

Intraoperative Hypotension and Related Risk Factors for Postoperative Mortality After Noncardiac Surgery in Elderly Patients: A Retrospective Analysis Report

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Background: Blood pressure fluctuation is very common during non-cardiac surgery in elderly. This retrospective study was to analyse whether intraoperative hypotension in elderly and other risk factors relate to the postoperative mortality.

Methods: A total of 118 cases (Observational group), who underwent noncardiac surgery in three medical centers between September 2014 and March 2017, and died in the hospital after the noncardiac surgery. With 1:2 ratio of propensity matching, 236 survival cases (Control group) were selected for comparison analyses with the death cases. Intraoperative blood pressure and perioperative parameters from both groups were collected from electronic anaesthesia charts. Data were analysed with univariate logistic regression analysis where variables with *p* values less than 0.05 were analysed with multivariate logistic regression analysis. The receiver operating characteristic (ROC) curve was constructed.

Results: There are five risk factors related to postoperative death in elderly patients: ASA grade, COPD, emergency surgery, general anesthesia, $60 < \text{MAP} \leq 65\text{mmHg}$ ($\text{OR} > 1$), and one factor may reduce the risk of postoperative mortality, which is PACU therapy ($\text{OR} < 1$). Compared with the Control group, the Observational group had a higher proportion of cerebral hernia, kidney injury and trauma ($p < 0.001$). The intraoperative blood transfusion volume and intraoperative blood loss volume were higher in the Observational group than the Control group ($p < 0.001$). The proportion of using vasoactive drugs was higher in the Observational group ($p < 0.001$), and there was more urine output during the operation in the Observational group ($p = 0.005$).

Conclusion: The intraoperative MAP of geriatric patients lower than 65mmHg is highly related to the postoperative mortality. Elderly patients with emergency surgery, high ASA grade and a history of COPD have an increased risk of postoperative mortality. General anesthesia is a risk factor for postoperative death in elderly patients, and the PACU therapy is a protective factor to avoid postoperative death.

Trial Registration: This study has been retrospectively registered in the Chinese Clinical Trials Registry (ChiCTR2000038912, 10/10/2020).

Keywords: intraoperative hypotension, geriatric patients, COPD, postoperative mortality, non-cardiac surgery

Introduction

With the acceleration of the social aging process and the increase of the elderly population, the proportion of elderly patients undergoing surgical treatment also increased gradually.¹ Due to the organs functional degradation in elderly patients, most of them

often suffer from various complications in the perioperative period, and the postoperative mortality rate is 4.4 times that of middle-aged and young patients.² Therefore, the risk factors associated with postoperative mortality in elderly patients are worth carefully analyzing.

Geriatric patients with decreased vascular self-regulation ability are prone to blood pressure fluctuations and hypotension. Besides, vital organs become more vulnerable for they are less able to tolerate hypotension.³ Many risk factors bring great challenges to the management of intraoperative anesthesia. Maintaining stable blood pressure is one of the core goals of intraoperative hemodynamic management.⁴ Although intraoperative hypotension is assumed to be associated with postoperative mortality, the threshold value of the blood pressure remains undefined, especially in the geriatric population.⁵ Therefore, it is of important clinical value to define the safe range of intraoperative blood pressure in elderly patients and regard it as one of the management objectives of hemodynamics. The purpose of this study is to investigate the correlation between the level and duration of intraoperative blood pressure and postoperative death in elderly patients and to explore other potential risk factors related to postoperative death in elderly patients.

Materials and Methods

Study Design and Population

This was a retrospective investigation study in three centers. The study was approved and the written informed consent was waived by the Ethics Committee of the First Affiliated Hospital (Southwest Hospital) of Army Medical University, PLA [KY2020182], by the Medical Ethics Committee of Second Affiliated Hospital (Xinqiao Hospital) of Army Medical University, PLA [2020-119-01] and by the Ethics Committee of Army Specialty Medical Center (Daping Hospital) of Army Medical University, PLA [2020.No.147] respectively. This study has been retrospectively registered in the Chinese Clinical Trials Registry (ChiCTR2000038912, 10/10/2020).

The inclusion criteria are as follows: the age of patients are ≥ 65 years old; the ASA Grade is I–IV; non-cardiac surgery; the operation time is more than 1 hour; the anesthetic methods are general anesthesia without combination with other anesthesia methods, spinal anesthesia or nerve block anesthesia; with definite clinical outcomes: cured, improved or died in the hospital. Died in the hospital after the operation within 60 days is defined as postoperative mortality. The transplantation surgery and adrenalectomy were excluded.

From September 2014 to March 2017, 31,654 cases met the inclusion and exclusion criteria. Having a postoperative mortality event, 118 cases were included in the Observational group, and the Control group was propensity matched by quantity from the survival cases. The number of surviving cases was twice as many as the Observational group in seven types of non-cardiac surgery (See Figure 1).

Collection and Collation of Blood Pressure Data

The intraoperative blood pressure data stored through the Mediston Anesthesia Information system (Mediston Medical Technology Company, Suzhou, China) were collected without distinction between noninvasive blood pressure and invasive blood pressure.

According to related studies, $MAP \leq 55\text{mmHg}$ and $SBP \leq 80\text{mmHg}$ can be regarded as intraoperative hypotension, while $MAP > 75\text{mmHg}$ and $SBP > 120\text{mmHg}$ are considered as nonhypotension thresholds.^{6,7} However, in order to explore a more accurate threshold of intraoperative hypotension in elderly patients, the researchers set the MAP analysis range to 5mmg and the SBP analysis range to 10mmHg for further study.

The blood pressure values, from anesthesia induction to the end of anesthesia, were collected and analyzed. According to the research scope of blood pressure above, the total number of minutes of SBP and MAP in the different range were calculated. The duration of SBP and MAP were recorded every 5 minutes and the invasive blood pressure was preferably selected for analysis. The blood pressure value at some time point, if obviously wrong or missing, would be replaced by the value recorded at the previous time point. In addition, the proportions of duration to the total operation time were measured.

To better realize the Logistic regression analysis, with reference of the ratio of blood pressure duration in each section to the total operation time, intraoperative blood pressure was defined as sort variables.

By analyzing the proportion of the duration of each blood pressure scope to the total operation time in the two groups, respectively, ratios of 0% and 10% were selected as the critical points. The ratio of 0% was defined as the variable 1, the ratio greater than 10% was defined as the variable 3, and the ratio between 0% and 10% was defined as the variable 2. These two critical points define intraoperative blood pressure as three categorical variables, and the distribution of

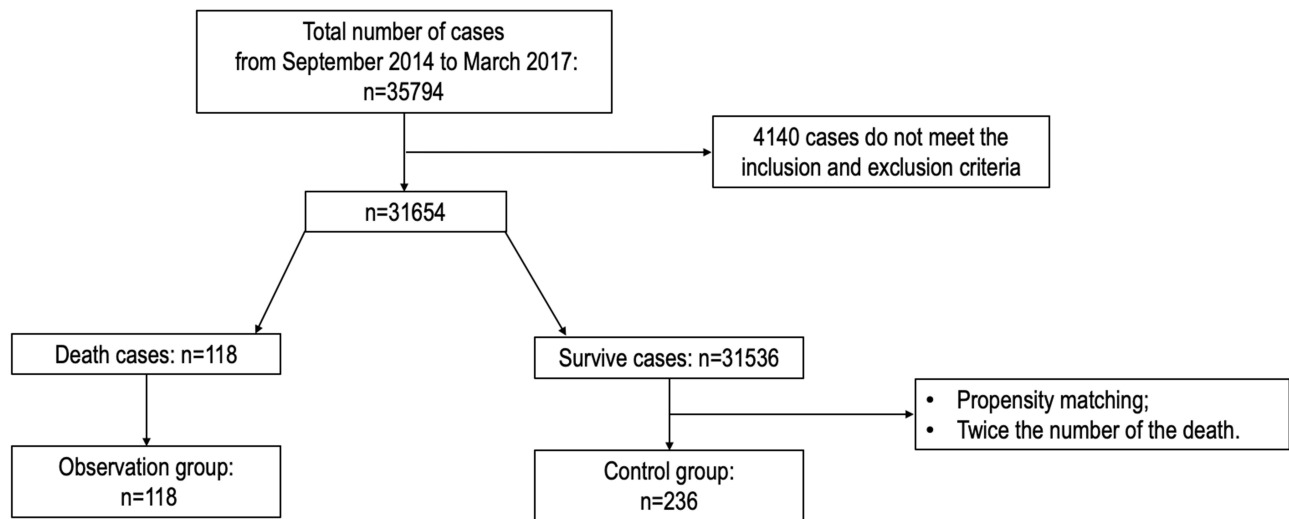


Figure 1 The flow chart of patient enrollment. Control group (n=236), Observational group (n=118).

the three variables is roughly balanced, which can better achieve multi-factor logistics regression.

Collection and Collation of Other Clinical Data

The clinical information, including blood pressure (MAP and SBP), gender, age, body mass index (BMI), ASA grade, NYHA heart function, operation type, operation time, anesthesia method, co-morbidity (hypertension, coronary heart disease, cerebral hernia, arrhythmia, diabetes, renal dysfunction, chronic obstructive pulmonary disease (COPD), anemia, trauma, malignant tumor), Charlson index, PACU therapy, intraoperative use of vasoactive drugs, intraoperative blood loss volume, Intraoperative urine output volume, intraoperative blood transfusion volume, intraoperative crystal liquid infusion, intraoperative colloidal fluid infusion, total intraoperative infusion, intraoperative resuscitation, and postoperative information were objectively extracted from the medical record system. Among these variables, Age, BMI, intraoperative blood loss volume, intraoperative urine output volume, intraoperative blood transfusion volume, intraoperative crystal liquid infusion, intraoperative colloidal fluid infusion, and total intraoperative infusion were regarded as continuous variables, the other variables were taken as sort variables. ASA grade, NYHA heart function were defined as 4 classification variables in accordance with existing classification standards. The mode of anesthesia is defined as two classification variables, including general anesthesia and non-general anesthesia. The other parameters were all seen as 2 classification variables: used or

unused. Postoperative information were not used for univariate or multiple logistic regression analysis.

Statistical Analysis

SPSS 18.0 software (SPSS, Chicago, Illinois, USA) was used to complete statistical analysis. The counting data were expressed using frequency and percentage, and the Chi-square test is used to compare between the two groups. Measurement data is represented by the median and quartile spacing [M (P25 P75)], Mann–Whitney *U*-test is used to compare between the two groups.

The blood pressure and the perioperative clinical indicators of elderly patients were analyzed by univariate logistic regression analysis firstly, and then the indicators with $p < 0.05$ were further analyzed by multiple logistic regression (stepwise regression method, entry: $P=0.05$; removal $P=0.10$). The linear model was used to diagnose the collinearity of the variables.

At last, the receiver operating characteristic (ROC) curve was established through multivariate regression analysis, which was used to analyze the sensitivity and specificity and determine the critical value.

Result

The Postoperative Mortality of Elderly Patients Undergoing Noncardiac Surgery in Three Hospitals

From September 2014 to March 2017, the postoperative mortality rate of the elderly patients in the three

Table I Clinical Characteristics of Patients

	Control Group (n=236)	Observational Group (n=118)	Statistic	p value
Demographics				
Age [M (P25, P75)]	71 (68–77)	72 (68–79)	–1.75	0.079
Male, n (%)	135 (57.20)	78 (66.10)	2.60	0.107
BMI [M (P25, P75)]	22.9 (21.0–25.4)	22.3 (19.8–24.8)	–1.77	0.076
ASA classification, n (%)				
I	9 (3.81)	1 (0.85)	82.65	< 0.001*
II	173 (73.31)	37 (31.36)		
III	51 (21.61)	54 (45.76)		
IV	3 (1.27)	26 (22.03)		
NYHA cardiac function, n (%)				
I	29 (12.29)	10 (8.47)	5.08	0.079
II	161 (68.22)	73 (61.86)		
III	46 (19.49)	35 (29.66)		
Co-morbidity				
Hypertension, n (%)	102 (43.22)	60 (50.85)	1.84	0.175
CHD, n (%)	38 (16.10)	24 (20.34)	0.98	0.323
Cerebral hernia, n (%)	1 (0.42)	8 (6.78)	12.83	< 0.001*
Arrhythmia, n (%)	28 (11.86)	11 (9.32)	0.52	0.471
Diabetes, n (%)	39 (16.53)	23 (19.49)	0.48	0.489
Renal dysfunction, n (%)	0 (0.00)	8 (6.78)	13.44	< 0.001*
COPD, n (%)	8 (3.39)	15 (12.82)	11.42	0.001*
Anemia, n (%)	19 (8.05)	6 (5.08)	1.05	0.304
Trauma, n (%)	3 (1.28)	14 (11.86)	19.21	< 0.001*
Malignant tumor, n (%)	67 (28.39)	31 (26.27)	0.18	0.675
Charlson index	1 (0, 2)	2 (1, 2)	–2.796	0.005*
Surgery information				
Intraoperative Crystal liquid infusion (mL) [M (P25, P75)]	1250 (800, 1787.5)	1500 (675, 2162.5)	–0.839	0.401
Intraoperative Colloidal fluid infusion (mL) [M (P25, P75)]	500 (500, 500)	800 (500, 800)	–6.667	< 0.001*
Total intraoperative infusion (mL) [M (P25, P75)]	1800 (1400, 2400)	2375 (1575, 3200)	–3.899	< 0.001*
Intraoperative blood loss (mL) [M (P25, P75)]	20 (100, 200)	200 (100, 400)	–5.181	< 0.001*
Intraoperative blood transfusion volume (mL) [M (P25, P75)]	0 (0, 0)	0 (0, 732.5)	–4.896	< 0.001*
Intraoperative urine volume (mL) [M (P25, P75)]	200 (0, 500)	300 (100, 600)	–2.779	0.005*
Emergency operation, n (%)	21 (8.90)	58 (49.15)	73.53	< 0.001*
Operation time (min) [M (P25, P75)]	172.5(120, 238.75)	220 (143.75, 331.25)	–4.199	< 0.001*
PACU therapy, n (%)	157(66.53)	51(43.22)	17.63	< 0.001*
The use of vasoactive drugs, n (%)	68(28.81)	58(49.15)	14.20	< 0.001*
General anesthesia, n (%)	171(72.46)	112 (94.92)	24.75	< 0.001*
Intraoperative resuscitation, n (%)	1 (0.42)	5 (4.24)	4.768	0.029*
Type of surgery, n (%)				
Neurosurgery	36 (15.25)	18 (15.25)	0.000	> 0.999
General surgery	32 (13.56)	16 (13.56)		
Hepatobiliary Surgery	34 (14.41)	17 (14.41)		
Urological surgery	28 (11.86)	14 (11.86)		
Thoracic surgery	36 (15.25)	18 (15.25)		
Orthopedic surgery	38 (16.10)	19 (16.10)		
Obstetrics and Gynecology Surgery	32 (13.56)	16 (13.56)		
Postoperative information				
Postoperative ICU therapy, n (%)	4 (1.69)	23 (19.49)	32.883	< 0.001*
Postoperative hospitalization (days) [M (P25, P75)]	18 (11, 24)	9 (3, 23.5)	–4.952	< 0.001*
Second post-operative surgery, n (%)	6 (2.54)	11 (9.32)	7.909	0.005*

(Continued)

Table 1 (Continued).

	Control Group (n=236)	Observational Group (n=118)	Statistic	p value
Postoperative organ dysfunction, n (%)				
Postoperative AKI	2 (0.85)	9 (7.62)	9.863	0.002*
Postoperative ARDS	2 (0.85)	5 (4.24)	3.079	0.079
Postoperative delirium	1 (0.42)	3 (2.54)	1.549	0.213
Postoperative stroke	0 (0.00)	4 (3.39)	8.881	0.012*

Note: * $p < 0.05$ was statistically significant.

Abbreviations: BMI, body mass index; ASA, American Society of Anesthesiologists; NYHA, New York Heart Association; CHD, coronary heart disease; COPD, chronic obstructive pulmonary disease; PACU, postanesthesia care unit; ICU, intensive care unit; AKI, acute kidney injury; ARDS, acute respiratory distress syndrome.

hospitals was 3.3%. Compared with the Control group, the proportion of comorbidity with cerebral hernia, renal failure and trauma was 6.78%, 6.78% and 11.86% respectively in the Observational group, which are higher than the ration in the Control group ($p < 0.001$), accordingly the Charlson index is higher in the Observation group than the Control group ($p=0.005$). The proportion of using vasoactive drugs and general anesthesia was also higher in the Observational group, the ration is 49.15% and 94.92%, respectively ($p < 0.001$). There was also significant difference in the amount of intraoperative blood transfusion and blood loss volume between the two groups ($p < 0.001$). Compared with the Control group, the Observational group had more intraoperative colloidal fluid infusion ($p<0.001$), total intraoperative infusion ($p<0.001$) and intraoperative urine output volume ($p=0.005$). After the operation, more patients in the Observation group were transmitted to the ICU ($p<0.001$) and suffered the second postoperative surgery ($p=0.005$). There were more patients in the Observation group diagnosed with postoperative AKI ($p=0.002$) and postoperative

stroke ($p=0.012$). Compared with the Observation group, patients in the Control group had experienced a longer postoperative hospitalization ($p<0.001$) (See Table 1).

The Blood Pressure in Each Section Between Two Groups

The duration of MAP and SBP in different blood pressure ranges was calculated respectively, and their proportion to the total operation time was expressed as the median and quartile spacing M (P25, P75) (see Tables 2 and 3). Compared with the Observational group, the median blood pressure duration ratio of the Control group was higher at the blood pressure interval of $\text{MAP} \geq 75\text{mmHg}$.

The Result of Univariate Logistic Regression Analysis

Univariate logistic regression analysis was used to analyze the potential relationship among intraoperative blood pressure, other clinical indicator, and postoperative mortality. The results indicated that some blood pressure sections, including MAP ($\text{MAP} \leq 55\text{mmHg}$, $55\text{mmHg} < \text{MAP} \leq$

Table 2 Percentage of Cumulative Duration of MAP in Total Operation Time [% , M (P25, P75)]

Group	$\leq 55 \text{ mmHg}$	$> 55 \sim 60 \text{ mmHg}$	$> 60 \sim 65 \text{ mmHg}$	$> 65 \sim 70 \text{ mmHg}$	$> 71 \sim 75 \text{ mmHg}$	$> 75 \text{ mmHg}$
Observational group	0 (0,1.84)	0 (0,4.13)	4.92 (0,10.78)	8.82 (1.44, 15.94)	12.27 (4.42, 19.34)	63.78 (44.23,90.38)
Control group	0 (0, 0)	0 (0,1.58)	0 (0,5.88)	4.23 (0,14.49)	8.45 (0,17.17)	79.67 (55.02, 98.24)

Table 3 Percentage of SBP Cumulative Duration in Total Operation Time [% , M (P25, P75)]

Group	$\leq 80 \text{ mmHg}$	$> 80 \sim 90 \text{ mmHg}$	$> 90 \sim 100 \text{ mmHg}$	$> 100 \sim 110 \text{ mmHg}$	$> 110 \sim 120 \text{ mmHg}$	$> 120 \text{ mmHg}$
Observational group	0 (0,2.42)	0 (0, 6)	5.82 (0,16.80)	14.04 (4.82, 22.43)	15.21 (9.2, 25.86)	50.78 (25.27, 69.14)
Control group	0 (0,0.41)	0 (0,3.30)	4.51 (0,15.31)	12.29 (1.72, 25)	17 (7.14, 27.92)	50 (22.73, 81.7)

60mmHg, 60mmHg < MAP ≤ 65mmHg, 65mmHg < MAP ≤ 70mmHg, 70mmHg < MAP ≤ 75mmHg) and SBP (SBP ≤ 80mmHg, 80mmHg < BP ≤ 90mmHg), might increase the postoperative mortality (OR>1). Other factors might be related to postoperative death of elderly patients ($p < 0.05$), such as ASA classification (OR=5.276, 95% CI, 3.504 to 7.946), cardiac function classification (OR=1.546, 95% CI, 1.042 to 2.295), method of general anesthesia (OR=7.096, 95% CI, 2.974 to 16.929), COPD (OR=4.191, 95% CI, 1.772 to 10.199), Emergency operation (OR=9.897, 95% CI, 5.567 to 17.594), severe trauma (OR=10.41, 95% CI, 2.929 to 37.003), intraoperative use of vasoactive drugs (OR=2.388, 95% CI, 1.511 to 3.775), intraoperative blood loss volume (OR=19.104, 95% CI, 8.716 to 41.875) and blood transfusion (OR=1.98, 95% CI, 1.151 to 3.404). MAP > 75mmHg (OR=0.346, 95% CI, 0.123 to 0.973) and PACU therapy (OR=0.383, 95% CI, 0.243 to 0.603) may be seen as a protective factor (See Table 4).

The Result of Multivariate Logistic Regression Analysis

According to the statistically significant variables in the results of univariate analysis, six related factors affecting the postoperative death of elderly patients were obtained by multivariate Logistic regression analysis (See Table 5).

Five variables: emergency surgery, general anesthesia, COPD, ASA classification, and 60mmHg < MAP ≤ 65mmHg were considered as risk factors for postoperative death in elderly patients (OR > 1). Among them, the risk of emergency surgery is 2.2 times higher than that of elective surgery, the risk increases by 2.96 times for every increase in ASA classification level and patients with a history of COPD would have a 4.2 times higher risk for postoperative death. 60mmHg < MAP ≤ 65mmHg is a high-risk factor for postoperative death in elderly patients. When the proportion of MAP between 60mmHg and 65mmHg exceeds 10%, the risk of postoperative death would increase by 1.5 to 2.25 times. Correspondingly, PACU therapy is a protective factor for postoperative death in elderly patients. The risk of postoperative death for patients observed by PACU was reduced by 80%. (OR < 1).

ROC Curve and the Determination of the Optimal Critical Value of the Regression Model

According to the results of multivariate regression analysis in Table 3, the regression equation is: $Y = -5.893 + 0.815X_1$

$+ 1.086X_2 + 2.761X_3 + 0.409X_4 + 1.415X_5 - 1.472X_6$.

According to the above regression equation, P value of prediction probability of postoperative death of elderly patients is calculated, $P = 1/1 + e^{-Y}$. The ROC curve of P value is established (See Figure 2). The area under the curve (AUC) is 0.850, and the 95% confidence interval (CI) is (0.805,0.894). The optimal critical P value on the ROC curve is 0.338, and the corresponding sensitivity and specificity are 73.5% and 82.2% respectively.

Discussion

In this logistic regression analysis study, we reported that MAP between 60mmHg and 65mmHg had a strong relationship with postoperative mortality in elderly patients. Additionally, ASA classification, emergency surgery, the method of general anesthesia, history of COPD were considered as high-risk factors (OR > 1). By contrast, PACU therapy was considered as a protective factor. The area under the ROC curve is 0.850, indicating that these clinical indicators are of great value in predicting postoperative mortality in elderly patients.

The result of this study on the lower threshold of intraoperative blood pressure in elderly patients is the most important achievement. The decrease in blood pressure during anesthesia and operation is inevitable. After the induction of general anesthesia, the circulatory inhibitory effects will make the peripheral vascular dilation and reduce peripheral vascular resistance; in spinal and regional anesthesia, the peripheral vascular will dilate as well. When the blood pressure decreases to some degree, it can be seen as intraoperative hypotension. In adult noncardiac surgery, some previous studies have suggested as follows: the morbidity of acute kidney injury and acute myocardial injury increased significantly when intraoperative MAP < 55 mmHg, even for a shorter duration;^{8,9} when the intraoperative MAP was 30% lower than the basal blood pressure level, the incidence of postoperative stroke would increase;³ continuous MAP < 64 mmHg was closely associated with stroke during cardiopulmonary bypass;⁷ in elderly patients with hypertension, the self-adjusting ability of the brain and kidneys may be impaired, causing the organs more prone to ischemia during hypotension.^{10,11} In current guidelines for noncardiac surgery, the American College of Cardiology (ACC) and the American Heart Association (AHA) recommends individualized management of surgical patients with preoperative complications.¹² These studies and consensus highlight the importance of blood pressure management in reducing

Table 4 Univariate Logistic Regression Analysis of Postoperative Mortality in Elderly Patients

	Regression Coefficient (B)	The Standard Deviation (SE)	Wald Value	P	The OR Value	95% CI
Age	0.197	0.151	1.710	0.191	1.218	0.906 ~ 1.637
Gender	0.378	0.235	2.587	0.108	1.459	0.921 ~ 2.311
Body mass index (BMI)	0.066	0.035	3.458	0.063	0.936	0.873 ~ 1.004
ASA classification	1.663	0.209	63.411	< 0.001*	5.276	3.504 ~ 7.946
NYHA cardiac function	0.436	0.202	4.675	0.031*	1.546	1.042 ~ 2.295
Hypertension	0.307	0.226	1.839	0.175	1.359	0.872 ~ 2.117
CHD	0.285	0.289	0.974	0.324	1.330	0.755 ~ 2.345
Cerebral hernia	2.839	1.067	7.078	0.008*	17.091	2.111 ~ 138.338
Arrhythmia	0.270	0.375	0.516	0.472	0.764	0.366 ~ 1.593
Diabetes	0.201	0.291	0.478	0.489	1.223	0.691 ~ 2.164
COPD	1.433	0.454	9.975	0.002*	4.191	1.722 ~ 10.199
Anemia	0.491	0.483	1.037	0.309	0.612	0.238 ~ 1.575
Trauma	2.343	0.647	13.109	< 0.001*	10.41	2.929 ~ 37.003
Malignant tumor	0.107	0.254	0.176	0.675	0.899	0.546 ~ 1.479
Blood pressure						
MAP ≤ 55 mmHg	0.557	0.201	7.696	0.006*	1.745	1.178 ~ 2.587
55 < MAP ≤ 60 mmHg	0.357	0.179	3.996	0.046*	1.430	1.007 ~ 2.030
60 < MAP ≤ 65 mmHg	0.537	0.150	12.809	< 0.001*	1.712	1.275 ~ 2.297
65 < MAP ≤ 70 mmHg	0.381	0.134	8.013	0.005*	1.463	1.124 ~ 1.905
70 < MAP ≤ 75 mmHg	0.373	0.145	6.600	0.010*	1.452	1.092 ~ 1.929
MAP > 75 mmHg	1.061	0.527	4.050	0.044*	0.346	0.123 ~ 0.973
SBP ≤ 80 mmHg	0.413	0.205	4.069	0.044*	1.511	1.012 ~ 2.258
80 < SBP ≤ 90 mmHg	0.342	0.156	4.849	0.028*	1.408	1.038 ~ 1.910
90 < SBP ≤ 100 mmHg	0.094	0.133	0.500	0.480	1.098	0.847 ~ 1.425
100 < SBP ≤ 110 mmHg	0.252	0.144	3.058	0.080	1.286	0.970 ~ 1.705
110 < SBP ≤ 120 mmHg	0.168	0.165	1.027	0.311	1.183	0.855 ~ 1.635
SBP > 120 mmHg	0.198	0.229	0.748	0.387	1.219	0.778 ~ 1.912
General anesthesia	1.959	0.444	19.507	< 0.001*	7.096	2.974 ~ 16.929
Emergency operation	2.292	0.294	60.969	< 0.001*	9.897	5.567 ~ 17.594
PACU therapy	0.960	0.231	17.195	< 0.001*	0.383	0.243 ~ 0.603
The use of vasoactive drugs	0.871	0.234	13.889	< 0.001*	2.388	1.511 ~ 3.775
Intraoperative urinary volume	0.062	0.248	0.062	0.803	0.940	0.578 ~ 1.528
Intraoperative blood loss	2.950	0.400	54.278	< 0.001*	19.104	8.716 ~ 41.875
Intraoperative blood transfusion volume	0.683	0.277	6.093	0.014*	1.980	1.151 ~ 3.404

Note: * $p < 0.05$ was statistically significant.

Abbreviations: BMI, body mass index; ASA, American Society of Anesthesiologists; NYHA, New York Heart Association; CHD, coronary heart disease; COPD, chronic obstructive pulmonary disease; PACU, postanesthesia care unit; ICU, intensive care unit; AKI, acute kidney injury; ARDS, acute respiratory distress syndrome.

Table 5 Multivariate Logistic Regression Analysis of Postoperative Mortality in Elderly Patients

Variable	Regression Coefficient (B)	The Standard Deviation (SE)	Wald Value	P	The OR Value	95% CI
X ₁ Emergency operation	0.815	0.403	4.089	0.043	2.260	1.025~4.982
X ₂ ASA classification	1.086	0.278	15.276	< 0.001	2.963	1.719~5.109
X ₃ General anesthesia	2.761	0.570	23.428	< 0.001	15.819	5.171~48.391
X ₄ 60mmHg < MAP ≤ 65mmHg	0.409	0.197	4.305	0.038	1.505	1.023~2.215
X ₅ COPD	1.415	0.569	6.186	0.013	4.117	1.350~12.557
X ₆ PACU therapy	-1.472	0.363	16.404	< 0.001	0.230	0.113~0.468
Constant	-5.893	0.911	41.828	< 0.001	0.003	

Note: $p < 0.05$ was statistically significant.

postoperative mortality and improving prognosis in elderly patients. However, there is no existing consensus on the maintenance range of blood pressure in elderly patients. From this point of view, our study provides evidence for clarifying the goal of blood pressure management in elderly patients. Our study regards the MAP between 60 and 65 mmHg as a risk factor for postoperative mortality in elderly patients. Although the results of our study did not show the correlation between MAP below 60mmHg and the postoperative mortality, combined with the theory of ischemia-reperfusion injury caused by hypotension, it can be inferred that MAP below 60mmHg may also affect postoperative mortality of elderly patients. A recent meta-analysis indicated that intraoperative hypotension was associated with an increased risk of postoperative AKI,

myocardial injury and 30-day mortality in non-cardiac surgery and suggested that intraoperative MAP < 60 mmHg should be avoided.¹³ So maybe MAP = 65mmHg can be regarded as the lower threshold of intraoperative blood pressure in elderly patients undergoing noncardiac surgery. When intraoperative MAP ≤ 65mmHg, it can be called an intraoperative hypotension event, which is a risk factor of postoperative mortality in elderly patients. It has been reported in adult noncardiac surgery that the duration of MAP < 49mmHg more than 5 minutes may increase the outcome of 30-day postoperative mortality.⁵ Our results of this study suggest that the threshold of intraoperative hypotension in elderly patients is 65mmHg, which is higher than that of 49mmHg. The reason behind this difference may be that the vascular system of elderly patients is more fragile and the ability of self-regulation is reduced. It is known that renal blood flow and glomerular filtration rate can keep stable when the MAP is in the range of 75-170 mmHg,¹⁴ however these indicates will become pressure-dependent once MAP beyond the range. Our study also indicates that the Observation group has a higher incidence of postoperative AKI, which may be related to the postoperative mortality rate. Therefore, MAP > 65mmHg should be maintained as far as possible in the perioperative circulation management of elderly patients.

The definition of intraoperative hypotension has not yet been unified. The absolute and relative definition of intraoperative hypotension have all been used in researches.⁸ Our study did not choose to use a relative intraoperative hypotension which may have a 20–40% drop in intraoperative blood pressure compared with preoperative blood pressure, based on the following considerations: Firstly, this study is a retrospective study, it is difficult to determine the accurate preoperative blood pressure; secondly, the relative intraoperative hypotension has not yet been unified and determined; thirdly, studies

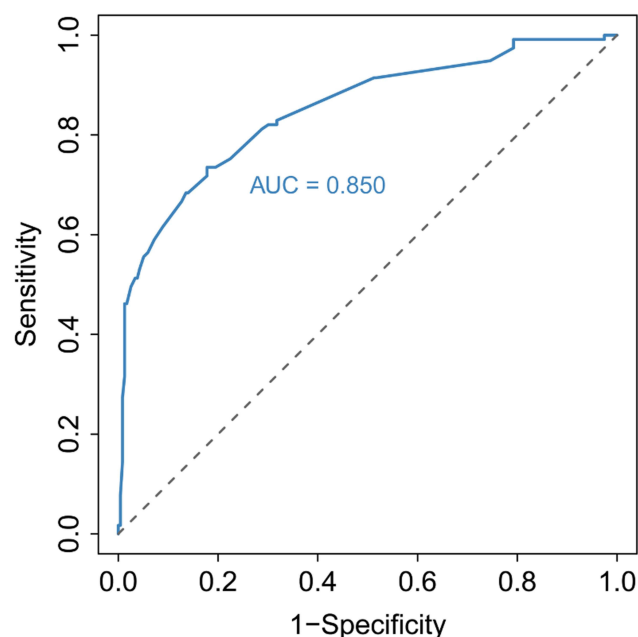


Figure 2 The ROC curve of P value predicted by logistic regression mode and the area under the curve (AUC).

have found that it would be feasible and beneficial to use the absolute value as the lower limit of intraoperative hypotension, it would be more easily for anesthesiologists to master and to incorporate into their clinical decision systems.

The results of our study about the high-risk factor, ASA grade, emergency surgery, as well as the protective factor, PACU therapy, in elderly patients are consistent with the current point of view and practice. According to the definition of ASA grade, the higher the ASA grade, the higher the anesthetic risk and postoperative mortality. Emergency surgery has been reported as an independent risk factor for postoperative mortality. And the significance of PACU therapy is to pay close attention to postoperative patients reducing the incidence of adverse prognosis.

COPD is estimated to be the third leading cause of death worldwide by 2020.¹⁵ Recent studies demonstrated that pneumonia and COPD individually affected the 30-day mortality and total mortality,¹⁶ which is consistent with our study. Elderly patients not only have a high incidence of COPD but also have a higher risk of postoperative mortality than young COPD patients.¹⁷ COPD is characterized by persistent airflow limitation, which in turn affects gas exchange, in severe cases, can lead to pulmonary hypertension and develop into cor pulmonale. Anesthetic drugs and anesthetic methods will inevitably affect the respiratory function of patients, not to mention the effects of surgical posture and other factors on the respiratory function of patients.¹⁸ It has been reported that postoperative pulmonary complications in patients with COPD are increased, and the occurrence of postoperative pulmonary complications is a high-risk factor for postoperative death.^{19,20} This is consistent with our findings that preoperative COPD is a high-risk factor for postoperative death in elderly patients. At present, the understanding of anesthetic management of elderly patients with COPD is still insufficient, our study highlights the necessity of paying attention to perioperative elderly patients with COPD and provides ideas for further research.

Our results demonstrate that general anesthesia is a risk factor about the postoperative mortality, so sufficient vigilance should be maintained in the choice of anesthetic methods for elderly patients. The choice and the safety of different anesthesia methods are still controversial. For geriatric patients undergoing hip surgery, nerve blocking

anesthesia was associated with fewer odds of adverse outcomes than general anesthesia.²¹ Some studies have shown that the use of regional anesthesia in patients with COPD is associated with lower incidences of composite morbidity, pneumonia, prolonged ventilator dependence, and unplanned postoperative intubation.^{22,23} From the results of our research, On the premise of meeting the surgical needs, avoiding general anesthesia for elderly patients may reduce postoperative mortality and improve the outcome. The results of our study also provide objective evidence for anesthesiologists to choose the method of anesthesia in the face of elderly patients.

Our study also has some limitations. Firstly, six related factors were obtained by multi-factor logistic analysis, which shows that the sample size is sufficient. However, a large number of variables are included in the univariate logistic regression analysis, which indicates that the sample size may be a little small. Secondly, there is a difference in the duration of operation between the Observation group and the Control group. The analysis of blood pressure is based on the proportion of the duration of each blood pressure segment, and the threshold of the duration of intraoperative hypotension can not be obtained. Thirdly, we explored the threshold of hypotension in elderly patients, but the impact of hypotension on the function of important organs is not confirmed by laboratory indicators.

Conclusions

Elderly patients are vulnerable groups, combined with our research results, attention should be paid to patients undergoing emergency surgery, with high ASA grade, and with a history of COPD. The mode of anesthesia should be prudently chosen and the MAP should be maintained above 65 mmHg intraoperatively. Optimizing the perioperative management of elderly patients is the key point to reduce the mortality of geriatric patients.

Abbreviations

95% CI, 95% confidence interval; ASA, American Society of Anesthesiologists; AUC, Area under the curve; IOH, intraoperative hypotension; COPD, chronic obstructive pulmonary disease; PACU, postanesthesia care unit; ROC, Receiver operating characteristic; RR, Risk ratio; MAP, mean arterial pressure; SBP, systolic blood pressure; MODS, multiple organs dysfunction syndrome; OR, odds ratio; ACC, American College of Cardiology; AHA, American Heart

Association. BMI, Body Mass Index; NYHA, New York Heart Association; CHD, Coronary Heart Disease; ICU, Intensive Care Unit; AKI, Acute Kidney Injury; ARDS, Acute Respiratory Distress Syndrome.

Data Sharing Statement

The datasets used and/or analysed during the current study are available from the corresponding author, Jianteng Gu, on reasonable request.

Ethics Approval and Consent to Participate

The study was approved and the written informed consent was waived by the Ethics Committee of the First Affiliated Hospital (Southwest Hospital) of Army Medical University, PLA [KY2020182], by the Medical Ethics Committee of Second Affiliated Hospital (Xinqiao Hospital) of Army Medical University, PLA [2020-119-01] and by the Ethics Committee of Army Specialty Medical Center (Daping Hospital) of Army Medical University, PLA [2020.No.147] respectively. This study has been retrospectively registered in the Chinese Clinical Trials Registry (ChiCTR2000038912, 10/10/2020). All methods performed in this study were in accordance with the relevant guidelines and regulations. All procedures performed in this study involving human participants were in accordance with the Ethical Standards of the Institutional Ethics Committee and with the 1964 Helsinki Declaration and its later amendments or comparable ethical standards.

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Disclosure

The authors report no conflicts of interest in this work.

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