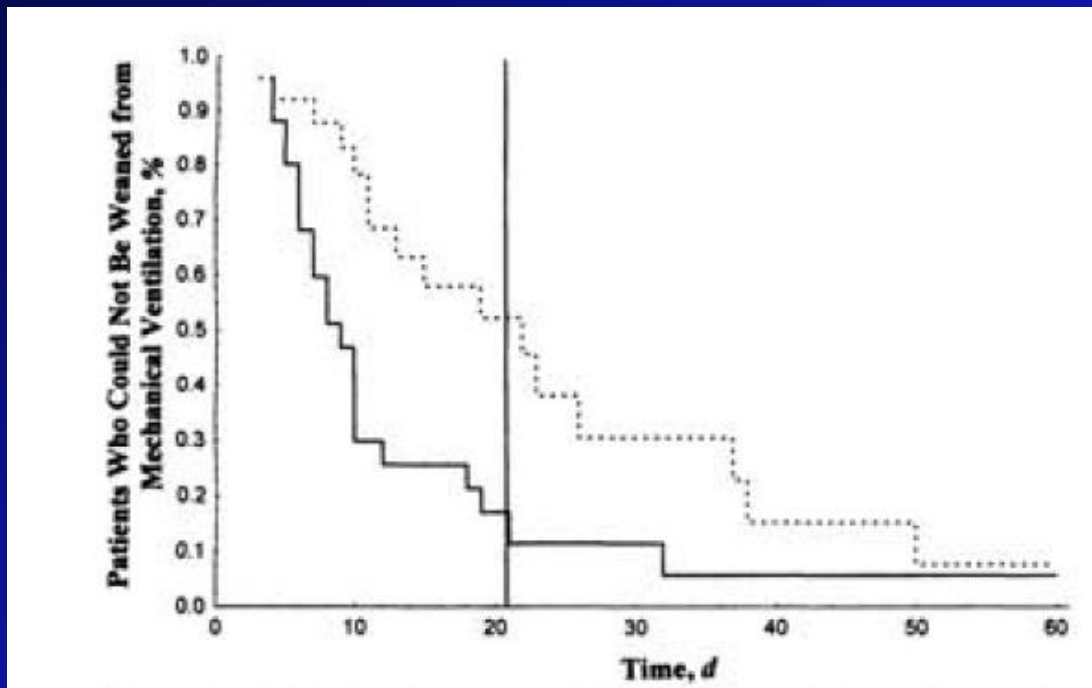
- IPSV

- PS  $17.6 \pm 2.1$  cm H<sub>2</sub>O
- Target RR ≤ 25/min
- PS decrease daily
- Intermittent T-piece trial

# NIV in weaning of patients with Respiratory Failure due to COPD - Results

- 50 enrolled, 25 in each group
- NIV vs. IPSV
  - Fewer days on ventilator: 10.2 vs 16.6 ( $p = 0.021$ )
  - Shorter ICU stay: 15.1 vs. 24 ( $p = 0.005$ )
  - Higher successful weaning at 21 days ( $p = 0.003$ )
  - No patient had VAP in NIV group (28% in IPSV)
  - Higher 60 days survival (92% vs 72%,  $p = 0.009$ )

# KM curves - weaning



**Figure 2. Kaplan-Meier curves for patients who could not be weaned from mechanical ventilation (defined as weaning failure or death linked to mechanical ventilation) in the two groups.**

The probability of weaning failure was significantly lower for the noninvasive ventilation group (cumulative probability for 60 days,  $P < 0.01$  by the log-rank test). The vertical line represents day 21, usually considered the threshold between weanable and unweanable patients. The solid line represents noninvasive pressure support ventilation; the dashed line represents invasive pressure support ventilation.

# Weaning of intubated COPD patients

NIV may facilitate weaning from ETMV in selected COPD patients

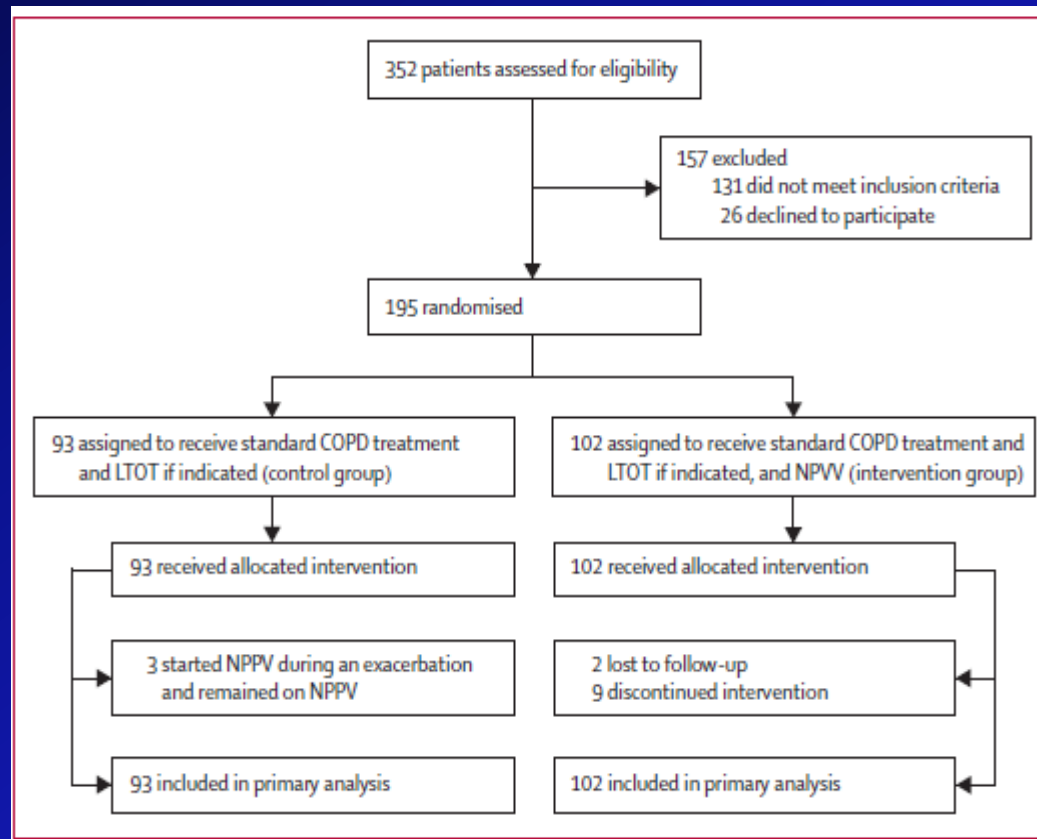




Home NIV for COPD

# Non-invasive positive pressure ventilation for the treatment of severe stable chronic obstructive pulmonary disease: a prospective, multicentre, randomised, controlled clinical trial

Thomas Köhnlein, Wolfram Windisch, Dieter Köhler, Anna Drabik, Jens Geiseler, Sylvia Hartl, Ortrud Karg, Gerhard Laier-Groeneveld, Stefano Nava, Bernd Schönhofer, Bernd Schucher, Karl Wegscheider, Carl P Crieé, Tobias Welte



# Subjects

- Stable COPD
- Stage IV disease with chronic hypercapnia
- NIV group – achieve 20% reduction in  $\text{PaCO}_2$
- Mean IPAP 21.6 cm  $\text{H}_2\text{O}$ ; EPAP 4.8 cm  $\text{H}_2\text{O}$ ; RR 16.1/min

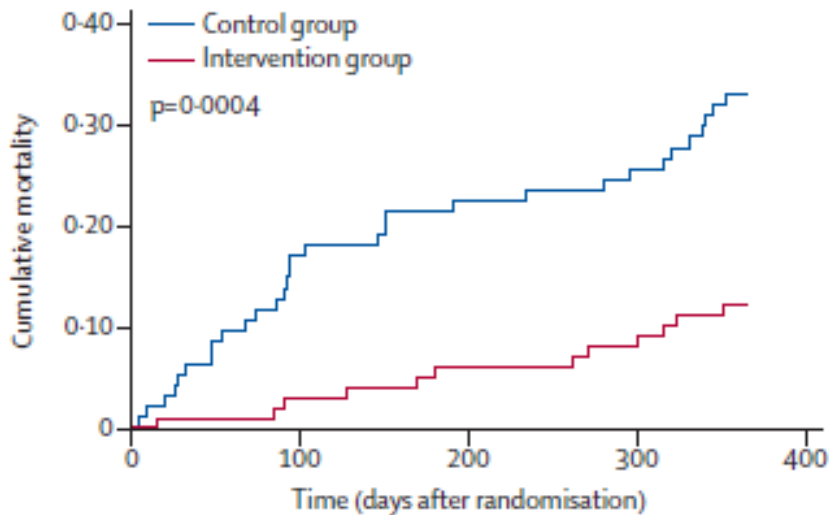
	Control group (n=93)	Non-invasive positive pressure ventilation group (n=102)
Age, years	64.4 (8.0)	62.2 (8.6)
Male, n (%)	56 (60%)	65 (64%)
Body-mass index, $\text{kg/m}^2$	24.5 (5.8)	24.8 (5.8)
FVC, % predicted	53.3% (13.8)	50.4% (13.3)
FEV <sub>1</sub> , % predicted	27.5% (8.9)	26% (11.0)
FEV <sub>1</sub> /FVC, %	41.2% (11.4)	40.4% (11.5)
Residual volume/total lung capacity, %	72.7% (8.9)	73.0% (8.5)
pH	7.39 (0.05)	7.39 (0.04)
$\text{PaCO}_2$ , kPa	7.7 (0.7)	7.8 (0.8)
$\text{PaO}_2$ , kPa*	8.7 (1.9)	8.6 (2.1)
$\text{SaO}_2$ , %*	90.8% (5.9)	90.3% (6.2)
$\text{HCO}_3^-$ , mmol/L	33.9 (4.1)	34.3 (4.0)
Base excess, mmol/L	8.0 (3.9)	7.8 (3.8)
6-min walk distance, m	249.6 (145.3)	226.7 (121.2)
Long-term oxygen treatment, n (%)	60 (65%)	67 (66%)

Data are mean (SD), unless otherwise stated. FVC=forced vital capacity. FEV<sub>1</sub>=forced expiratory volume in 1 s.  $\text{PaCO}_2$ =arterial carbon dioxide pressure.  $\text{PaO}_2$ =arterial oxygen pressure.  $\text{SaO}_2$ =arterial oxygen saturation.  $\text{HCO}_3^-$ =bicarbonate. \*In patients with long-term oxygen treatment, oxygen was applied via nasal cannula at the previously prescribed flow rate.

**Table 1: Baseline demographic and clinical characteristics of patients**

# Outcomes

- 1-yr mortality
  - NIV: 11.8%
  - Control: 33.3%
- Hazard ratio 0.24
- QoL – 6.2 point improvement in SGRQ,  $p = 0.029$



Number at risk  
Control group 93 77 72 69  
Intervention group 102 95 92 90



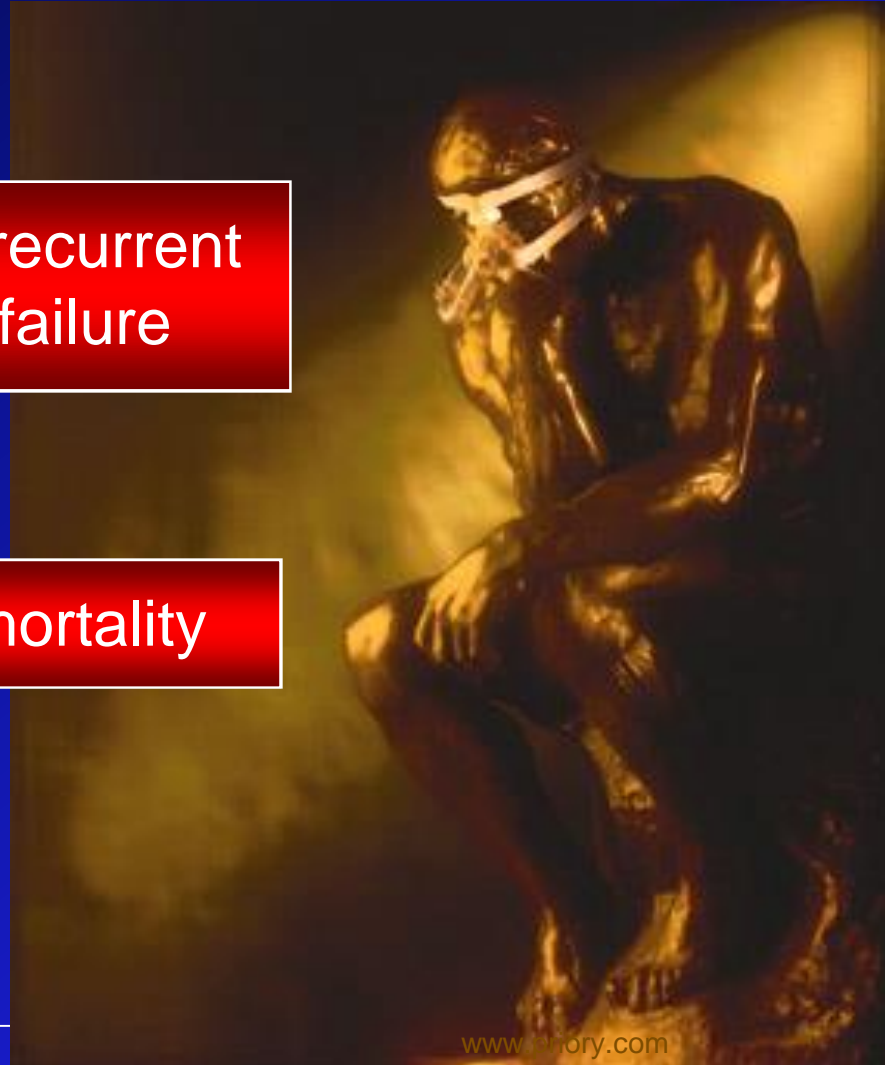
# Home NIV in survivors of COPD after ARF

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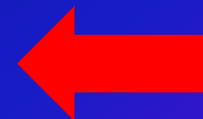
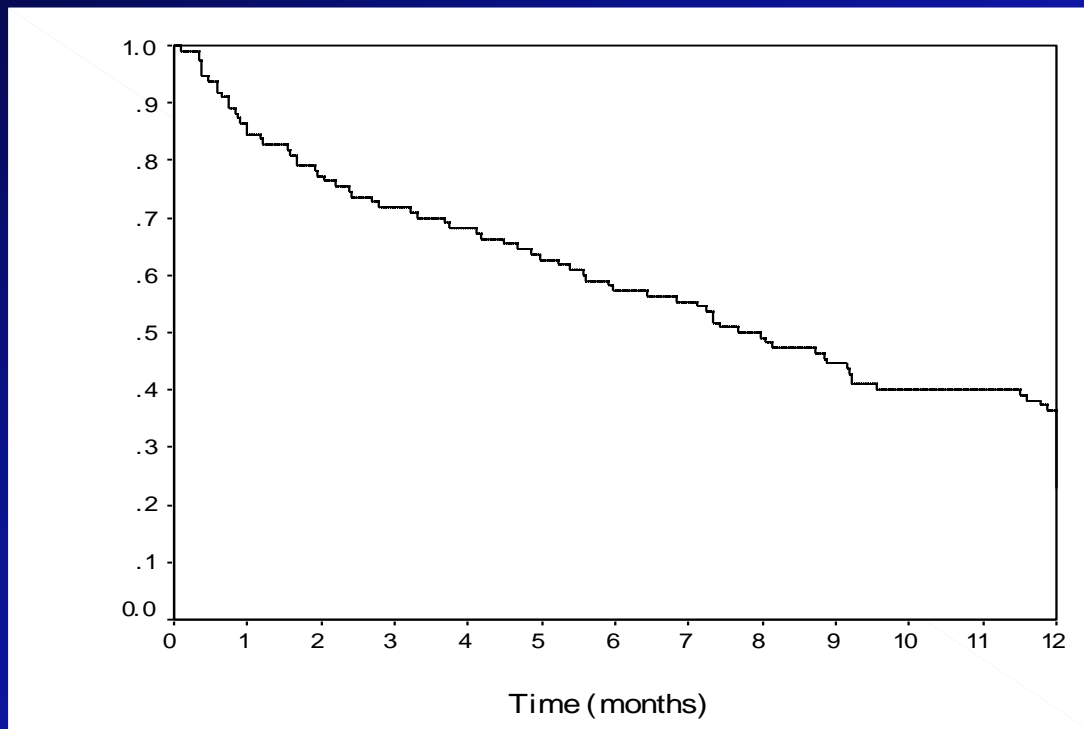
# Problem of the Survivor of ARF in COPD

High rates of recurrent respiratory failure

High mortality

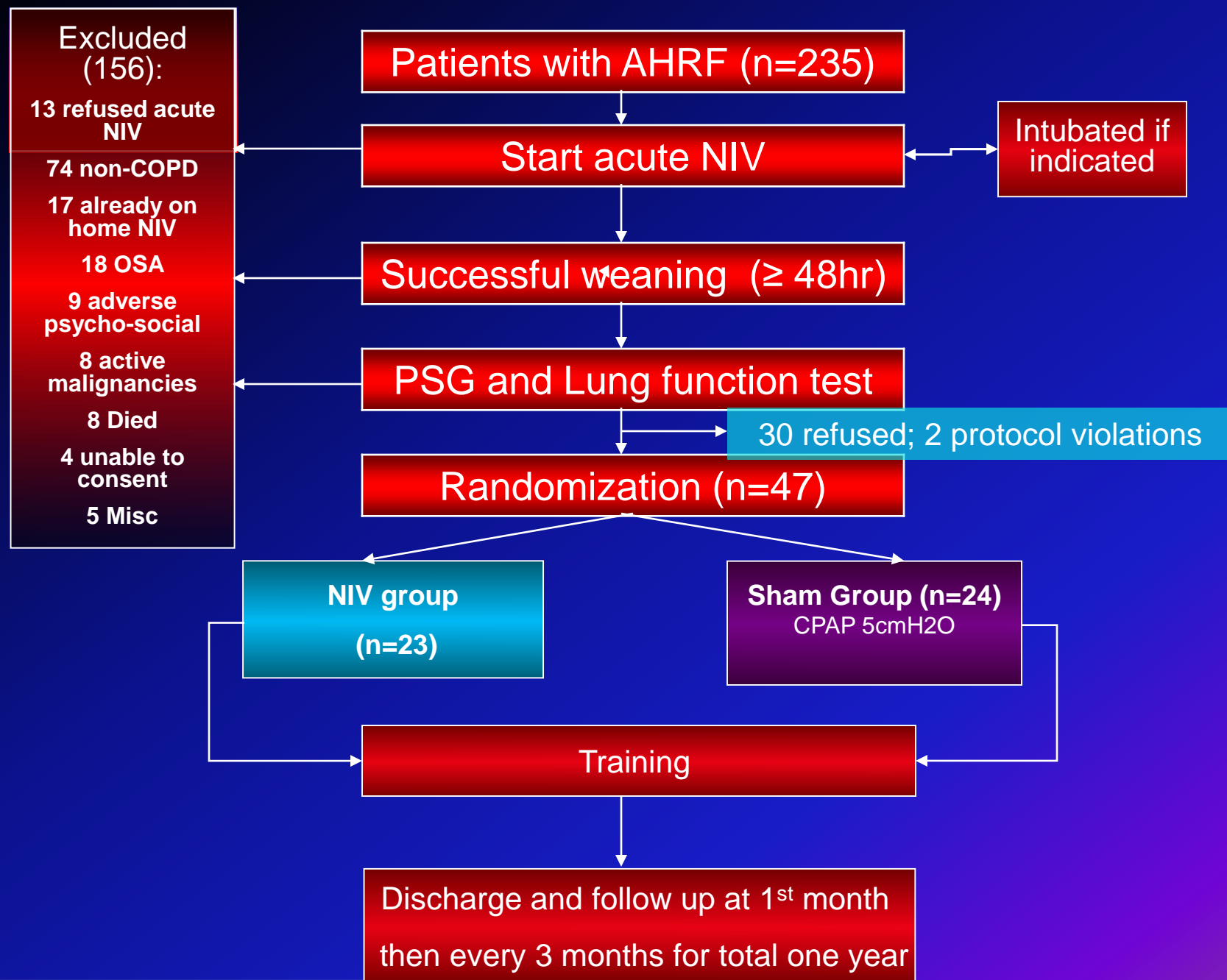


# Probability of Event-free Survival



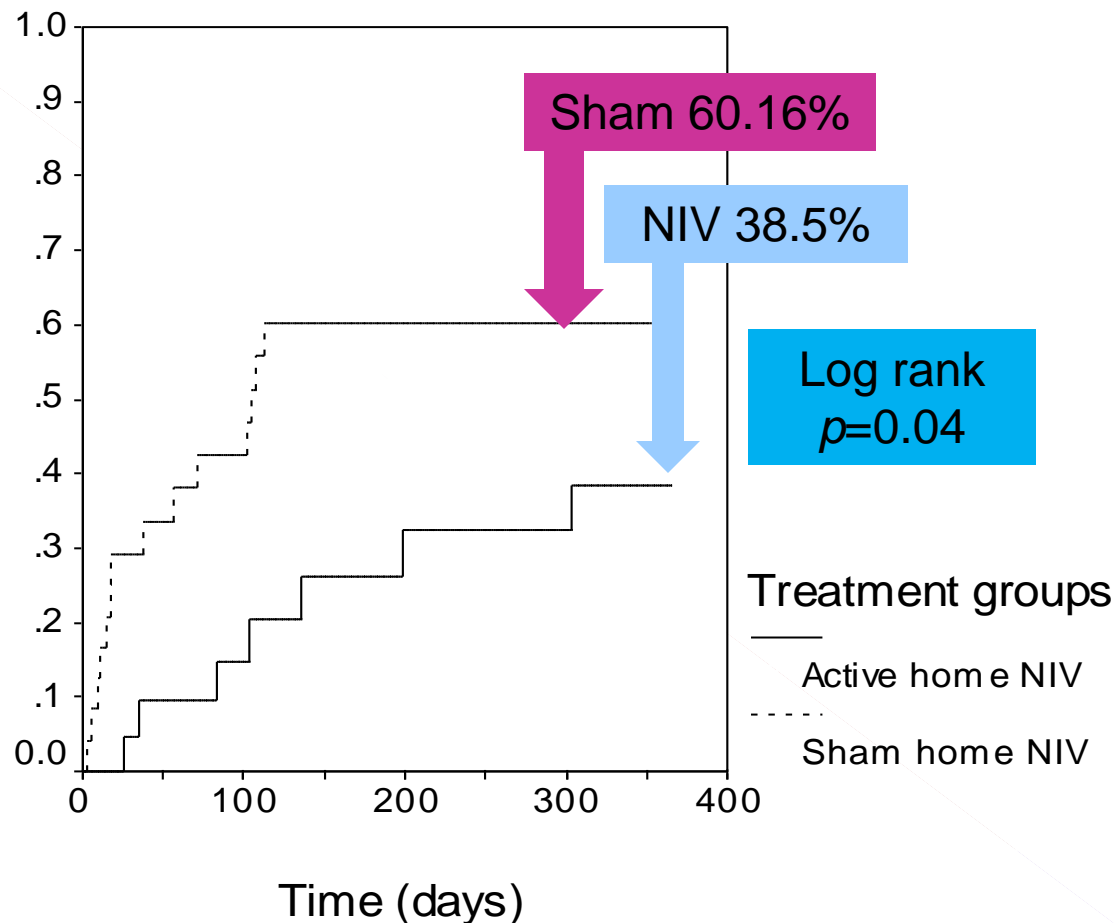
Recurrent  
AHRF: 63%  
in 1 year

Death ~ 50%





# Primary Study Outcome - Recurrent AHRF



# HMV vs HOT after AE-COPD

Murphy P, et al.

JAMA. 2017 Jun 6; 317(21):  
2177–2186.

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# Inclusion

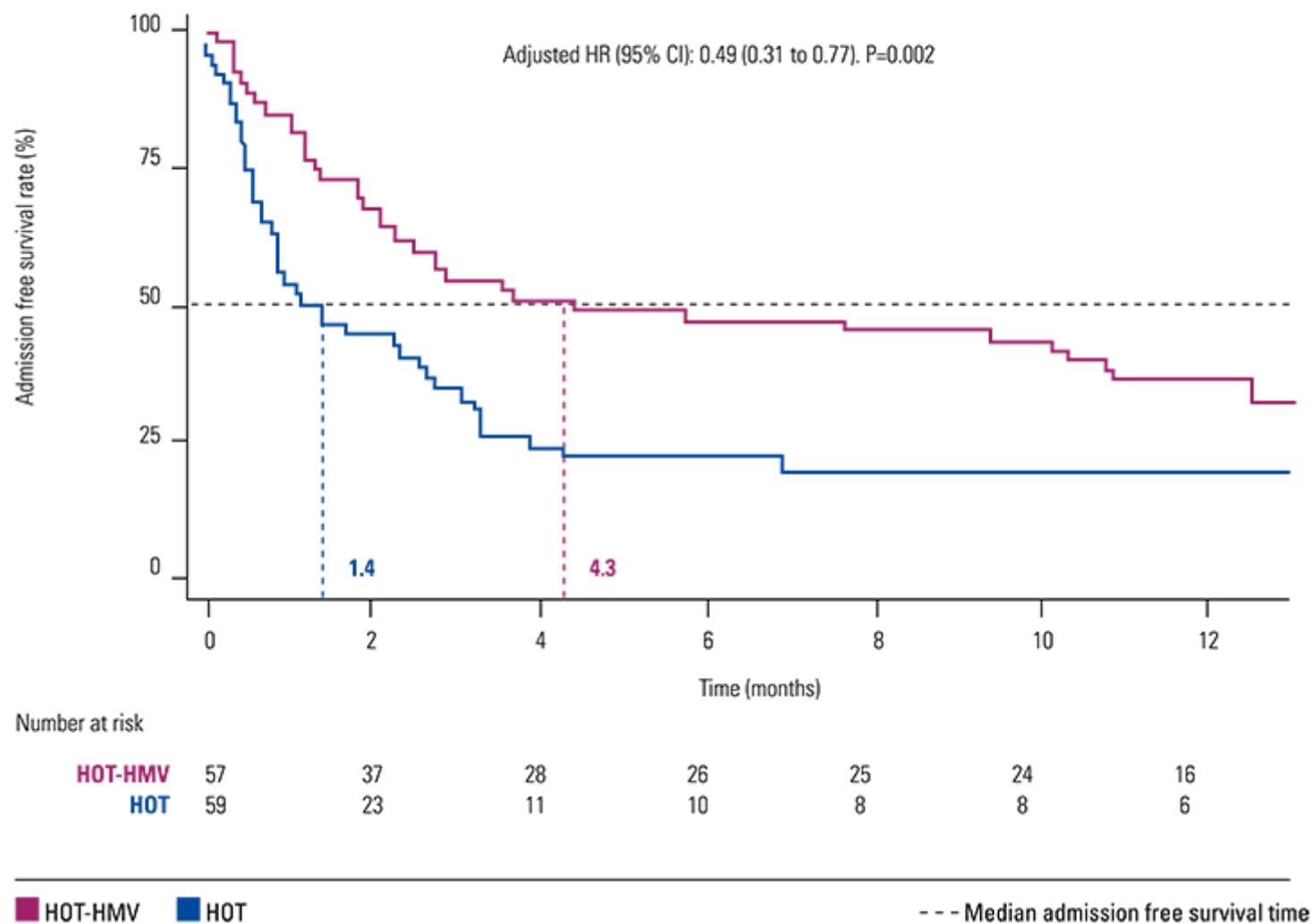
- FEV1 <50% of predicted
  - In patient admission with AE-COPD.
  - Persistent hypercapnia 2 to 4 weeks after the resolution of the hypercapnic acidosis.
  - Chronic hypoxia PaO<sub>2</sub> <55 mmHg or <60 mmHg complications
  - Smoking > 20 pack-years.
-

# Results

1. HMV combined with HOT can significantly reduce the risk of hospital readmission or death by 51% within 12 months in hypercapnic COPD patients.
  2. Improved QoL
  3. Health economics analysis underway
-

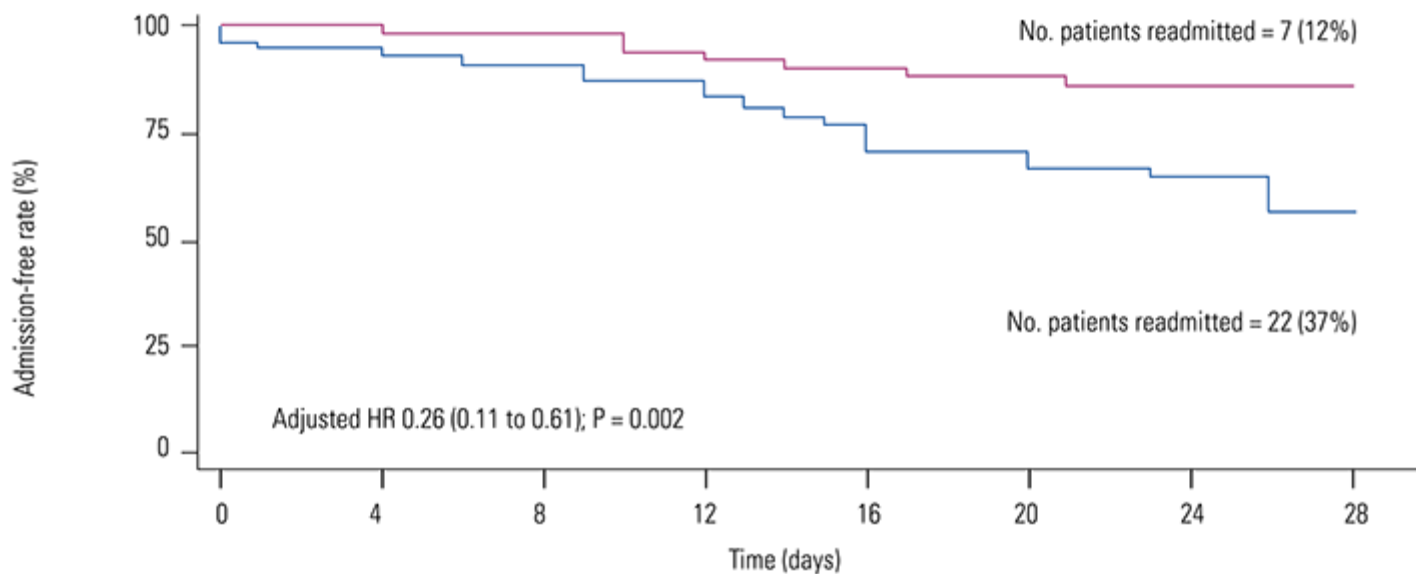
## Time to readmission or death from randomisation to follow-up at 1 year

Intention to treat analysis





# Time to hospital readmission by treatment arm



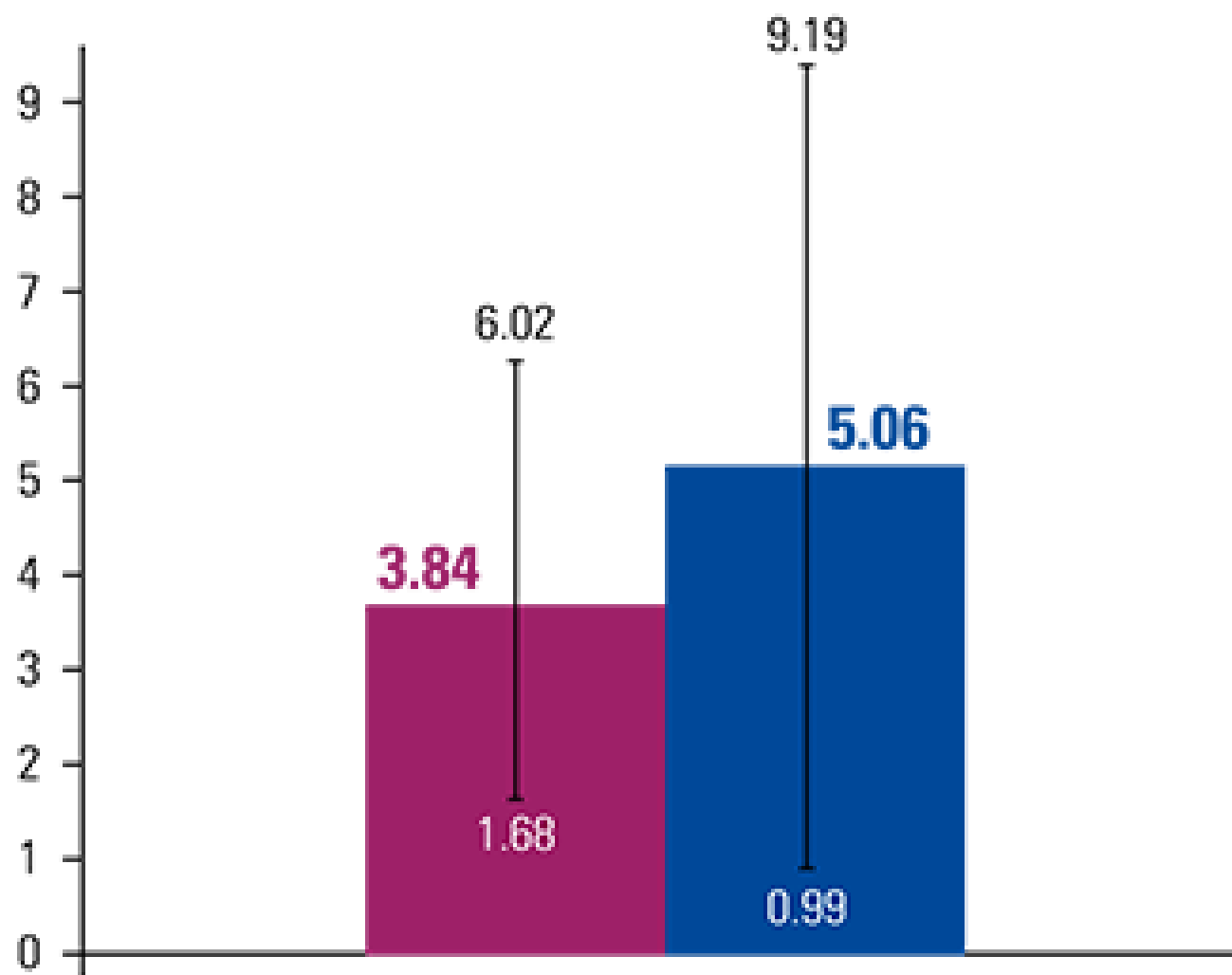
Number at risk

<b>HOT-HMV</b>	57	52	50	48	46	45	44	44
<b>HOT</b>	59	49	46	44	38	35	32	28

**HOT-HMV** **HOT**

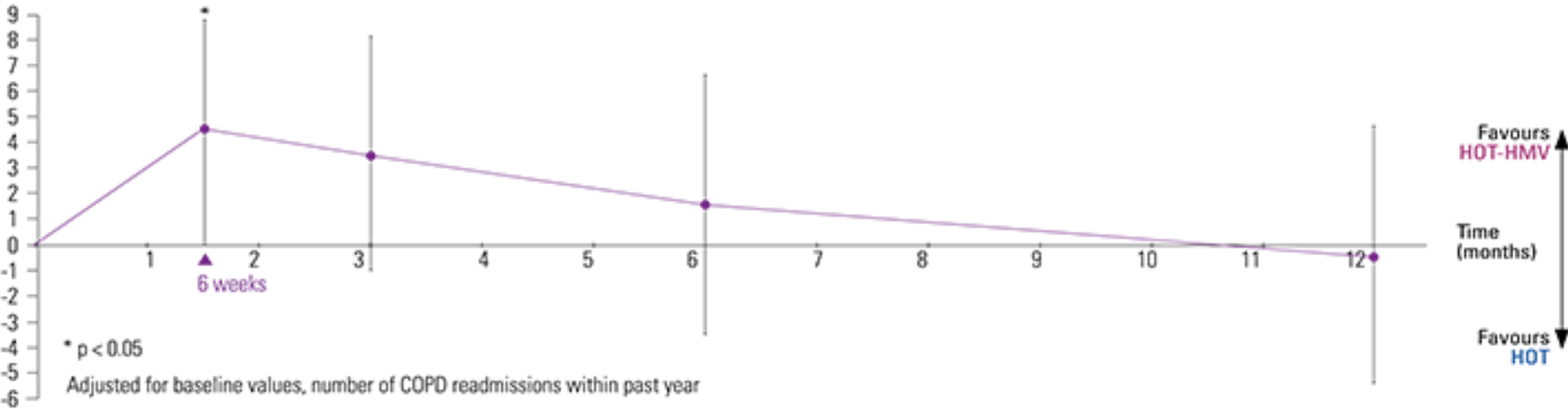
# Median exacerbation rate over one year (25<sup>th</sup> to 75<sup>th</sup> percentile)

■ H0T-HMV  
■ H0T



Adjusted rate ratio (95% CI): 0.66 (0.46 to 0.95). P = 0.03

SRI questionnaire



# Home NIV for COPD

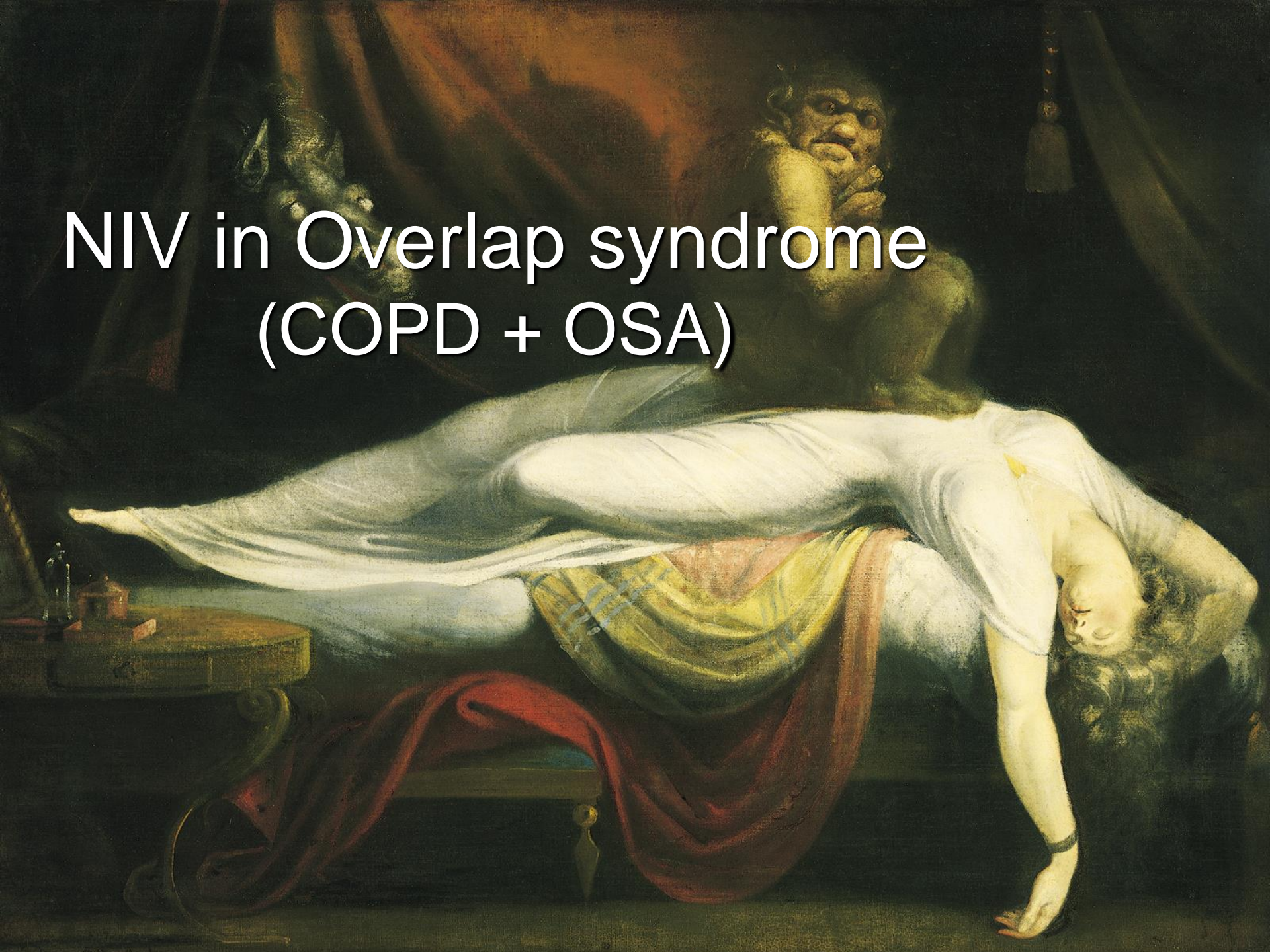
## Stable hypercapnia

- Improves survival(1 RCT)
- Improves QoL
- Maximal tolerated pressure

## Following ARF

- Persistent hypercapnia > 2 weeks
- Improved survival
- ? Recurrent ARF
- More RCTs needed

# NIV in Overlap syndrome (COPD + OSA)



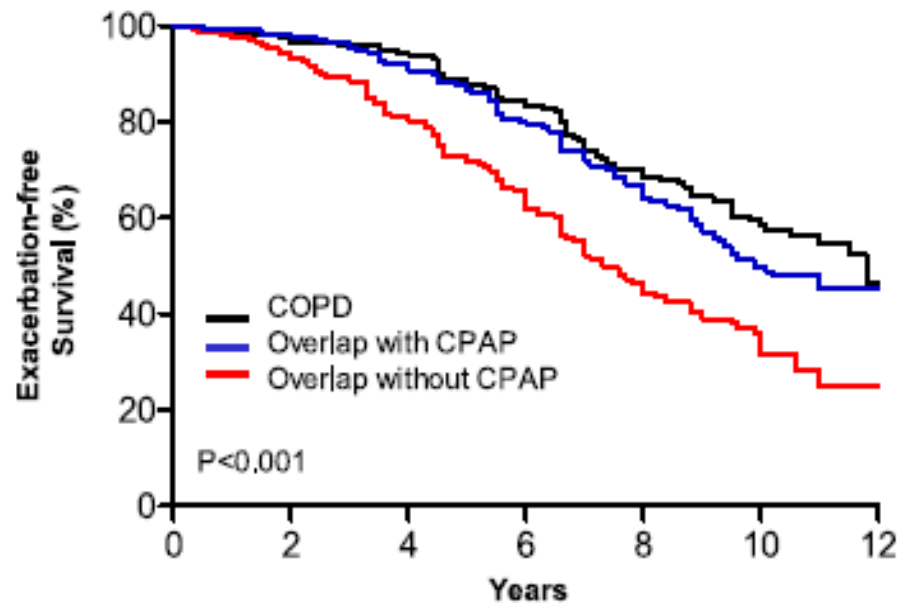
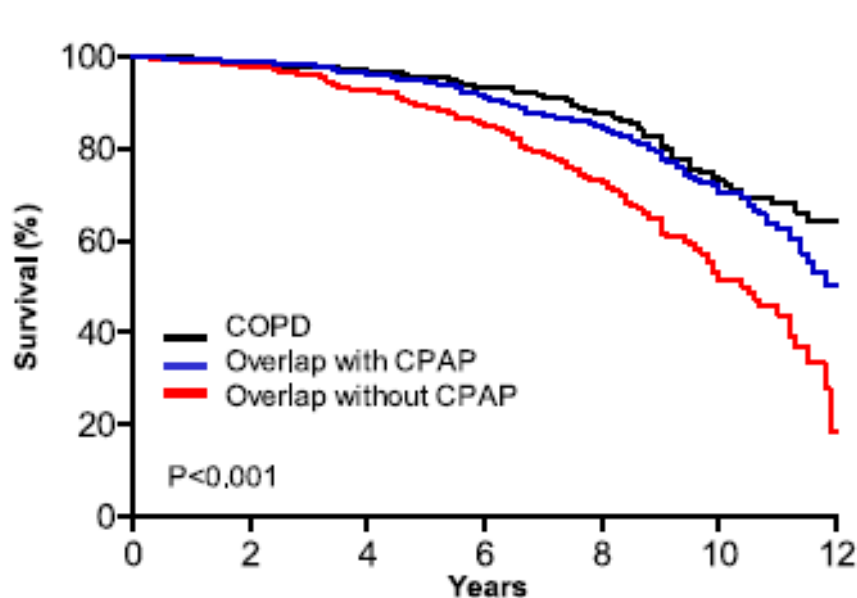


## II. Overlap syndrome (COPD + OSA)

Worse prognosis vs. either condition alone

- More nocturnal desaturation
  - Worse hypoxemia and hypercapnia
  - More nocturnal arrhythmias
  - Pulmonary HT, RHF
  - Lower 5-yr survival
-

# Survivals and exacerbation requiring hospitalisation



# CPAP in overlap syndrome

Adjusted for age, sex, BMI, smoking, alcohol, comorbidities, FEV1 and sleepiness scores:

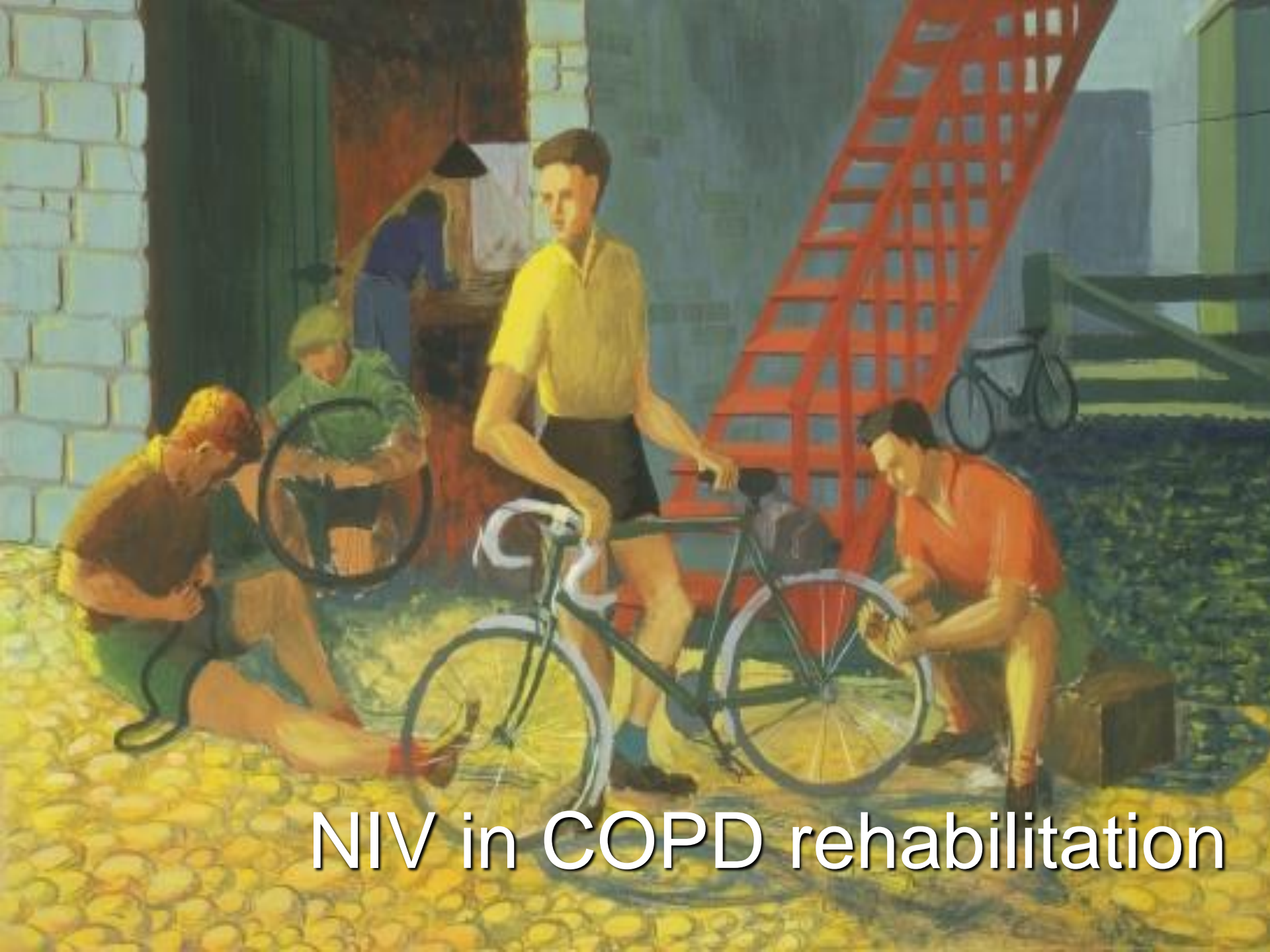
- Untreated overlap syndrome vs. COPD
  - Higher risk of mortality
  - Higher risk of hospitalisation for COPD exacerbation
- CPAP-treated overlaps have same mortality and exacerbation risk vs. COPD

*(Not RCT because of ethical constraints)*

# NIV in Overlap syndrome

CPAP improves survival and  
reduces hospitalisation for  
COPD/OSA

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NIV in COPD rehabilitation



# III. NIV for COPD rehabilitation

## NIV during exercise training

- Inspiratory pressure support improves ET and SOB [Keilty SE, et al. Thorax 1994]
- PAV+CPAP improves endurance time during cycling [Dolmage TE, et al. Chest 1997]
- IPS sustains exercise induced lactataemia (13.6 min vs. 5.5 min) – training effect? [Polkey MI et al. Thorax 2000]
- Systematic review: exertional dyspnoea and exercise endurance favour NIV [van't Hul A, et al. J Cardiopulm Rehab 2002]
- Long term benefit?

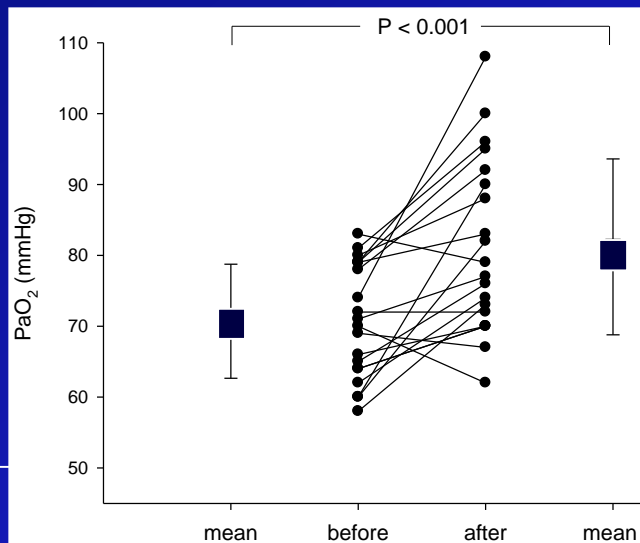
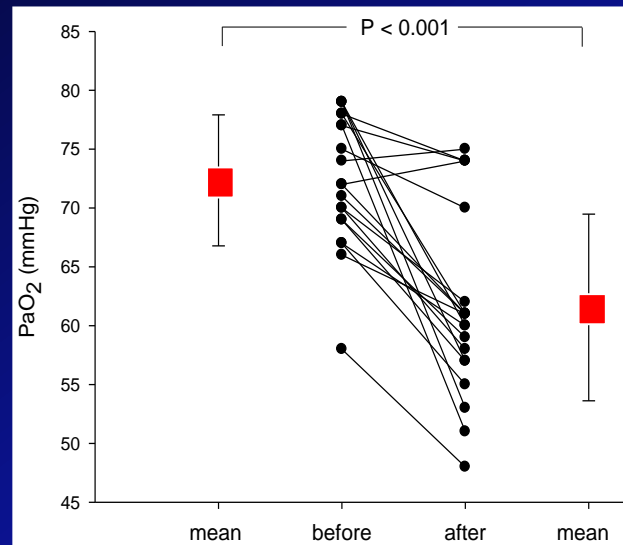
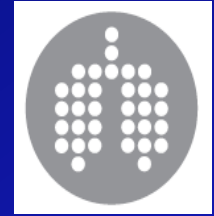
## Nocturnal NIV + PR

- NIV + PR improves fatigue, MRF score, cognition, PaCO<sub>2</sub>, daily step count in hypercapnic COPD [Duiverman ML et al. Thorax 2008]
- NIV + PR improves 6-min walk and longest distance walked, FEV<sub>1</sub>, ABG, SF-36 and hyperinflation in GOLD IV COPD [Köhnlein T et al. Resp Med 2009]



# Noninvasive ventilation during walking in patients with severe COPD: a randomised cross-over trial

M. Dreher, J.H. Storre and W. Windisch



50

P < 0.001

53

6

209

PaCO<sub>2</sub>  
[mmHg]

Borg  
Dyspnea  
Scale  
P < 0.001

Walking  
distance  
[m]  
P < 0.05

50

n.s.

51

4

252

# NIV in COPD rehabilitation

NIV maybe a useful adjunct in  
rehabilitation for severe COPD

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# NIV in palliative care?

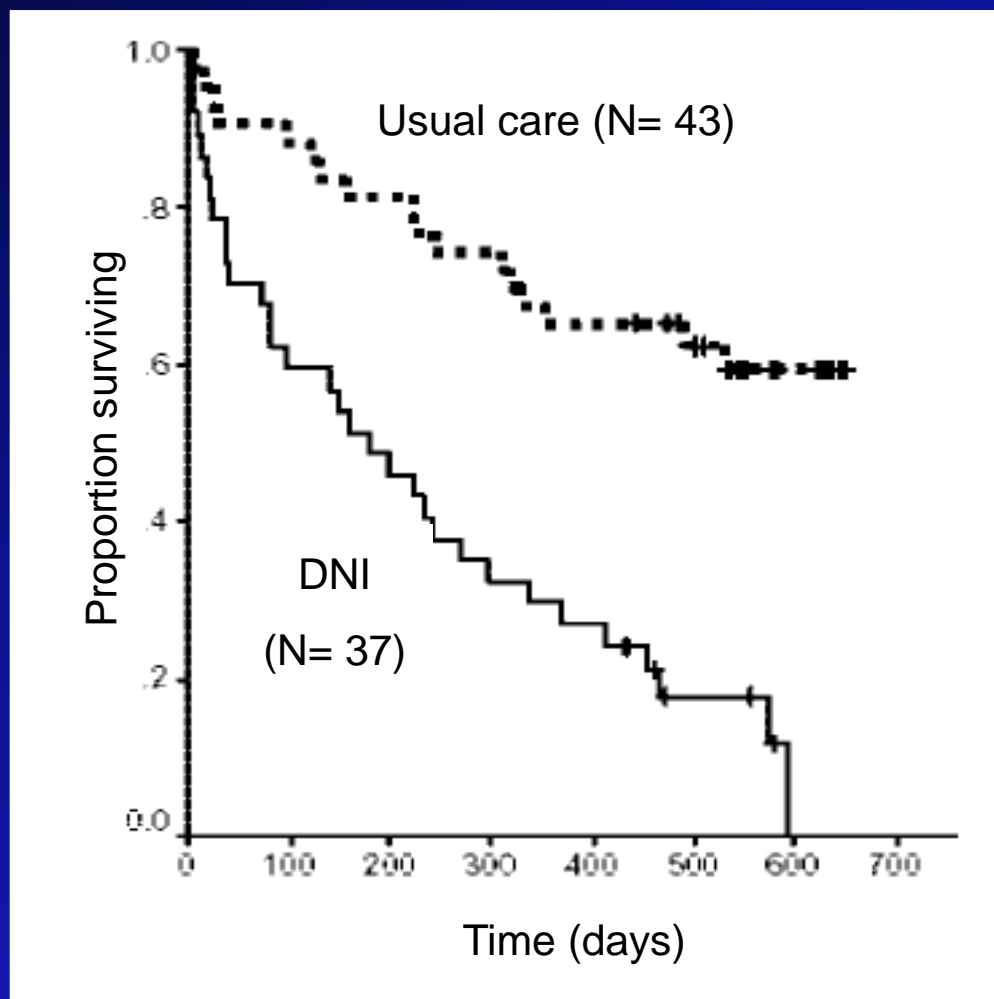
(Courtesy of Dr Jeff Ng, PC unit,  
Haven of Hope Hospital)



# Consideration near the end of life

Dimensions	Potential Benefits	Potential Burdens
Physical	<ol style="list-style-type: none"> <li>1. Prolong survival</li> <li>2. Relief in dyspnoea</li> </ol>	<ol style="list-style-type: none"> <li>1. Prolong the process of dying</li> <li>2. Discomfort over face (e.g. pressure narcosis)</li> <li>3. Building up of bronchial mucus</li> <li>4. Noise, especially from alarm of NIV machine</li> <li>5. Demanding in cooperation by patient</li> </ol>
Psychological		Causing anxiety (including claustrophobia)
Social	<ol style="list-style-type: none"> <li>1. Improvement in alertness in communication</li> <li>2. Buying time to say goodbye and to achieve "closure"</li> </ol>	<ol style="list-style-type: none"> <li>1. Hindrance in communication by the mask</li> <li>2. Discharge at home impossible without a ventilator</li> </ol>
Others		<ol style="list-style-type: none"> <li>1. Difficulty in withdrawal</li> <li>2. Training required for care-givers</li> <li>3. Cost</li> </ol>

# NIV for DNR COPD – poor survival



# Palliative use of NIV in COPD

No RCT evidence

Individualise

Need to define specific therapeutic goal



# Conclusions

AE-COPD with ARF	
Weaning from ET-MV	
Home NIV	
COPD/OSA	
Pulmonary Rehabilitation	
Palliative care	

# Conclusions

AE-COPD with ARF	Trial of NIV in majority
Weaning from ET-MV	
Home NIV	
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# Conclusions

AE-COPD with ARF	Trial of NIV in majority
Weaning from ET-MV	NIV may facilitate weaning in selected patients
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# Conclusions

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COPD/OSA	
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Pulmonary Rehabilitation	Useful adjunct Nocturnal use/Training use
Palliative care	



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COPD/OSA	CPAP/NIV reduced mortality/hospitalisation
Pulmonary Rehabilitation	Useful adjunct Nocturnal use/Training use
Palliative care	Individualised Burdensome

Thank you

