



Clinical characteristics and outcomes of mechanically ventilated elderly patients in intensive care units: a Chinese multicentre retrospective study

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Background: In recent years, the number of elderly patients receiving mechanical ventilation (MV) in intensive care units (ICUs) has increased. However, the evidence on the outcomes of elderly mechanically ventilated patients is scant in China. Our objective was to evaluate the characteristics and outcomes in elderly patients (≥ 65 years) receiving MV in the ICU.

Methods: We performed a multicentre retrospective study involving adult patients who were admitted to the ICU and received at least 24 hours of MV. Patients were divided into three age groups: under 65, 65–79, and ≥ 80 years. The primary outcome was hospital mortality. We performed univariate and multivariate logistic regression analysis to identify factors associated with hospital mortality.

Results: A total of 853 patients were analysed. Of those, 61.5% were ≥ 65 years of age, and 26.0% were ≥ 80 years of age. There were significant differences in the principal reason for MV among the three age groups ($P < 0.001$). Advanced age was significantly associated with total duration of MV, ICU length of stay (LOS), and ICU costs (all $P < 0.001$), but not with hospital LOS and hospital costs ($P > 0.05$). In addition, mortality rates in the ICU, hospital, and at 60 days significantly increased with age (all $P < 0.001$). In the age group of 80 years and older, the mortality rates were 47.7%, 49.5%, and 50.0%, respectively. Multivariate logistic regression analysis had found that age, Acute Physiology and Chronic Health Evaluation (APACHE) II score, partial pressure of oxygen in arterial blood/fraction of inspired oxygen ($\text{PaO}_2/\text{FiO}_2$) ratio, total duration of MV, ICU LOS, and the decision to withhold/withdraw life-sustaining treatments were independent influence factors for mortality rates.

Conclusions: Mechanically ventilated elderly patients (≥ 65 years) have a higher ICU and hospital mortality, but the hospital LOS and hospital costs are similar to younger patients. Advanced age should be considered as a significant independent risk factor for hospital mortality of mechanically ventilated ICU patients.

Keywords: Elderly; mechanical ventilation (MV); intensive care unit; outcomes; mortality

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Introduction

It is well-known that population aging is already taking place in most countries and regions of the world. The proportion of the world's population aged 60 years or older increased from 8% in 1950 to 12% in 2013. The older population is also aging itself; the proportion of persons aged 80 years or older within the older population increased from 7% in 1950 to 14% in 2013. Twenty-three million persons aged 80 years or over were living in China in 2013 (1).

As the population ages and life expectancy increases, the number of elderly patients admitted to intensive care units (ICUs) is rapidly increasing in many countries (2-5) and has been estimated to rise considerably (3,4). Elderly patients currently account for 42–52% of ICU admissions and for almost 60% of all ICU days (3,6-8). A retrospective analysis in Australia and New Zealand found an annual increase of 5.6% in the number of very elderly (age ≥ 80 years) patients (3). The main reasons for ICU admission in elderly patients (age ≥ 65 years) are cardiovascular issues (23–24%) (7,9), respiratory issues (26–52%) (10-16), trauma (29%) (17), or sepsis (22–32%) (3,18-21). Invasive mechanical ventilation (MV) has become one of the most common treatments used in elderly patients admitted to the ICU. About half or more of elderly patients in the ICU need MV (3,7,9-11,14-16,18,21-23). With our aging population, the number of patients with MV will steadily increase, with a projected 80% increase by 2026 when compared to 2000 (24).

Although many clinical studies have described the clinical characteristics and outcomes of elderly patients in the ICU, there is still a lack of information on the prognosis of mechanically ventilated elderly patients in the ICU. This information is important in clinical practice when deciding whether to transfer the patient to an ICU, and whether to initiate or withdraw MV. The aim of this study is to describe clinical characteristics and outcomes of mechanically ventilated elderly patients in the ICU.

We present the following article in accordance with the STROBE reporting checklist (available at <http://dx.doi.org/10.21037/jtd-20-2748>).

Methods

Study setting and design

This retrospective observational cohort study was carried out in fourteen ICUs of thirteen tertiary teaching hospitals in Beijing between January 2012 and June 2013.

Among the fourteen participating ICUs, ten were medical-surgical ICUs, two were surgical ICUs, one was a respiratory ICU, and one was a medical ICU. The number of ICU beds ranged from 8 to 20 during the study period. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). This retrospective study was approved by the Research Ethics Boards of all participating institutions (approval number: 2019FXHEC-KY167), with a waiver of informed consent.

Study population

Patients who were admitted to the ICU and received at least 24 hours of invasive MV within the first 48 hours of ICU stay were eligible. Patients were excluded if they were less than 18 years old, had incomplete data sets, were diagnosed with neuromuscular disease, required chronic MV prior to hospital admission, or were transferred from other facilities and had already been intubated or tracheotomised. A patient was considered as one case if they were admitted to the ICU several times during the study period, and only data from the first ICU admission was analysed.

In our study, patients aged 65 years and older at the time of hospital admission, were defined as the elderly, and the cohort was divided into three age groups for analysis: under 65, 65–79, and 80 years and older. Those under 65 years of age were used as a reference population.

Data collection

For every enrolled patient, the following data were recorded: demographic and epidemiological characteristics; reason for ICU admission; comorbidities; severity of illness; primary reason for MV; MV parameters and settings; sedatives, analgesics, and neuromuscular blockers used during MV; total duration of MV; ventilator-free days (25); the occurrence of successful weaning within 28 days; ICU and hospital length of stay (LOS); ICU and hospital costs; complications of MV; the occurrence of withholding or withdrawing life-sustaining treatments; and discharge destination. To obtain survival data after hospital discharge, we contacted survivors, their relatives, or their general practitioner or nursing homes by phone.

Severity of illness was assessed using the Acute Physiology and Chronic Health Evaluation (APACHE) II score (26). The severity of illness as rated by the APACHE II score was given with and without age points to illustrate the impact that age has on the score.

Outcomes of interest

The primary outcome was hospital mortality. Secondary outcomes included the total duration of MV, ICU and hospital LOS, and ICU and 60-day mortality rates.

Statistical analysis

Statistical analyses were carried out using SPSS 21.0 (SPSS Inc., Chicago, Illinois, USA). Data were expressed as the mean \pm standard deviation (SD) for normally distributed continuous variables, the median [interquartile range (IQR)] for non-normally distributed variables, and the number (percentage) for categorical variables. Continuous variables were compared using the Kruskal-Wallis test or the Mann-Whitney U test. Categorical variables were compared using the chi-square or Fisher's exact tests. Logistic regression analysis was used to determine risk factors for the ICU, hospital, and 60-day mortality rates. Variables were introduced into the model based on clinical and statistical significance (P value <0.2 on univariate analysis). We then performed three separate multivariate logistic regression analyses using a forward selection procedure to determine risk factors for ICU mortality (first analysis), hospital mortality (second analysis), and 60-day mortality (third analysis). We defined cut-off points for ICU LOS and total duration of MV to create a binary dependent variable: seven days for ICU stay and five days for total duration of MV. These cut-off points were chosen because they were the median values for the variables. A two-sided P value <0.05 was considered statistically significant.

Results

Baseline characteristics

A total of 853 patients were included in this study. Of these, there were 328 patients under the age of 65 years (38.5%), 303 patients aged 65–79 years (35.5%), and 222 patients aged 80 years and older (26.0%). *Table 1* presents baseline demographics and clinical characteristics of mechanically ventilated patients in the ICU. There were statistically significant differences in body mass index (BMI) and smoking history among the three age groups. However, significant differences were not found among the three age groups due to gender, past history of surgery, and non-invasive mechanical ventilation (NIMV) before ICU admission.

With advancing age, patients admitted to the ICU were

more likely to be from the medical ward (from 13.1% in the age group under 65 years to 32.4% in the age group of 80 years and older), while the proportion from the surgical ward decreased (54.0% in the age group under 65 years to 30.6% in the age group of 80 years and older).

With increasing age, the prevalence of some pre-existing comorbidities such as hypertension, chronic renal failure, chronic heart failure, chronic obstructive pulmonary disease (COPD), cancer, and stroke significantly increased. The prevalence of having two or more comorbidities also increased with age (P <0.001). In the two older age groups, more than 50% of patients had two or more comorbidities.

ICU hospitalization characteristics

Table 2 presents the hospitalization characteristics of mechanically ventilated patients in the ICU. The APACHE II scoring system was used to assess the severity of illness in patients admitted to the ICU, both with and without age points. There was a significant difference among the three age groups with median APACHE scores of 13 points (IQR, 8–19) in the age group under 65 years, 18 points (IQR, 12–24) in the age group of 65–79 years, and 19 points (IQR, 13–24) in the age group 80 years and older (P <0.001). When compared with patients aged less than 65 years, the two older age groups had higher APACHE scores (both P <0.001). After filtering out the age points, there was still a significant difference in APACHE scores between the age group under 65 years and the age group of 80 years and older [11 points (IQR, 6–17) *vs.* 14 points (IQR, 8–19), P=0.034]; however, there was not a significant difference between the age group under 65 years and the age group of 65–79 years (P=0.114).

We found significant differences among the three age groups (P <0.001) in the principal reason for MV: pneumonia (31.1%) was the most prevalent reason for the initiation of MV in the age group of 80 years and older, followed by postoperative status (23.4%), as well as COPD or asthma (10.4%) and congestive heart failure (10.4%). However, the most prevalent reason for MV in the age group under 65 years old and the age group between 65 and 79 years old was postoperative status (48.2% and 37.0%, respectively), followed by pneumonia (11.9% and 17.2%, respectively) and acute respiratory distress syndrome (ARDS; 9.8% and 10.9%, respectively).

Arterial blood gas analysis before MV in the three age groups showed statistical differences in partial pressure of oxygen in arterial blood/fraction of inspired oxygen (PaO₂/FiO₂)

Table 1 Baseline characteristics of mechanically ventilated patients in ICU

Variable	<65 years (n=328)	65–79 years (n=303)	≥80 years (n=222)	P value ^b
Age (years), MD (IQR)	50 (38, 58)	74 (70, 77)*	84 (82, 87) [#]	<0.001
Female, n (%)	120 (36.6)	117 (38.6)	77 (34.7)	0.650
BMI (kg/m ²)	24 (21, 26)	23 (21, 25)	22 (19, 24) [#]	<0.001
History of smoking, n (%)	118 (36.0)	123 (40.6)	60 (27.0) [#]	0.005
Past history of surgery, n (%)	87 (26.5)	69 (22.8)	59 (26.6)	0.478
NIMV before ICU admission	41 (12.5)	42 (13.9)	19 (8.6)	0.168
ICU admission source, n (%)				<0.001
Medical ward	43 (13.1)	70 (23.1)*	72 (32.4)*	<0.001
Surgical ward	177 (54.0)	152 (50.2)	68 (30.6) [#]	<0.001
Emergency department	94 (28.7)	75 (24.8)	78 (35.1) [#]	0.034
Others	14 (4.3)	6 (2.0)	4 (1.8)	0.126
Comorbidities, n (%)				
Hypertension	77 (23.5)	183 (60.4)*	143 (64.4)*	<0.001
Diabetes	42 (12.8)	113 (37.3)*	78 (35.1)*	<0.001
Chronic renal failure	25 (7.6)	36 (11.9)	36 (16.2)*	0.007
Chronic heart failure	9 (2.7)	34 (11.2)*	37 (16.7)*	<0.001
COPD	8 (2.4)	38 (12.5)*	39 (17.6)*	<0.001
Cirrhosis	7 (2.1)	2 (0.7)	2 (0.9)	0.249
Pulmonary fibrosis	10 (3.0)	9 (3.0)	6 (2.7)	0.971
Cancer	26 (7.9)	44 (14.5)*	34 (15.3)*	0.010
Brain stroke	31 (9.5)	71 (23.4)*	75 (33.8) [#]	<0.001
No. of comorbidities ^a , n (%)				<0.001
None	182 (55.5)	52 (17.2)	31 (14.0)	
1	85 (25.9)	93 (30.7)	57 (25.7)	
≥2	61 (18.6)	158 (52.1)	134 (60.4)	

^a, comorbidities included hypertension, insulin-dependent diabetes, chronic renal failure, chronic heart failure [New York Heart Association (NYHA) functional classification III–IV], chronic obstructive pulmonary disease, cirrhosis, pulmonary fibrosis, cancer, and brain stroke;

^b, comparison among the three age groups; *, $P < 0.05$, significantly different from the group of patients aged under 65 years; [#], $P < 0.05$, significantly different from the group of 65–79 years. ICU, intensive care unit; MD, median; IQR, interquartile range; BMI, body mass index; MV, mechanical ventilation; NIMV, non-invasive mechanical ventilation; APACHE II, Acute Physiology and Chronic Health Evaluation II; COPD, chronic obstructive pulmonary disease.

ratio ($P = 0.026$), partial pressure of carbon dioxide (PaCO_2 , $P < 0.001$), and pulse oxygen saturation (SpO_2 , $P < 0.001$), but this was not observed in arterial pH ($P = 0.126$).

The combination of synchronized intermittent mandatory ventilation and pressure support mode (SIMV + PSV) was the most preferred mode of MV in the three age groups (39.0%, 43.6%, and 34.2% in the under 65 years age

group, 65–79 years old group, and 80 years and older group, respectively). However, the MV mode was not significantly different among the three age groups ($P = 0.404$).

Regarding the management of patients during MV, significant differences were found in tidal volume (VT; $P < 0.001$) and the use of analgesics ($P < 0.001$) among the three age groups. In the whole cohort, more than 70%

Table 2 Hospitalization characteristics of mechanically ventilated patients in ICU

Variable	<65 years (n=328)	65–79 years (n=303)	≥80 years (n=222)	P value ^a
APACHE II score, MD (IQR)	13 (8, 19)	18 (12, 24)*	19 (13, 24)*	<0.001
APACHE II without age points, MD (IQR)	11 (6, 17)	13 (7, 19)	14 (8, 19)*	0.023
Primary reason for MV, n (%)				<0.001
Acute respiratory failure				
ARDS	32 (9.8)	33 (10.9)	12 (5.4)	0.080
Postoperative	158 (48.2)	112 (37.0)*	52 (23.4)*#	<0.001
Congestive heart failure	9 (2.7)	16 (5.3)	23 (10.4)*	0.001
Aspiration	3 (0.9)	4 (1.3)	12 (5.4)*#	0.001
Pneumonia	39 (11.9)	52 (17.2)	69 (31.1)*#	<0.001
Sepsis	22 (6.7)	17 (5.6)	9 (4.1)	0.416
Trauma	9 (2.7)	2 (0.7)	0 (0.0)*	0.012
Cardiac arrest	8 (2.4)	14 (4.6)	5 (2.3)	0.196
Other	17 (5.2)	12 (4.0)	5 (2.3)	0.226
Acute on chronic respiratory failure				
COPD or asthma	4 (1.2)	23 (7.6)*	23 (10.4)*	<0.001
Chronic pulmonary disease	8 (2.4)	7 (2.3)	2 (0.9)	0.397
Coma	19 (5.8)	11 (3.6)	10 (4.5)	0.434
Arterial blood gas analysis prior to MV, MD (IQR)				
pH	7.37 (7.27, 7.42)	7.36 (7.25, 7.42)	7.34 (7.23, 7.41)	0.126
PaO ₂ /FiO ₂	182 (117, 315)	159 (110, 239)*	180 (120, 268)	0.026
PaCO ₂ (mmHg)	38 (33, 45)	40 (34, 55)*	42 (34, 65)*	<0.001
SpO ₂ (%)	97 (90, 100)	95 (88, 99)*	93 (88, 99)*	<0.001
Mode and parameter setting at the beginning of MV				
Mode of MV, n (%)				0.404
VCV	71 (21.6)	62 (20.5)	42 (18.9)	
PCV	31 (9.5)	36 (11.9)	33 (14.9)	
PSV	73 (22.3)	57 (18.8)	54 (24.3)	
SIMV	20 (6.1)	12 (4.0)	14 (6.3)	
SIMV + PSV	128 (39.0)	132 (43.6)	76 (34.2)	
Others	5 (1.5)	4 (1.3)	3 (1.4)	
Ventilator's parameter setting, MD (IQR)				
VT (mL/kg predicted bodyweight)	7.38 (6.60, 8.25)	7.28 (6.37, 8.03)	6.87 (6.07, 7.91)*#	0.001
Applied PEEP (cmH ₂ O)	5 (5, 8)	6 (5, 8)	6 (5, 8)	0.429
Peak pressure (cmH ₂ O)	22 (18, 28)	24 (20, 29)*	23 (20, 28)	0.010
Plateau pressure (cmH ₂ O)	18 (14, 20)	18 (14, 22)	16 (12, 20)	0.087

Table 2 (continued)

Table 2 (continued)

Variable	<65 years (n=328)	65–79 years (n=303)	≥80 years (n=222)	P value ^a
Use of sedatives, n (%)	280 (85.4)	253 (83.5)	177 (79.7)	0.219
Use of analgesics, n (%)	239 (72.9)	185 (61.1)*	106 (47.7)*#	<0.001
Use of neuromuscular blockers, n (%)	13 (4.0)	15 (5.0)	6 (2.7)	0.429

^a, comparison among the three age groups; *, P<0.05, significantly different from the group of patients aged under 65 years; #, P<0.05, significantly different from the group of 65–79 years. ICU, intensive care unit; APACHE II, Acute Physiology and Chronic Health Evaluation II; MD, median; IQR, interquartile range; MV, Mechanical ventilation; ARDS, acute respiratory distress syndrome; COPD, chronic obstructive pulmonary disease; PaO₂, partial pressure of oxygen in arterial blood; FiO₂, fraction of inspired oxygen; PaCO₂, partial pressure of carbon dioxide in arterial blood; SpO₂, pulse oxygen saturation; VCV, volume-control ventilation; PCV, pressure-control ventilation; SIMV, synchronized intermittent mandatory ventilation; PSV, pressure support ventilation; PEEP, positive end-expiratory pressure.

(71.9%) of patients were ventilated with a VT <8 mL/kg of predicted bodyweight, and in the third age group, the proportion of patients with a VT <8 mL/kg of predicted bodyweight was the highest (76.1%). The proportion of patients using analgesics in the three age groups decreased with age (72.9% in the age group under 65 years to 47.7% in the age group of 80 years and older). There were no significant differences in the median levels of positive end expiratory pressure (PEEP) and platform pressure among the three age groups, or in the use of sedatives and neuromuscular blockers (all P>0.05).

Clinical courses and outcomes

Table 3 presents the clinical courses and outcomes in each of the three age groups. There were statistically significant differences in the total duration of MV (P<0.001), ventilator-free days (P<0.001), and successful weaning rate within 28 days (P<0.001) among the three age groups. As shown in Table 3, it can be clearly seen that compared with patients in the group aged 65–79 years and the group aged under 65 years, the total MV time of patients in the age group of 80 years and older increased significantly, while the ventilator-free days, and successful weaning rate within 28 days decreased significantly. Only the total duration of MV between the two older age groups had no statistically significant difference [135 (IQR, 59–280) vs. 165 (IQR, 67–404) hours, P=0.101].

There was no statistically significant difference in the hospital LOS among the three age groups (P=0.749). However, the difference in the ICU LOS did differ significantly (P<0.001). In the group of patients aged 80 and older, the ICU LOS was 13 days (IQR, 6–24 days), as compared with 8 days (IQR, 4–19 days) for patients aged

65–79 years (P=0.005) and 7 days (IQR, 4–14 days) for patients aged under 65 years (P<0.001).

Mortality rates in the ICU, hospital, and at 60 days significantly increased with age. In the age group of 80 years and older, the hospital mortality rate was 49.5%, which was significantly different from the 19.8% rate for the age group under 65 years (P<0.001) and the 35.0% rate for the age group of 65–79 years (P=0.001). There was also a significant difference in the hospital mortality rate between the age group 65–79 years and the age group under 65 years (P<0.001). The same was true for ICU mortality and 60-day mortality rates.

There was no statistically significant difference in hospital costs among the three age groups (P=0.084). However, ICU costs did differ significantly (P<0.001). The ICU costs were higher in the age group of 65–79 years (with a median of 68,000 CNY) and the age group of 80 years and older (with a median of 89,000 CNY) than those in the age group under 65 years (with a median of 57,000 CNY; P=0.030 and P<0.001, respectively), but there was no significant difference between the two elderly groups (P=0.061).

Table 3 also shows the rate of events that occurred over the course of MV. The proportion of patients who required reintubation tended to be higher in the age group of 80 years and older. Furthermore, the rates of subsequent ventilator-associated pneumonia and the proportion of patients who underwent prolonged MV were significantly higher in the age group of 80 years and older (all P<0.01). In addition, the decision to withhold or withdraw life-sustaining treatments tended to be much more often in the age group of 80 years and older (19.4%). A considerable proportion of the 572 patients who survived the hospital were discharged home. Additionally, the proportion of

Table 3 Clinical course and outcomes of mechanically ventilated patients in ICU

Variable	<65 years (n=328)	65–79 years (n=303)	≥80 years (n=222)	P value ^d
Total duration of MV (hours), MD (IQR)	96 (47, 202)	135 (59, 280)*	165 (67, 404)*	<0.001
Ventilator-free days (days), MD (IQR)	24 (5, 28)	20 (0, 27)*	2 (0, 26)*#	<0.001
Successful weaning within 28 days ^a , n (%)	237 (72.3)	180 (59.4)*	98 (44.1)*#	<0.001
ICU LOS (days), MD (IQR)	7 (4, 14)	8 (4, 19)	13 (6, 24)*#	<0.001
Hospital LOS (days), MD (IQR)	23 (12, 37)	23 (14, 40)	24 (13, 35)	0.749
ICU mortality, n (%)	61 (18.6)	96 (31.7)*	106 (47.7)*#	<0.001
Hospital mortality, n (%)	65 (19.8)	106 (35.0)*	110 (49.5)*#	<0.001
60-day mortality, n (%)	82 (25.0)	114 (37.6)*	111 (50.0)*#	<0.001
ICU costs (10,000 CNY), MD (IQR)	5.7 (2.7, 10.6)	6.8 (3.4, 13.7)*	8.9 (4.6, 15.6)*	<0.001
Hospital costs (10,000 CNY), MD (IQR)	10.6 (5.8, 16.7)	11.3 (6.6, 19.0)	11.7 (6.8, 19.3)	0.084
Complications of MV, n (%)				
Barotrauma ^b	7 (2.1)	6 (2.0)	1 (0.5)	0.263
VAP	43 (13.1)	51 (16.8)	60 (27.0)*#	<0.001
Self-extubation	2 (0.6)	3 (1.0)	5 (2.3)	0.233
Reintubation	10 (3.0)	15 (5.0)	20 (9.0)*	0.009
Tracheotomy	53 (16.2)	38 (12.5)	46 (20.7)#	0.042
PMV ^c	28 (8.5)	35 (11.6)	45 (20.3)*#	<0.001
Withhold/withdraw life-sustaining treatments, n (%)	38 (11.6)	48 (15.9)	43 (19.4)*	0.040
Discharge destination, n (%)				
Home	207 (78.7)	167 (84.8)	86 (76.8)	
Respiratory care ward	8 (3.0)	5 (2.5)	10 (8.9)	
Nursing home	2 (0.8)	8 (4.1)	3 (2.7)	
Other hospital	46 (17.5)	17 (8.6)	13 (11.6)	

^a, successful weaning from MV was defined as complete respiratory autonomy for at least 48 h; ^b, Barotrauma refers to the development of at least one of the following: interstitial emphysema, pneumothorax, pneumomediastinum, pneumoperitoneum or subcutaneous emphysema; ^c, PMV was defined as the need for mechanical ventilation for more than 21 days; ^d, comparison among the three age groups; *, P<0.05, significantly different from the group of patients aged under 65 years; #, P<0.05, significantly different from the group of 65–79 years. ICU, intensive care unit; MV, Mechanical ventilation; MD, median; IQR, interquartile range; LOS, length of stay; CNY, Chinese yuan; VAP, ventilator-associated pneumonia; PMV, prolonged mechanical ventilation.

patients that survived discharge and returned home in all three age groups was similar. However, in the age group of 80 years and older, nearly one-ninth (8.9%) of survivors were transferred to a respiratory care ward.

Univariate and multivariate logistic regression analyses

Table 4 presents the statistical results of univariate and multivariate analyses for ICU mortality, hospital mortality,

and 60-day mortality. Multivariate logistic regression analysis had found that age, APACHE II score, PaO₂/FiO₂ ratio, total duration of MV, ICU LOS, and the decision to withhold/withdraw life-sustaining treatments were independent influence factors for ICU mortality, hospital mortality, and 60-day mortality. In addition, the number of comorbidities was also an independent risk factor for ICU mortality. Multivariate logistic regression analysis also showed that when compared with the reference group

Table 4 Results of the logistic regression analyses

Variable	ICU mortality		Hospital mortality		60-day mortality	
	OR (95% CI)	P value	OR (95% CI)	P value	OR (95% CI)	P value
Univariate analysis						
Age groups (vs. <65)						
65–79	2.030 (1.404–2.935)	<0.001	2.177 (1.519–3.120)	<0.001	1.810 (1.286–2.546)	0.001
≥80	4.000 (2.727–5.867)	<0.001	3.974 (2.723–5.800)	<0.001	3.000 (2.087–4.312)	<0.001
BMI (kg/m ²)	0.922 (0.886–0.958)	<0.001	0.925 (0.891–0.961)	<0.001	0.933 (0.899–0.968)	<0.001
History of smoking	1.160 (0.857–1.568)	0.337	1.144 (0.850–1.540)	0.373	1.159 (0.866–1.551)	0.320
ICU admission source: Medical ward	3.373 (1.209–9.427)	0.020	2.581 (1.848–3.604)	<0.001	2.857 (2.047–3.988)	<0.001
No. of comorbidities (vs. none)						
1	2.145 (1.408–3.269)	<0.001	1.941 (1.296–2.907)	0.001	1.784 (1.209–2.631)	0.004
≥2	3.280 (2.238–4.807)	<0.001	2.854 (1.979–4.115)	<0.001	2.658 (1.869–3.780)	<0.001
APACHE II score	1.106 (1.084–1.129)	<0.001	1.101 (1.079–1.123)	<0.001	1.098 (1.076–1.119)	<0.001
Cause of MV: pneumonia	2.082 (1.462–2.964)	<0.001	2.154 (1.517–3.058)	<0.001	2.329 (1.643–3.302)	<0.001
PaCO ₂ (mmHg)	1.005 (0.998–1.012)	0.143	1.005 (0.998–1.011)	0.173	1.004 (0.997–1.010)	0.278
PaO ₂ /FiO ₂	0.998 (0.997–0.999)	0.001	0.998 (0.997–0.999)	0.002	0.998 (0.997–0.999)	0.001
Total duration of MV ≥5 days	3.378 (2.473–4.615)	<0.001	2.984 (2.208–4.033)	<0.001	2.953 (2.203–3.960)	<0.001
ICU LOS ≥7 days	1.287 (0.951–1.742)	0.102	1.334 (0.991–1.797)	0.058	1.379 (1.030–1.845)	0.031
Withhold/withdraw life-sustaining treatments	10.775 (6.941–16.727)	<0.001	11.567 (7.326–18.262)	<0.001	19.394 (11.202–33.577)	<0.001
Multivariate analysis						
Age groups (vs. <65)						
65–79	1.348 (0.838–2.171)	0.218	1.442 (0.908–2.290)	0.121	1.312 (0.832–2.068)	0.242
≥80	2.799 (1.714–4.573)	<0.001	2.519 (1.558–4.072)	<0.001	1.843 (1.141–2.977)	0.012
BMI (kg/m ²)	0.957 (0.910–1.005)	0.081	0.960 (0.914–1.008)	0.099	0.964 (0.918–1.011)	0.134
ICU admission source: Medical ward	1.211 (0.774–1.895)	0.401	1.356 (0.872–2.106)	0.176	1.540 (1.001–2.370)	0.050
No. of comorbidities (vs. none)						
1	1.441 (0.822–2.527)	0.202	1.409 (0.816–2.432)	0.218	1.260 (0.739–2.148)	0.395
≥2	1.835 (1.065–3.162)	0.029	1.667 (0.981–2.835)	0.059	1.438 (0.854–2.421)	0.172
APACHE II score	1.085 (1.058–1.113)	<0.001	1.079 (1.053–1.106)	<0.001	1.069 (1.043–1.095)	<0.001
Cause of MV: pneumonia	0.958 (0.596–1.540)	0.859	1.036 (0.649–1.655)	0.881	1.245 (0.781–1.985)	0.357
PaCO ₂ (mmHg)	0.996 (0.987–1.004)	0.308	0.995 (0.987–1.003)	0.245	0.994 (0.985–1.002)	0.144
PaO ₂ /FiO ₂	0.999 (0.998–1.000)	0.027	0.999 (0.998–1.000)	0.046	0.999 (0.998–1.000)	0.016
Total duration of MV ≥5 days	5.046 (2.653–9.598)	<0.001	3.761 (2.086–6.782)	<0.001	3.449 (1.951–6.097)	<0.001
ICU LOS ≥7 days	0.253 (0.131–0.491)	<0.001	0.358 (0.195–0.658)	0.001	0.369 (0.205–0.666)	0.001
Withhold/withdraw life-sustaining treatments	7.828 (4.710–13.009)	<0.001	8.741 (5.210–14.666)	<0.001	14.323 (7.882–26.028)	<0.001

OR, odds ratio; ICU, intensive care unit; BMI, body mass index; APACHE II, Acute Physiology and Chronic Health Evaluation II; MV, Mechanical ventilation; PaCO₂, partial pressure of carbon dioxide in arterial blood; PaO₂, partial pressure of oxygen in arterial blood; FiO₂, fraction of inspired oxygen; LOS, length of stay.

(under 65 years) and adjusted for other variables, the odds ratios of ICU death, hospitalization death and death within 60 days in patients aged 80 years and older were 2.799 (95% CI: 1.714–4.573), 2.519 (95% CI: 1.558–4.072) and 1.843 (95% CI: 1.141–2.977), respectively.

Discussion

In this multicentre retrospective study, our main findings were that the elderly account for a major proportion of adult patients receiving MV in ICU. Mechanically ventilated elderly patients have a lower ICU, hospital and 60-day survival, but the hospital LOS and hospital costs are similar to younger patients. Advanced age should be considered as an important independent risk factor for mortality rates of patients receiving MV in the ICU.

In recent decades, a few studies have shown that the proportion of elderly patients receiving MV treatment in ICUs reaches 25.4–50.5% (27–30). In our study, we found that this proportion of elderly patients was up to 61.5%, and the proportion of very elderly patients was up to 26.0%, which was significantly higher than in the previous studies. There are several reasons why this might be the case, including the following: first, it is well-known that population aging is taking place much more rapidly now in developing countries than it had in developed countries in the past. In particular, China is one of the fastest aging countries in the world (1). This demographic trend is bound to increase the demand for health care resources. Secondly, there is still a lack of well-defined thresholds for elderly patients, and the definition of the term “elderly” has varied from 65 to 75 years old (7,10,22,23,27,29–36). Therefore, the research results may differ.

As reported by others, acute respiratory failure was the most frequent indication for MV (27,28,30,37), accounting for 87.5% of the principal reason for MV for our total study group. There were statistically significant differences in the principal reason for MV among the three age groups ($P<0.001$). Compared with the other two age groups, very elderly (age ≥ 80 years) patients were more likely to receive MV for medical indications (pneumonia, aspiration, and congestive heart failure) and less likely to receive MV for postoperative reasons and trauma. Esteban *et al.* (27) also found that older patients (age >70 years) were ventilated because of cardiac diseases (heart failure and cardiac arrest) and fewer were ventilated because of coma, ARDS and trauma. The reasons for this may be explained as follows: first, we found that with age, patients admitted to the ICU

were more likely to come from the medical ward, while the proportion coming from surgical wards declined. Secondly, in our study, we found that the prevalence of having two or more comorbidities increased with age in patients with MV in the ICU ($P<0.001$). Having several organ dysfunctions and failures, such as cardiovascular and respiratory dysfunction and failure increased the relative risk of receiving MV. Multivariate analysis revealed that factors such as increasing age, New York Heart Association (NYHA) functional classification III or IV, neurologic and respiratory reasons for admission, emergency surgery; pneumonia, and lower Glasgow Coma Score were associated with the use of MV (38).

Unlike other studies (28,38), our study shows that SIMV + PSV was the main invasive ventilatory mode in the whole group as well as in all three age groups, followed by assist-control ventilation and PSV. In fact, this may reflect heterogeneity in the selection of ventilatory modes in different ICUs and countries (37,38). However, there was no statistically significant difference in ventilatory modes among the three age groups ($P=0.404$). In our cohort, the median VT setting was 7.26 mL/kg of predicted bodyweight (IQR, 6.37–8.11), a value lower than that described in some studies (28,37,38). More than 70% (71.9%) of patients were ventilated with a VT <8 mL/kg of predicted bodyweight, and this proportion increased with age. This value is greater than the 48.4% reported by Metnitz *et al.* in a multinational cohort (38). This may be related to the median $\text{PaO}_2/\text{FIO}_2$ ratios being <200 in the three age groups. Metnitz *et al.* found that patients with the lowest $\text{PaO}_2/\text{FIO}_2$ ratios were ventilated more often with tidal volumes <8 mL/kg of body weight, than patients with higher $\text{PaO}_2/\text{FIO}_2$ ratios (38).

Age has always been considered to be a factor in increased ICU resource utilization. Our findings may support this. In this study, we found statistically significant differences in duration of MV, length of ICU stay, and ICU costs among the three age groups. However, this does not mean that the length of hospital stays and hospital costs of elderly patients have increased significantly. The association between age and hospital costs in patients receiving MV was studied by Chelluri *et al.* (39). They found that daily and total hospital costs were lower in older patients. They suggest that the lower resource use for older patients may be related to a preference for less aggressive care by elderly patients and their families or by healthcare providers. We also support this view because we found that the decision to withhold or withdraw life-sustaining treatments tended to be much more often with aging. Moreover, Turnbull *et al.* (40)

have also shown a higher rate of limiting life support among older patients.

Our results suggest that ICU mortality, hospital mortality, and 60-day mortality of ventilated elderly patients in the ICU are significantly elevated. Age was strongly associated with mortality rates of mechanically ventilated elderly patients in the ICU, and remained a strong predictor of mortality even after adjusting for other variables. Our study adopted the most widely used criteria to define an elderly population, and analyzed the relationship between age and ICU mortality, hospital mortality and 60-day mortality in the three age intervals. We present the results of a quantitative assessment of increased risk of death associated with specific age intervals (≥ 80 years).

To our knowledge, this is the first study in China to evaluate the characteristics and outcomes in elderly patients (≥ 65 years) receiving MV in the ICU. However, we also are aware of several limitations of our study. First, this study is a retrospective study, and the data were obtained in 2012–2013, which may impose temporal limitations on the applicability of this data set. Second, we only evaluated ICU mortality, hospital mortality, and 60-day mortality, without discussing long-term mortality, activities of daily living (ADL), and quality of life (QOL) after discharge, because we did not collect relevant data. Future studies should focus on long-term mortality, physical rehabilitation, and QOL of elderly patients discharged from the hospital. Third, we did not collect data on clinical and laboratory variables during the period of MV, such as changes in blood pressure, heart rate, respiratory rate, pH, PaO₂, PaCO₂, SpO₂, and ventilator parameters. Thus, we cannot further assess the effectiveness of MV.

Conclusions

In conclusion, the results obtained in the present study indicate that the elderly patients constitute a major proportion of adult patients receiving MV in the ICU.

The elderly mechanically ventilated patients (≥ 65 years) have a higher ICU and hospital mortality, but the hospital LOS and hospital costs are similar to younger patients. Advanced age should be considered as a significant independent risk factor for hospital mortality of mechanically ventilated ICU patients. Evaluating the clinical characteristics and outcomes of elderly mechanically ventilated patients has become important in terms of the effective use of limited medical resources and making decisions for clinicians, patients and their families.

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Footnote

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Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). This retrospective study was approved by the Research Ethics Boards of all participating institutions (approval number: 2019FXHEC-KY167), with a waiver of informed consent.

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