

RICS
Rotterdam, The Netherlands
April 5, 2023; 13:30–14:00 HH



Future of Automated Ventilation

patient vs. healthcare provider



Disclosures

- Xenios/Fresenius, Germany
- Hamilton Medical AG, Switzerland



Agenda

- history of ventilation
- history of automated ventilation
- benefits of automated ventilation
- need for automated ventilation?
- wrap-up



Agenda

- history of ventilation
- history of automated ventilation
- benefits of automated ventilation
- need for automated ventilation?
- wrap-up



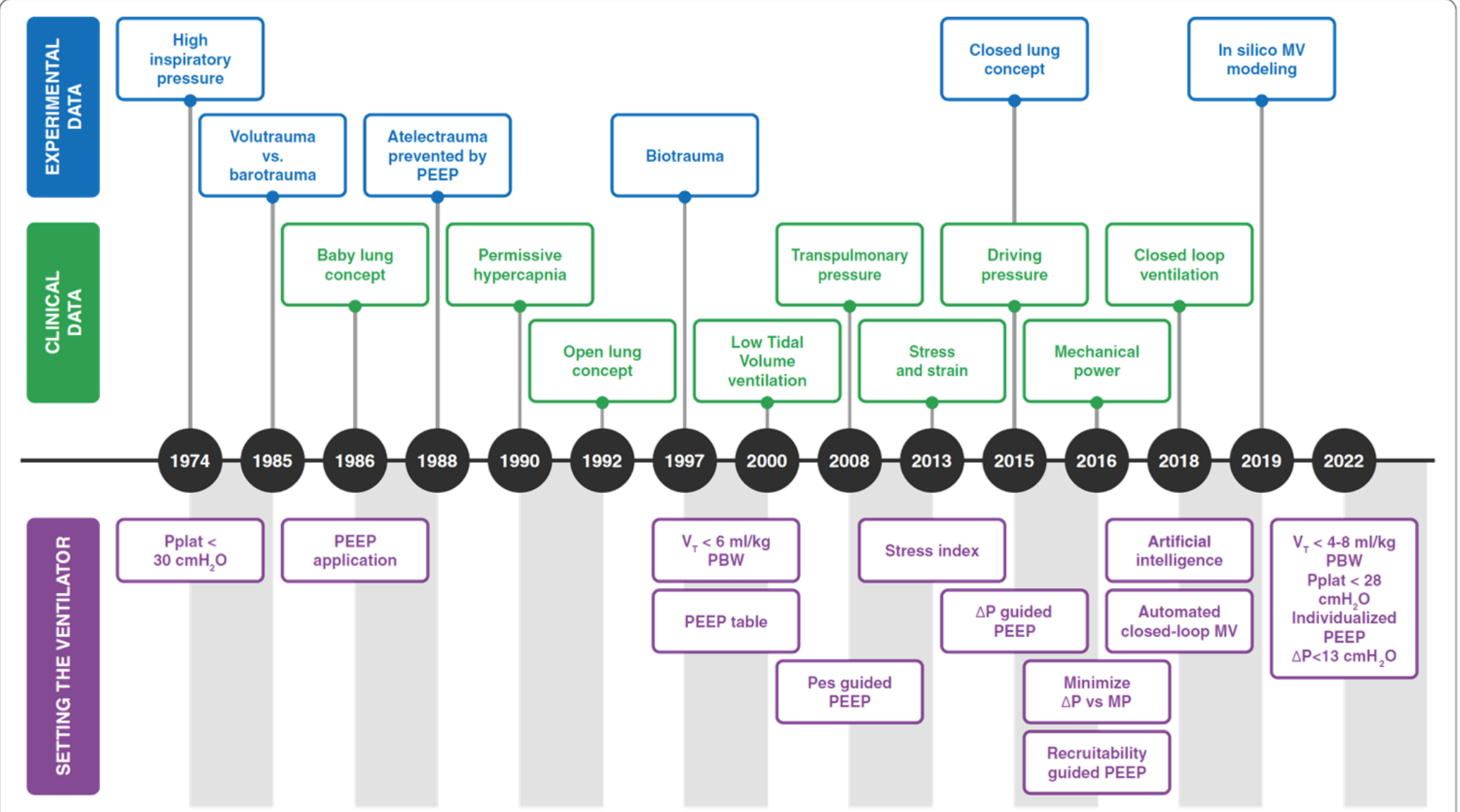


Fig. 1 Contributions over time to the knowledge of how to apply protective ventilation

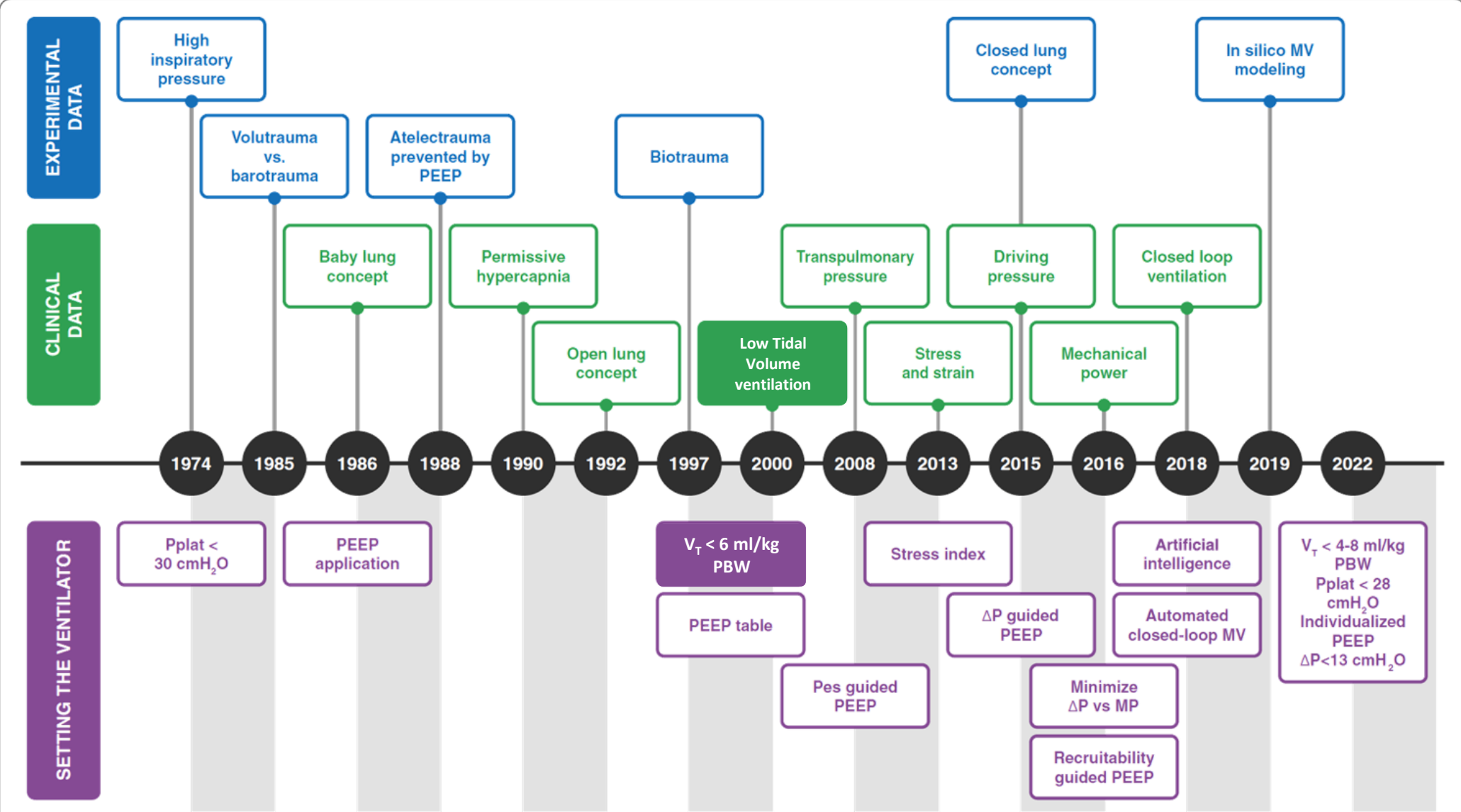


Fig. 1 Contributions over time to the knowledge of how to apply protective ventilation



QUESTION Does the use of a lower tidal volume (V_T) with mechanical ventilation affect important clinical outcomes in ARDS patients?

CONCLUSION Ventilation with a lower V_T than is traditionally used results in decreased mortality and increases the number of days without ventilator use.

POPULATION



344 Women **516 Men**

patients with ARDS

mean age: **52** years

LOCATION

ICUs in the
USA



VENTILATION STRATEGIES

861 patients with mild,
moderate or severe ARDS

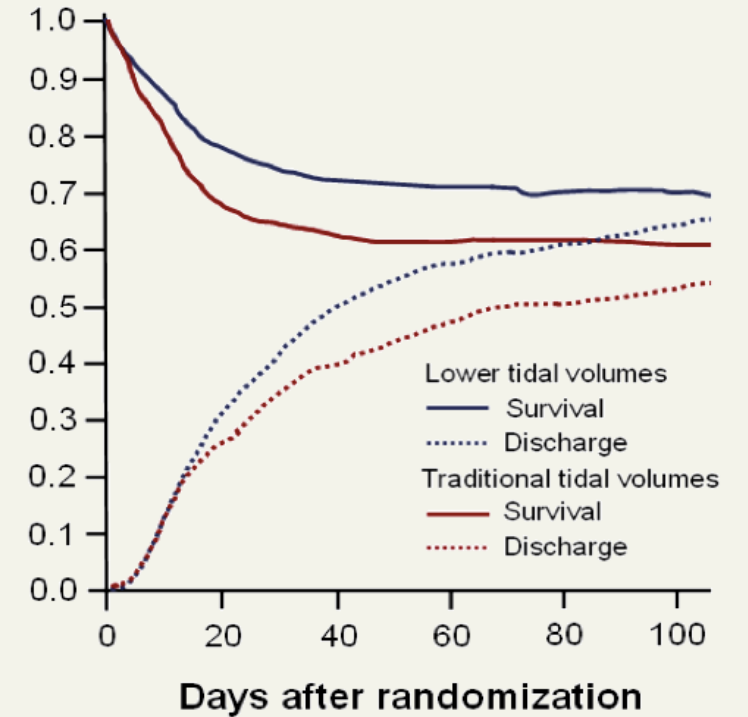
432 patients
lower V_T
(6 ml/kg)
[~ 400 to 500 ml]

429 patients
traditional V_T
(12 ml/kg)
[~1000 to 1200 ml]

(PRIMARY) OUTCOME

hospital mortality and duration of invasive
ventilation

FINDINGS



QUESTION For patients in the ICU who are ventilated for reasons other than ARDS, is low tidal volume superior to intermediate tidal volume?

CONCLUSION Among ICU patients receiving invasive ventilation, a strategy with a low tidal volume was not superior to using intermediate tidal volume.

POPULATION



621 Men 340 Women

ICU patients without ARDS expected to be intubated for more than 24 hours

Median age: 68 years (IQR, 59-76)

LOCATIONS

6 ICUs in the Netherlands



INTERVENTIONS

961 Patients randomized

477 Randomized
475 Analyzed

484 Randomized
480 Analyzed

Low tidal volume

Started at tidal volume of 6 mL/kg; tidal volume then decreased in steps of 1 mL/kg predicted body weight

Intermediate tidal volume

Started at tidal volume of 10 mL/kg; if plateau pressure exceeded 25 cm H₂O, tidal volume was decreased in steps of 1 mL/kg predicted body weight

PRIMARY OUTCOME

Number of ventilator-free days and alive at day 28

FINDINGS



QUESTION What is the impact of mechanical power on mortality in patients with ARDS as compared with that of primary ventilator variables such as the ΔP , V_T , and RR?

CONCLUSION Mechanical power was associated with mortality during controlled mechanical ventilation in ARDS, but a simpler model using only the ΔP and RR was equivalent.

POPULATION



1728 Women 2821 Men

Patients with ARDS

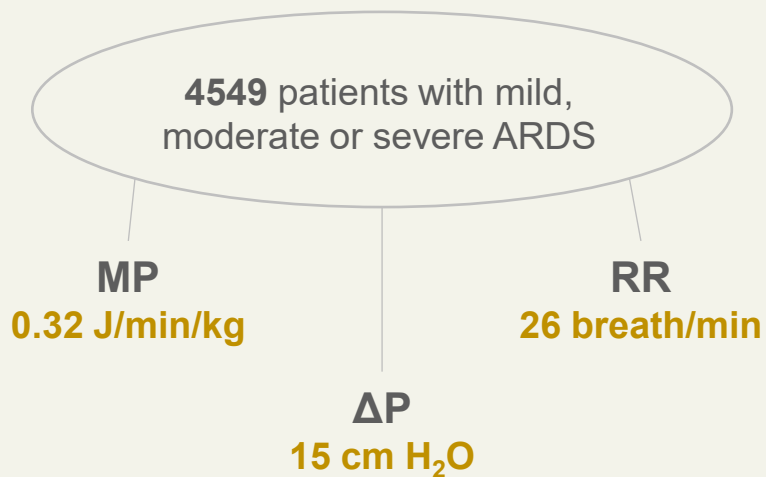
Mean Age: 55 years

LOCATION

6 RCTs and 1 observational study



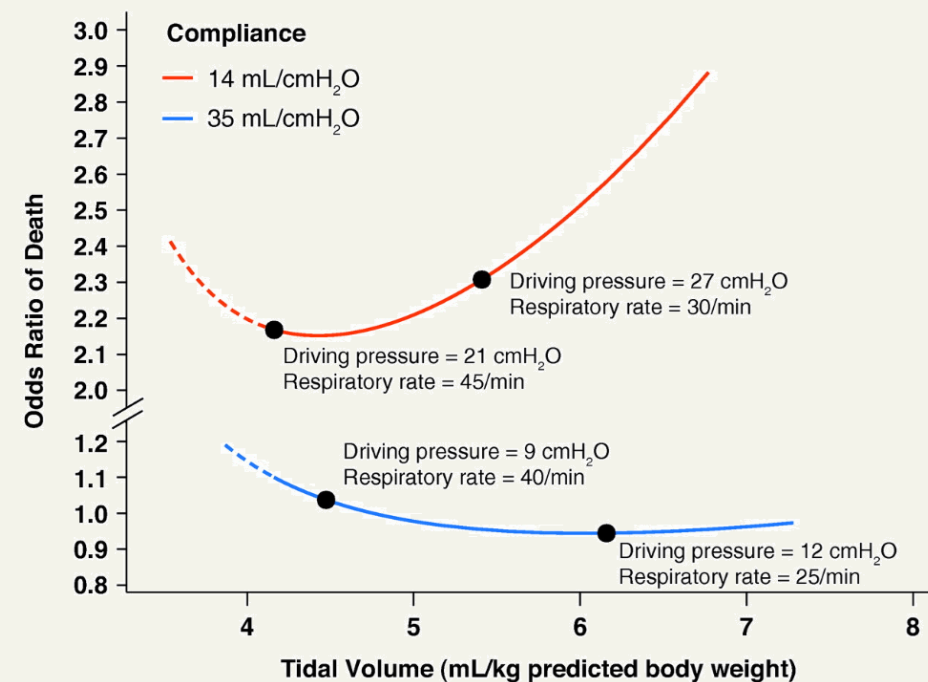
VENTILATION PARAMETERS



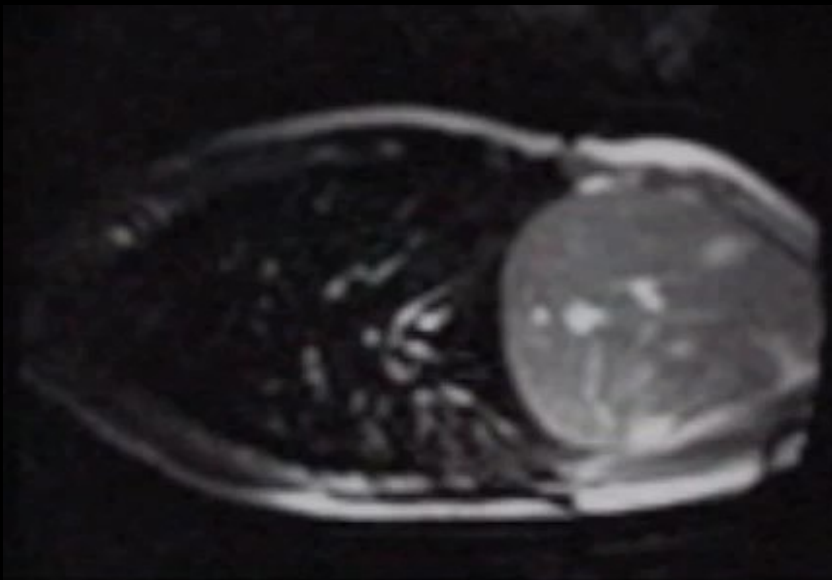
(PRIMARY) OUTCOME

Mortality at 28 or 60 days

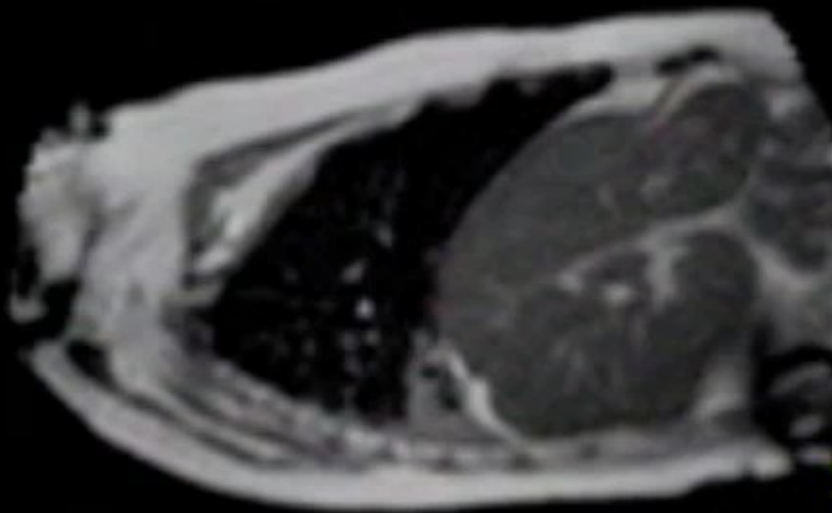
FINDINGS



ACTIVE



PASSIVE



during spontaneous vs non-spontaneous ventilation, note:

- the position of diaphragm
- the aerated lung size
- presence of atelectasis
- presence of overdistension

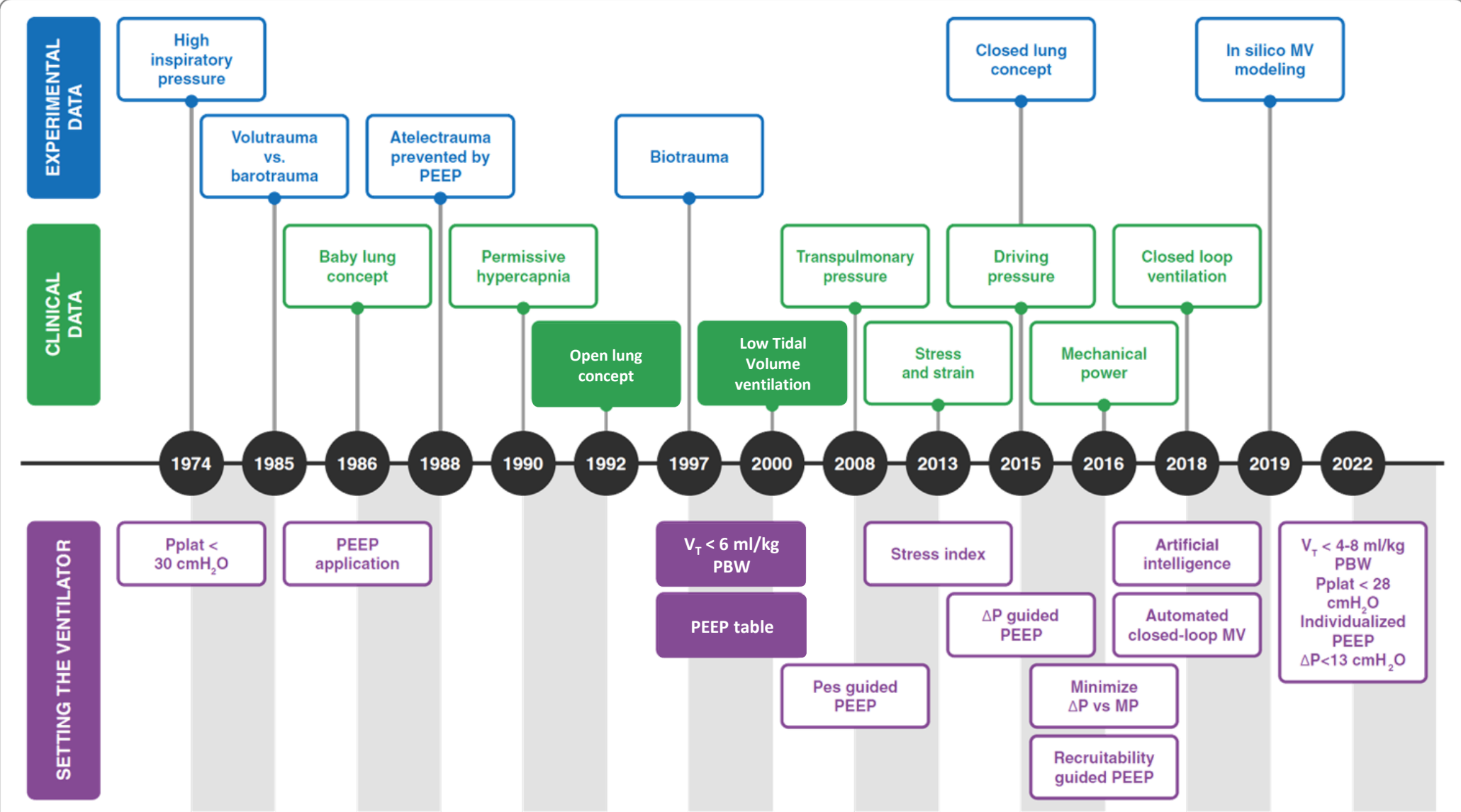


Fig. 1 Contributions over time to the knowledge of how to apply protective ventilation

QUESTION Does use of a lung recruitment maneuver associated with PEEP titration according to the best respiratory-system compliance reduce 28-day mortality of patients with moderate to severe ARDS, compared with a conventional low-PEEP strategy?

CONCLUSION A strategy using a lung recruitment maneuver and titrated PEEP increased mortality of patients with moderate to severe ARDS.

POPULATION



379 Women 631 Men

consecutive patients with moderate to severe ARDS

Mean Age: 51 years

LOCATION

120 ICUs from 9 countries



VENTILATION STRATEGIES

1010 patients with moderate or severe ARDS

501 patients titrated (high) PEEP [>15 cm H₂O] with RM

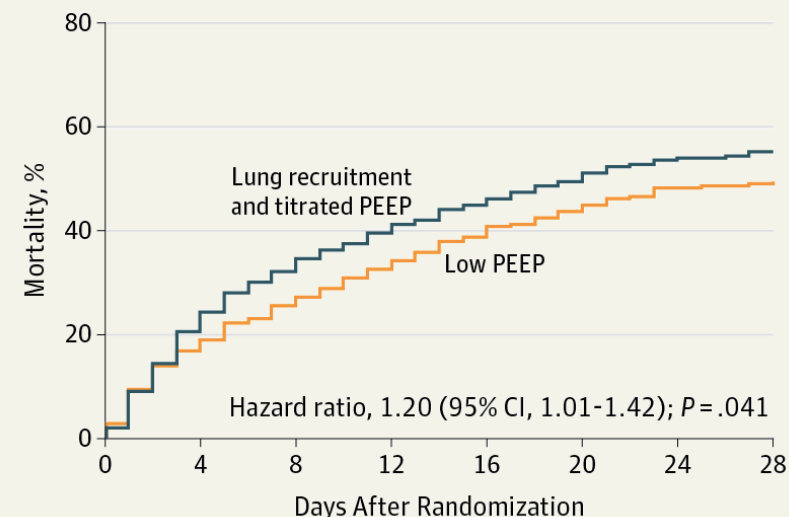
509 patients standard (low) PEEP [< 12 cm H₂O] without RM

(PRIMARY) OUTCOME

28-day mortality; length of ICU and hospital stay; VFD28; pneumothorax requiring drainage or barotrauma within 7 days

FINDINGS

better oxygenation, but worse outcomes, and more pneumothorax and barotrauma with high PEEP



QUESTION Is there heterogeneity in treatment effects in patients enrolled in the ART, using a machine learning approach?

CONCLUSION Recruitment maneuvers and titrated PEEP may be harmful in ARDS patients with pneumonia or requiring vasopressor support. Driving pressure appears to modulate the association between the ART study intervention, etiology of ARDS, and mortality.

POPULATION



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consecutive patients with moderate to severe ARDS

Mean Age: **51** years

LOCATION

120 ICUs
from **9** countries



VENTILATION STRATEGIES

1010 patients with moderate or severe ARDS

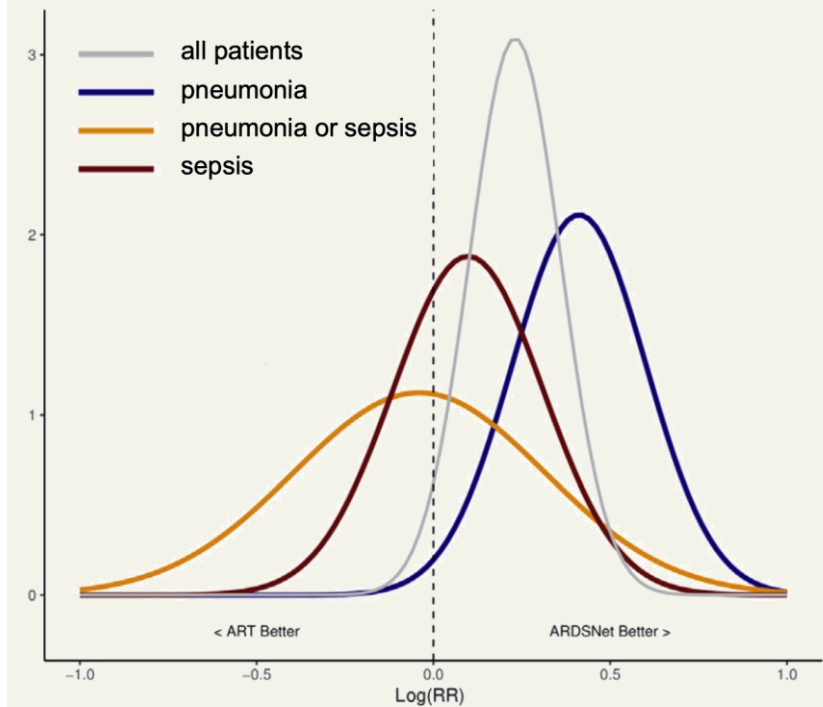
501 patients
titrated (high) PEEP
[>15 cm H₂O] with RM

509 patients
standard (low) PEEP
[< 12 cm H₂O] without RM

OUTCOME OF THE BAYESIAN ANALYSIS

28-day mortality

FINDINGS



QUESTION Does a mechanical ventilation strategy that is personalized to individual patients' lung morphology improve the survival of patients with ARDS when compared with standard of care?

CONCLUSION Personalization of ventilation decreased mortality in patients with ARDS [in the posthoc analysis]; a ventilator strategy misaligned with lung morphology substantially increases mortality.

POPULATION



114 Women 286 Men

patients with ARDS for less than 12 hours

Median Age: 62 years

LOCATION

20 ICUs in France



INTERVENTION

400 patients with moderate to severe ARDS

204
standard care

V_T 6 ml/kg PBW
low PEEP

196
personalized care

non-focal

V_T 6 ml/kg PBW
high PEEP
with RM

focal

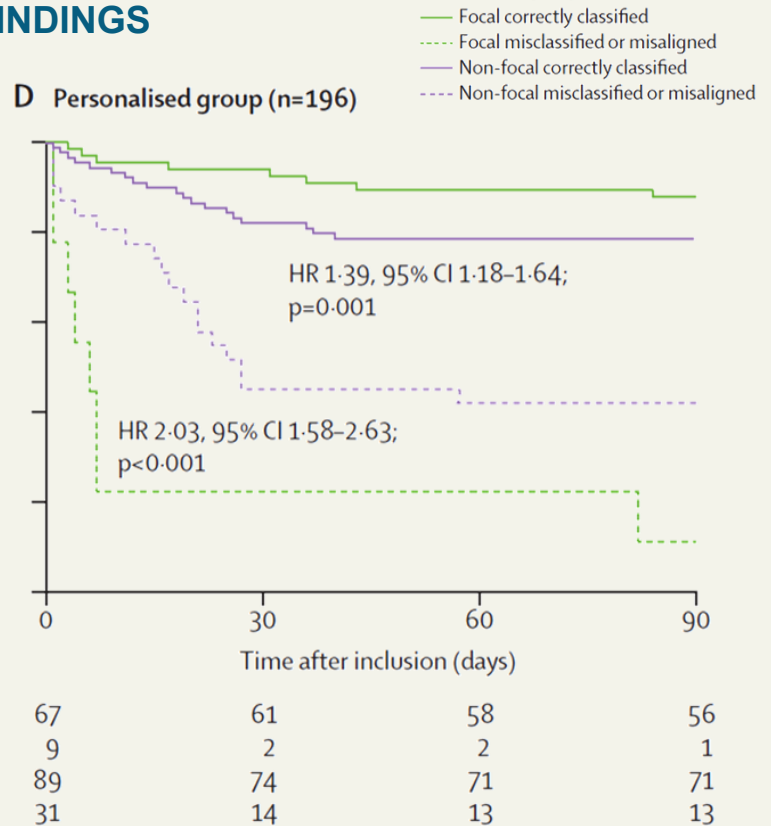
V_T 8 ml/kg PBW
low PEEP and
prone positioning

(PRIMARY) OUTCOME

mortality at day 90; ventilator-free days, ARDS resolution; LOS in ICU; barotrauma

FINDINGS

D Personalised group (n=196)



QUESTION In ICU patients who received invasive ventilation for reasons other than acute respiratory distress syndrome (ARDS), is a strategy with lower positive end-expiratory pressure (PEEP) noninferior to higher PEEP with respect to ventilator-free days at day 28?

CONCLUSION This clinical trial found that among ICU patients receiving invasive ventilation, a strategy with lower PEEP was noninferior to a strategy using higher PEEP for the outcome of ventilator-free days, supporting the use of lower PEEP in patients without ARDS.

POPULATION



623 Men 346 Women

Adults without ARDS expected not to be extubated within 24 hours of intubation

Median age: 66 years

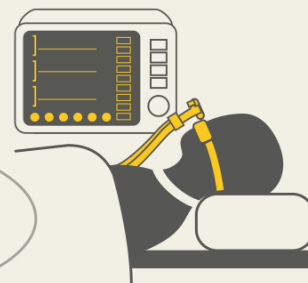
LOCATIONS

8 ICUs in the Netherlands



INTERVENTION

980 Patients randomized
969 Patients analyzed

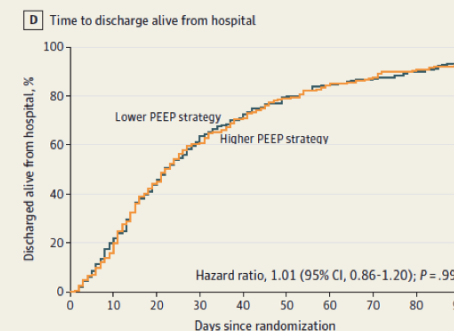
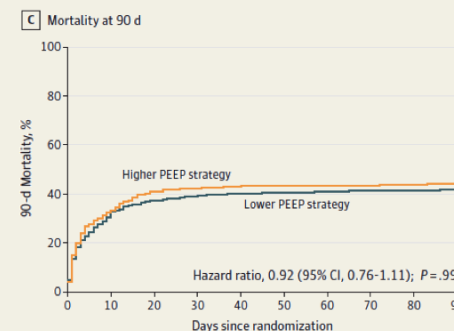
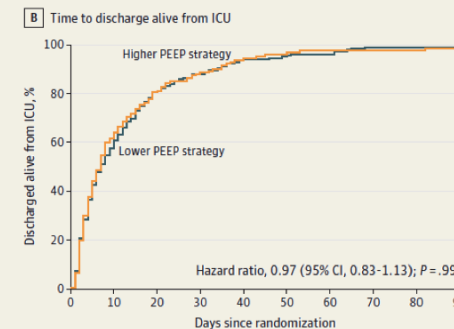
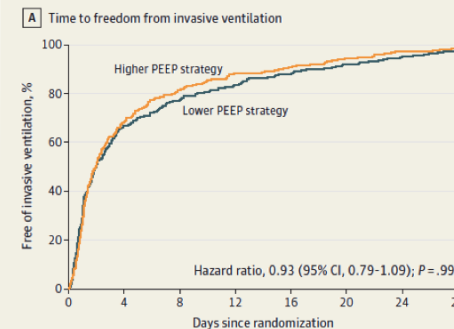


476 Lower PEEP strategy
Lowest level between 0-5 cm H₂O

493 Higher PEEP strategy
Lowest level of 8 cm H₂O

PRIMARY OUTCOME

Number of ventilator-free days by study day 28 (noninferiority margin of -10%)



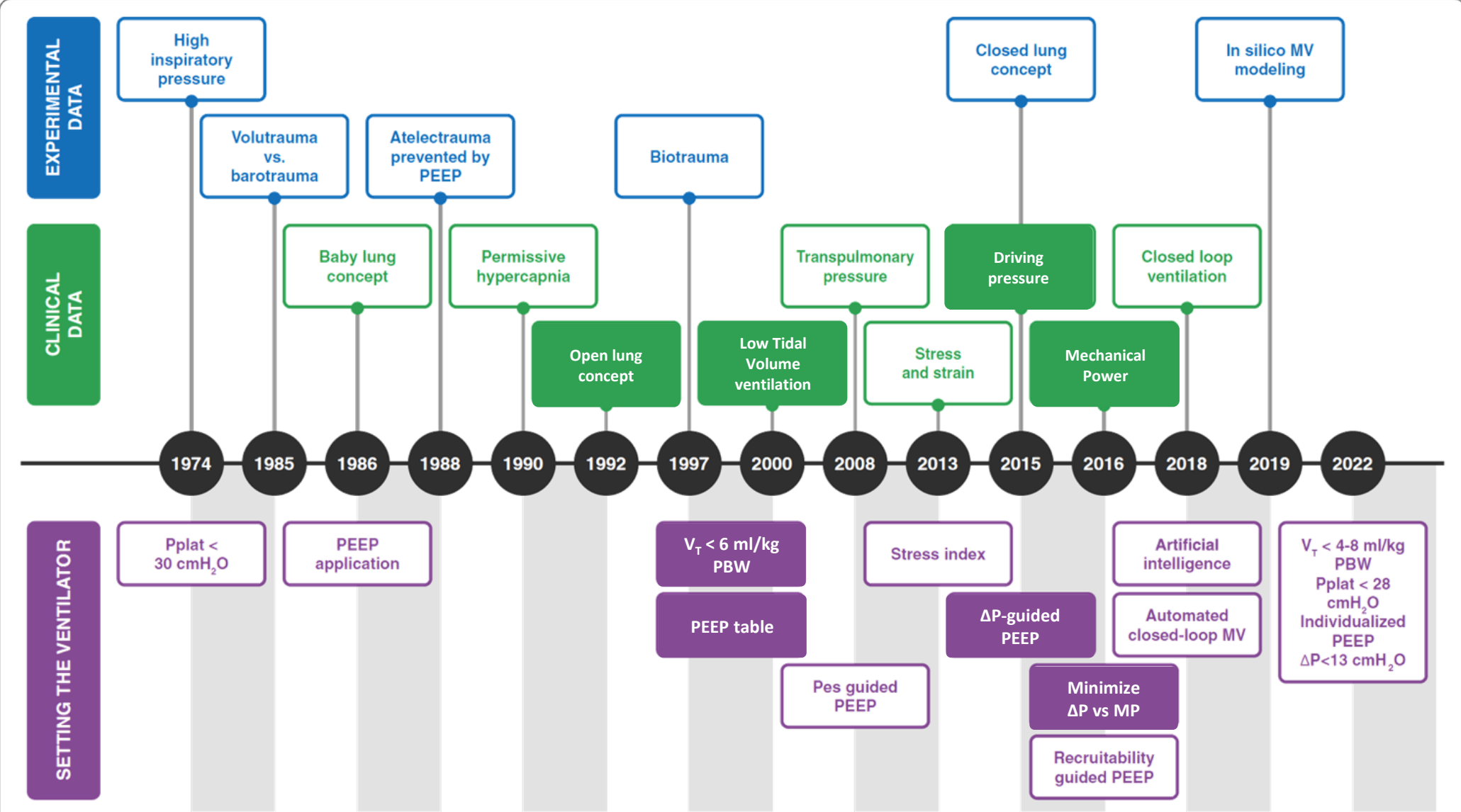


Fig. 1 Contributions over time to the knowledge of how to apply protective ventilation



QUESTION Is ΔP an index more strongly associated with survival than V_T or PEEP in patients who are not actively breathing?

CONCLUSION ΔP is the ventilation variable that best stratified risk; decreases in ΔP owing to changes in ventilator settings may be strongly associated with increased survival.

POPULATION



~40% Women ~60% Men

patients with ARDS included in RCTs

Mean Age: from 34 to 60 years

SOURCE

9 trials worldwide



METHODS

multilevel mediation analysis of individual patient data from 3562 patients

prediction model
univariate
multivariate

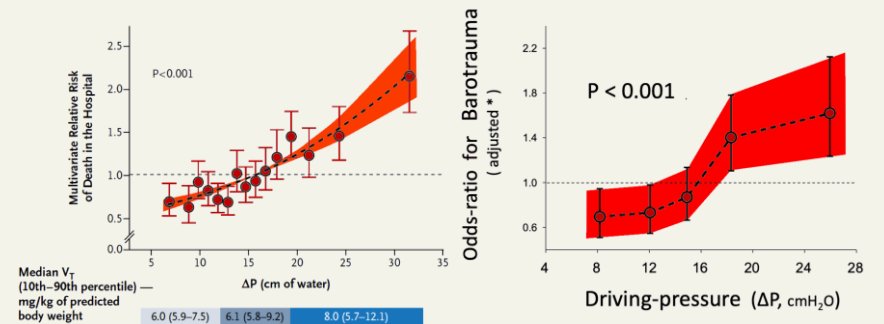
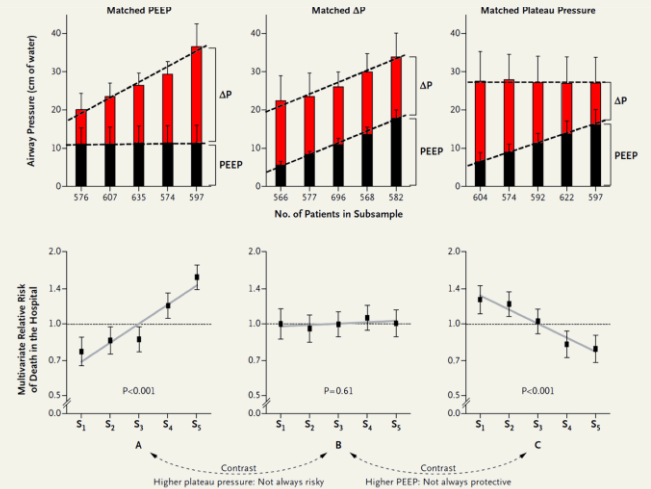
mediation analysis

risk priority of ΔP , V_T and PEEP

(PRIMARY) OUTCOME

60-day mortality

FINDINGS



QUESTION What is the association between exposure to different intensities of mechanical ventilation over time and intensive care unit (ICU) mortality in patients with acute respiratory failure?

CONCLUSION Cumulative exposure to higher intensities of mechanical ventilation was harmful, even for short durations.

POPULATION



5141 Women 8267 Men

patients receiving ventilation
for 4 hours or more

Median Age: 62 years

LOCATION

9 ICUs in
Toronto, Canada

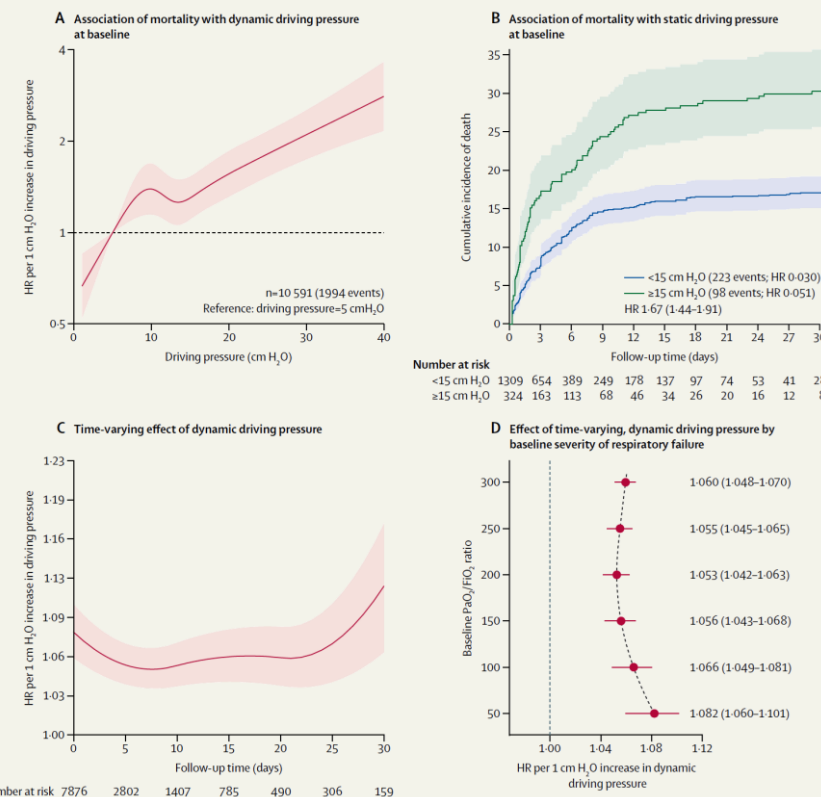


FINDINGS

	Exposure to high driving pressure		Exposure to high mechanical power	
	HR estimate (95% CrI)	p value	HR estimate (95% CrI)	p value
Baseline variables				
PaO ₂ /FiO ₂ , mm Hg	0.945 (0.896–0.994)	0.026	0.977 (0.930–1.031)	0.38
Age, years	1.108 (1.048–1.160)	<0.0001	1.128 (1.080–1.182)	<0.0001
APACHE III score	1.602 (1.526–1.680)	<0.0001	1.591 (1.524–1.669)	<0.0001
APACHE pH	0.832 (0.809–0.859)	<0.0001	0.840 (0.820–0.864)	<0.0001
Time-varying variables				
Days with driving pressure ≥15 cm H ₂ O	1.049 (1.023–1.076)	<0.0001
Days with mechanical power ≥17 J/min	1.069 (1.047–1.092)	<0.0001

1622 (20.6%) of 7876 patients died; 64 281 daily observations were recorded. HRs were the adjusted HRs associated with a 1-SD increment in the given variable. Values higher than 1 indicate increased mortality. The values used for SDs were as follows: PaO₂/FiO₂ ratio 119; pH 0.11; age 17 years; and APACHE III score 29. The effects of the number of days with either driving pressure greater than or equal to 15 cm H₂O or mechanical power greater than or equal to 17 J/min were estimated using Quasi-Poisson models in the joint model analyses. HR=hazard ratio. CrI=credible interval. PaO₂=partial pressure of oxygen. FiO₂=fraction of inspired oxygen. APACHE=Acute Physiology and Chronic Health Evaluation.

Table 3: Cumulative effect on HRs of exposure to high intensities of mechanical ventilation for 7876 patients with available data



QUESTION Does the intensity of ventilation, reflected by the mechanical power of ventilation (MP), has an association with outcome in invasively ventilated patients without ARDS.

CONCLUSION In ICU patients without ARDS, MP has an independent association with mortality. This finding suggest that MP holds an added predictive value over its individual components, making MP an attractive parameter to monitor and target in these patients.

POPULATION



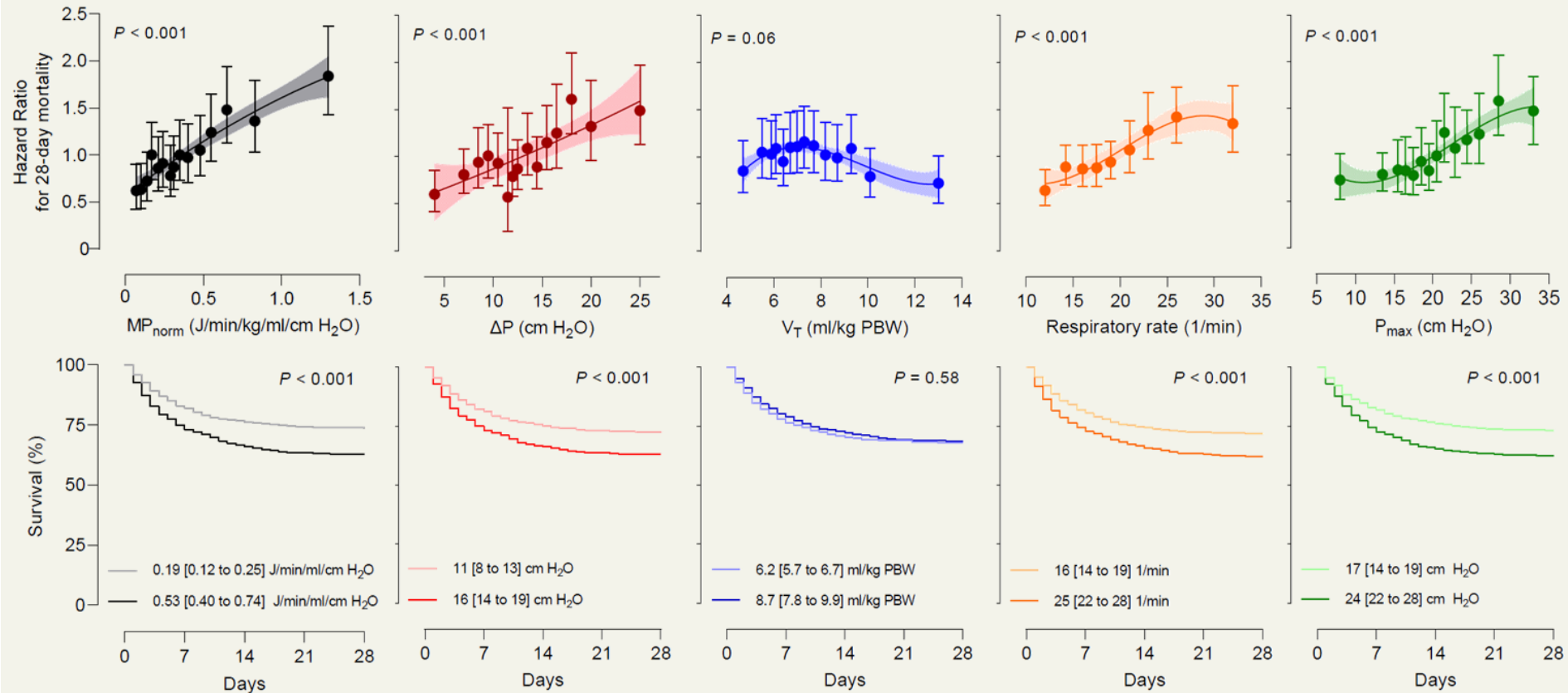
1962 Patients

ICU patients without ARDS, expected to need invasive ventilation > 24 hours

Median Age: 67 years

LOCATION

8 ICUs in the Netherlands



Agenda

- history of ventilation
- history of automated ventilation
- benefits of automated ventilation
- need for automated ventilation?
- wrap-up



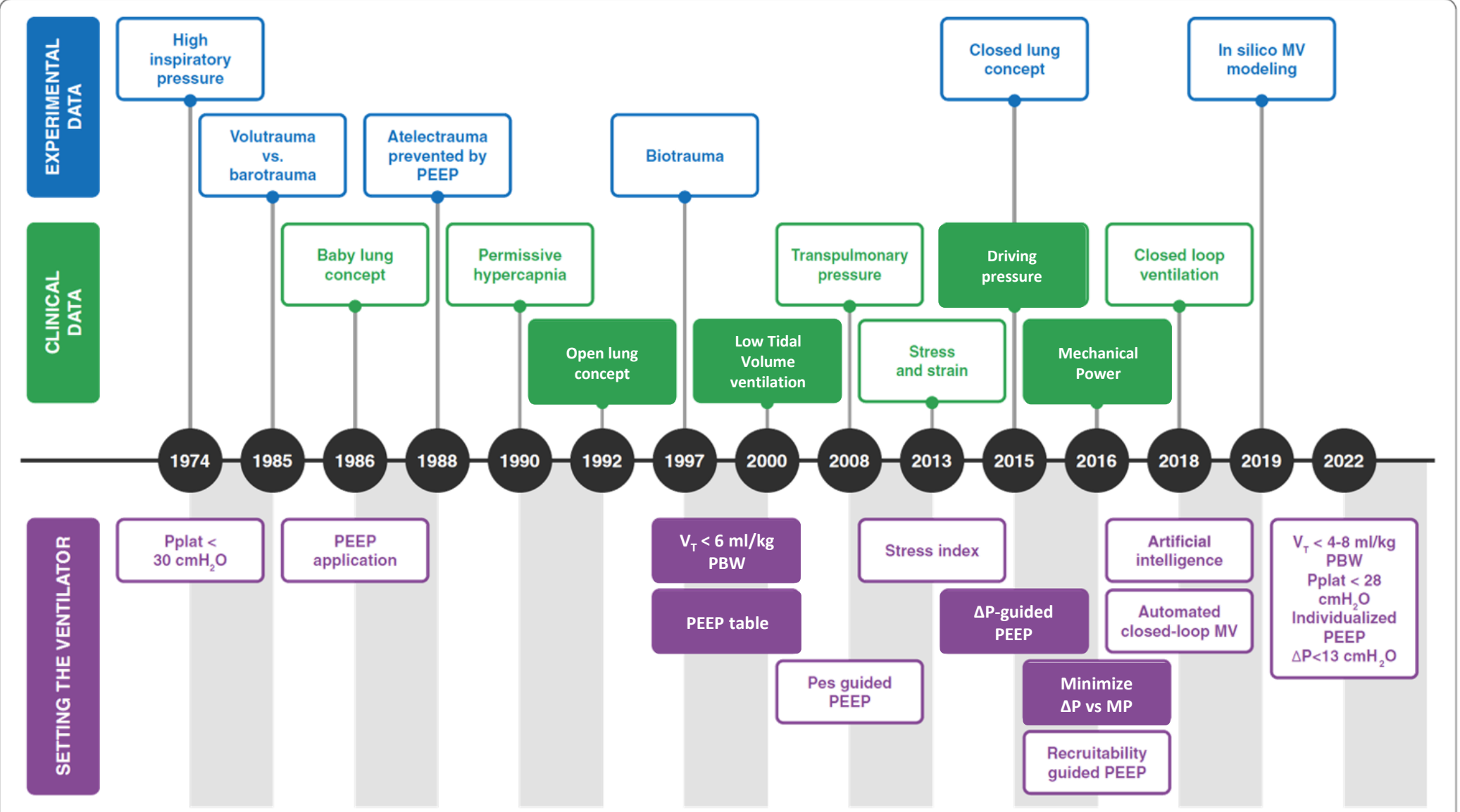


Fig. 1 Contributions over time to the knowledge of how to apply protective ventilation

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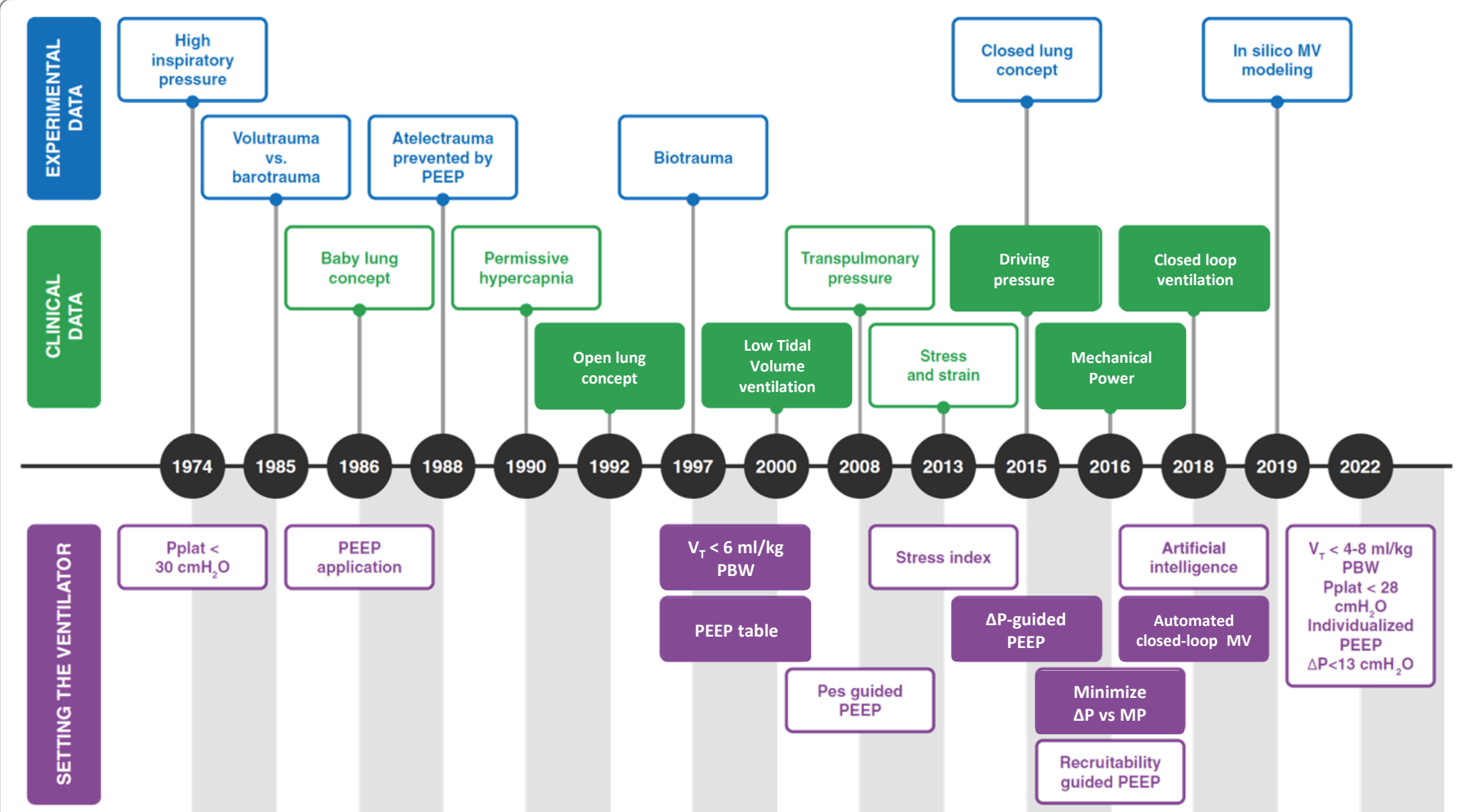
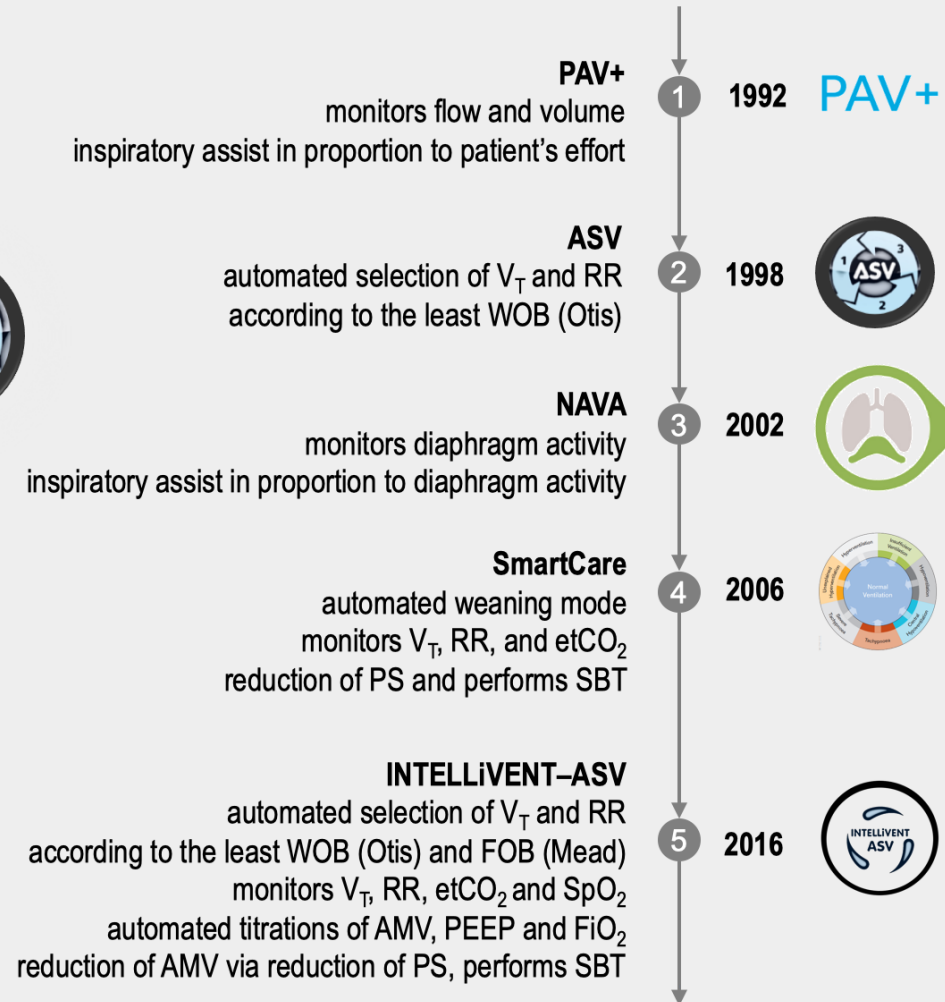


Fig. 1 Contributions over time to the knowledge of how to apply protective ventilation

A History of Automated Ventilation

Despite years of research in mechanical ventilation many settings remained to be set by hand
Here is when the ventilators became smarter



QUESTION Is the amount of mechanical power of ventilation (MP) under adaptive support ventilation (ASV) different from that under nonautomated pressure-controlled ventilation?

CONCLUSION This study suggests ASV may have benefits compared with pressure-controlled ventilation with respect to the MP transferred from the ventilator to the respiratory system in passive invasively ventilated critically ill patients.

POPULATION



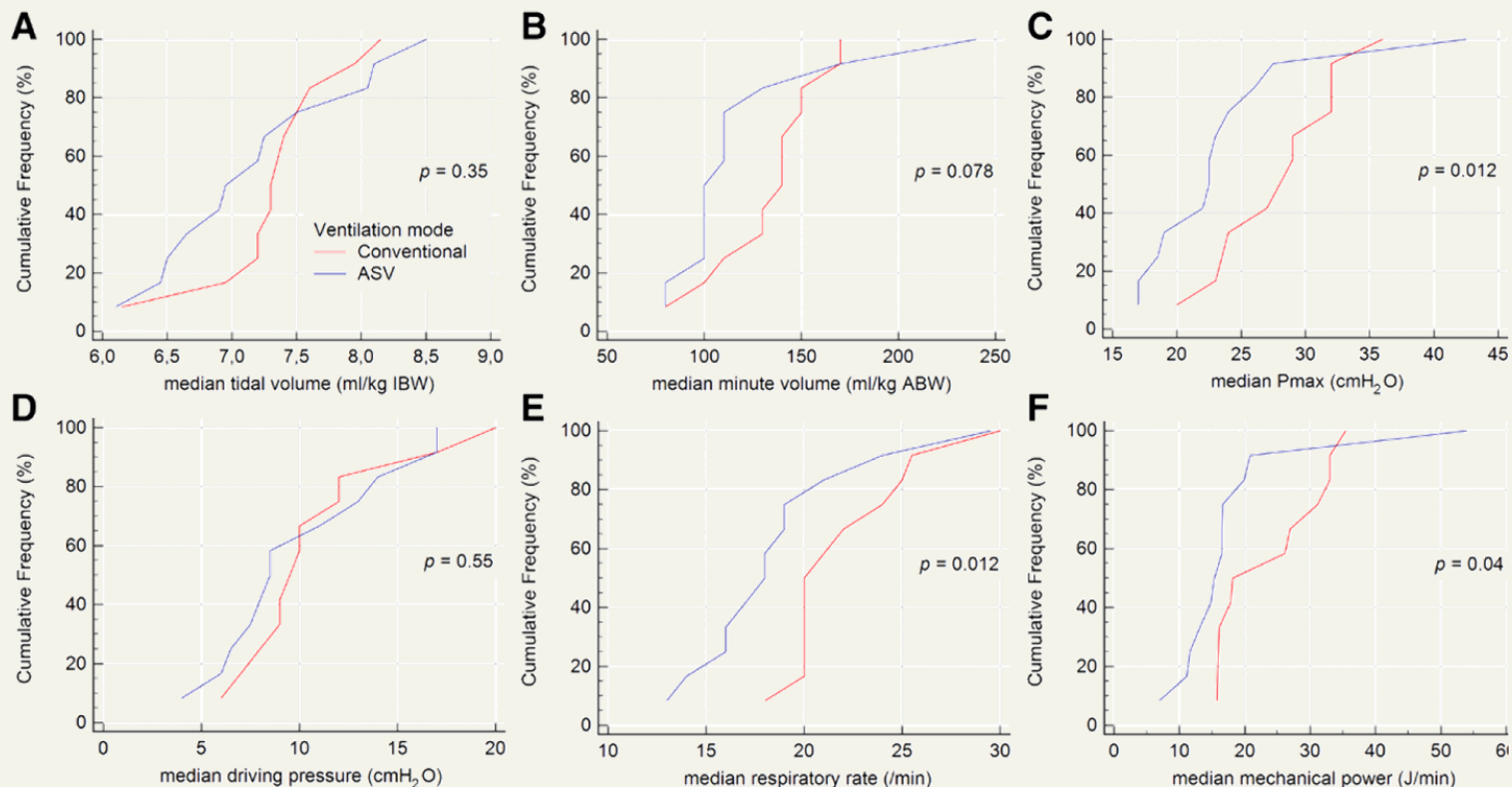
7 Women 15 Men

patients expected to need invasive ventilation for the next 24 hours

Median Age: 67 years

LOCATION

1 ICUs in The Netherlands



QUESTION In patients receiving post-operative ventilation after cardiac surgery, does INTELLiVENT-ASV improve the quality of breathing compared with conventional ventilation?

CONCLUSION Fully automated ventilation in patients after cardiac surgery optimized lung-protective ventilation during postoperative ventilation, with fewer episodes of severe hypoxaemia and an accelerated resumption of spontaneous breathing.

POPULATION



67 Women 153 Men

patients after uncomplicated cardiac surgery

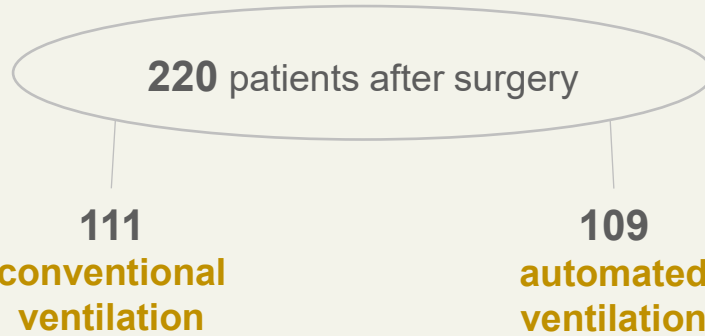
Median Age: 62–76 years

LOCATION

1 ICU in the Netherlands



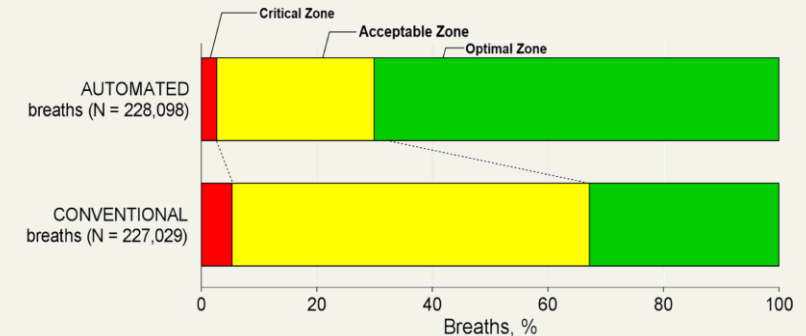
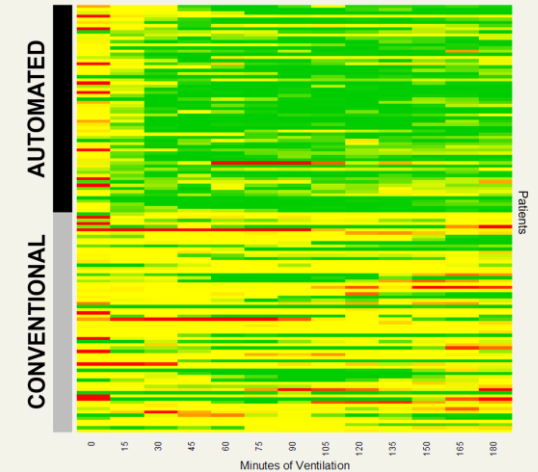
INTERVENTION



(PRIMARY) OUTCOME

proportion of breath within predefined optimal, acceptable, and critical ranges (V_T , P_{max} , SpO_2 and $etCO_2$); severe hypoxaemia ($SpO_2 < 85\%$) and resumption of spontaneous breathing

FINDINGS



QUESTION In COVID-19 patients with ARDS, does INTELLiVENT-ASV reduce the driving pressure and mechanical power of ventilation compared with conventional ventilation?

CONCLUSION INTELLiVENT-ASV reduces the intensity of ventilation in COVID-19 patients with ARDS.

POPULATION



12 Women 39 Men

COVID-19 with moderate to severe ARDS

Median Age: 63 years

LOCATION

2 ICUs in the Netherlands



INTERVENTION

51 patients intubated in the ICU for acute hypoxemia



conventional ventilation

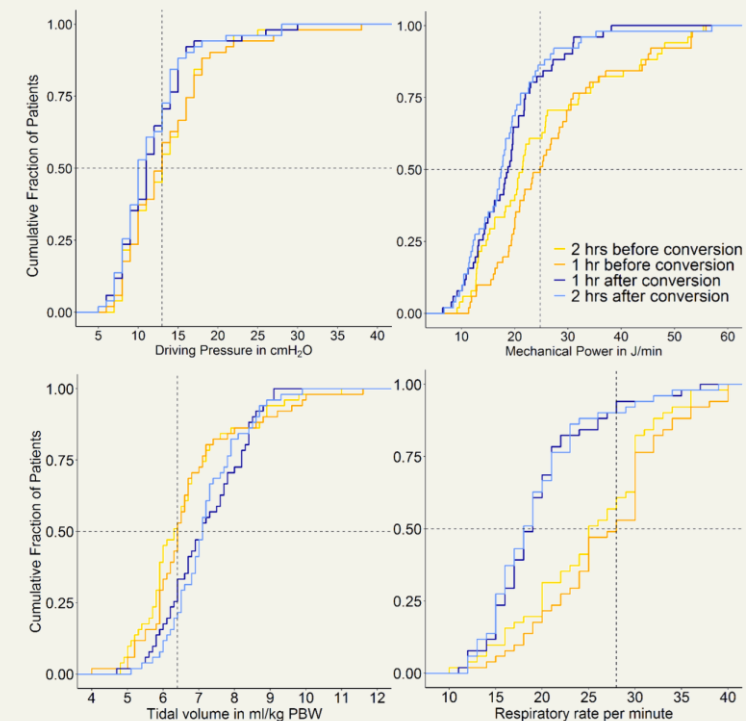
crossover

automated ventilation

(PRIMARY) OUTCOME

ΔP and MP before and after converting from conventional ventilation to INTELLiVENT-ASV

FINDINGS





QUESTION In COVID-19 patients with ARDS, does INTELLiVENT-ASV reduce the driving pressure and mechanical power of ventilation compared with conventional ventilation?

CONCLUSION INTELLiVENT-ASV reduces the intensity of ventilation in COVID-19 patients with ARDS.

POPULATION



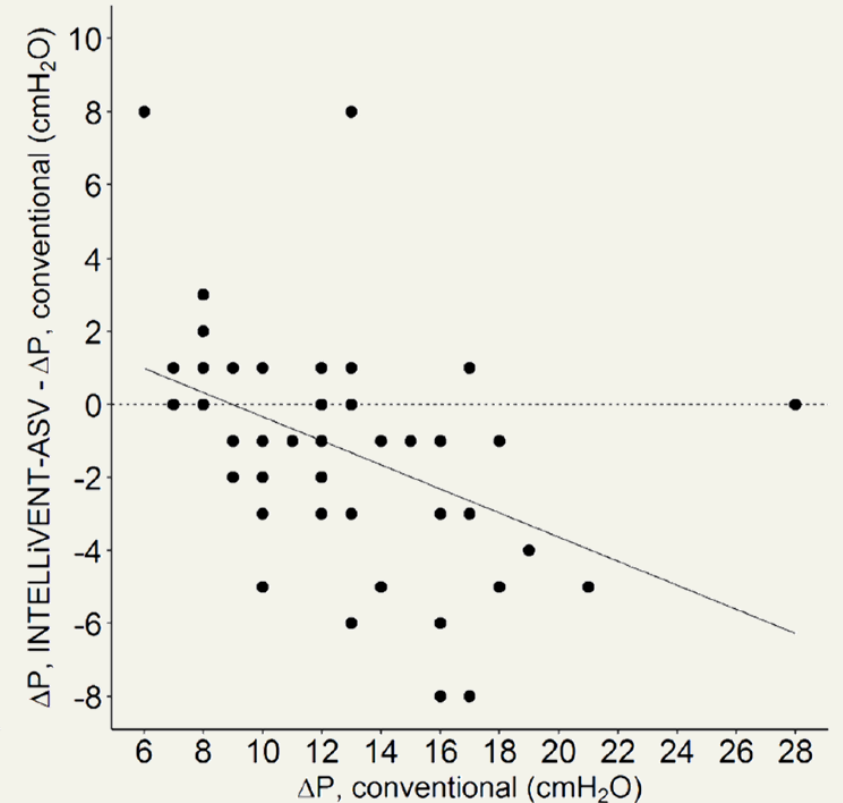
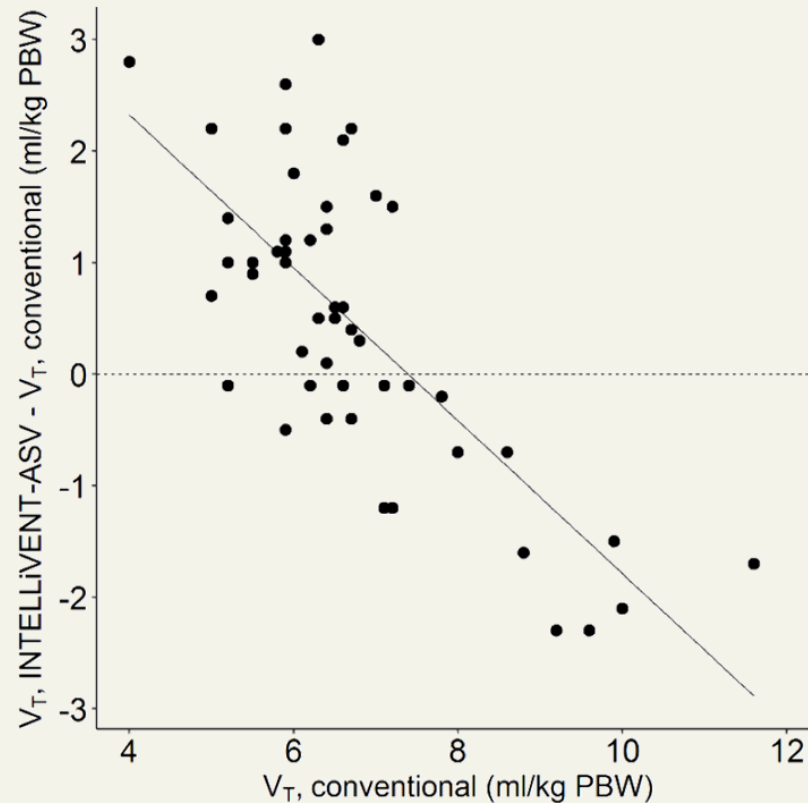
12 Women 39 Men

COVID-19 with moderate to severe ARDS

Median Age: 63 years

LOCATION

2 ICUs in the Netherlands



QUESTION What is the effect of automated closed-loop oxygen control, compared to automated ventilation with manual oxygen titrations, on time spent in predefined pulse oximetry (SpO₂) zones in pediatric critically ill patients?

CONCLUSION In this randomized crossover trial in pediatric critically ill patients under invasive ventilation with ASV, the percentage of time spent within in optimal SpO₂ zones increased with the use of closed-loop oxygen control.

POPULATION

37 Pediatric Patients



children with or without ARDS under invasive ventilation

Median Age: 1 year

LOCATION

1 ICU in Turkey



INTERVENTION

37 patients under ASV for acute hypoxemia

2 hours manual or automated FiO₂ control

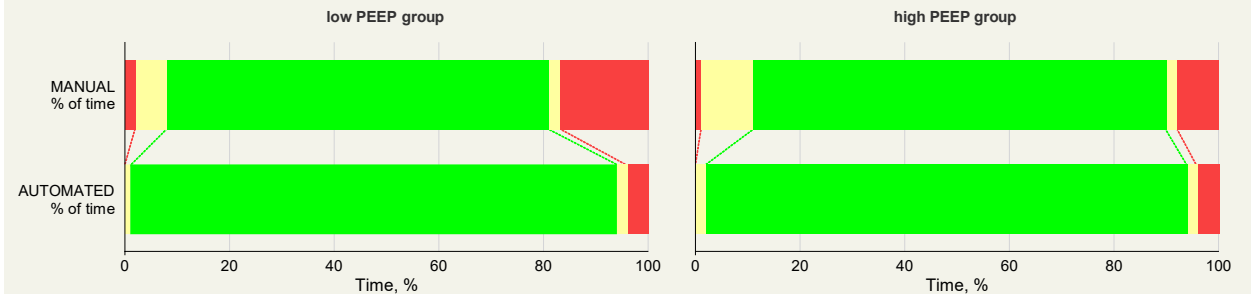
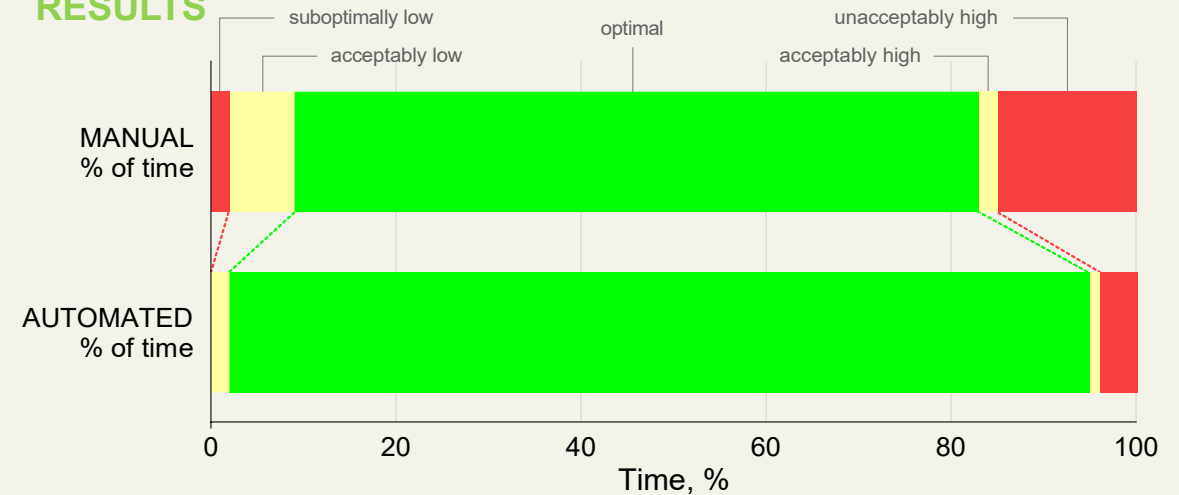
crossover

2 hours automated or manual FiO₂ control

(PRIMARY) OUTCOME

FiO₂ settings and SpO₂ readings

RESULTS



QUESTION What is the efficacy of a closed-loop oxygen control in critically ill patients with moderate to severe acute hypoxemic respiratory failure (AHRF) treated with high flow nasal oxygen (HFNO).

CONCLUSION Closed-loop oxygen control improves oxygen administration in patients with moderate-to-severe AHRF treated with HFNO, increasing the percentage of time in the optimal oxygenation range and decreasing the workload of healthcare personnel.

POPULATION

45 patients under HFNO



patients with moderate to severe ARF, including patients with COVID-19

Median Age: 49 year

LOCATION

1 ICU in Spain



INTERVENTION

45 patients under HFNO

4 hours manual or automated FiO₂ control

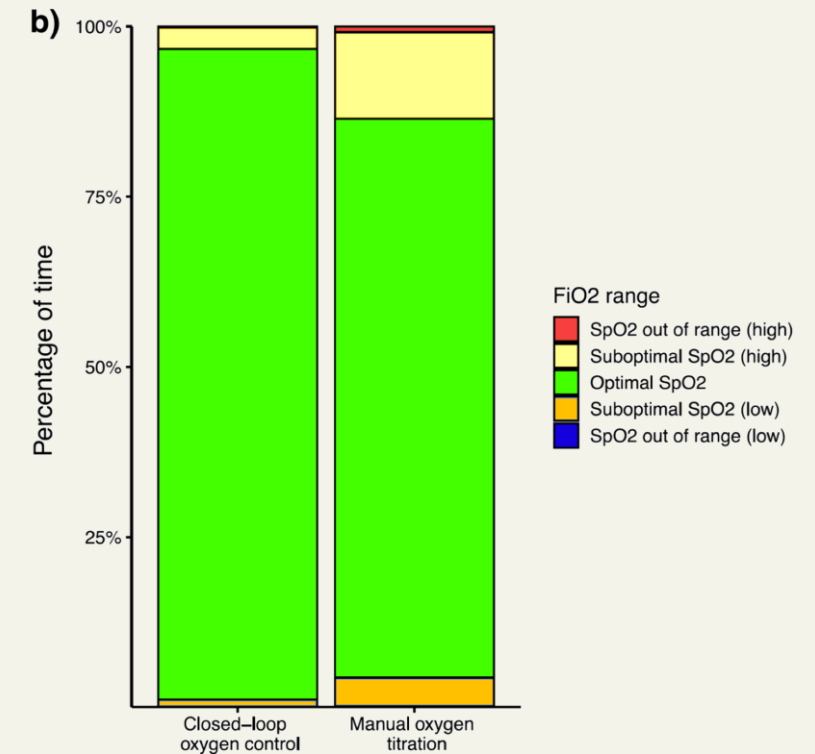
crossover

4 hours automated or manual FiO₂ control

(PRIMARY) OUTCOME

percentage of time spent in the individualized optimal SpO₂ ranges

RESULTS



QUESTION What is the effect of HFNO with closed-loop control of the fraction of inspired oxygen (FiO₂), compared to HFNO with manual titrations of the FiO₂, on time spent in predefined pulse oximetry (SpO₂) zones in pediatric critically ill patients?

CONCLUSION In this randomized crossover trial in pediatric critically ill patients under HFNO, the percentage of time spent within in optimal SpO₂ zones increased with the use of closed-loop FiO₂ control.

POPULATION

23 Pediatric Patients



children with acute hypoxemic respiratory failure under HFNO

Median Age: 1 year

LOCATION

3 ICUs in Turkey



INTERVENTION

23 patients under HFNO for acute hypoxemia

manual or automated FiO₂ control

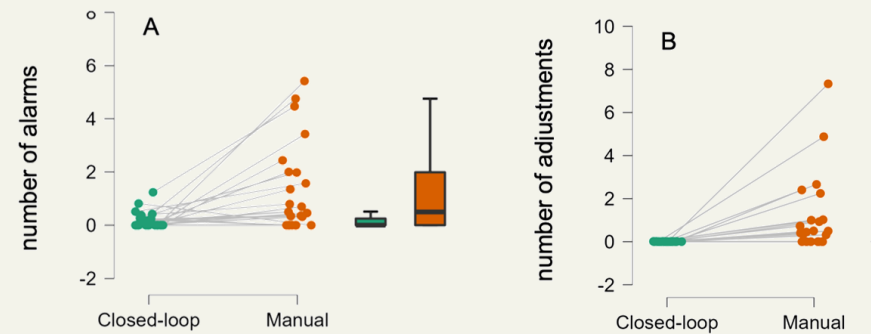
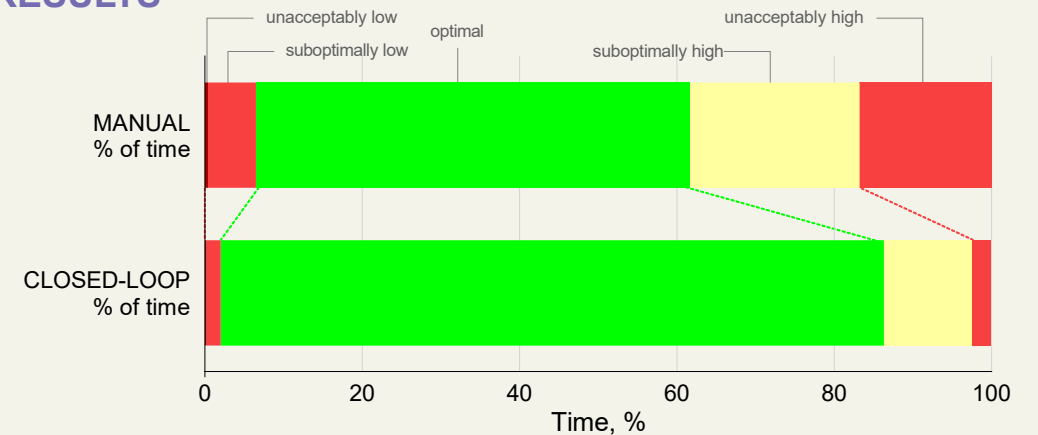
crossover

automated or manual FiO₂ control

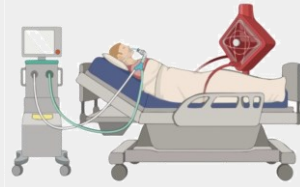
(PRIMARY) OUTCOME

FiO₂ settings and SpO₂ readings; alarms and manual adjustments

RESULTS



Lung Protective Ventilation during ECMO



how to set the ventilator (and adjust ECMO) by hand and the potential role of automation

cannulation for ECMO

initiation

start on time, to prevent unnecessarily long use of harmful ventilation

1

blood flow

use sufficiently high blood flows, to provide optimal oxygenation

2

initial sweep gas flow

use sufficiently high gas flows, to provide good decapneization

3

sweep gas flow

adjust to allow use of even lower V_T and RR, to keep the energy transfer low

4

decannulation



tracheal intubation

tidal volume

target a low V_T , from 4 to 8 ml/kg PBW

1

positive end-expiratory pressure

use PEEP levels that prevent too much alveolar collapse (with ECMO PEEP is suggested to be set > 10 cm H_2O)

2

respiratory rate

target an alveolar minute volume that fits patient's need, and use permissive hypercapnia

3

driving pressure

adjust V_T and titrate PEEP, so that ΔP stays low

4

mechanical power

titrate ventilator settings and use permissive hypercapnia, so that the energy transferred to the lung stays low

5

extubation

automated modes (but studies that support their use during ECMO remain lacking)



PAV+

inspiratory assist in proportion to patient's effort



ASV

automated selection of V_T and RR according to the least work of breathing



NAVA

inspiratory assist in proportion to diaphragm activity



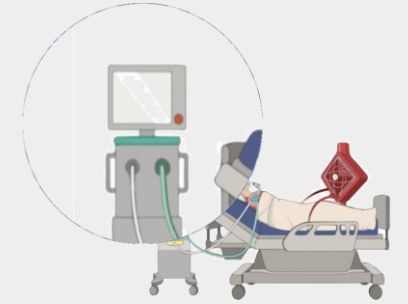
SmartCare

automated weaning: reduction of support and performs spontaneous breathing trials



INTELLiVENT-ASV

automated selection of V_T and RR according to the least work of breathing and force of breathing (power), automated titrations of AMV, PEEP and FiO_2 , and automated weaning: reduction of support, performs spontaneous breathing trials



Agenda

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- history of automated ventilation
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- need for automated ventilation?
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Agenda

- effectiveness
- safety
- efficacy?

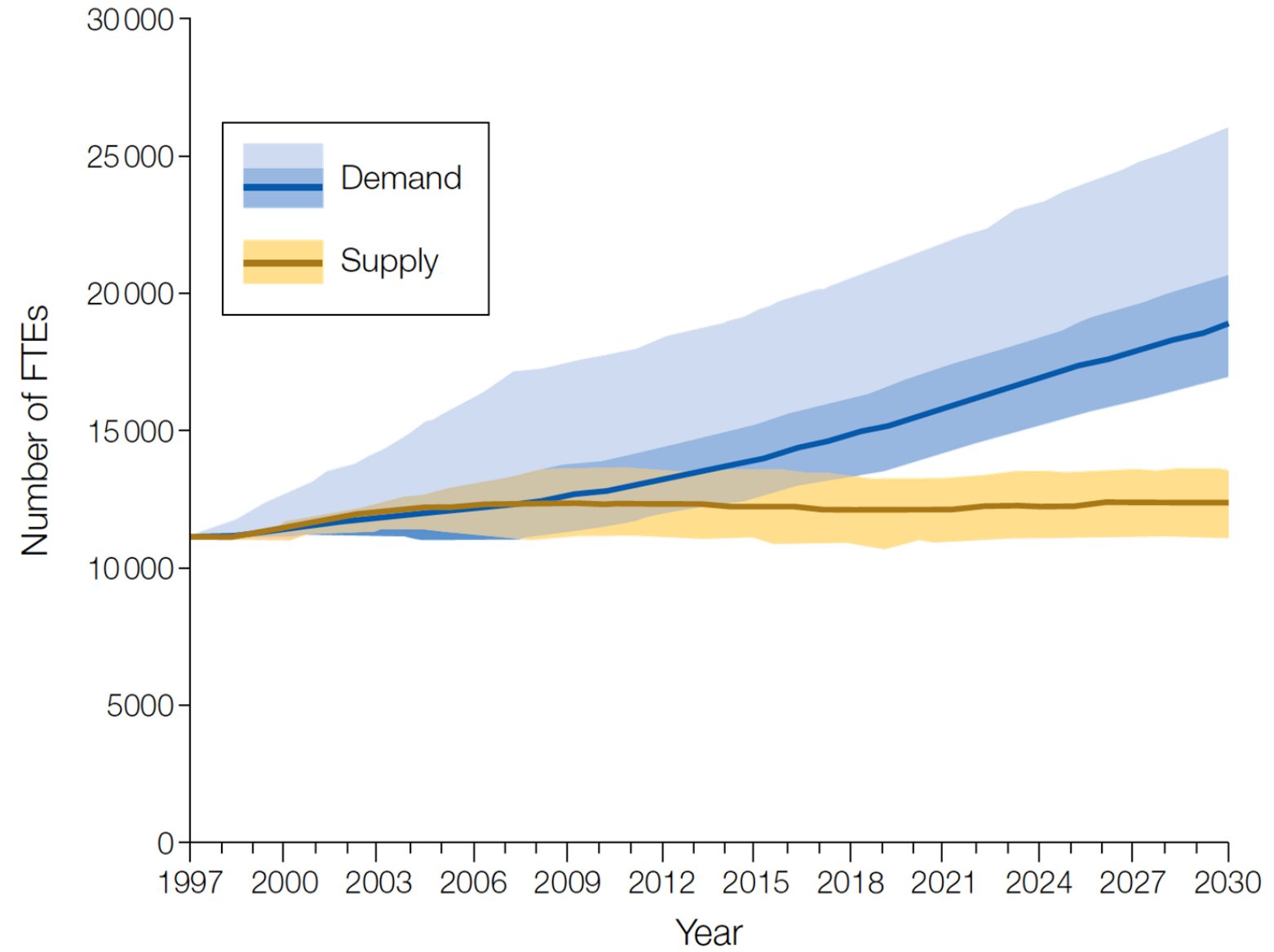


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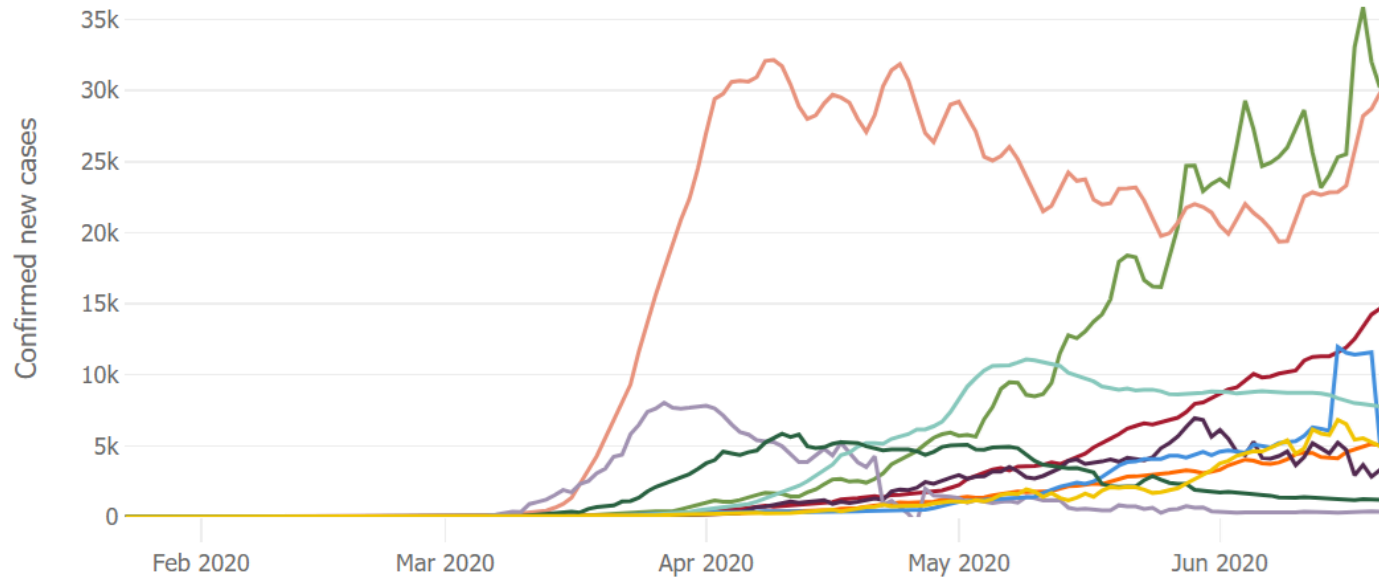


A Intensivists



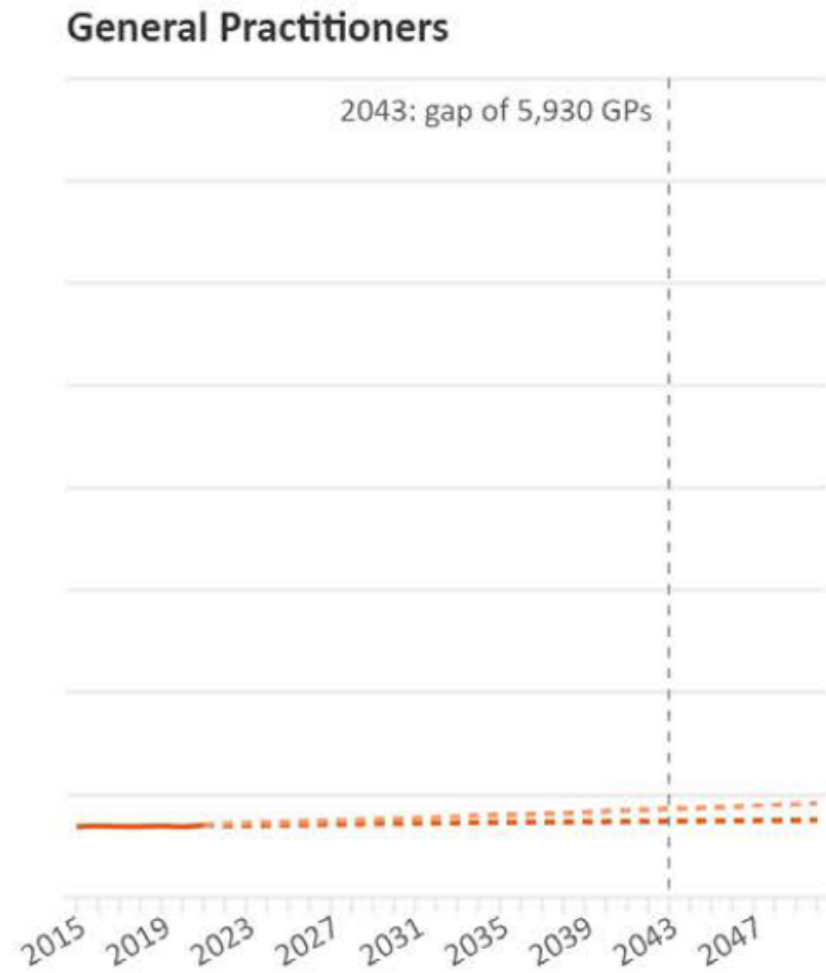
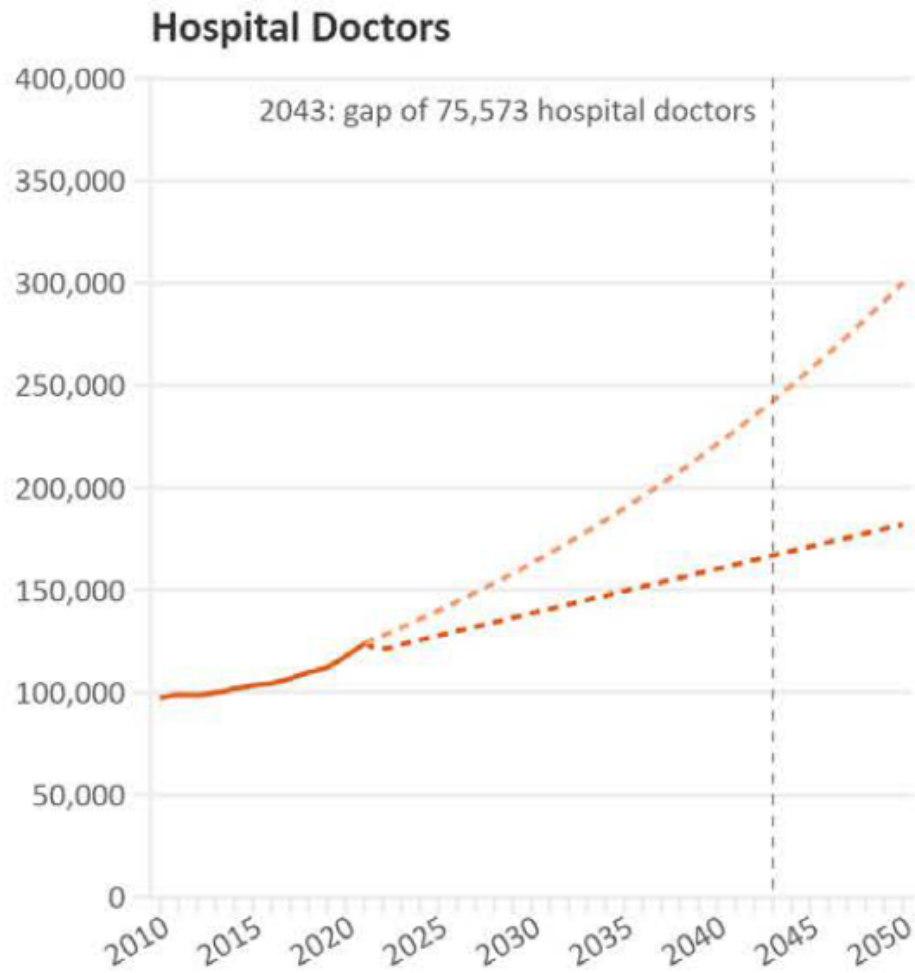
Daily confirmed new cases (5-day moving average)

Outbreak evolution for the current 10 most affected countries



Click any country below to hide/show from the graph:

- Brazil
- Mexico
- US
- India
- Peru
- Spain
- Chile
- Russia
- United Kingdom
- Pakistan



Potential gap between supply and demand of NHS doctors (FTE)

Gap between **projected trend growth** and **required growth** to meet expected growth in activity level

QUESTION What is the precise amount of oxygen consumption with high-flow nasal oxygen (HFNO) and with mechanical ventilation in patients with acute hypoxemic respiratory failure due to COVID-19?

CONCLUSION Actual oxygen consumption, hourly oxygen consumption, and total oxygen consumption are substantially higher in COVID-19 patients that start with HFNO compared with patients that start with mechanical ventilation.

POPULATION



275 Patients

COVID-19 patients with acute hypoxemic failure receiving respiratory support

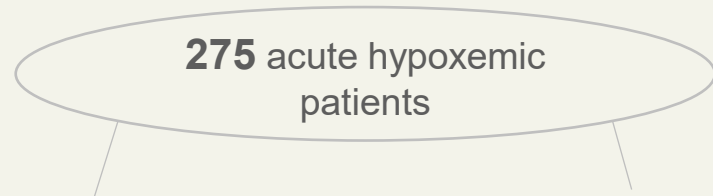
Median Age: **63** year

LOCATION

2 ICUs in Europe



COMPARISON



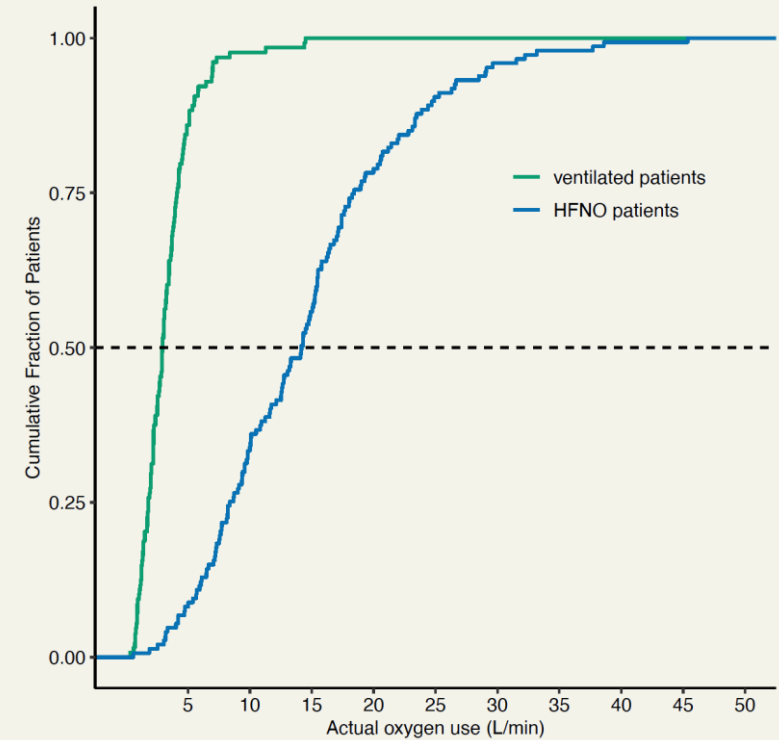
147
started with
HFNO

128
started with
mechanical ventilation

(PRIMARY) OUTCOME

oxygen consumption per minute during the first 2 full calendar days of ICU admission; oxygen consumption per hour and total oxygen consumption in the same time frame

FINDINGS



QUESTION What is the safety, efficacy and workload for the health care team of INTELLiVENT–ASV versus conventional modes over a 48–hour period?

CONCLUSION INTELLiVENT–ASV requires less manual intervention and delivered more variable PEEP, while delivering ventilation safe and effective ventilation in terms of V_T , RR, SpO_2 and $etCO_2$.

POPULATION



86 Patients

patients expected to need ventilation for more than 48 hours

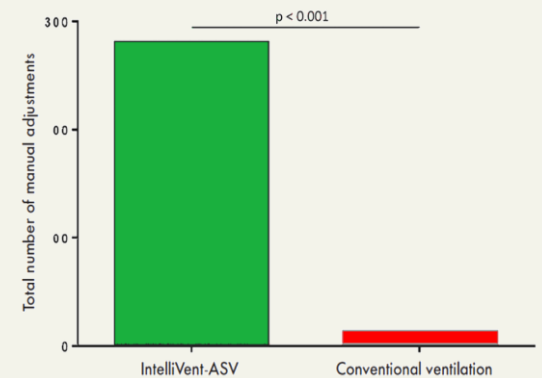
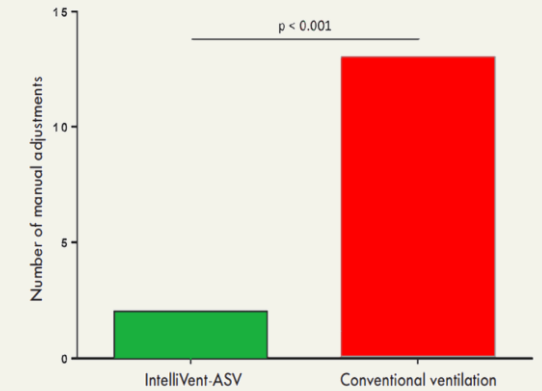
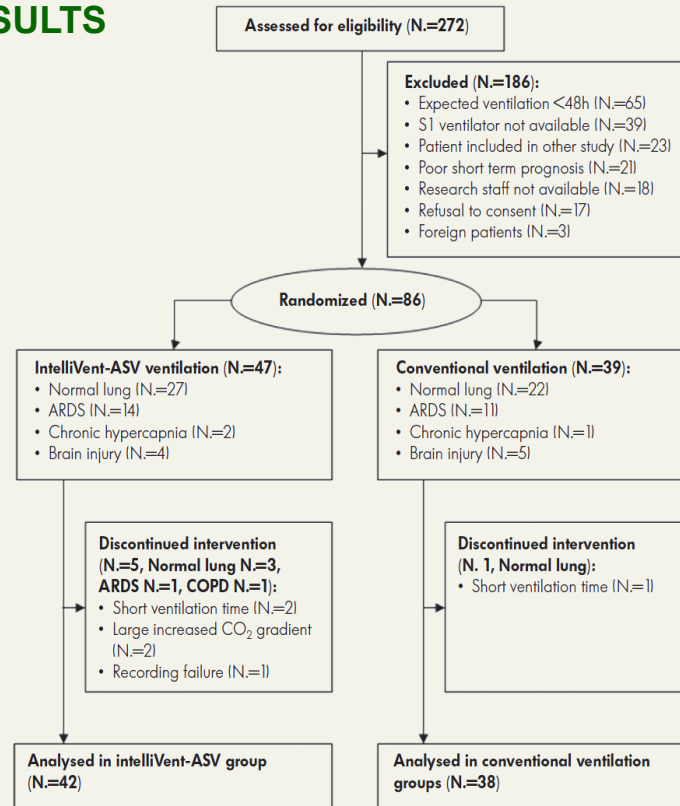
Median Age: 59 year

LOCATION

1 ICU in Belgium



RESULTS



QUESTION Does INTELLiVENT–ASV reduce the number of manual ventilator setting changes compared to conventional ventilation modes like volume assist control and pressure support in ICU patients?

CONCLUSION For mechanically ventilated ICU patients, INTELLiVENT–ASV significantly reduces the number of manual ventilator setting changes with the same number of arterial blood gas analysis and sedation dose, and is easier to use.

POPULATION



60 Patients

patients with an expected duration of ventilation of at least 48 hours

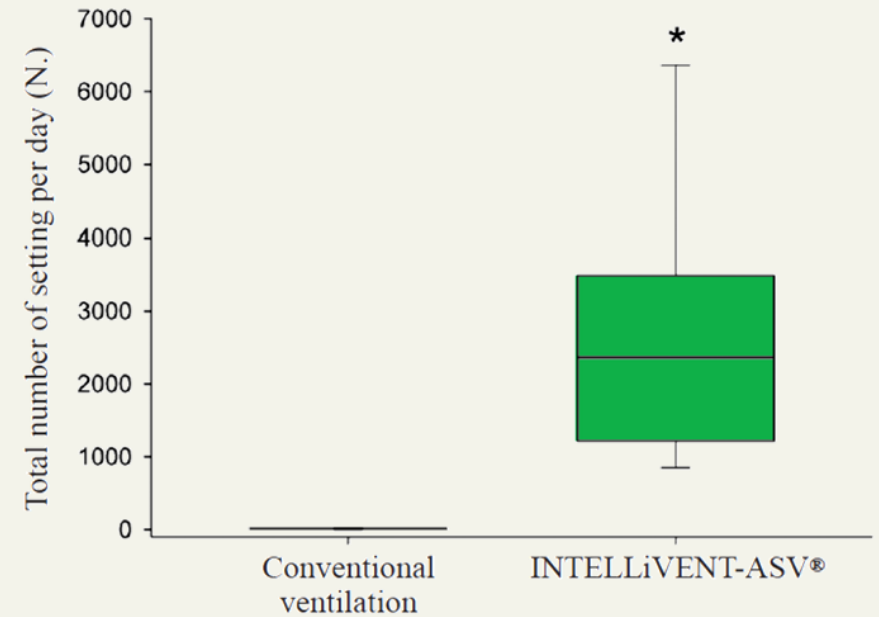
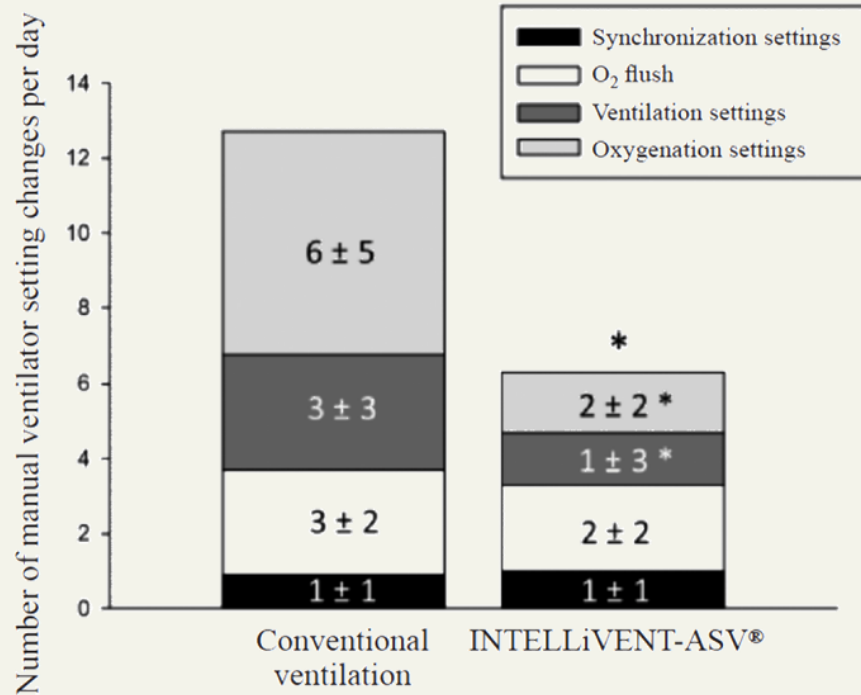
Median Age: **65** year

LOCATION

1 ICU
in France



RESULTS



QUESTION How does INTELLiVENT–ASV performs, in terms of lung–protective ventilation, compared to conventional mechanical ventilation in the resource–constrained setting of the COVID–19 pandemic?

CONCLUSION During an early highpoint of the pandemic, mechanical ventilation using INTELLiVENT–ASV was associated with a higher degree of lung–protective ventilation than was conventional mechanical ventilation.

POPULATION



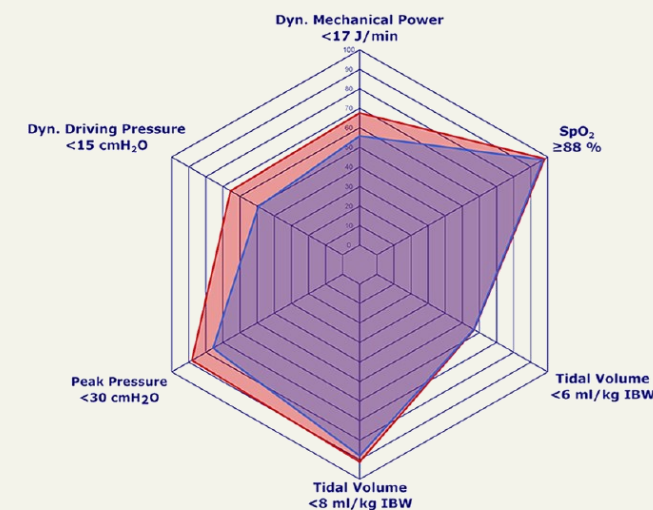
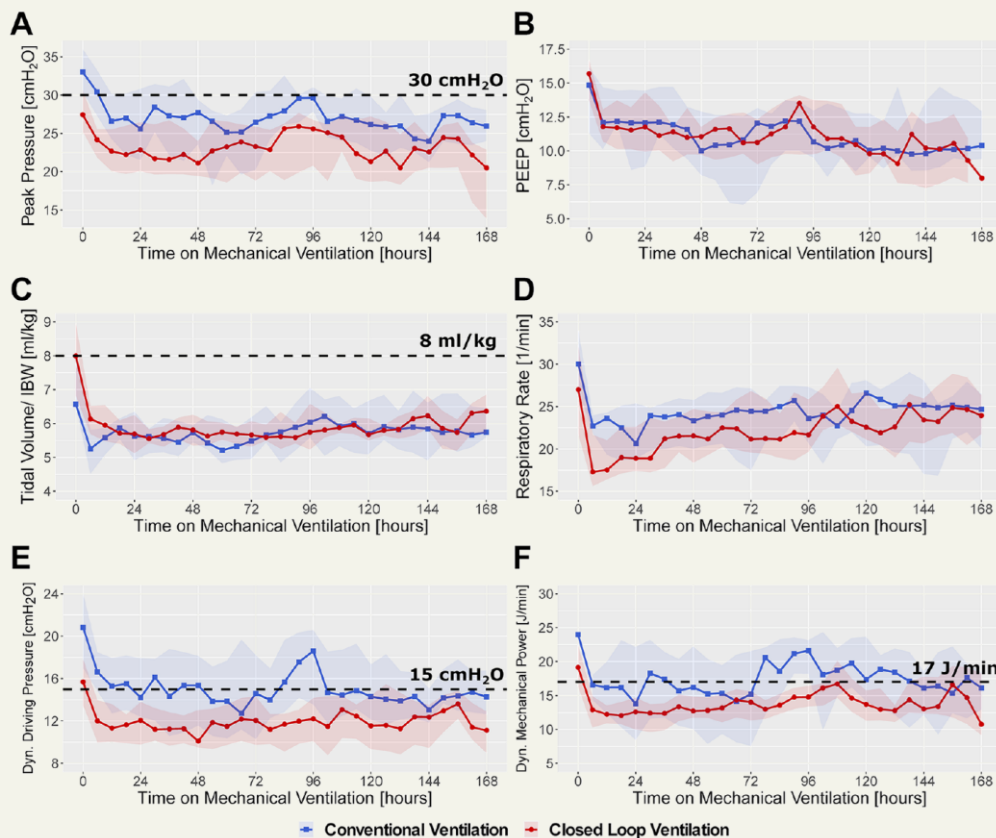
40 Patients

COVID–19 ARDS patients receiving mechanical ventilation

Median Age: 63 year

LOCATION

1 ICU in Switzerland



2.8 [2.3–3.6] automatic adjustments *per minute*
 4 [2–7] vs. 7 [3–12] manual adjustments *per hour*

Wrap-up

- ventilation can be complex and time-consuming
- automated modes are increasingly available
- effectiveness, safety, effectivity
- workload reductions

