ORIGINAL ARTICLE

Non-invasive ventilation as a first-line treatment for acute respiratory failure: "real life" experience in the emergency department

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Objective: To describe our experience with non-invasive ventilation (NIV) for patients with acute respiratory failure (ARF) in the emergency department (ED).

Methods: A prospective/retrospective, observational study on 190 patients with ARF (mean $\pm$ SD age 72.2 ± 12.9 years, mean APACHE II score 18.9 ± 5.9), who received 200 NIV trials in an ED. We analysed the NIV register data (prospectively collected) and medical records (retrospective data abstraction) and evaluated clinical indications for NIV, patient outcomes, and predictive factors for success and death. NIV success was defined as tolerance of the procedure and no need for endotracheal intubation (ETI).

Results: Main indications to NIV were cardiogenic pulmonary oedema (CPE) (70 trials), acute exacerbation of COPD (39), both CPE and acute exacerbation of COPD (11), pneumonia (48), decompensation of obesity/hypoventilation (6), other conditions (26). The procedure was successful in 60.5% of trials. Global mortality was 34.5%, similar to the APACHE II predicted mortality of 32%. ETI rates were 6.5% and tracheostomy rates 1%. The improvement of pH within six hours after NIV initiation was predictive of survival in the hypercapnic group.

Conclusions: Our results confirm the global efficacy of NIV in an ED setting, and show that, in spite of lower success rate in "real practice" in comparison with RCTs, an intermediate care unit can represent an appropriate and less expensive setting to perform this technique. The low rate of ETI seems to be because of the high number of patients for whom NIV was used as "ceiling" treatment.

Non-invasive ventilation (NIV) consists of mechanical breaths delivered by a tightly fitting nasal or facial mask or a helmet instead of an artificial airway, such as an endotracheal or tracheostomy tube. A number of prospective randomised controlled trials (RCTs)¹⁻⁷ and meta analyses⁸⁻¹⁰ has demonstrated that NIV is effective and safe for selected patients with acute respiratory failure (ARF). When ARF is because of acute exacerbations of chronic obstructive pulmonary disease (COPD), NIV is now considered standard of treatment.^{11, 12} In patients with hypoxemic cardiogenic pulmonary oedema (CPE) resistant to standard medical treatment, continuous positive airway pressure (CPAP) (which is here considered a mode of NIV even if it is not assisted ventilation) has been shown to reduce intubation rate and accelerate the recovery.^{13, 14} Non-invasive pressure support ventilation seems to reduce intubation rate in the subgroup of CPE patients with hypercapnia.¹⁵ Patients with ARF because of other diseases and conditions have been treated with NIV with some benefit,¹⁶⁻¹⁸ so extending the indications for this ventilatory technique. However, RCTs show that NIV has beneficial effects in the setting of a research study, which are not automatically reproducible in the real clinical practice, where less selected populations of patients are treated and less intensive care and monitoring can be provided, except for an intensive care unit (ICU). In addition, although in randomised trials reductions of intubation and death rates are considered primary clinical goals, these may not always be considered realistic objectives in everyday clinical practice. Actually, palliation, comfort, and relief of dyspnoea can represent acceptable end points of NIV for patients with poor prognosis or who refuse advanced life support. For these patients, NIV can be utilised as "ceiling" treatment. In this study, we describe the experience of our

emergency department (ED) with NIV as first-line treatment in patients with ARF and analyse the outcomes in relation to the clinical goals previously established for each individual patient.

METHODS

This study was conducted from January 2002 to March 2003 in an ED of a university-affiliated urban hospital with 1200 beds, which includes a 19 bed high dependency unit (HDU). The unit provides acute care and monitoring for critically ill or high risk patients who are admitted from the emergency ward and who do not immediately require or are not otherwise candidates for invasive mechanical ventilation. It provides an acute NIV service, which is also available for inpatients who develop ARF while staying in surgical or medical wards for other reasons. Emergency physicians manage the unit, rotating between it and the emergency ward approximately every three months. The medical HDU staff are divided in two teams, each composed of one full time senior physician, three full time junior doctors, and two residents, each rotating over two 6 hours and one 12 hour shifts. All physicians have been formally trained in the delivery of NIV and in airway management, and a rapid access to endotracheal intubation (ETI) and invasive

Abbreviations: APACHE II, Acute Physiology and Chronic Health Evaluation II score; ARF, acute respiratory failure; CAP, community acquired pneumonia; COPD, chronic obstructive pulmonary disease; CPAP, continuous positive airway pressure; CPE, cardiogenic pulmonary oedema; ECG, electrocardiogram; ED, emergency department; EPAP, expiratory positive airway pressure; ETI, endotracheal intubation; HDU, high dependency unit; ICU, intensive care unit; IPAP, inspiratory positive airway pressure; NIV, non-invasive ventilation; PEEP, positive end expiratory pressure; RCT, randomised controlled trials

mechanical ventilation is guaranteed without delay 24 hours a day. However, the experience and expertise in NIV administration of medical staff are highly variable. In addition, because of a large shortage of nurses during 2002, nurse:patient ratio was 1:4 in the first shift, down to 1:6 or 1:9 in afternoon and nocturnal shifts, which sometimes reduces nursing care and monitoring capability to critical levels. No dedicated respiratory therapist was available in the unit during the period of study. Adherence to the practical guidelines of American Thoracic Society¹⁹ and British Thoracic Society²⁰ was generally maintained on NIV administration. Criteria for initiating NIV were the presence of moderate to severe dyspnoea plus either $p\text{CO}_2 > 45$ mm Hg with $\text{pH} < 7.35$ or $\text{PaO}_2 < 60$ mm Hg with FiO_2 up to 0.6. In patients with hypercapnic ARF because of chest wall deformity, neuromuscular disorder, morbid obesity, or decompensated obstructive sleep apnoea, a NIV trial could be started even in absence of respiratory distress, as these patients often present with depressed mental status associated to abnormal gas exchange. The critical pathway of our hospital allows patients presenting to the ED with ARF (or developing it while in hospital wards) to be evaluated by both an ICU specialist and an ED staff member, who together make a decision about the need for invasive or non-invasive ventilatory assistance and where to admit the patient: in ICU or in HDU. Usually, a NIV trial is offered to each patient with ARF who has no need for immediate ETI or who declines it or is not otherwise a candidate for ETI because of poor prognosis of underlying disease or advanced age. Whenever possible, a management plan is made before initiating a NIV trial about what to do in case of a NIV trial failure—that is, either to intubate and mechanically ventilate the patient or to consider the NIV trial as a “ceiling” treatment, taking into account the stage of underlying disease, the patient or family’s wishes about advanced life support, and the available beds in ICU. During the study, NIV was provided initially in the emergency room and continued there as long as a HDU bed became available. Because of the limited availability of HDU beds, the length of stay in the emergency room could prolong for many hours.

Most of the time, Ventilators Vision (Respironics, Murrysville, PA, USA) were used, which permitted delivery of known concentrations of oxygen and direct monitoring of pressure, volume, and flow during ventilation. Conventional ICU ventilators (Evita 2, Draeger, Lubeck, Germany or Horus Taema, France) were used when the synchronisation between patient and ventilator was difficult or when the clinician believed the patient to be at high risk of NIV failure. Facial masks were used either exclusively or in the period of stabilisation and adapting to NIV. Some patients who were able to breath with closed mouth could subsequently use nasal masks. The bilevel pressure support was the most commonly used mode of ventilation; CPAP was used for patients with pulmonary oedema without hypercapnia. In hypercapnic patients, NIV protocol consisted in setting inspiratory positive airway pressure (IPAP) and expiratory positive airway pressure (EPAP) at low levels at the beginning of procedure—for example, 8/3 or 10/4 cm H_2O —and then in increasing pressure support up to obtain a tidal volume of about 7 ml/kg, a respiratory rate less than 25 breaths/min, and patient comfort. EPAP was increased to counterbalance intrinsic positive end expiratory pressure (PEEP) and improve inspiratory triggering. In hypoxemic patients, CPAP was set initially at 5–6 cm H_2O and gradually increased up to obtain haemoglobin saturation $> 90\%$ with $\text{FiO}_2 < 60\%$. Generally, each patient was subjected to monitoring of electrocardiogram (ECG), blood pressure, heart rate, respiratory rate, SpO_2 , arterial blood gas levels, and clinical status during the first hours of ventilation and

subsequently depending on severity of ARF and availability of equipment. Central venous and peripheral artery accesses were also obtained for selected patients who required more intensive monitoring.

Two of us (CA and FM) analysed patients’ data, which were in part prospectively collected from the NIV register of the unit (demographic data, baseline physiological variables, arterial blood gas data, NIV mode) and in part retrospectively obtained from the medical records through data abstraction forms (indication to and duration of NIV, pressure and FiO_2 settings, outcomes such as need for endotracheal intubation, HDU and hospital mortality, HDU and hospital length of stay). The competent Hospital Authority approved access to patient medical records and use of data. Every effort was made to understand the clinical indications for NIV and the underlying disease(s), based on emergency physician’s notes, progressive notes of the HDU physician in charge, and final diagnosis on medical record at discharge. The success of NIV trial was defined as tolerance of the procedure and avoidance of ETI. Failure of a NIV trial was defined as either intolerance of the procedure or need for ETI, which was the presence of any of the following criteria within 14 days of admission: a) $\text{pH} < 7.20$; b) $\text{pH} 7.20\text{--}7.25$ on two occasions one hour apart; c) hypercapnic coma; d) $\text{PaO}_2 < 45$ mm Hg despite of maximal tolerated FiO_2 , and e) cardiorespiratory arrest.⁵ If any of these criteria was met, the patient could be offered ETI or continued medical treatment with or without NIV, depending on patient and family’s wishes and the attending physician and ICU specialist’s combined evaluation in relation to the goals of treatment.

We also tried to estimate the predictive value of some factors (chosen a priori), which other authors have suggested being associated with NIV success or failure.^{21 22 17}

STATISTICAL ANALYSIS

The descriptive analysis was performed for the overall NIV population and for single categories of disease. Results are presented as absolute values and frequencies, means and standard deviations, or medians and interquartile ranges. Predictive factors for response to NIV and for death were identified by univariate analysis: independent variables included Acute Physiology and Chronic Health Evaluation (APACHE) II score, arterial blood gas values (basal pH and $p\text{CO}_2$), and level of consciousness, adjusted for sex and age. As predictors of death, we have also considered the absence of improvement of $p\text{CO}_2$ and of pH within six hours after NIV initiation. Predictors are expressed as odd ratios (OR) and their 95% confidence intervals (CI). Variables that were univariately associated with either response to NIV or death ($p < 0.2$) were considered for the final stepwise logistic regression. Analysis was performed using STATA version 8.0 (Stata, College Station, Texas, USA, 2003).

RESULTS

Altogether, 190 patients (93 women) with a mean ($\pm$ SD) age of 72.2 ± 12.9 years and a mean APACHE II score of 18.9 ± 5.9 received a NIV trial for 200 episodes of ARF during the study period. Eight patients received two trials and one patient received three trials for distinct episodes of ARF. Baseline characteristics of global population on admission are shown in table 1.

Main clinical indications to NIV and characteristics of each indication group are reported in table 2.

The ventilation modes were bilevel pressure support (172 trials) and CPAP (25 trials). Two patients who started with CPAP were shifted to bilevel positive airway pressure ventilation because of unsuccessful CPAP. In one patient, the ventilation mode was not specified. Median pressure settings were IPAP 16 cm H_2O (interquartile range

Table 1 Baseline characteristics of 190 patients with acute respiratory failure (ARF) treated with non-invasive ventilation (NIV)

Characteristics	Number $\pm$ SD (or %)*
Mean age $\pm$ SD (years)	72.2 $\pm$ 12.9
Number (%) female	93 (48.9%)
Clinical parameters on admission:	
Mean APACHE II score on admission (n = 191 trials)	18.9 $\pm$ 5.9
Systolic blood pressure (mm Hg): mean $\pm$ SD	138.1 $\pm$ 32.1
Diastolic blood pressure (mm Hg): mean $\pm$ SD	82.1 $\pm$ 17.2
Heart rate (bpm): mean $\pm$ SD	103.9 $\pm$ 21.1
Respiratory rate (bpm): mean $\pm$ SD	33 $\pm$ 9
Mental status, number (%) of patients (n = 199 trials)	
Alert	137 (68.8%)
Slightly decreased level of consciousness	47 (23.6%)
Severely decreased level of consciousness	15 (7.5%)
Median basal arterial blood gas value (IQR)	
pH	
All patients (180/200 trials)	7.27 (7.20–7.34)
Hypercapnic patients (143/144 trials)	7.26 (7.18–7.30)
Non-hypercapnic patients (37/37 trials)	7.39 (7.32–7.44)
PaCO ₂ mmHg	
All patients (181/200 trials)	64 (47.4–75.6)
Hypercapnic patients (144/144 trials)	70 (57–78)
Non-hypercapnic patients (37/37 trials)	37 (32.8–41)
PaO ₂ mmHg	
All patients (178/200 trials)	57.4 (46–72)
Hypercapnic patients (141/144 trials)	57 (46–72)
Non-hypercapnic patients (37/37 trials)	60 (48–69.4)
PaO ₂ /FiO ₂	
Non-hypercapnic patients (17/37 trials)	193 (115–223)
Patients coming from	
Emergency room	157 (78.5%)
Medical ward	26 (13%)
Surgical ward	17 (8.5%)
Clinical indication to NIV, number (and %) of trials	
CPE	70 (35%)
COPD exacerbation	39 (19.5%)
CPE and COPD exacerbation	11 (5.5%)
Pneumonia and COPD exacerbation	21 (10.5%)
Pneumonia	27 (13.5%)
Obesity/hypoventilation decompensation	6 (3%)
Others†	26 (13%)

SD, standard deviation; IQR, interquartile range; COPD, chronic obstructive pulmonary disease; CPE, cardiogenic pulmonary oedema.

*Unless stated otherwise.

†Others: pleural effusion (4), amyotrophic lateral sclerosis (2), postoperative pulmonary failure (6), lung cancer (4), pulmonary embolism (4), cardiogenic shock (2), septic shock (2), hemophthisis (1), and multiple myeloma (1).

(IQR 14–18), EPAP 6 cm H₂O (IQR 5–6), and CPAP 10 cm H₂O (IQR 8–10). Median NIV duration was 24 hours (IQR 6–72) and median FiO₂ was 0.4 (IQR 0.34–0.5).

Based on the previous definition of success, 121 NIV trials (60.5%) were successful and 113 patients were discharged at home alive. Eight patients were considered to benefit from the technique, but successively died because of their

underlying diseases (four because of advanced neoplastic disease, two because of acute myocardial infarction, one because of cerebral stroke, and one died 79 days after the NIV trial for unknown cause). Seventy nine trials (39.5%) failed, 16 of which for intolerance and 63 for procedure inefficacy. Only two of the intolerant patients underwent invasive mechanical ventilation (one was intubated and the other one was subjected to tracheostomy): both were discharged at home alive, together with the other seven who were not intubated. Seven patients intolerant of procedure were not intubated for varied reasons (two for advanced neoplastic disease, three for advanced age, and two refused advanced life support) and died. Thirteen of 63 patients in whom NIV trial failed for inefficacy were intubated and eight survived, one of whom was tracheostomised. The remaining 50 patients of the failure group were not considered candidate for intubation and only two survived to discharge (fig 1).

Mortality in HDU was 27.4% and global mortality at hospital discharge was 34.5%, which was slightly higher than the death rate predicted by APACHE II (32%). Median length of stay in HDU was 6 days (IQR 3–10) and median length of stay in hospital was 15 days (IQR 8–26). Endotracheal intubation rate was 6.5% and tracheostomy was made in 1% of NIV trials (table 3).

Nasal bridge ulcer was the main complication of NIV, but we cannot report its rate because of missed data on medical records. However, in one patient only the nasal bridge ulcer was the cause of procedure withdrawal.

Success rates and observed and APACHE II predicted mortalities for each main disease category are reported in table 4.

We found that, of the determinants of success evaluated (APACHE II, mental status, basal pH and PaCO₂, and improvement of pH and PaCO₂ within six hours after NIV initiation), the APACHE II score only was a predictive factor of success and of death, both in the global population of patients and in the hypercapnic group. The improvement of pH within six hours (OR 0.14, CI 95% 0.03 to 0.62; $p = 0.009$), but not that of PaCO₂ (OR 0.33, CI 95% 0.97 to 1.13; $p = 0.07$) was predictive of survival in the hypercapnic group. Multivariate analysis for the previous factors showed that only APACHE II was an independent predictive factor for response to NIV both in overall population (OR 0.85, CI 95% 0.79 to 0.91; $p = 0.001$) and in hypercapnic group (OR 0.80, CI 95% 0.72 to 0.89; $p = 0.001$) (Table 5).

DISCUSSION

A main finding of our study is the feasibility of NIV in an intermediate care unit within the ED, where patients can receive NIV at an earlier stage and with minor utilisation of resource than in ICU. The overall success rate of NIV in our study (60.5%) is quite lower than that reported in RCTs, but compares favourably with that of other similar observational studies. Alsous *et al*²³ reported a global success rate of 63% in their retrospective series of 80 patients with ARF (80% with hypercapnic ARF, APACHE II score of 17.2), who, differently from our patients, were treated exclusively with bilevel positive airway pressure in ICU. Level of monitoring and nurse staffing in their setting, certainly superior to those available for our patients, probably helped obtain better outcomes than ours. In another retrospective study of NIV for patients admitted at mixed intermediate and intensive care units with ARF from various causes, Sinuff *et al* reported that NIV was stopped secondary to improvement in 48.4% of their patients.²⁴ In a successive study by the same authors, the implementation of a guideline for the use of NIV did not obtain better outcomes of care in terms of ETI and mortality rates.²⁵ Girault *et al* demonstrated the clinical efficacy of NIV applied for different indications in a medical ICU, with an

Table 2 Baseline characteristics of each non-invasive ventilation (NIV) indication group

	CPE total	Hypercapnic	Non-hypercapnic	Pneumonia total	Hypercapnic	Non-hypercapnic	COPD	CPE + COPD	Pneumonia + COPD	Hypercapnic pneumonia + COPD	Ob-hypoven
Number trials (number patients)	70* (69)	45 (44)	18 (18)	27* (26)	12 (12)	11 (11)	39 (37)	11	21* (20)	18 (17)	6 (6)
Age†	76.4±12.5	77.3±12	76±10.7	64.9±14.9	66±13.7	62.3±16.8	71.3±8.1	83.8±6.4	74.6±9.9	73.9±10.1	58.1±5.4
F/M	44/25	30/14	10/8	10/16	4/8	5/6	15/22	7/4	7/13	6/11	3/3
APACHE II†	18.3±5.8 (66/70)	18.5±5 (45/45)	18.7±6.9 (16/18)	18.6±5.1 (27/27)	20.8±4.4 (12/12)	17.7±5.4 (11/11)	19±5.3 (38/39)	19.4±6.5 (11/11)	20.9±4.5 (21/21)	21.1±4.8 (18/18)	13.5±5.8 (6/6)
pH‡	7.27 (7.19–7.31)	7.24 (7.16–7.29)	7.34 (7.29–7.41)	7.37 (7.3–7.42)	7.31 (7.16–7.36)	7.44 (7.39–7.47)	7.26 (7.21–7.3)	7.24 (7.19–7.28)	7.29 (7.22–7.34)	7.27 (7.21–7.33)	7.26 (7.22–7.3)
PaCO ₂ ‡	51 (42.6–61.1)	55 (49–67)	37.2 (30.5–40.7)	45 (37.6–59.5)	59.5 (53.7–65)	37 (34.1–41)	75.6 (70.8–86.3)	77 (65–78.5)	71 (60.7–93)	72.5 (64.7–97)	70.9 (68.4–72.7)
PaO ₂ ‡	56 (46–71.2)	56 (44.7–65.8)	62.5 (47.2–74.4)	54 (47.6–63)	49.5 (47–54.2)	60 (52.1–67)	61 (47–73)	65 (43.7–77.5)	52.1 (46–72.5)	50.3 (39.5–69)	64.5 (55.5–72.9)
PaO ₂ /FiO ₂ ‡	185 (134–242.2)	165.5 (129–231.5)	223 (204.2–244)	142 (91.5–196.7)	175.5 (111.7–237.7)	105.5 (86–173)	184 (154.5–213.5)	191 (116.5–257.2)	161 (144–240)	197.5 (146.2–243.7)	252 (218.5–295)
	(32/70)	(24/45)	(9/18)	(12/27)	(6/12)	(6/11)	(15/39)	(6/11)	(9/21)	(8/18)	(3/6)

APACHE, Acute Physiology and Chronic Health Evaluation; CPE, cardiogenic pulmonary oedema; COPD, chronic obstructive pulmonary disease; F/M, female/male; ob-hypoven, obesity-hypoventilation decompensation.

*Trials with available arterial blood gas data: CPE 63/70; pneumonia 23/27; pneumonia + COPD 20/21.

†Mean ± standard deviation.

‡Median, interquartile range (available data/number trials).

overall success rate of 64%.²⁶ The difference in success rate with respect to RCTs could be because of three principal reasons. The first one is related to the setting where our patients were treated: initially in an emergency ward and subsequently in an understaffed intermediate care unit. Monitoring levels and intensity of nursing care were lower than usually offered in ICUs, where most RCTs on NIV were performed. Second, the selection of our patients for NIV was not as strict as in the RCTs where patients not candidate for intubation were excluded—for example, in the study of Brochard *et al.*,² 69% of patients admitted with ARF for exacerbation of COPD were excluded from the NIV trial. In

our series, of 63 patients who failed the NIV trial for inefficacy, 50 were not considered eligible for intubation and 48 died. These patients received non-invasive ventilation as “ceiling treatment” for their ARF. The third reason for different outcomes involves the absence of double blinding in the RCTs on NIV. Because of the nature of the intervention (applying a mask to the patient), double blinding of treatment assignment was not possible for NIV studies, probably overestimating the treatment effect.²⁷

As for outcomes of single categories of disease, the higher success rate obtained in the subgroup of patients with CPE (78.6%) in comparison with others subgroups is most

Figure 1 Outcomes of non-invasive ventilation (NIV) trials. ETI, endotracheal intubation

Table 3 Global outcomes

Successful trials, number and %	121 (60.5%)
Failures, number and %	79 (39.5%)
Intolerance	16 (8%)
Inefficacy:	
Early*	52 (26%)
Late†	11 (5.5%)
Intubation, number and %	13 (6.5%)
Tracheostomy, number and %	2 (1%)
HDU mortality, number (and %)	52 (27.4%)
of patients	
Global mortality, number (and %)	68 (34.5%)
of patients	
Mortality predicted by APACHE II	32%
HDU length of stay, median days (IQR)	6 (3–10)
Hospital length of stay, median days (IQR)	15 (8–26)

APACHE, Acute Physiology and Chronic Health Evaluation; HDU, high dependency unit; IQR, interquartile range.

*Early: failure within 48 hours of NIV; †Late: failure after 48 hours of NIV.

probably because of the ready reversibility of heart failure with medical treatment, which can work quite fast and so reduce the time during which the patient needs to be ventilated. The median NIV duration in these patients was 7 hours versus 24 hours (IQ 6–72 h) in the global population. The low success rate obtained in patients with community acquired pneumonia (CAP) confirms the results of previous studies indicating the poor efficacy of NIV in this type of patients.²⁸ In a randomised trial in patients with severe CAP, Confalonieri *et al*¹⁷ found, at a post hoc analysis, that the CAP subgroup with COPD randomised to NIV had a significant reduction in intubation rate, duration of intermediate respiratory ICU stay, and mortality at two months from study entry. The group of CAP patients with COPD in our study had a mortality rate of 50% at hospital discharge, slightly lower than that of CAP patients without COPD (57%), but a lower success rate of NIV (28.6% *v* 37%, respectively). Both groups of CAP patients had a mortality rate higher than that predicted by APACHE II—that is, 38.6% in CAP patients with COPD and 30.1% in CAP patients without COPD. We believe the high death rate of CAP patients in our series to be related to their high comorbidity and high number of patients with do-not-intubate codes (44.4% in CAP patients and 42.8% in CAP patients with COPD).

Another finding in our study of “real clinical practice” is the presence of a high number of patients who are not

considered candidate for invasive mechanical ventilation and for whom NIV is delivered as “ceiling” treatment. Actually, in our series, the rate of ETI (6.5%) was relatively low, mainly because of the high number of patients who refused or were otherwise judged not candidate for invasive mechanical ventilation. Many of these patients received NIV for palliative reasons. Consequently, mortality rate at hospital discharge was 34.5% in our series (6.6% in NIV responders and 75.9% in NIV non-responders), which is much higher than in RCTs, where patients for whom ETI is not an option are not eligible to the study. In our series, those patients constituted a large part of global population (32%). With the limits of a retrospective study, these last results should induce a serious rethinking of the value and opportunity of NIV in patients who are not candidate for ETI.

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Table 4 Outcomes for categories of disease

	CPE			Pneumonia			COPD	CPE + COPD	Pneumonia + COPD	Hypercapnic pneumonia + COPD	Ob-hypoven
	Total	Hypercapnic	Non-hypercapnic	Total	Hypercapnic	Non-hypercapnic					
Number trials	70*	45	18	27*	12	11	39	11	21*	18	6
Number patients	69	44	18	26	12	11	37	11	20	17	6
Successes†	55/70 (78.6%)	38/45 (84.4%)	13/18 (72.2%)	10/27 (37%)	5/12 (41.6%)	4/11 (36.4%)	25/39 (64.1%)	6/11 (54.5%)	6/21 (28.6%)	5/18 (27.8%)	6/6 (100%)
Actual mortality†	16/69 (23.1%)	8/44 (18.2%)	6/18 (33.3%)	15/26 (57.7%)	8/12 (66.7%)	5/11 (45.4%)	11/37 (29.7%)	2/11 (18.2%)	10/20 (50%)	9/17 (52.9%)	0/6
APACHE II predicted mortality	21.9%			30.1%			24.8%	NA	38.9%		NA

APACHE, Acute Physiology and Chronic Health Evaluation; CPE, cardiogenic pulmonary oedema; COPD, chronic obstructive pulmonary disease; ob-hypoven, obesity-hypoventilation decompensation; NA, not applicable.

*number (%); †trials with available arterial blood gas data: CPE 63/70, pneumonia 23/27, pneumonia + COPD 20/21.

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Table 5 Predictive factors (adjusted for age and sex) of non-invasive ventilation (NIV) success and death

	Overall			Hypercapnic group		
	OR	CI 95%	p	OR	CI 95%	p
Age, years	0.99	0.97–1.01	0.499	0.99	0.96–1.02	0.132
Sex (male=1)	1.88	1.04–3.31	0.035	1.70	0.85–3.37	0.633
APACHE II	0.89	0.83–0.94	0.001	0.86	0.80–0.93	0.001
pH*	1.00	0.97–1.03	0.897	1.01	0.97–1.05	0.499
PaCO ₂ (mm Hg)	0.99	0.98–1.01	0.237	0.97	0.96–0.99	0.017
Decreased level of consciousness	0.75	0.48–1.20	0.231	0.70	0.41–1.20	0.193

	Overall			Hypercapnic group		
	OR	CI 95%	p	OR	CI 95%	p
Age, years	1.03	1.00–1.05	0.039	1.03	0.99–1.06	0.102
Sex (male=1)	0.58	0.32–1.06	0.075	0.49	0.24–1.02	0.057
APACHE II	1.09	1.03–1.16	0.003	1.12	1.04–1.20	0.003
pH*	1.01	0.98–1.04	0.640	0.98	0.94–1.02	0.315
PaCO ₂ (mm Hg)	1.00	0.98–1.01	0.612	1.01	0.99–1.03	0.410
Decreased level of consciousness	1.55	0.97–2.51	0.067	1.65	0.93–2.91	0.085
†PaCO ₂ 6 h	0.44	0.17–1.14	0.091	0.33	0.97–1.13	0.078
†pH 6 h	0.16	0.05–0.50	0.002	0.14	0.03–0.62	0.009

*pH values were multiplied for 100. The OR obtained is for pH increments of 0.01.

†Improvement of PaCO₂ and pH within 6 h after NIV initiation.