

급성 호흡부전 환자에서 산소치료

Pulmonology & Critical Care Medicine

Chonnam National University Hospital

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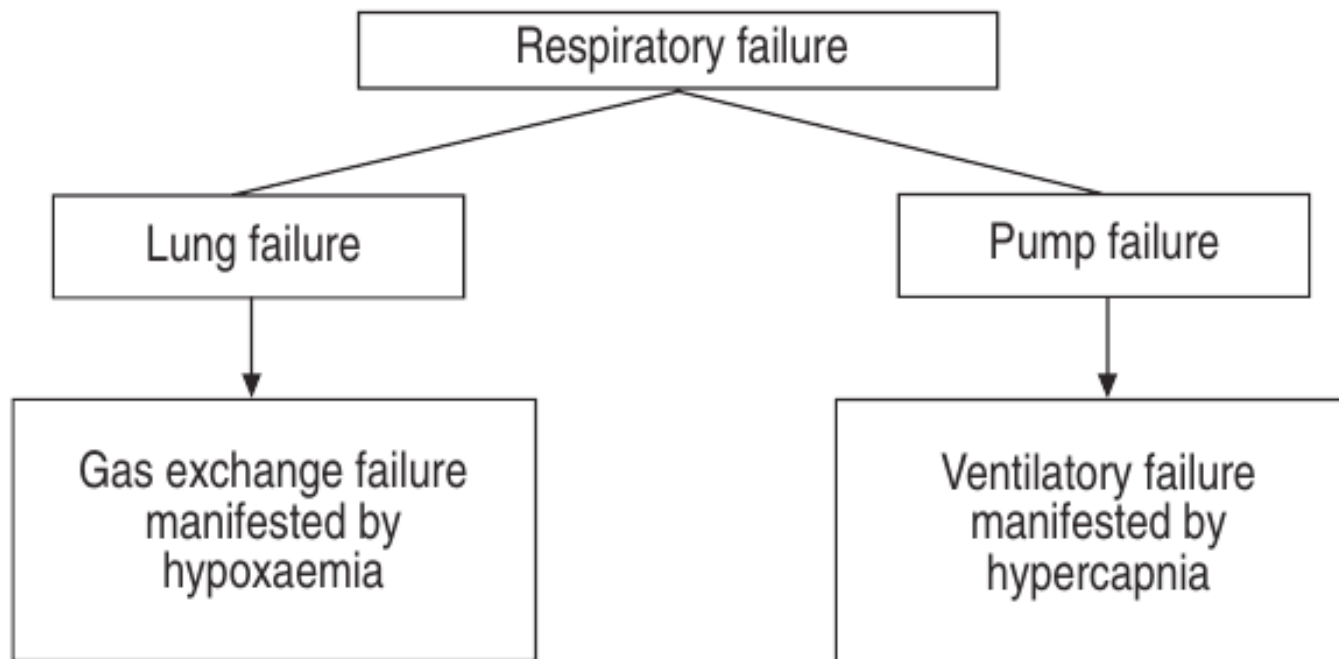
Contents

- 1) Respiratory failure
- 2) Acute hypoxemic respiratory failure
- 3) Acute hypercapnic respiratory failure

Definition of Respiratory Failure

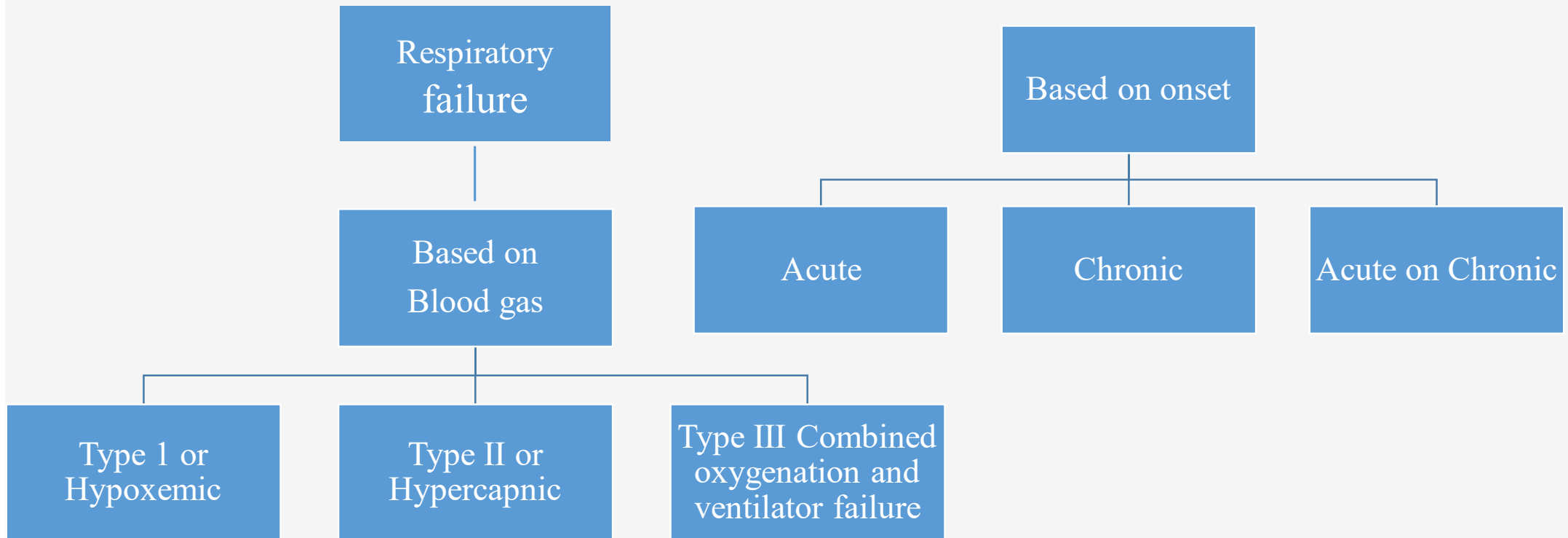
- Respiratory system fails in one of both of gas exchange functions
 - ✓ Oxygenation
 - ✓ Elimination of carbon dioxide
- Arterial oxygen tension (PaO_2) < 60mmHg
- Arterial carbon dioxide tension (PaCO_2) > 45mmHg

Types of Respiratory Failure



- Gas exchanging organ
 - Lung
 - Pulmonary vessels
- Ventilation organ
 - Chest wall
 - Respiratory muscle
 - Chest cage
 - CNS
 - Spinal, Peripheral nerve

Classification of Respiratory Failure

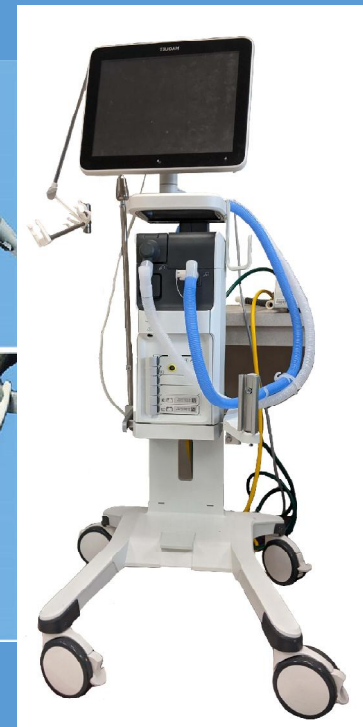
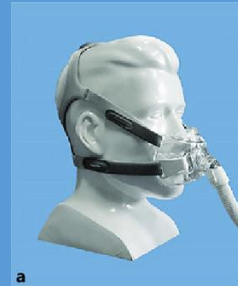


Methods of O₂ supply

Oxygen Supply



Ventilator



CASE 1

- 55세 여자 환자가 3일전부터 발생한 기침, 가래를 동반한 호흡곤란 증상으로 응급실로 내원함.
- Vital sign
 - BP: 100/50mmHg, HR:120/min, RR:34/min, BT:38.2°C, SaO₂ 78%
- ABGA
 - RA : pH 7.39, pCO₂ 26mmHg, PaO₂ 46mmHg, SaO₂ 81%

CASE 1



- No history of chronic pulmonary, cardiac disease
- Supply O₂ via mask 10L/min
- Vital sign after O₂ supply
 - ✓ BP: 100/50 mmHg, HR: 120/min, RR: 29/min
 - ✓ ABGA pH 7.45, pCO₂ 25mmHg, PaO₂ 70 mmHg, SaO₂ 92%



Goals of Oxygen Therapy

- The ability to supply the patient with the oxygen they need
- Comfort
- Avoid endotracheal intubation
- Improve mortality
- Prevention of adverse effect

Comparing HFNC and Conventional oxygen therapy

- 17 Patients with ARF requiring > 9L/min
- Face mask (9L/min – 15L/min)
- HFNC setting : FiO₂ 60%, initial Flow 40L/min

Table. Changes in Dyspnea and Respiratory Parameters Comparing Conventional Oxygen Therapy to High Flow Nasal Cannula

	H0	H + 15 min	H + 30 min	H + 60 min
Borg scale (<i>n</i> = 9)	6 (5–7)	4 (3–4)*	4 (2–4)†	3 (2–4)†
Visual analog scale (<i>n</i> = 9)	7 (5–8)	5 (2–6)*	4 (2–6)†	3 (1–5)‡
Respiratory rate, breaths/min (<i>n</i> = 17)	28 (25–32)	25 (23–30)*	25 (21–30)‡	25 (21–28)†
S _{pO₂} , % (<i>n</i> = 17)	90 (88.5–94)	96 (90–99)‡	95 (90–100)†	97 (92.5–100)†

Values are median (IQR).

* *P* < .05.

† *P* < .001.

‡ *P* < .01.

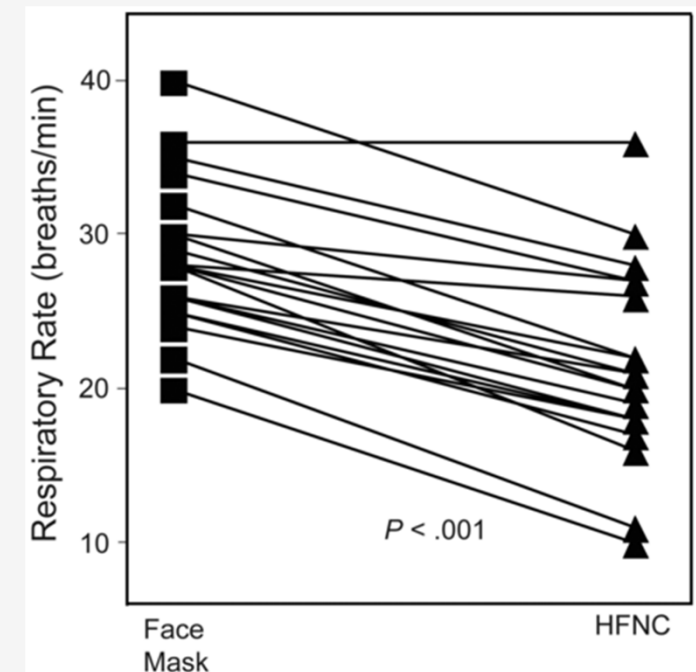
H0 = time just before switching from conventional oxygen therapy to high flow nasal cannula;

Comparing HFNC and Conventional oxygen therapy

- $\text{SaO}_2 < 96\%$ while receiving $\text{FiO}_2 \geq 0.5$ via face mask
- Comparing face mask(15L/min) and HFNC 30L/min

	Face Mask	HFNC	P
Subjective Evaluation			
Dyspnea	6.8 (4.1–7.9)	3.8 (1.3–5.8)	.001
Mouth dryness	9.5 (8.0–10.0)	5 (2.3–7.0)	< .001
Overall comfort	5.0 (2.3–6.8)	9.0 (8.0–10.0)	< .001
Respiratory and Gas-Exchange Variables			
Total oxygen flow (L/min)	15.0 (12.0–20.0)	30.0 (21.3–38.7)	< .001
Fraction of delivered oxygen	1.0 (0.8–1.0)	1.0 (0.8–1.0)	.32
Respiratory rate (breaths/min)	28 (25–32)	21 (18–27)	< .001
pH [†]	7.42 ± 7.38–7.47	7.44 (7.38–7.50)	.06
P_{aO_2} (mm Hg) [‡]	77 (64–88)	127 (83–191)	.002
P_{aCO_2} (mm Hg) [‡]	37 (33–45)	37 (32–43)	.51
HCO_3^- (mmol/L) [‡]	25.0 (22.1–28.5)	24.5 (22.2–29.1)	.09
Base excess (mmol/L) [‡]	1.0 (–2.3–4.8)	–1.0 (–2.3 to 5.3)	.055
S_{pO_2} (%)	95 (91–97)	98 (96–99)	.002
Hemodynamic Variables			
Mean arterial pressure (mm Hg)	87 (76–94)	86 (71–93)	.36
Heart rate (beats/min)	94 (77–112)	85 (73–108)	> .99

* All values are median and interquartile range.
[†] Only in patients with an arterial catheter.
 HFNC = high-flow nasal cannula
 S_{pO_2} = blood oxygen saturation measured via pulse oximetry



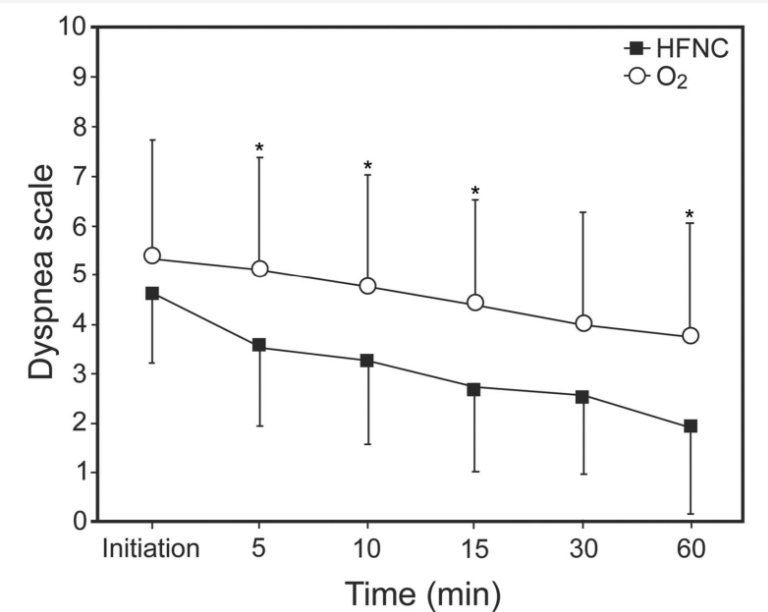
Comparing HFNC and Conventional Oxygen Therapy

- 40 Patients with ARF (RR > 24/min and SpO₂ < 94% on RA)
- HFNC (Flow 35 L/min, FiO₂ adjusted to SpO₂ ≥ 94%) vs. COT for 60 min
- CHF, Asthma attack, COPD c AE, Pneumonia
- Mean SpO₂ 86-88%

Table 2. Comparing Clinical and Physiologic Parameters for the HFNC and COT Groups at the End of the Study

Parameter	HFNC	COT
Dyspnea scale score, mean ± SD	2.0 ± 1.8	3.8 ± 2.3
Breathing frequency, mean ± SD breaths/min	26.0 ± 6.2	27.5 ± 4.9
Mean arterial pressure, mean ± SD mm Hg	88.7 ± 10.9	101.0 ± 24.8
Heart rate, mean ± SD beats/min	91.7 ± 19.3	101.6 ± 24.2
S _{pO₂} , mean ± SD %	96.8 ± 2.5	97.6 ± 2.0
Comfort scale score, mean ± SD	1.6 ± 1.7	3.7 ± 2.4

HFNC = high-flow nasal oxygen cannula
COT = conventional oxygen therapy



Comparing HFNC and Conventional Oxygen Therapy

- 303 Patients with ARF (RR $\geq$ 22/min and SpO₂ < 92% on RA)
- HFNC (Flow 40 L/min, FiO₂ adjusted to SpO₂ $\geq$ 94%) vs. COT
- Outcomes : conversion to NIV or IMV, Hospital LOS, 90-day mortality
- Cause of hypoxia : COPD, Pneumonia, CHF, Asthma

Outcome	HFNC (n = 165)	Standard O ₂ (n = 138)	P
Conversion to mechanical ventilation in emergency department, n, % (95% CI)	6, 3.6 (1.5–7.9)	10, 7.2 (3.8–13)	.16
90-d mortality, n, % (95% CI)	35, 21.2 (15.6–28.1)	24, 17.4 (11.9–24.6)	.40
Pneumothorax, n, % (95% CI)	0, 0 (0–3)	0, 0 (0–3)	
Subcutaneous emphysema, n, % (95% CI)	0, 0 (0–3)	0, 0 (0–3)	
Pressure sore, n, % (95% CI)	0, 0 (0–3)	0, 0 (0–3)	
Apnea, n, % (95% CI)	0, 0 (0–3)	1, 0.7 (0–4)	.45
Fall in GCS $\geq$ 2 points, n, % (95% CI)	1, 0.6 (0–3.7)	6, 4.3 (1.8–9.4)	.050
Fall in GCS due to CO ₂ retention,* n, % (95% CI)	0, 0 (0–3)	3, 2.2 (0.4–6)	.09
Intubated,* n, % (95% CI)	1, 0.6 (0–4)	3, 2.2 (0.5–6)	.33
Admitted to ICU,* n, % (95% CI)	8, 4.8 (2.3–9.4)	5, 3.6 (1.3–8.4)	.60
Mechanical ventilation within 24 h,* n, % (95% CI)	9, 5.5 (2.8–10.2)	16, 11.6 (7.2–18.1)	.053
In-hospital mortality,* n, % (95% CI)	15, 9.1 (5.5–14.6)	11, 8.0 (4.4–13.8)	.73
Emergency department LOS, median (IQR) h	4.5 (3.6–5.8)	4.9 (3.6–5.9)	.32
Hospital LOS, median (IQR) d	5.0 (2.8–8.3)	5.6 (2.8–9.3)	.43
Time to mechanical ventilation in emergency department,* median (IQR) min	6, 139.5 (70–214.5)	10, 126 (19.75–197.75)	.43
Time to mechanical ventilation in ward,* median (IQR) min	3, 1,040 (755.5–1,222.5)	6, 857 (395–1,116)	.71

Comparing HFNC and Conventional oxygen therapy

Intubation rate

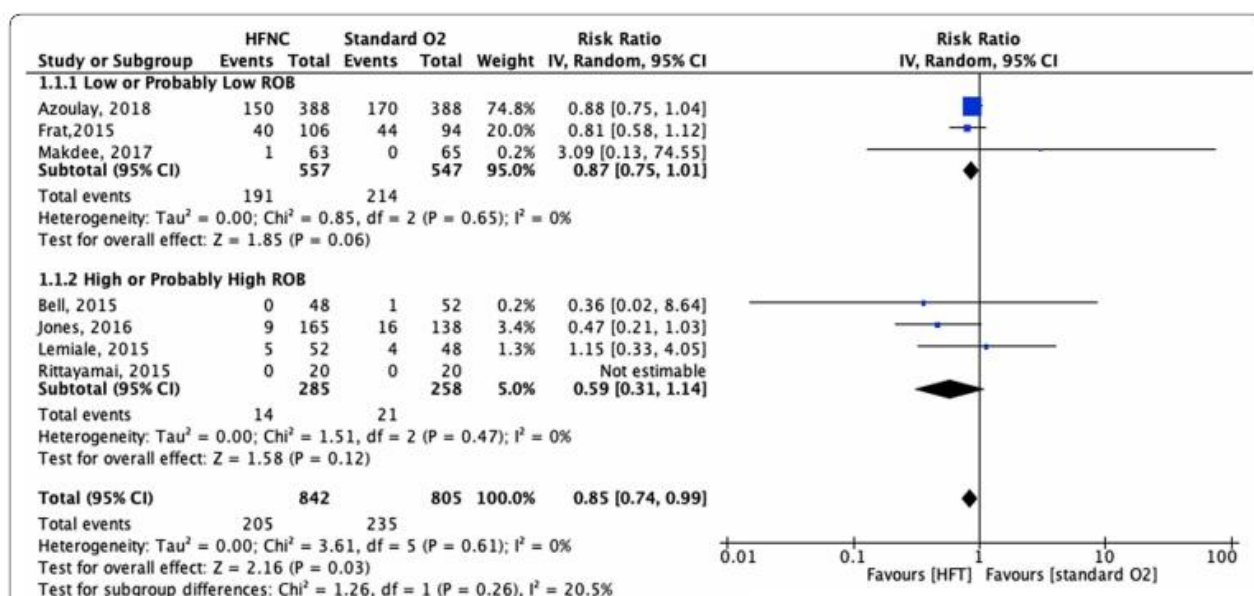


Fig. 3 Need for invasive mechanical ventilation

Mortality

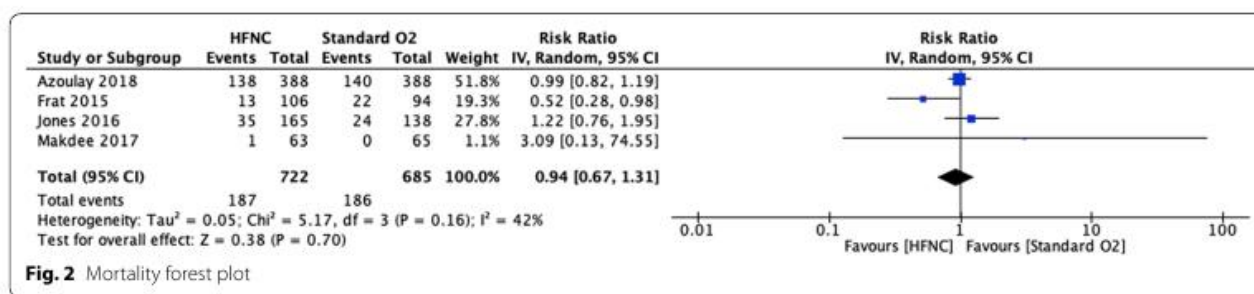


Fig. 2 Mortality forest plot

ERS clinical practice guidelines: HFNC in acute respiratory failure

HFNC for hypoxaemic acute respiratory failure

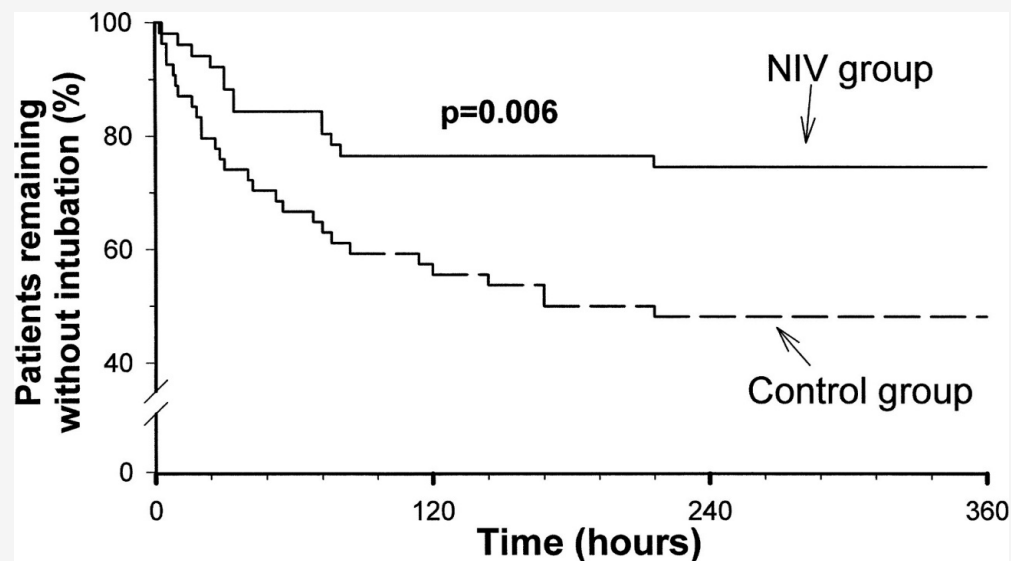
PICO question 1: Should HFNC or COT be used in patients with acute hypoxaemic respiratory failure?

Recommendation 1

We suggest the use of HFNC over COT in adults with acute hypoxaemic respiratory failure (conditional recommendation, moderate certainty of evidence).

Comparing NIV and Conventional oxygen therapy

- 105 Patients with ARF (P/F ratio ≤ 120)
- Venturi mask (54) vs. NIV (51)



Exclusion Criteria

- (1) hypercapnia (PaCO₂ of more than 45 mm Hg) on admission
- (2) need for emergency intubation
- (3) recent esophageal, facial, or cranial trauma or surgery
- (4) severely decreased consciousness (a Glasgow coma score ≤ 11)
- (5) severe hemodynamic instability despite fluid repletion and use of vasoactive agents
- (6) a lack of cooperation
- (7) tracheotomy or other upper airway disorders
- (8) severe ventricular arrhythmia or myocardial ischemia
- (9) active upper gastrointestinal bleeding
- (10) an inability to clear respiratory secretions
- (11) ≥ 1 severe organ dysfunction in addition to respiratory failure.

Comparing NIV and Conventional oxygen therapy

Intubation rate

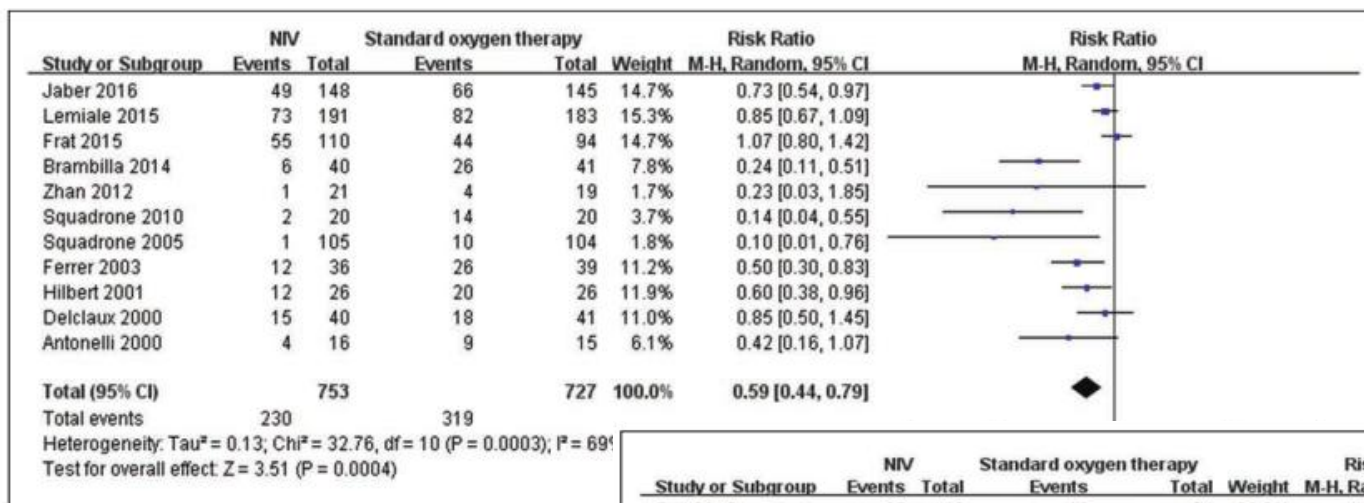


Figure 3. Intubation rate in acute hypoxemic nonhypercapnic respiratory failure patients randomized to noninvasive ventilation (NIV) versus standard oxygen therapy. M-H = Mantel-Haenszel.

Mortality

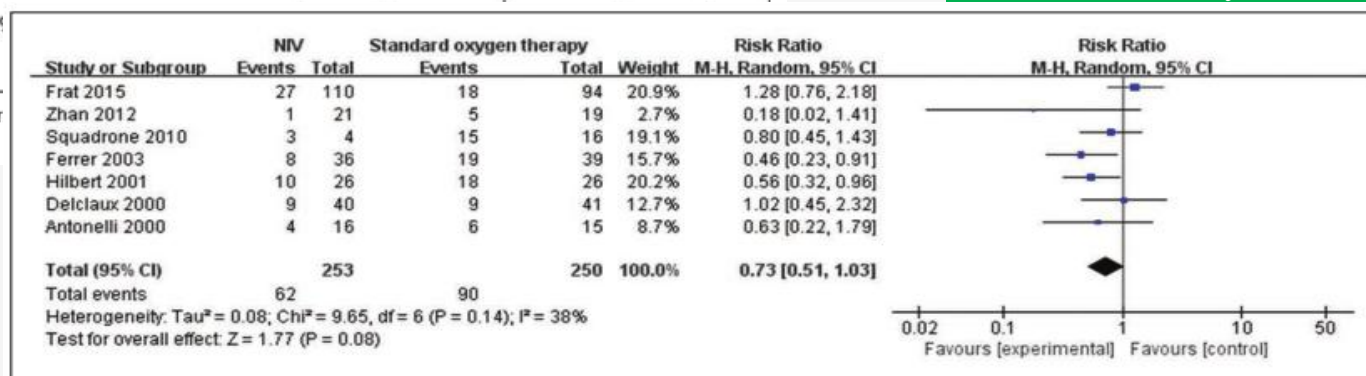


Figure 4. ICU mortality in acute hypoxemic nonhypercapnic respiratory failure patients randomized to noninvasive ventilation (NIV) versus standard oxygen therapy. M-H = Mantel-Haenszel.

Benefit of NIV in acute hypoxemic respiratory failure

➤ Patients likely to benefit

- ✓ Acute cardiogenic pulmonary edema

➤ Patients likely to benefit

- ✓ Hypoxemic nonhypercapnic respiratory failure not due to ACPE
(Inability to cooperate, protect airway, or clear secretions)
- ✓ Acute respiratory failure due to asthma exacerbation
- ✓ Post-operative or chest trauma induced acute respiratory failure

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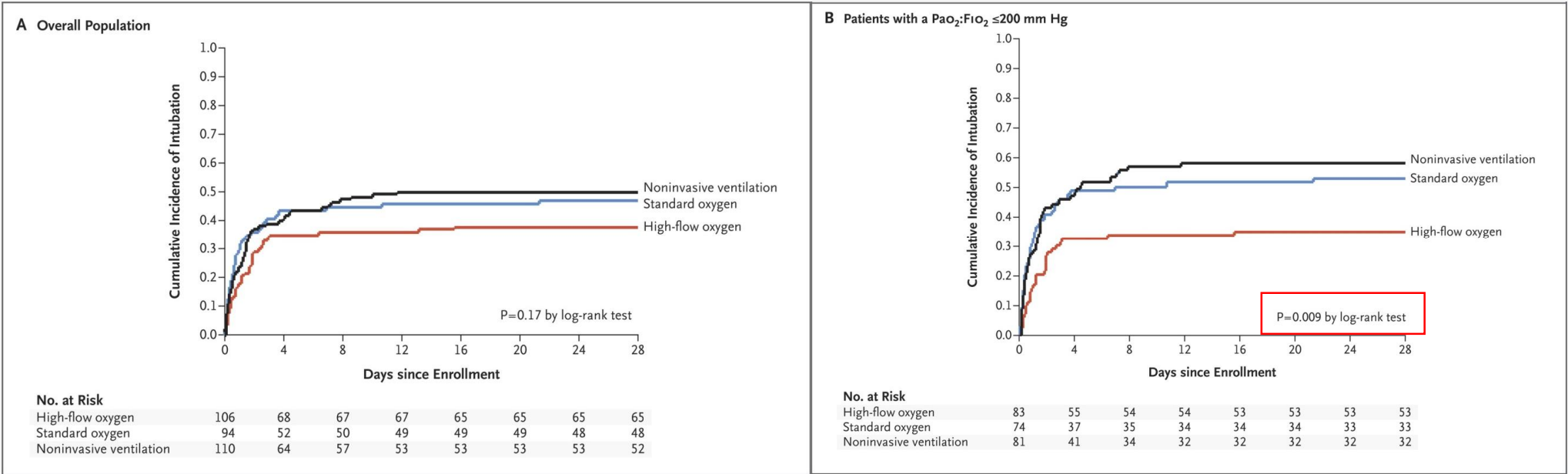
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High-Flow Oxygen through Nasal Cannula in Acute Hypoxemic Respiratory Failure

- Multi-center Randomized controlled trial, 23 ICUs in France and Belgium
- 310 patients with acute hypoxemic respiratory failure, HFNC(106) vs. SOT(94) vs. NIV(110)
- Acute hypoxemic respiratory failure : P/F ratio < 300
- Outcome
 - ✓ Primary outcome : Intubation rate at day 28
 - ✓ 2ndary outcome : All cause mortality in ICU and at 90 days

Intubation rate



HFNC(38%) vs. COT(47%) vs. NIV(50%)

HFNC(35%) vs. COT(53%) vs. NIV(58%)

All cause mortality

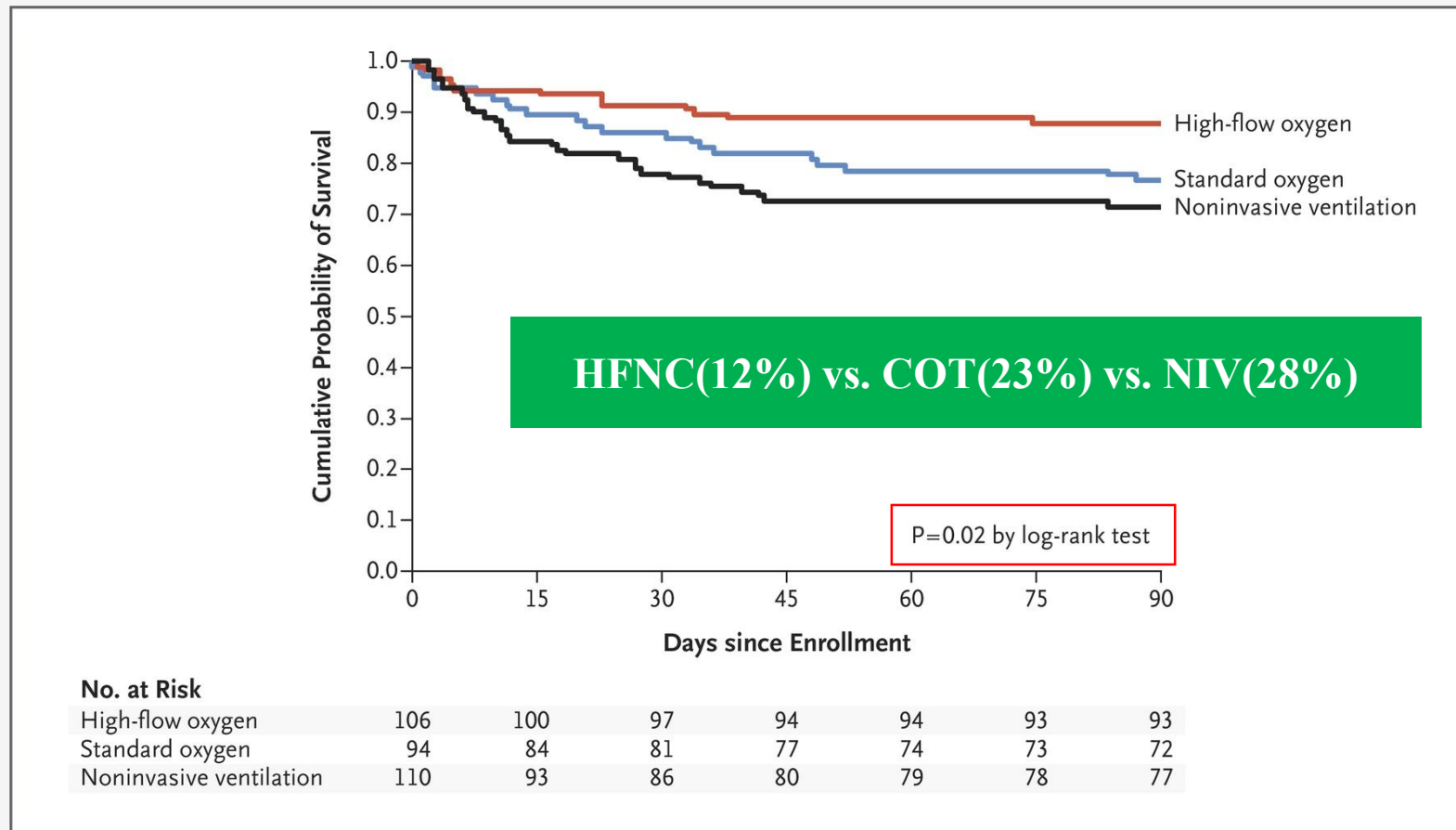


Table 1. Characteristics of the Patients at Baseline, According to Study Group.*

Characteristic	High-Flow Oxygen (N = 106)	Standard Oxygen (N = 94)	Noninvasive Ventilation (N = 110)
Age — yr	61±16	59±17	61±17
Male sex — no. (%)	75 (71)	63 (67)	74 (67)
Body-mass index†	25±5	26±5	26±6
SAPS II‡	25±9	24±9	27±9
Current or past smoking — no. (%)	34 (32)	36 (38)	40 (36)
Reason for acute respiratory failure — no. (%)			
Community-acquired pneumonia	71 (67)	57 (61)	69 (63)
Hospital-acquired pneumonia	12 (11)	13 (14)	12 (11)
Extrapulmonary sepsis	4 (4)	5 (5)	7 (6)
Aspiration or drowning	3 (3)	1 (1)	2 (2)
Pneumonia related to immunosuppression	6 (6)	4 (4)	10 (9)
Other	10 (9)	14 (15)	10 (9)
Bilateral pulmonary infiltrates — no. (%)	79 (75)	80 (85)	85 (77)
Respiratory rate — breaths/min	33±6	32±6	33±7

Adverse effect of NIV



Incidence of side effects	Any grade	Grade 3-4
Eye redness	76 (17%)	17 (4%)
Rhinorrhea	137 (31%)	28 (6%)
Mouth dryness	329 (75%)	116 (27%)
Hourseness	74 (27%)	11 (3%)
Bloating	89 (20%)	21 (3%)
Interface related pain	377 (86%)	19 (4%)
Pressure sore	209 (48%)	4 (1%)
Noticeable leaks	267 (61%)	65 (15%)
Deventilation dyspnoea	83 (19%)	21 (5%)
Perceived patient ventilator-asynchrony	75 (17%)	23 (5%)

NIV Interfaces



Full face mask



face mask

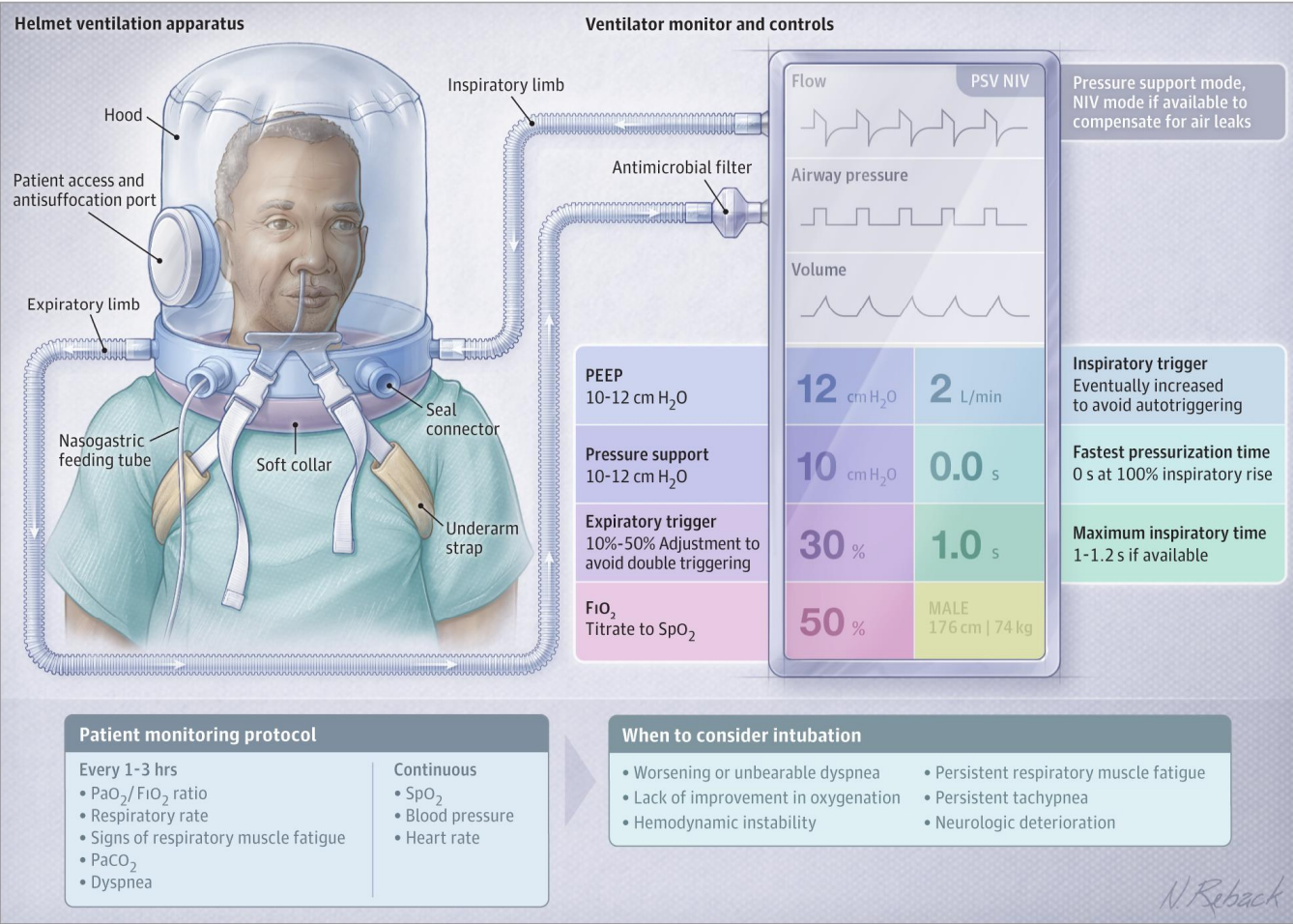


Nasal mask



Helmet mask

Helmet NIV

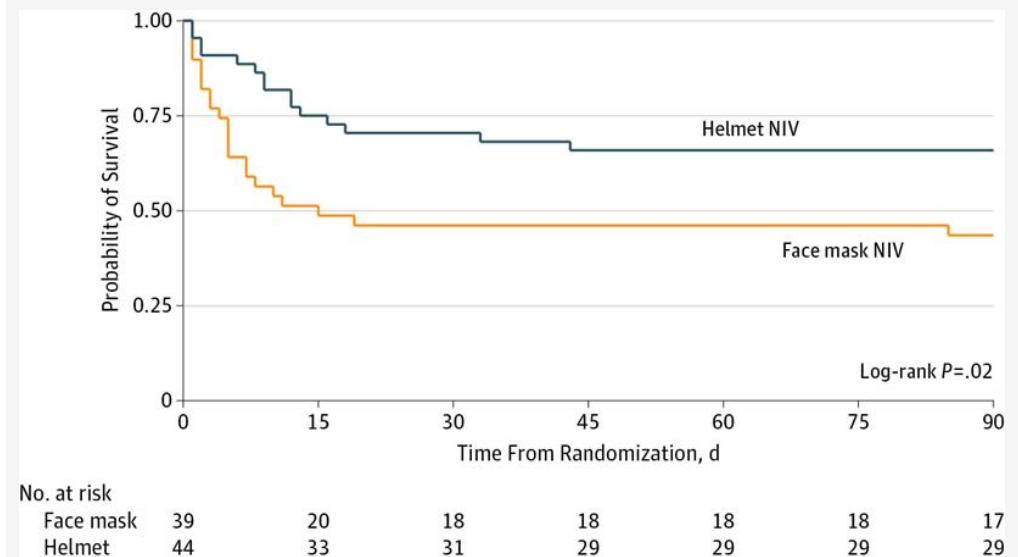


Comparing between NIV interfaces

- 83 Patients with ARDS, P/F ratio 90-220, Shock (25%)
- Helmet device (44) vs. Face mask device (39)

Table 2. Primary and Secondary Outcomes and Adverse Events

	Face Mask (n = 39)	Helmet (n = 44)	Absolute Difference (95% CI)	P Value
Primary outcome, No. (%)				
Endotracheal intubation	24 (61.5)	8 (18.2)	-43.3 (-62.4 to -24.3)	<.001
Reason for intubation				
Respiratory failure	20 (83.3)	3 (37.5)	-45.3 (-82.5 to -9.1)	.01
Circulatory failure	3 (12.5)	0 (0)	-12.5 (-25.7 to 0.7)	.55
Neurologic failure	1 (4.2)	5 (62.5)	58.3 (24.8 to 92.8)	.001
Secondary outcomes, median (IQR), d				
Ventilator-free days	12.5 (0.49-28)	28 (13.7-28)	8.4 (13.4 to 3.4)	<.001
ICU length of stay	7.8 (3.9-13.8)	4.7 (2.5-8.7)	-2.76 (-6.07 to 0.54)	.04
Hospital length of stay	15.2 (7.8-19.7)	10.1 (6.5-15.9)	-2.92 (-8.47 to 2.63)	.16
Mortality, No. (%)				
Hospital	19 (48.7)	12 (27.3)	-21.4 (-41.9 to -1.0)	.04
90 d ^a	22 (56.4)	15 (34.1)	-22.3 (-43.3 to -1.4)	.02
Adverse events				
Mask deflation	0 (0)	2 (4.5)		
Skin ulceration	3 (7.6)	3 (6.8)		



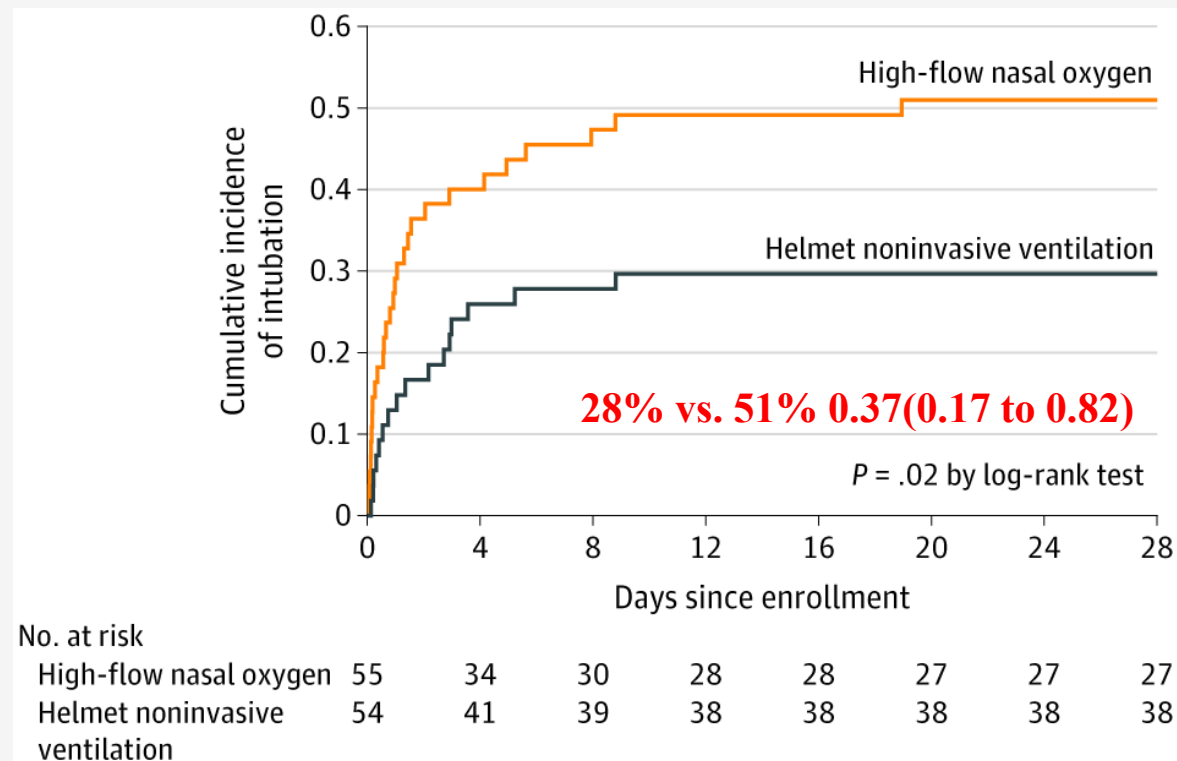
Comparing between NIV interfaces

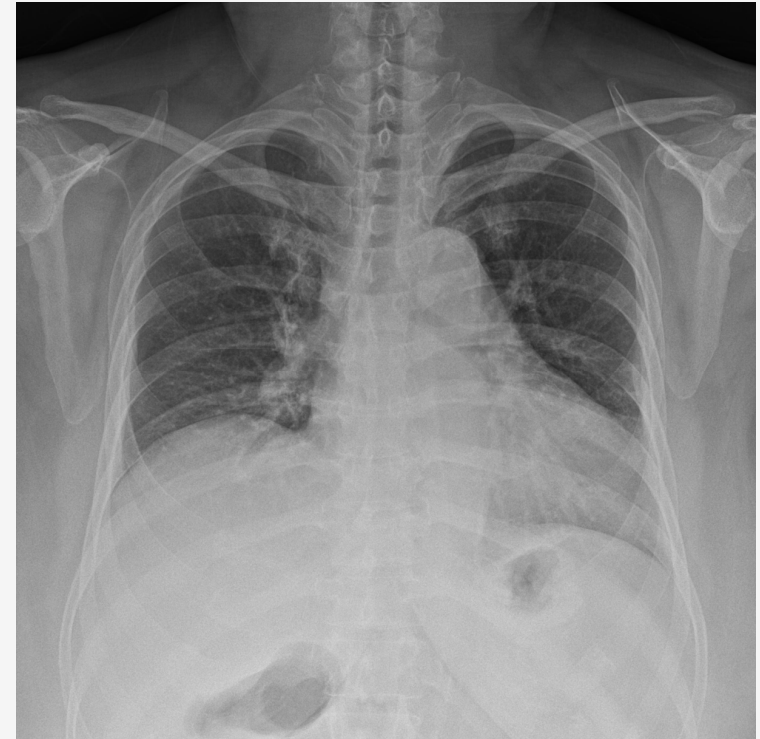
Table 3. Level of Respiratory Support and Physiologic Parameters During Noninvasive Ventilation

	Noninvasive Ventilation, Median (IQR)		<i>P</i> Value
	Face Mask (n = 39)	Helmet (n = 44)	
Respiratory support with NIV ^a			
Duration of NIV, h	26.4 (7.0-60.0)	19.8 (8.4-45.6)	.68
PEEP, cm H ₂ O	5.1 (5.0-8.0)	8 (5.0-10.0)	.006
Pressure support, cm H ₂ O	11.2 (10.0-14.5)	8 (5.6-10.0)	<.001
FIO ₂ , %	60 (50.0-68.6)	50 (40.0-60.0)	.02
SpO ₂ , %	95.3 (92.3-96.7)	96.2 (94.8-98.4)	.13
Respiratory rate, breaths/min			
Baseline	28.3 (22.1-34.4) ^b	27.7 (21.5-34.6) ^b	
After randomization	29.1 (22.1-37.6)	24.5 (20.4-30.5)	

Helmet NIV vs. HFNC in COVID-19

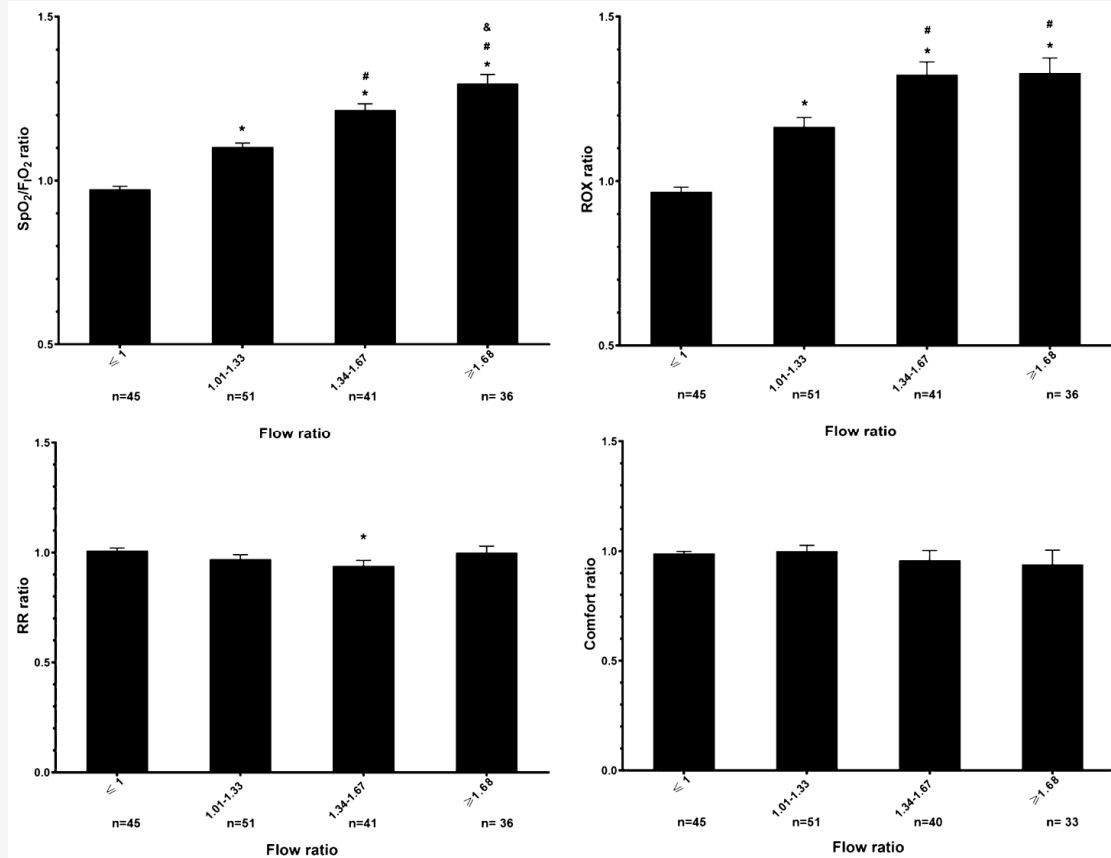
- 109 Patients with ARF ($\text{PaO}_2/\text{FiO}_2 \leq 200$), HFNC vs. Helmet NIV
- Outcomes : respiratory support free days, intubation rate, mortality





Response to different flow ratios.

- HFNC and required a minimum FiO_2 of 0.4 to maintain pulse oximetry (SpO_2) of 90–97%
- Peak tidal inspiratory flow
 - ✓ Mean : 34 ± 9
 - ✓ Median : 31 [27-42]

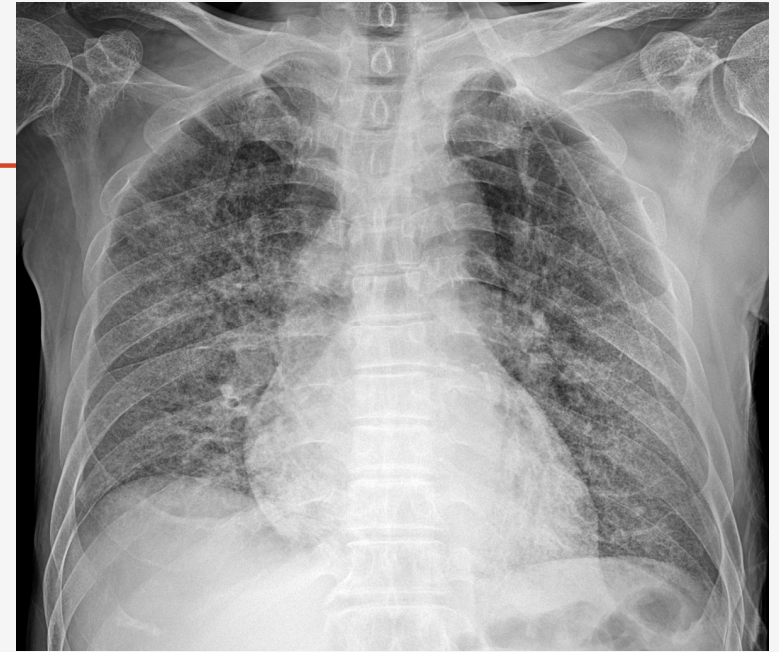


CASE 2

- 60/M
- C/C dyspnea (onset : 3 days ago)
- fever, chill, cough, sputum after greenhouse work

- Vital sign
 - BP: 100/60mmHg, HR:79/min, **RR:24/min**, BT:38.1°C

- ABGA via Mask 8L/min
 - RA : pH 7.44, pCO₂ 33mmHg, **PaO₂ 65mmHg, SaO₂ 93%**





	Flow (L/min)	FiO2	SaO2	RR
Day 1	60	0.7	94%	24
Day 2	60	0.5	96%	20
Day 3	60	0.8	95%	24
Day 4	60	0.8	93%	28
Day 5	60	0.9	91%	34

Is HFNC safe to all patients?

- Restrospective, HFNC failure
- 175 patients, Timing of intubation before 48hr(early) and after 48hr(late)

Variables	Crude		Propensity-adjusted ^a		Propensity-matched ^b	
	Odds ratio (95 % CI)	P value ^c	Odds ratio (95 % CI)	P value ^c	Odds ratio (95 % CI)	P value ^c
Primary outcome						
Overall ICU mortality	0.323 (0.158–0.658)	0.002	0.317 (0.143–0.700)	0.005	0.369 (0.139–0.984)	0.046
Secondary outcomes						
Extubation success	3.284 (1.361–7.923)	0.008	3.091 (1.193–8.013)	0.020	2.057 (0.746–5.672)	0.163
Ventilator-weaning	3.056 (1.470–6.351)	0.003	3.380 (1.492–7.656)	0.004	2.495 (1.039–5.991)	0.041
Ventilator-free days to day 28	0.542 (0.383–0.768) ^d	0.001 ^e	0.516 (0.349–0.763) ^d	0.001 ^e	0.639 (0.431–0.946) ^d	0.026 ^e
14-Day mortality from HFNC application	0.949 (0.455–1.977)	0.888	0.712 (0.312–1.622)	0.418	0.608 (0.231–1.606)	0.316
14-Day mortality from intubation	0.653 (0.325–1.311)	0.231	0.482 (0.218–1.067)	0.072	0.447 (0.168–1.184)	0.105
28-Day mortality from HFNC application	0.820 (0.416–1.616)	0.566	0.680 (0.318–1.457)	0.322	0.896 (0.440–1.824)	0.763
28-Day mortality from intubation	0.571 (0.287–1.138)	0.111	0.557 (0.258–1.198)	0.134	0.802 (0.380–1.692)	0.563
Length of ICU stay	0.827 (0.586–1.169) ^f	0.282 ^g	0.830 (0.552–0.800) ^f	0.372 ^g	1.329 (0.598–2.952) ^f	0.485 ^g

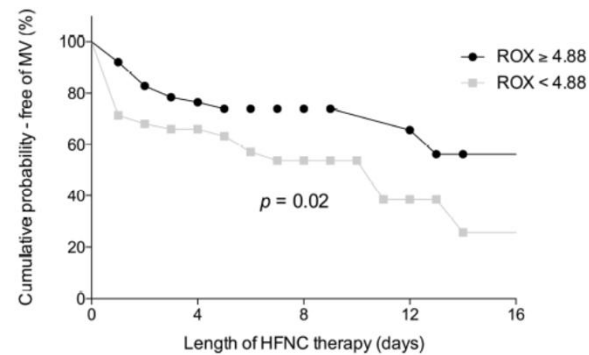
- The early intubated patients had better overall ICU mortality

ROX index – Predict outcome of HFNC

➤ Acute respiratory failure d/t Pneumonia., 191 patients

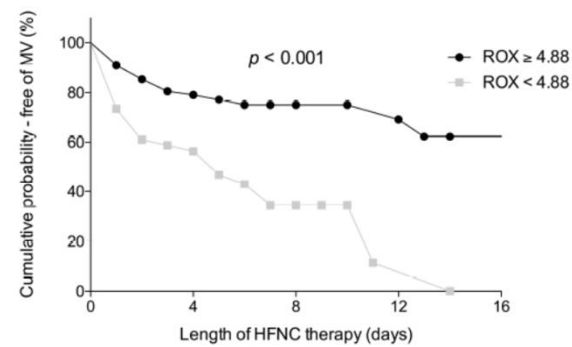
➤ ROX index
$$\frac{\text{SaO}_2 / \text{FiO}_2}{\text{Respiratory rate}}$$

Survival proportions according to ROX index at 2h of HFNC



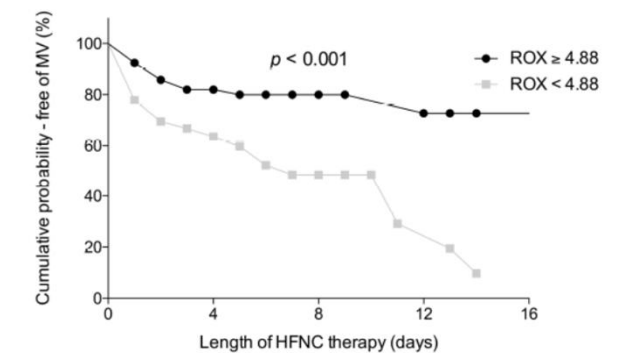
Number at risk					
ROX ≥ 4.88	104	40	11	8	1
ROX < 4.88	73	26	11	4	1

Survival proportions according to ROX index at 6h of HFNC



Number at risk					
ROX ≥ 4.88	126	53	17	12	3
ROX < 4.88	48	16	8	1	0

Survival proportions according to ROX index at 12h of HFNC

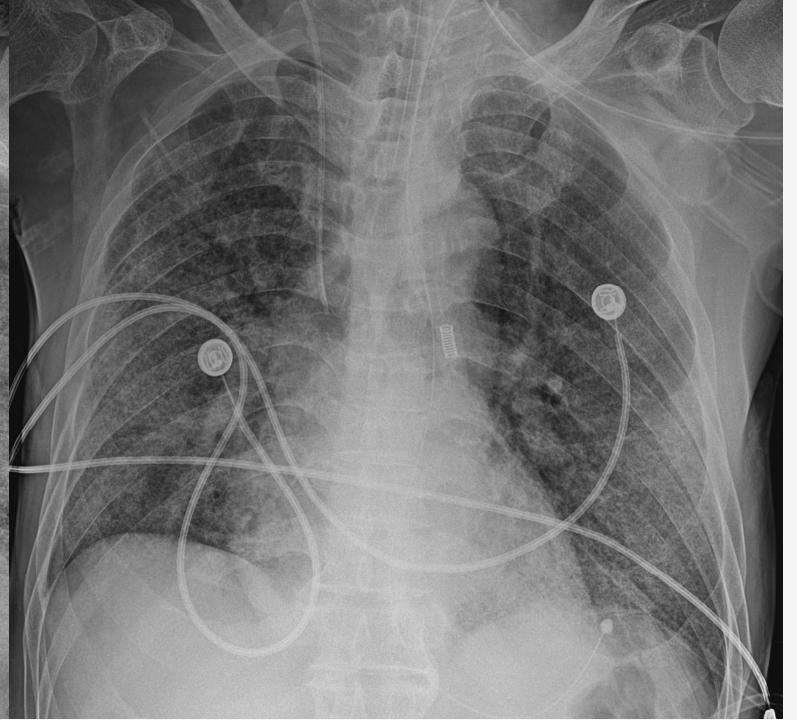
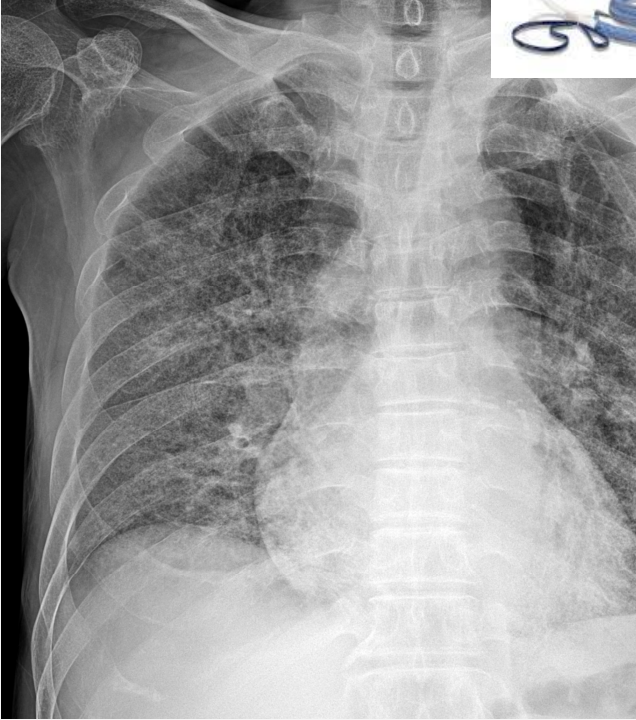


Number at risk					
ROX ≥ 4.88	119	51	13	10	3
ROX < 4.88	38	20	10	3	0

	Flow (L/min)	FiO2	SaO2	RR	ROX index
Day 1	60	0.7	94%	24	2.74
Day 2	60	0.5	96%	20	9.60
Day 3	60	0.8	95%	24	3.16
Day 4	60	0.8	93%	28	2.65
Day 5	60	0.9	91%	34	2.40



Intubation



Risk factor of HFNC failure

- 180 patients, HFNC (n=87)
- First line Treatment of ARDS (n=45)

	All Subjects With ARDS (N = 45)	HFNC Success (n = 27)	HFNC Failure (n = 18)	P
Age, median (IQR), y	57.9 (38.7–74.2)	46.1 (39–75.2)	62.8 (37.1–72.6)	.89
Males, n (%)	22 (49)	13 (48)	9 (50)	.86
At least one comorbid condition, n (%)	27 (60)	14 (52)	14 (78)	.08
Chronic heart failure	5 (11)	2 (7)	3 (17)	.33
COPD	5 (11)	3 (11)	2 (11)	> .99
Neurodegenerative disease	4 (9)	3 (11)	1 (6)	.52
Reason for HFNC oxygen therapy, n (%)				
Pneumonia	36 (80)	23 (85)	13 (72)	.44
ALI of extra-pulmonary origin	4 (9)	2 (7)	2 (11)	.67
Toxic ARDS	5 (11)	2 (7)	3 (17)	.33
At least one associated organ failure, n (%)	20 (44)	7 (26)	13 (76)	.002
Hemodynamic	11 (24)	2 (7)	9 (50)	.001
Kidney	12 (27)	6 (22)	6 (33)	.59
Neurological	4 (9)	0	4 (22)	.01
SAPS II score, median (IQR)	36 (24–44)	29 (22–37)	46 (29–61.5)	.001
Highest breathing frequency, median (IQR), breaths/min	34 (30–40)	33 (30–40)	37 (29–40)	.57
Initial P _a O ₂ /F _{IO₂} , median (IQR)	137 (88.5–208.5)	145.3 (97.5–223.5)	115.3 (84–177.1)	.26
Lowest P _a O ₂ /F _{IO₂} , median (IQR)	108.6 (73–137.1)	124 (93–217)	91.5 (64–129.5)	.02
HFNC oxygen therapy duration of < 24 h, n (%)	22 (49)	10 (37)	12 (67)	.05
Duration of therapy, median (IQR), h	24 (12.5–50)	32 (16–53)	20 (12–31)	.16
ICU stay, median (IQR), d	4 (3–12.5)	3 (2–5)	13.5 (5.5–19)	.001
Alive at ICU discharge, n (%)	35 (78)	26 (96)	9 (50)	.001
Alive at day 28, n (%)	33 (71)	24* (89)	9 (50)	.003

Delayed intubation was associated with higher mortality

- 449 patients with hypoxemia who were receiving NIV

Table 5 Early versus late intubation in patients with a HACOR score of >5 at 1 h of noninvasive ventilation

NIV time points and hospital mortality	Intubation at ≤ 12 h (N = 88)	Intubation at >12 h (N = 175)	p
HACOR score at NIV initiation	8.9 ± 3.9	7.7 ± 3.0	<0.01
HACOR score at 1 h of NIV	11.4 ± 4.0	8.8 ± 3.1	<0.01
HACOR score before intubation	12.2 ± 4.7	11.4 ± 5.0	0.22
Time from NIV initiation to intubation (h)	5 (2–10)	53 (23–132)	<0.01
Hospital mortality	58 (66%)	138 (79%)	0.03

Values in table are presented as the mean $\pm$ SD, the median with the IQR in parenthesis, or as a number with the percentage in parenthesis, as appropriate

HACOR heart rate, acidosis, consciousness, oxygenation, and respiratory rate, NIV noninvasive ventilation

- **NIV failure rate : 58%**
- **Mortality of patients with NIV failure : 75%**
- **Hospital Mortality : Early intubation (66%) vs. Late intubation (79%) p=0.03**

Risk factor of NIV failure

NIV Failure Rate (%)

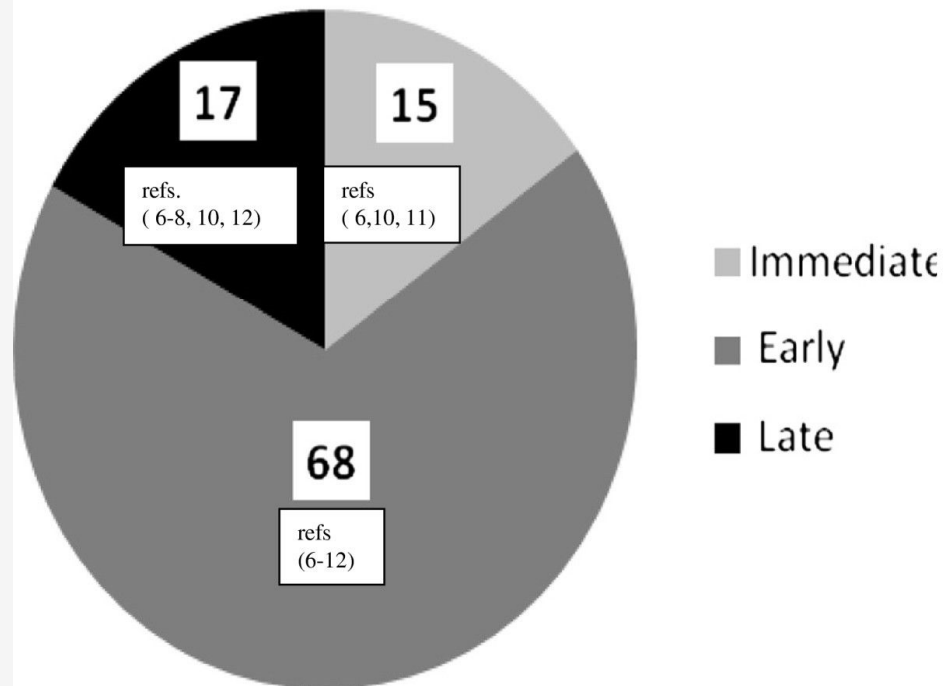


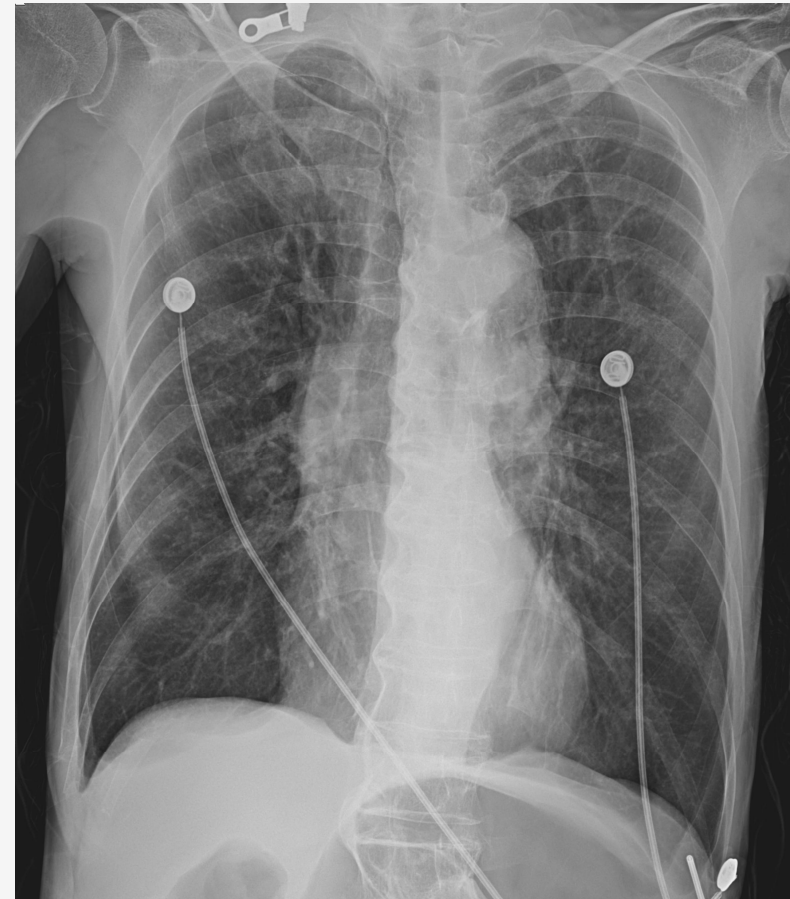
Table 3. Risk Factors for NIV Failure in Patients With Acute Hypoxemic Respiratory Failure

Diagnosis of ARDS or pneumonia
Age > 40 y
Hypotension (systolic blood pressure < 90 mm Hg)
Metabolic acidosis (pH < 7.25)
Low oxygenation index (P_{aO_2}/F_{IO_2})
Simplified Acute Physiology Score II > 34
Failure to improve oxygenation within first hour of NIV ($P_{aO_2}/F_{IO_2} > 175$ mm Hg)

ARDS = acute respiratory distress syndrome
 F_{IO_2} = fraction of inspired oxygen
(Based on data in References 23-25.)

CASE 3

- C/C dyspnea (O : 2 days ago)
- COPD로 외래 추적 중 – ICS/LABA
- URI history (+)
- Wheezing(+)
- Alert mental state



CASE 3

pH	7.23
PCO2	62
PO2	55
Hct	55
Na+	143
K+	3.8
Ca++	1.11
Hgb	17.1
BE-ECF	-1.6
BE-B	-3.2
SBC	21.9
HCO3	26.0
TCO2	27.9
O2 SAT	82
O2 CT	.
nCa++	.
An. gap	.
유산	1.9
포도당(blo	109

➤ Diagnosis

- Acute exacerbation of COPD
- Acute hypercapnic respiratory failure



Indications for NIV in AECOPD

Indications for Noninvasive Mechanical Ventilation (NIV)

Table 5.7

At least one of the following:

- Respiratory acidosis ($\text{PaCO}_2 \geq 6.0 \text{ kPa}$ or 45 mmHg and arterial $\text{pH} \leq 7.35$)
- Severe dyspnea with clinical signs suggestive of respiratory muscle fatigue, increased work of breathing, or both, such as use of respiratory accessory muscles, paradoxical motion of the abdomen, or retraction of the intercostal spaces
- Persistent hypoxemia despite supplemental oxygen therapy

➤ ABGA

✓ **pH 7.23, pCO_2 62mmHg, PO_2 55mmHg, SaO_2 82%**

Comparing NIV and Convention oxygen Therapy

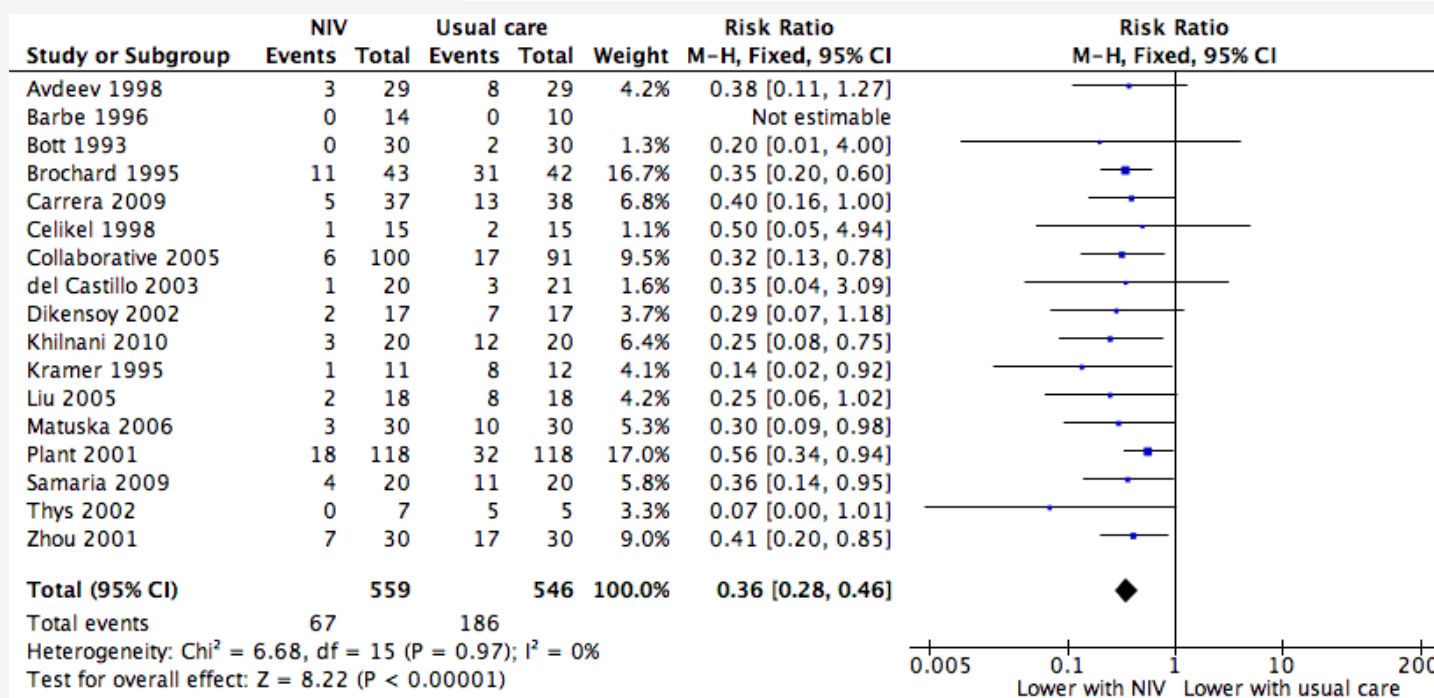
- Prospective multicenter RCT, 14 UK hospitals
- 236 patients, COPD c AE, pH 7.25-35, PaCO₂ > 45 mmHg
- NIV vs. COT (Target SaO₂ 85-90%)

	Standard treatment (n=118)	NIV (n=118)		Standard	NIV	p
Mean age (SD)	69 (8)	69 (7)	Intention-to-treat			
Sex (M/F)	63/55	54/64	Failed	32/118 (27%)	18/118 (15%)	0.02
Respiratory rate per min (range)	28 (24-40)	28 (24-36)	Died	24/118 (20%)	12/118 (10%)	0.05
pH (range)	7.31 (7.26-7.35)	7.32 (7.25-7.35)	Subgroup analysis			
PaCO ₂ kPa (SD)	8.65 (1.70)	8.82 (1.51)	pH<7.30			
PaO ₂ kPa (range)	7.00 (4.71-12.31)	6.88 (4.50-13.8)	Failed	16/38 (42%)	13/36 (36%)	0.64
Ranges in parentheses are median data with 5th and 95th centiles.			Died	13/38 (34%)	8/36 (22%)	0.31
			pH>=7.30			
			Failed	16/80 (20%)	5/82 (6%)	0.01
			Died	11/80 (14%)	4/82 (5%)	0.06

Table 2: **Primary outcome and in-hospital mortality**

Non-invasive ventilation for the management of acute hypercapnic respiratory failure due to exacerbation of chronic obstructive pulmonary disease (Review)

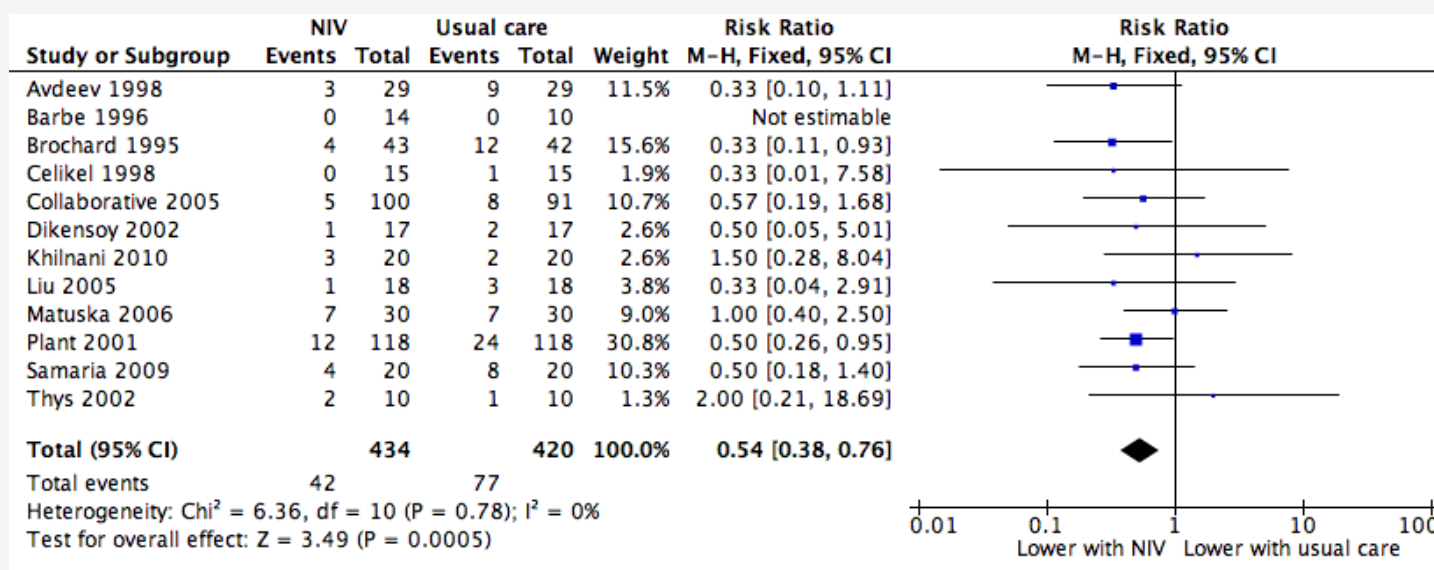
Osadnik CR, Tee VS, Carson-Chahhoud KV, Picot J, Wedzicha JA, Smith BJ



Intubation rate

Non-invasive ventilation for the management of acute hypercapnic respiratory failure due to exacerbation of chronic obstructive pulmonary disease (Review)

Osadnik CR, Tee VS, Carson-Chahhoud KV, Picot J, Wedzicha JA, Smith BJ



Mortality

CASE 2

pH	7.23
PCO2	62
PO2	55
Hct	55
Na+	143
K+	3.8
Ca++	1.11
Hgb	17.1
BE-ECF	-1.6
BE-B	-3.2
SBC	21.9
HCO3	26.0
TCO2	27.9
O2 SAT	82
O2 CT	.
nCa++	.
An. gap	.
유산	1.9
포도당(blo	109



pH	7.32
PCO2	43
PO2	74
Hct	48
Na+	147
K+	2.9
Ca++	1.02
Hgb	14.9
BE-ECF	-3.9
BE-B	-3.9
SBC	21.7
HCO3	22.2
TCO2	23.5
O2 SAT	93
O2 CT	.
nCa++	.
An. gap	.
유산	1.4
포도당(blo	193

CASE 2

2023-01-20 18:25

Sedation 위해 Midazolam 3mg I.V done .

[중재] Intubation함 (E-tube size : 7.5, 삽입길이 : 24 cm,

Ultian 5mg+P/S 50cc I.V infusion start (3cc/hr)

Dextomin 200mcg+P/S 48cc I.V infusion start (3cc/hr). Ventilator applied (CMV mode, FIO2 0.7, RR 17회, VT 360ml, PF 41ml, PEEP 5cmH2O).

2023-01-20 18:20

환자 RR 35~40 회 이상의 tachypnea

보이며 숨쉬기 힘들다고 호소하여 담당의에게 보고함

2023-01-20 16:20

환자 RR 지속적으로 39회 이상 check

며 숨을 몰아쉬는 모습 보임. 모니터상 HR 130 회 이상의 tachycardia 보이며 숨 쉬
게 보고함

2023-01-20 16:00

[중재] 환자 상태 확인함. 의식상태 확인함 (변화 : 없음, 상태 : Alert).

BIPAP유지중임 (FIO2 0.3, IT 0.85, IPAP 21, EPAP 6).

[중재] SpO2 확인함 (SpO2 : 92 %).

[사정] Dyspnea 있음 (증상 정도 : moderate, RR : 37 회/min). 부착물 유지중임 (종류 : Foley cath, A-line(Rt.radial), Lt.arm & Rt.leg Pph I.V line). 욕창중재 활동을 시행함.

[중재] 낙상의 가능성을 증가시킬 수 있는 환경을 사정하고 조정함. 낙상의 가능성이 큰 환자의 신체적 결손을 파악하고 사정함.

Monitoring of NIV failure

Table 4. Monitoring Capabilities Recommended for NIV in Acute Respiratory Failure

Continuous observation from central monitoring area

Frequent checks of:

Comfort

Tolerance

Mask fit

Air leak

Patient-ventilator synchrony

Vital signs, especially respiratory rate

Accessory muscle use

Ventilator tidal volume (aim for 6–7 mL/kg)

Continuous telemetry

Electrocardiogram trace

Oximetry

Blood gas values at baseline, after 1–2 h, and as indicated

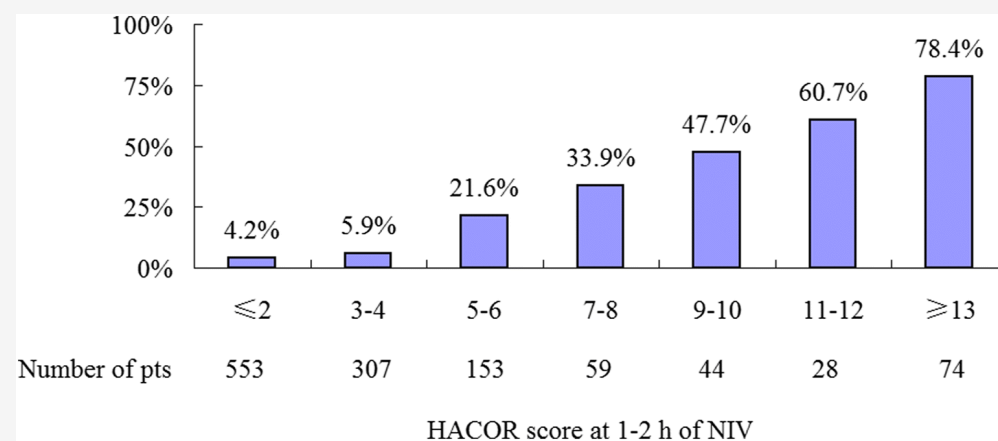
NIV = noninvasive ventilation

Signs of treatment failure of NIV

- Worsening or lack of improvement in gas exchange
- Signs of respiratory muscle fatigue
- Feeling of unbearable dyspnea
- Development of respiratory acidosis
- Presence of unmanageable tracheal secretions
- Hemodynamic instability

Prediction of NIV failure

Variable	Value	Score
HR	≤ 120	0
	≥ 121	1
pH	≥ 7.35	0
	7.30-7.34	2
	7.25-7.29	3
	< 7.25	4
Glasgow	15	0
	13-14	2
	11-12	5
	≤ 10	10
PaO ₂ /FiO ₂	>201	0
	176-200	2
	151-175	3
	126-150	4
	101-125	5
	≤ 100	6
RR	≤ 30	0
	31-35	1
	36-40	2
	41-45	3
	≥ 46	4



Prediction of in-hospital mortality in AHRF requiring NIV

NIVO score	Points
Chest radiograph consolidation	1
Glasgow Coma Scale ≤ 14	1
Atrial fibrillation	1
pH < 7.25	1
Time to acidaemia > 12 h	2
eMRCD 5a	2
eMRCD 5b	3
Total	9

TABLE 4 In-hospital and 90-day mortality by Noninvasive Ventilation Outcomes (NIVO) score increment and risk category

	Patients n	In-hospital mortality	90-day mortality
NIVO score			
0	67	0	10.4
1	79	8.9	20.3
2	133	5.3	15.8
3	152	15.1	26.3
4	116	19.0	40.5
5	97	35.1	46.4
6	54	53.7	59.3
7	26	65.4	76.9
8	8	87.5	87.5
9	1	100	100
Total	733	20.1	32.2
Risk category			
Low (0–2)	279	5.0	15.8
Medium (3–4)	268	16.8	32.5
High (5–6)	151	41.2	50.1
Very high (7–9)	35	71.4	80.0

CASE 4

- 기저로 COPD를 가지고 있는 80세 남자 환자가 호흡곤란을 주소로 내원하였다.
내원시 양폐로 천명음(wheezing)이 들렸다. 의식은 명료하였다.
- ABGA
 - RA state : pH 7.38 pCO₂ 72mmHg, PO₂ 43mmHg, SaO₂ 78%
- IV steroid 및 IV antibiotics 치료를 시작하였다.
- 다음으로 선택할 수 있는 Oxygen Therapy는?



Indications for NIV in AECOPD

Indications for Noninvasive Mechanical Ventilation (NIV)

Table 5.7

At least one of the following:

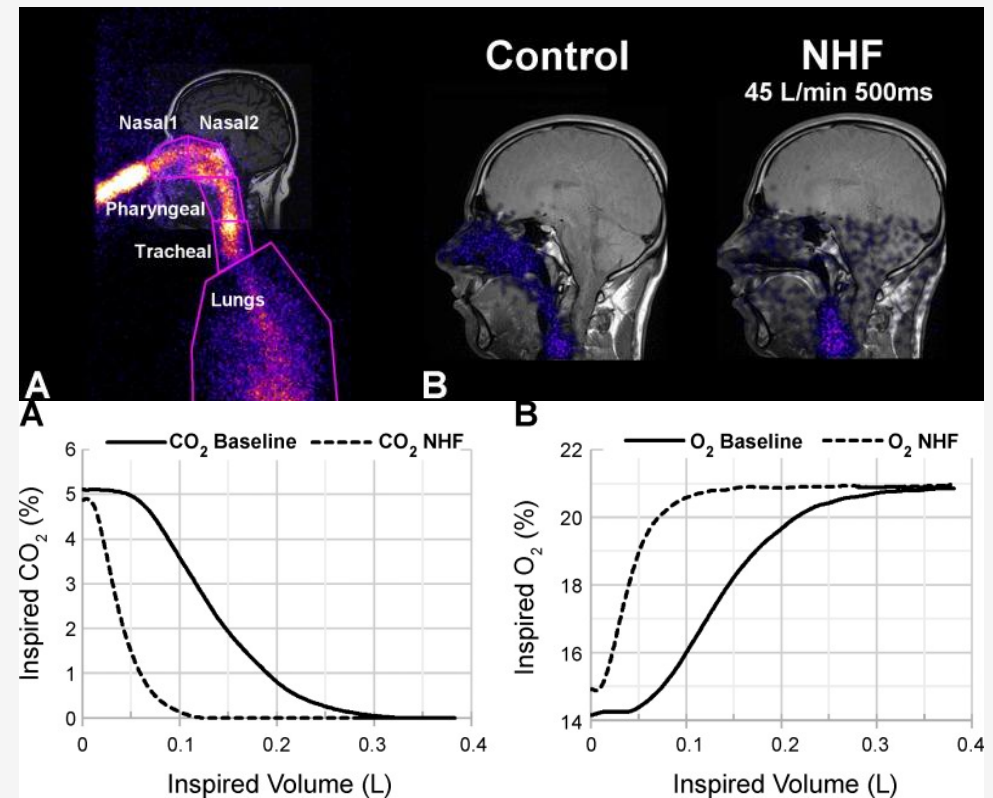
- Respiratory acidosis ($\text{PaCO}_2 \geq 6.0 \text{ kPa}$ or 45 mmHg and arterial $\text{pH} \leq 7.35$)
- Severe dyspnea with clinical signs suggestive of respiratory muscle fatigue, increased work of breathing, or both, such as use of respiratory accessory muscles, paradoxical motion of the abdomen, or retraction of the intercostal spaces
- Persistent hypoxemia despite supplemental oxygen therapy

➤ ABGA

- **pH 7.38 pCO₂ 72mmHg**, PO₂ 43mmHg, SaO₂ 78%

Mechanism of HFNC in COPD patients

- Improve mucociliary clearance
- Reduce dead space
- Deliver low level of airway pressure
- Comfort



HFNC may be useful in COPD

	Total patients (n = 81)	Hypercapnia group (n = 46)	Nonhypercapnia group (n = 35)
Males, n (%)	46 (56.8)	18 (39.1)	28 (80.0)
Age, y ($\pm$ SD)	69.5 ($\pm$ 14.7)	69.7 ($\pm$ 13.6)	69.2 ($\pm$ 16.3)
Diabetes mellitus, n (%)	20 (24.7)	10 (21.7)	10 (28.6)
Hypertension, n (%)	29 (35.8)	17 (37.0)	12 (34.3)
Chronic lung diseases, n (%)	49 (60.5)	35 (76.1)	14 (40.0)
Chronic obstructive pulmonary disease	42 (85.7)	33 (94.3)	9 (64.3)
Lung cancer	3 (6.1)	0 (0)	3 (21.4)
Asthma	2 (4.1)	2 (5.7)	0 (0)
Interstitial lung disease	2 (4.1)	0 (0)	2 (14.3)
Causes of respiratory failure			
Acute exacerbation of chronic obstructive pulmonary disease	38 (46.9)	30 (65.2)	8 (22.9)

Table 2
Arterial blood gas analysis an

	Hypercapnia group (n = 46)		
	Before-HFNC ABG	After-HFNC ABG	P
Paco ₂ , mm Hg ($\pm$ SD)	73.2 ($\pm$ 20.0)	67.2 ($\pm$ 23.4)	.02
Pao ₂ , mm Hg ($\pm$ SD)	71.6 ($\pm$ 40.0)	85.0 ($\pm$ 34.7)	.02
pH ($\pm$ SD)	7.28 ($\pm$ 0.08)	7.31 ($\pm$ 0.08)	<.01
HCO ₃ ⁻ , mmol/L ($\pm$ SD)	32.9 ($\pm$ 7.0)	32.0 ($\pm$ 7.9)	.21
SpO ₂ , % ($\pm$ SD)	82.5 ($\pm$ 17.2)	91.9 ($\pm$ 7.4)	<.01
Respiratory rate	24.7 ($\pm$ 5.8)	23.6 ($\pm$ 5.2)	.03

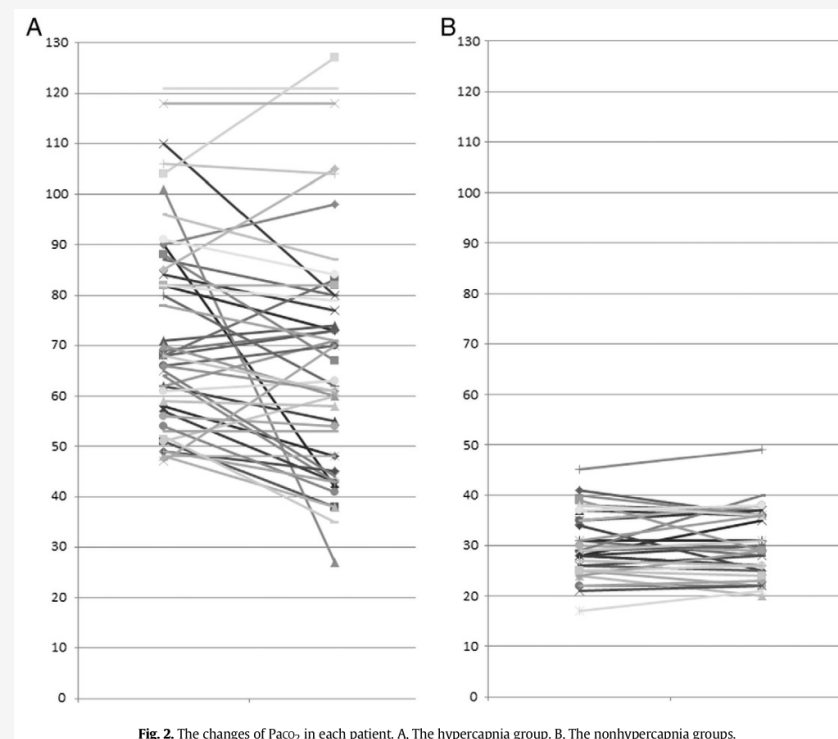
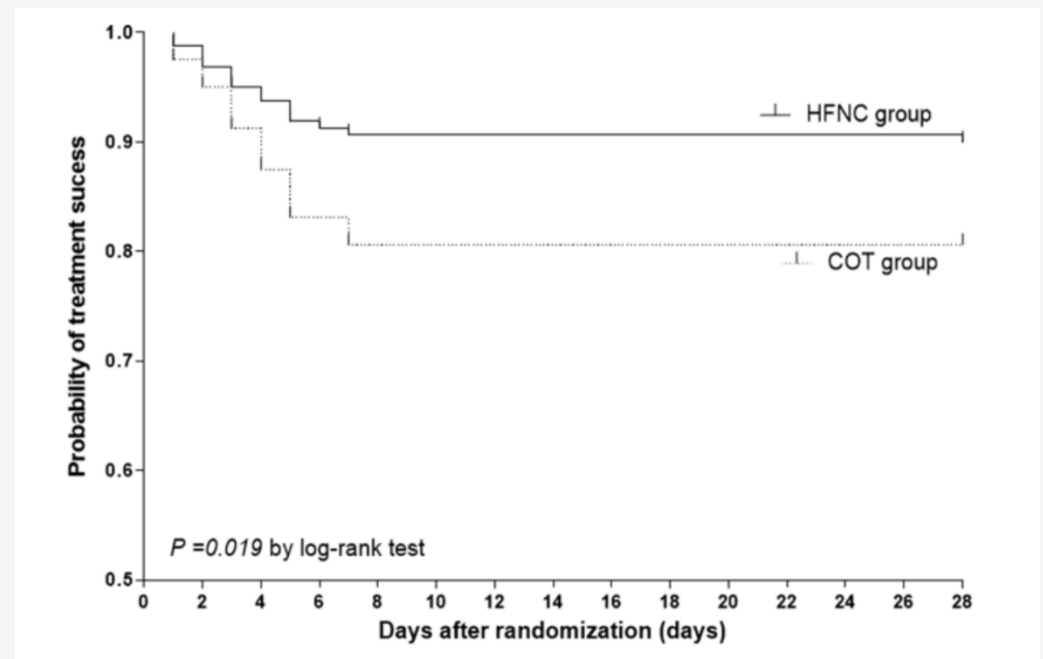


Fig. 2. The changes of Paco₂ in each patient. A, The hypercapnia group. B, The nonhypercapnia groups.

HFNC may reduce the need to intubation

- Prospective RCT, 320 patients, China
- Patients with AECOPD with $\text{pH} \geq 7.35$, $\text{PaO}_2 < 60$ mmHg, and $\text{PaCO}_2 > 45$ mmHg

Outcome	HFNC	COT	P
Treatment failure	16 (10.0%)	31 (19.4%)	0.026
Patients received NIV	13 (8.1%)	26 (16.3%)	0.039
CAT scores at discharge	12.0 ± 7.6	17.0 ± 7.3	0.002
Subjective discomfort score	2.3 ± 1.1	3.9 ± 1.6	<0.001
Reason for NIV			0.050
Respiratory acidosis	12 (7.5)	24 (15.0)	
Obvious dyspnea	4 (2.5)	7(4.4)	



HFNC may be useful in AECOPD

- Not tolerated NIV, COPD c AE, $\text{pH} \leq 7.38$ and $\text{PaCO}_2 > 45\text{mmHg}$
- 38 patients, Initiation setting : Flow 50L/min

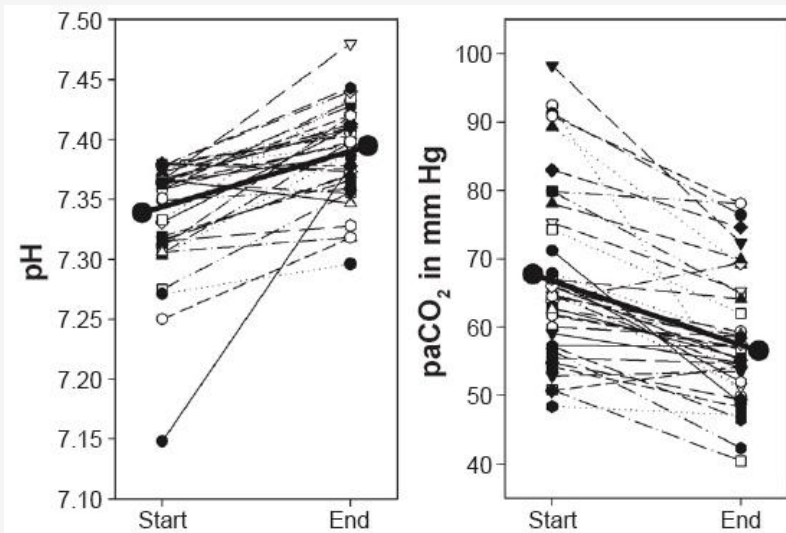


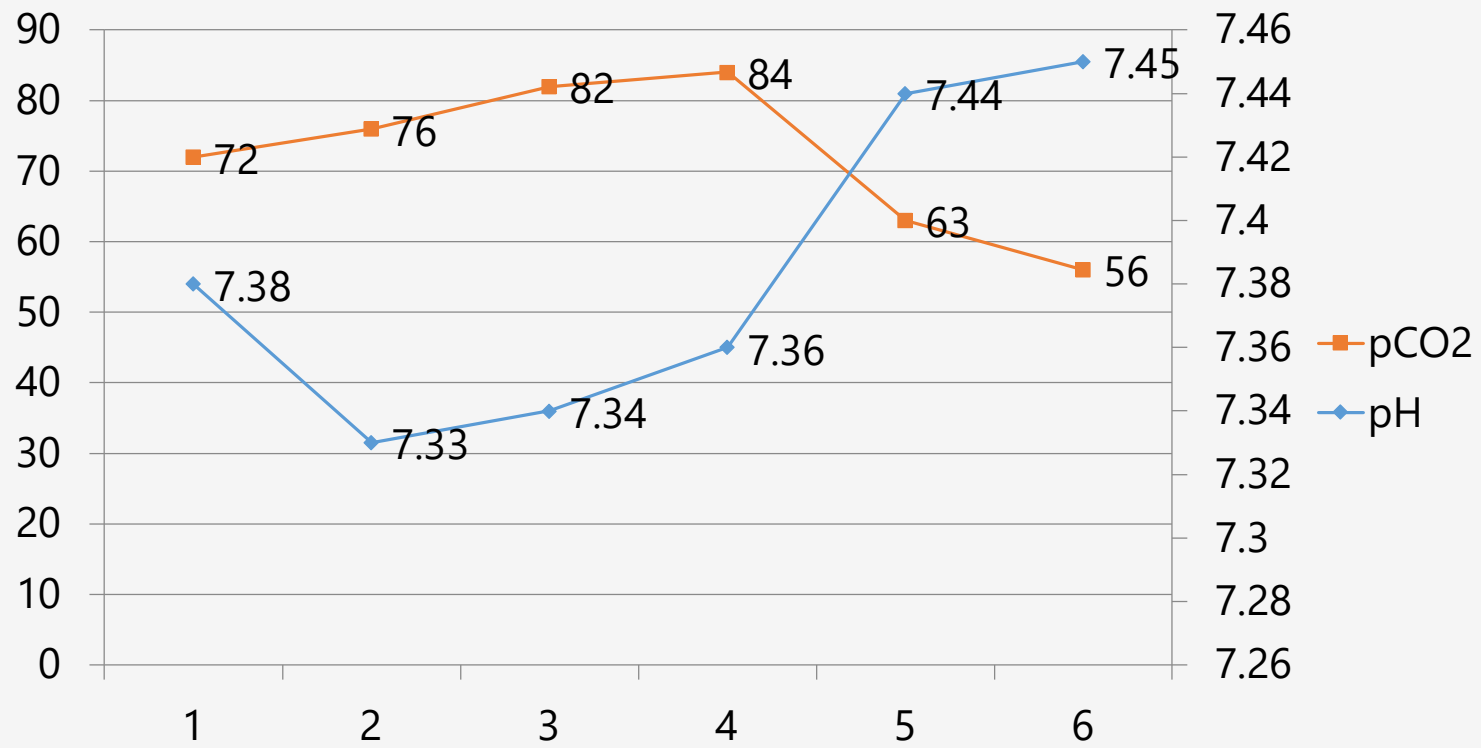
Figure 1 Changes in pH and PaCO_2 in all patients; the bold line marks mean values; $P < 0.05$.

Table 1 Parameters during NHF treatment

Parameters	All patients ($\text{pH} \leq 7.38$; $n=38$)			
	Baseline (mean $\pm$ SD)	End (mean $\pm$ SD)	Change (mean $\pm$ SD)	P-value
pH	7.339 $\pm$ 0.041	7.392 $\pm$ 0.048	0.052 $\pm$ 0.048	0.000
pCO_2 mmHg	67.6 $\pm$ 12.9	58.5 $\pm$ 9.7	-9.1 $\pm$ 8.8	0.001
pO_2 mmHg	58.3 $\pm$ 15.2	58.3 $\pm$ 17.6	-	0.983
SO_2 %	85.3 $\pm$ 9.5	86.2 $\pm$ 11.3	-	0.798
HCO_3 mmol/L	30.8 $\pm$ 5.1	31.6 $\pm$ 5.3	-	0.636
Base excess mmol/L	6.9 $\pm$ 4.9	7.7 $\pm$ 5.4	-	0.633
NHF flow L/min	-	25.8 $\pm$ 8.2	-	-
Treatment time min	-	195 $\pm$ 231	-	-
Patients with $\text{pH} < 7.35$ ($n=17$)				
pH	7.298 $\pm$ 0.046	7.379 $\pm$ 0.051	0.082 $\pm$ 0.060	0.000
pCO_2 mmHg	73.7 $\pm$ 13.7	59.6 $\pm$ 11.2	-14.2 $\pm$ 10.6	0.002
pO_2 mmHg	56.9 $\pm$ 14.8	55.6 $\pm$ 20.7	-	0.845
SO_2 %	84.8 $\pm$ 10.1	84.3 $\pm$ 11.9	-	0.904
HCO_3 mmol/L	31.6 $\pm$ 5.5	32.5 $\pm$ 5.6	-	0.661
Base excess mmol/L	7.4 $\pm$ 5.3	8.5 $\pm$ 5.7	-	0.591
NHF flow L/min	-	26.3 $\pm$ 7.9	-	-
Treatment time min	-	252 $\pm$ 251	-	-

Notes: Changes in blood gas analyses data, usage time and flow rates during NHF therapy in all patients and in the subgroup with $\text{pH} < 7.35$. All data are organized as mean $\pm$ SD. "-" indicates no significant differences to report.

Abbreviation: NHF, nasal high-flow.



HFNC (Flow 60L/min, FiO₂ 60%)

Risk factor Failure of HFNC in COPD patients

Table 4 Comparison of Physiological Parameters Between HFNC Success and Failure Groups

Characteristic	HFNC Group	Baseline	24 h	48 h	72 h	120 h	p ^a
pH	Success	7.38±0.05	7.40±0.04	7.40±0.04	7.41±0.04	7.41±0.04	0.253
	Failure	7.40±0.04	7.38±0.04	7.39±0.02	7.38±0.06	7.37±0.07	0.197
	p ^c	0.281	0.334	0.756	0.018	0.030	0.050 ^b
PaO ₂ (mmHg)	Success	54.8±4.8	72.1±8.7	70.6±14.0	71.3±15.5	71.2±16.4	0.000
	Failure	54.9±5.0	71.2±11.7	68.1±10.3	68.9±15.2	78.2±25.1	0.487
	p ^c	0.518	0.786	0.649	0.629	0.384	0.878 ^b
PaCO ₂ (mmHg)	Success	54.6±6.9	52.4±8.0	51.5±5.6	52.2±7.2	50.9±8.9	0.000
	Failure	58.0±8.9	59.0±11.8	55.8±5.6	63.8±11.0	68.7±13.2	0.144
	p ^c	0.086	0.006	0.012	0.006	0.000	0.000 ^b
SaO ₂ (%)	Success	89.1±4.7	93.3±2.7	93.3±2.2	93.1±3.3	93.7±2.6	0.000
	Failure	90.5±4.8	92.9±1.9	92.3±2.5	92.2±4.1	95.4±4.2	0.160
	p ^c	0.295	0.592	0.237	0.460	0.179	0.628 ^b
Respiratory rate (bpm)	Success	20.1±1.5	21.3±2.3	22.4±5.9	21.8±2.8	21.4±2.4	0.013
	Failure	22.4±2.5	22.7±2.6	22.0±2.8	23.0±2.4	21.5±1.9	0.896
	p ^c	0.032	0.059	0.829	0.205	0.936	0.784 ^b

Risk factor of HFNC failure in AECOPD

TABLE 2.

Changes in Respiratory Rate, $Paco_2$, and pH After 1 Hour of High-Velocity Nasal Insufflation According to the Result of the Therapy

Variable	Success			Failure		
	Baseline	1 hr HVNI	<i>p</i>	Baseline	1 hr HVNI	<i>p</i>
Respiratory rate (breaths/min) ^a	29 (28–31)	22 (20–24)	< 0.001	30 (25–31)	23 (22–29)	0.379
$Paco_2$ (mm Hg) ^a	56 (51–67) →	51 (44–59)	< 0.001	66 (56–77) →	72 (68–78)	0.392
pH ^{a,b}	7.32 (7.28–7.34)	7.37 (7.34–7.40) ^b	0.006	7.31 (7.29–7.32)	7.31 (7.28–7.35) ^b	0.56

HVNI = high-velocity nasal insufflation.^aValues expressed as median (interquartile range).^b*p* = 0.022 for pH at the time of HVNI therapy (Mann-Whitney *U* test).

HFNC vs. NIV in COPD Patients

- 65 patients
- NIV IPAP 10-20 cmH₂O FiO₂ 1.0
- HFNC 35 L/min, FiO₂ 1.0

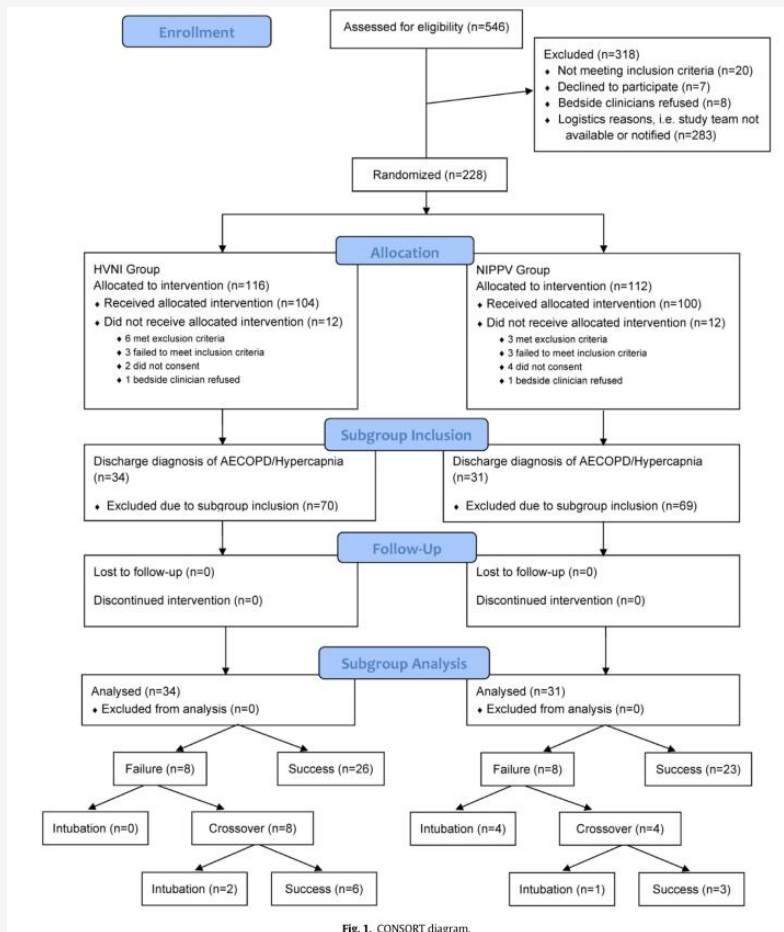


Fig. 1. CONSORT diagram.

Table 2
Therapy failure.

	HVNI, n = 34	NIPPV, n = 31	p-value
Intubation at 72 h, n (%)	2 (5.9%)	5 (16.1%)	0.244
Arm failure at 72 h, n (%)	8 (23.5%)	8 (25.8%)	1.000
Crossover at 72 h, n (%)	8 (23.5%)	4 (12.9%)	0.346

Data is reported as number (%). Statistical significance ($p < 0.05$) determined by Fisher's Exact Test.

Indications for Invasive mechanical ventilation in AECOPD

Indications for Invasive Mechanical Ventilation

Table 5.8

- Unable to tolerate NIV or NIV failure
- Status post-respiratory or cardiac arrest
- Diminished consciousness, psychomotor agitation inadequately controlled by sedation
- Massive aspiration or persistent vomiting
- Persistent inability to remove respiratory secretions
- Severe hemodynamic instability without response to fluids and vasoactive drugs
- Severe ventricular or supraventricular arrhythmias
- Life-threatening hypoxemia in patients unable to tolerate NIV

Summary

- Respiratory failure : lung failure, pump failure
- Acute hypoxemic respiratory failure
 - ✓ HFNC > COT or NIV
 - ✓ NIV : Acute cardiogenic pulmonary edema
- Acute hypercapnic respiratory failure
 - ✓ NIV > HFNC, COT
 - ✓ HFNC : hypercapnic AECOPD without respiratory acidosis

가...감사합니다!

넘죽

