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The value of cardiopulmonary comorbidity in patients with acute large vessel occlusion stroke undergoing endovascular thrombectomy: a retrospective, observational cohort study

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Abstract

Background Chronic lung and heart diseases are more likely to lead an intensive end point after stroke onset. We aimed to investigate characteristics and outcomes of endovascular thrombectomy (EVT) in patients with acute large vessel occlusion stroke (ALVOS) and identify the role of comorbid chronic cardiopulmonary diseases in ALVOS pathogenesis.

Methods In this single-center retrospective study, 191 consecutive patients who underwent EVT due to large vessel occlusion stroke in neurological intensive care unit were included. The chronic cardiopulmonary comorbidities and several conventional stroke risk factors were assessed. The primary efficacy outcome was functional independence (defined as a mRS of 0 to 2) at day 90. The primary safety outcomes were death within 90 days and the occurrence of symptomatic intracranial hemorrhage(sICH). Univariate analysis was applied to evaluate the relationship between factors and clinical outcomes, and logistic regression model were developed to predict the prognosis of ALVOS.

Results Endovascular therapy in ALVOS patients with chronic cardiopulmonary diseases, as compared with those without comorbidity, was associated with an unfavorable shift in the NHISS 24 h after EVT [8(4,15.25) versus 12(7.5,18.5), P = 0.005] and the lower percentage of patients who were functionally independent at 90 days, defined as a score on the modified Rankin scale of 0 to 2 (51.6% versus 25.4%, P = 0.000). There was no significant between-group difference in the frequency of mortality (12.1% versus 14.9%, P = 0.580) and symptomatic intracranial hemorrhage (13.7% versus 19.4%, P = 0.302) or of serious adverse events. Moreover, a prediction model showed that existence of cardiopulmonary comorbidities (OR=0.456, 95%CI 0.209 to 0.992, P = 0.048) was independently associated with functional independence at day 90.

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Conclusions EVT was safe in ALVOS patients with chronic cardiopulmonary diseases, whereas the unfavorable outcomes were achieved in such patients. Moreover, cardiopulmonary comorbidity had certain clinical predictive value for worse stroke prognosis.

Keywords Cerebral infarction, Acute ischemic stroke, Endovascular thrombectomy, Prognosis, Comorbidity

Background

Acute ischemic stroke (AIS) is a central nervous system (CNS) disease leading the second cause of death worldwide and is proved to be influenced by many factors [1]. Although limited by treatment time window and classification of cerebral infarction, the efficacy and safety of endovascular thrombectomy (EVT) for eligible patients with acute large vessel occlusion stroke (ALVOS) have been fully confirmed by positive results of large randomized clinical trials in recent years [2–4]. Treatment futility criteria for EVT—characteristics defining patients' subgroups that do not benefit from EVT—still remain elusive [5].

Recent data suggest that distinctive factors [6–9] may impact functional outcome in ALVOS patients. Chronic lung and heart diseases cause discomfort in activities of daily living (ADLs) or exercise [10] and are more likely to lead an intensive end point after stroke onset [11, 12]. Previous studies have focused on the impact of comorbidity on stroke, but none have considered whether comorbidity affect endovascular therapy and it is uncertain whether EVT could help retard functional deterioration. A better understanding of the association of comorbidity with stroke severity, intraoperative manipulation and outcomes is therefore required to understand apparent associations with poststroke functional independence. Therefore, our study was designed to test the hypothesis that cardiopulmonary comorbidities would lead to worse neurological functional outcomes by comparing the characteristics and outcomes between patients who underwent endovascular therapy with and without cardiopulmonary comorbidities.

Methods

Participant

This study was a retrospective analysis. We enrolled patients with anterior or posterior circulation LVOS who underwent EVT from January 2018 to September 2022 in our stroke center. The research protocol was reviewed and approved by the Ethics Committee (approval number: K2023-021-01). Written informed consent was obtained from all patients or their legal representatives. We have adhered to the STROBE statement (Supplementary material).

Patients were eligible if they met the following inclusion criteria. (a) Signs and symptoms consistent with the diagnosis of an acute ischemic stroke according to the criteria of Chinese guidelines for diagnosis and treatment of acute ischemic stroke 2018 [13]. (b) Age \geq 18 years. (c) Endovascular therapy can be initiated (femoral puncture) in 24 h of stroke onset. Stroke onset is defined as the time the patient was last known to be at their neurologic baseline. (d) Occlusion of the internal carotid artery, proximal segment of the middle cerebral artery (M1/M2), the anterior cerebral artery (A1/A2), or basilar artery (BA) can be confirmed by digital subtraction angiography (DSA) or computed tomographic angiography (CTA). The exclusion criteria includes: (a) intracranial hemorrhage; (b) massive brain infarction (anterior circulation infarction > 1/2 MCA territory, posterior circulation infarction > 2/3 Pontine or midbrain volume); (c) known allergy to iodine that precludes an endovascular procedure; (d) known hereditary or acquired hemorrhagic diathesis, coagulation factor deficiency; (e) previous severe disability [modified Rankin Score (mRS) $[14] \ge 3$]; (f) severe renal failure. The treatment protocol and methods have been published before [15]. The flow chart of the inclusion of the study population is displayed in Fig. 1.

Clinical assessment

The demographics characteristics and medical history of all patients were collected and recorded. Chronic cardiopulmonary comorbidity is a clear diagnosis of various cardiac diseases [rheumatic heart disease or various cardiomyopathy or coronary atherosclerotic heart disease (acute coronary syndrome or after coronary artery stent implantation or coronary artery bypass grafting)] with a New York Heart Association (NYHA) cardiac function grade III~IV or cardiac ultrasound showing left ventricular ejection fraction (LVEF) less than 45%; and/or a clear diagnosis of chronic obstructive pulmonary disease or pulmonary heart disease, with type I or type II respiratory failure before the stroke onset. Other clinical assessments were performed at baseline; assessments included the score on the modified Rankin scale (mRS) [14], the National Institutes of Health Stroke Scale (NIHSS) [16], the Trial of ORG 10,172 in Acute Stroke Treatment (TOAST) classification [17] and ASPECTS [18] or pc-ASPECTS [19]. The treatment-related information was



Fig. 1 Flow chart of the inclusion of the study population. mRS = modified Rankin Scale, OTP = symptoms onset to groin puncture time

recorded by a neurointerventionalist during thrombectomy therapy, including intravenous thrombolysis, occlusion site, tandem lesion, stroke onset to door time(ODT), door to puncture time (DPT), puncture to reperfusion time (PRT), anesthesia type, the thrombectomy approach (stent-retrievers or suction thrombectomy systems), rescue therapy (balloon dilation or stent implantation), times of pass through, modified thrombolysis in cerebral infarction (mTICI) score [20].

Outcome

The primary efficacy outcome was functional independence (defined as a score on the modified Rankin scale of 0 to 2) at day 90. The secondary efficacy outcomes were the NIHSS right after and 24 h after endovascular therapy. The primary safety outcomes were death within 90 days and the occurrence of symptomatic intracranial hemorrhage(sICH), defined as an increase of at least 4 points in the NIHSS score that was associated with parenchymal, subarachnoid or intraventricular hemorrhage on CT imaging within 36 h after thrombectomy. The outcomes were collected by telephone or outpatient consultation at day 90 after EVT.

Treatment

Patients with 6 to 24 h of onset were scanned by CT perfusion (CTP), and evaluated whether there was mismatch area. Anesthesia type depended on the situation, and femoral artery puncture was performed after anesthesia. Aspiration thrombectomy or stent-retriever thrombectomy was performed with approved thrombectomy device, at the discretion of the neurointerventionalist. If the thrombectomy did not successfully recanalize after 4 operations, balloon angioplasty or stent implantation could be used for rescue therapy. If the arteries were still blocked after rescue therapy, the treatment was considered to be failed. Permanent stenting and ballon angioplasty could be combined. All patients were admitted to neurological intensive care unit (NICU) for at least 72 h after the operation. Other perioperative medical therapy was carried out according to the guidelines [13].

Statistical analysis

Python 3.10 software was used for all the statistical analysis. The Kolmogorov-Smirnov method was used to test the normal distribution of the continuous variables. Then the mean \pm sd of the measurement data was used for the normal distribution, and the student test was used for the comparison between the groups. Metrical data that did not conform to normal distribution was expressed as median (interquartile range) [M (P25, P75)]; and Mann-Whitney U test was used for comparison between groups. The Categorical variables were expressed by the number of cases and percentage (%), and the comparison between groups was carried out by Chi-square test or Fisher's exact test as appropriate. Statistically significant differences were defined as P < 0.05. Moreover, the prognosis model for primary efficacy outcomes was calculated by logistic regression, in which the variables were tested in the univariate analysis. Statistically significant differences were defined as P < 0.1 in this case. And due to the unbalanced outcomes, the research participants were stratified randomly divided into a training set and an internal validation set at a ratio of 7:3 to better the logistic regression model. A receiver operating characteristic (ROC) curve was drawn, and the area under the curve (AUC) was calculated to evaluate the predictive efficacy of the logistic regression model.

Result

General characteristic

A total of 191 ALVOS patients who received EVT were included in this study. The median age was 70(59, 80) years. Among them, there were 67 patients with cardiopulmonary comorbidity (CC group) and 124 patients with non-cardiopulmonary comorbidity (NCC group). The demographics and baselines characteristics of the overall patients are shown in Table 1. The characteristics were not balanced. Patients with cardiopulmonary complications were more likely to be female (27.4% versus 56.7%, P=0.000), older [67(57, 77) versus 76 (67.5, 82.5), P=0.000], and to have atrial fibrillation (35.5% versus 71.4%, P=0.000) and coronary heart disease (14.5%)

versus 50.7%, P=0.000). Unexpectedly, the CC group was significantly less likely to smoke (24.2% versus 10.4%, P=0.022). There was difference in Toast classification of cerebral infarction between the two groups (P=0.000). Cardioembolism (46.8%) and large-artery atherosclerosis (43.5%) were the main causes in the NCC group, while large-artery atherosclerosis (80.6%) was the main cause in the CC group. The NIHSS scores of the whole cohort at admission ranged from 0 to 40, with an average of 17.53 ± 7.08. There was no difference in baseline NIHSS scores (P=0.293) and ASPECT (P=0.073) scores. Before stroke onset, the distribution of mRS scores had a significant difference between the two groups (P=0.023). Patients with cardiopulmonary complications were less likely to have pre-stroke mRS score at 0 (82.3% versus 67.2%).

Clinical characteristic

The clinical characteristics of the overall patients are shown in Table 2. The times of pass through was 2(1, 4) in the CC group and more than 1 (1, 3) in the NCC group, which were significantly different (P=0. 001). The proportion of balloon angioplasty (33.1% versus 11.9%,

 Table 1
 Demographics and baseline characteristics stratified by cardiopulmonary comorbidities

	0.000*
	0.000*
Age, median(IQR) 70 (59,80) 67 (57,77) 76 (67.5,82.5) 2782.000 ^b	*
Male, n(%) 119 (62.3%) 90(72.6%) 29 (43.3%) 15.896 ^c	0.000
Medical history, n(%)	
lschemic stroke 51 (26.7%) 29 (23.4%) 22 (32.8%) 1.984 ^c	0.159
Diabetes mellitus 75 (39.3%) 46 (37.1%) 29 (43.3%) 0.698 ^c	0.403
Hypertension 129 (67.5%) 82 (66.1%) 47 (70.1%) 0.321 ^c	0.571
Hyperlipidemia 74 (38.7%) 42 (33.9%) 32 (47.8%) 3.536 ^c	0.060
Atrial fibrillation 92 (48.2%) 44 (35.5%) 48 (71.4%) 22.778 ^c	0.000*
Coronary heart disease 52 (27.2%) 18 (14.5%) 34 (50.7%) 28.817 ^c	0.000*
Smoking 37 (19.4%) 30 (24.2%) 7 (10.4%) 5.262 ^c	0.022*
Drinking 19 (9.9%) 16 (12.9%) 3 (4.5%) 3.447 ^c	0.063
TOAST classification, n(%) 24.728 ^c	0.000*
Cardioembolism 70 (36.6%) 58 (46.8%) 12 (17.9%)	
Large-artery atherosclerosis 108 (56.5%) 54 (43.5%) 54 (80.6%)	
Other 13 (6.8%) 12 (9.7%) 1 (1.5%)	
Baseline NIHSS, mean ± SD 17.53 ± 7.08 17.14 ± 7.44 18.27 ± 6.35 -1.054 ^a	0.293
Pre-stroke mRS score, n(%) 6.002 ^c	0.0497*
0 147 (77.0%) 102 (82.3%) 45 (67.2%)	
1 26 (13.6%) 14 (11.3%) 12 (17.9%)	
2 18 (9.4%) 8 (6.5%) 10 (14.9%)	
ASPECTS 8.52±1.67 8.68±1.56 8.22±1.82 1.802 ^a	0.073

IQR Interquartile range, SD Standard deviation, TOAST The Trial of ORG 10,172 in Acute Stroke Treatment, NIHSS The National Institutes of Health Stroke Scale, mRS Modified Rankin Scale, ASPECTS Alberta stroke program early CT score or pc-ASPECTS

* for *P* < 0.05

 $^a\,$ for t-value, b for Z-value, $^c\,$ for $\chi^2\,$ value

Characteristics	All patients (n = 191)	NCC group (<i>n</i> = 124)	CC group (n=67)	Test value	P-value
Intravenous thrombolysis, n(%)	75 (39.3%)	52 (41.9%)	23 (34.3%)	1.055 ^c	0.304
General anesthesia, n(%)	109 (57.05%)	70 (56.5%)	39 (58.2%)	0.055 ^c	0.815
Tandem lesion, n(%)	44 (23.0%)	32 (25.8%)	12 (17.9%)	1.530 ^c	0.216
Times of thrombectomy, median(IQR)	2 (1,3)	1 (1,3)	2 (1,4)	2984 ^b	0.001*
Balloon angioplasty, n(%)	49 (25.6%)	41 (33.1%)	8 (11.9%)	10.177 ^c	0.001*
Stent implantation, n(%)	27 (14.1%)	23 (18.5%)	4 (6.0%)	5.670 ^c	0.017*
Recanalization (mTICI 2b/3), n(%)	167 (87.4%)	110 (88.7%)	57 (85.1%)	0.523 ^c	0.470
Intraoperative embolization, n(%)	24 (12.6%)	14 (12.9%)	10 (14.9%)	0.523 ^c	0.470
Location of occlusion, n(%)				4.054 ^c	0.132
Anterior circulation	162 (84.3%)	105 (84.7%)	57 (85.1%)		
Posterior circulation	27 (14.1%)	19 (15.3%)	8 (11.9%)		
Both anterior and posterior	2 (1%)	0 (0%)	2 (3.0%)		
ODT (min), mean ± SD	260.47±225.07	269.95 ± 219.14	242.93 ± 236.33	0.794 ^a	0.428
DPT (min), mean ± SD	91.04±57.93	94.67±65.03	84.33±41.33	1.182 ^a	0.239
PRT (min), mean ± SD	81.05 ± 57.39	78.43 ± 55.80	85.91 ± 60.34	-0.861ª	0.390

 Table 2
 clinical characteristics stratified by cardiopulmonary comorbidities

IQR Interquartile range, SD Standard deviation, mTICI Modified thrombolysis in cerebral infarction, NIHSS The National Institutes of Health Stroke Scale, ODT Onset to door time, DPT Door to puncture time, PRT Puncture to recanalization time

* for *P*<0.05

 $^a\,$ for t-value, $^b\,$ for Z-value, $^c\,$ for $\chi^2\,$ value

P=0.001) and that of stent implantation (18.5% versus 6.0%, P=0.017) were lower in the CC group. There was no significant difference between the two groups in anesthesia type, tandem lesions, vascular recanalization, embolism occurred during operation, sites of occlusion, time from onset to hospital, door to puncture time (DPT), puncture to recanalization time (PRT), and NIHSS score after EVT (P > 0.05).

Outcome

Efficacy and safety outcomes are presented in Table 3. The NIHSS at 24 h after EVT was higher in the CC group, and the difference was statistically significant [8(4,15.25) versus 12(7.5,18.5), P=0.005]. At day 90, the good functional outcomes were achieved in 17 (25.4%) patients in the CC group, which was significantly less (P=0.000) than those in the NCC group. The mortality rate was numerically lower in the NCC group, but the difference was not significant (12.1% versus 14.9%, P=0.580). Out of the total number of 25 mortality cases, 10 cases were in the CC group, including 5 cases of cerebral hernia or hemorrhage, 1 case of heart failure, 2 cases of myocardial infarction, 1 case of pulmonary infection and 1 case of renal failure, while the other 15 cases were in the NCC group, including 10 cases of cerebral hernia or hemorrhage, 1 case of gastrointestinal hemorrhage, 1 case of

Table 3 Outcomes stratified by cardiopulmonary comorbidities

Characteristics	All notionts	NCC aroun		Test value	<i>P</i> -value
Characteristics	(n=191)	(<i>n</i> = 124)	(<i>n</i> =67)	lest value	
90d mRS 0~2, n(%)	81 (42.4%)	64 (51.6%)	17 (25.4%)	12.262 ^c	0.000*
NIHSS after EVT, mean \pm SD	14.03 ± 9.51	13.51 ± 9.78	15.01 ± 8.97	-1.045 ^a	0.297
NIHSS at 24 h, median (IQR)	10 (4,17.5)	8(4,15.25)	12 (7.5,18.5)	3210.50 ^b	0.005*
Mortality, n(%)	25 (13.1%)	15 (12.1%)	10 (14.9%)	0.306 ^c	0.580
Hemorrhage transformation, n(%)	30 (15.7%)	17 (13.7%)	13 (19.4%)	1.065 ^c	0.302
Brain herniation, n(%)	19 (9.9%)	12 (9.7%)	7 (10.4%)	0.029 ^c	0.658
Decompressive hemicraniectomy, n(%)	10 (5.2%)	7 (5.6%)	3 (4.5%)	0.1200 ^c	0.73

IQR Interquartile range, SD Standard deviation

* for *P*<0.05

 $^a\,$ for t-value , $\,^b\,$ for Z-value , $\,\,^c$ for $\chi^2\,$ value

pulmonary embolism and 2 cases of respiratory failure. There was no difference in the proportion of any postoperative complications, including incidence of ISCHs, brain herniation and decompressive hemicraniectomy. To verify whether the age made the outcomes worse, we did a subgroup analysis between 22 elderly and 45 young patients in the CC group. The result showed that there was no significant difference in 90d good functional outcomes [26.7%(12/45) versus 29.4%(5/22), χ 2=0.121, *P*=0.728].

Prognosis analysis

Then, a univariate analysis of all the patients is shown in Table 4. The good functional outcomes at day 90 (mRS score 0-2) were achieved in 81 (42.4%) patients. Among the variables that could be accessed to before EVT, the following was identified as predictors of a good outcome at 3 months (P < 0.1): Age, Ischemic stroke, Diabetes mellitus, Hyperlipidemia, Coronary heart disease, cardiopulmonary complication, intravenous thrombolysis, Baseline NIHSS. When predictors were selected for the logistic regression analysis (Table 5; Fig. 2), history of diabetes mellitus (OR=0.459, 95%CI 0.219 to 0.960, P=0.039), higher baseline NIHSS (OR=0.880, 95%CI 0.820 to 0.935, P=0.000) and existence of cardiopulmonary comorbidities (OR=0.456, 95%CI 0.209 to 0.992, P=0.048) were associated with a poor prognosis. The AUC of the logistic regression model was 0.75 (Fig. 3), the weighed F1 score was 0.73. It indicated that cardiopulmonary comorbidity was an independent risk factor for the prognosis.

Discussion

We studied the characteristics and outcomes of thrombectomy in patients with cardiopulmonary diseases. Our retrospective study finds that baseline characteristics are not balanced, there are more female, older, higher proportion of atrial fibrillation and coronary heart disease, and smoking. These variables are risk factors of chronic lung [21] and cardiac [22, 23], as well as stroke. A number of retrospective studies have shown that patients with congestive heart failure is associated with atrial fibrillation (AF) [22, 24], which is consistent with the result in our study. Although ALVOS patients in the CC group have more AF, fewer stroke onset was caused by cardioembolism. It is speculated that long-term chronic cardiopulmonary dysfunction may result in an increase in the proportion of large-vessel atherosclerosis infarction.Chronic heart failure reduced cerebral blood flow and abnormal auto-regulation might result in impaired perfusion and arterial stiffness in the brain [25, 26]. Previous study showed poor pulmonary dysfunction could cause cerebral small vessel diseases [21, 27]. The lungs and heart are irrevocably linked in their oxygen and carbon dioxide transport functions [28]. Cardiopulmonary comorbidity reduces oxygen supply to the brain, which could lead to endothelial damage and accelerate the process of atherosclerosis, then aggravate the negative effect on pathogenesis of ischemia lesions.

Previous studies have shown that age is one of the factors of poor outcomes, but few have studied whether these populations comorbid with cardiopulmonary diseases. Our study shows that chronic cardiopulmonary comorbidity is a risk factor to the poor prognosis after EVT. Then, we performed a subgroup analysis of the CC group, predicting that the process of comorbidity leading to poor outcomes is not related to the age. On the contrary, the poorer cardiopulmonary function may account for the poor outcomes in elderly patients with ALVOS.

In terms of clinical characteristics between groups, patients with cardiopulmonary comorbidities were more likely to have higher mRS before stroke onset, while there was no difference in the baseline NIHSS score between the two groups. Therefore, we suspect that chronic cardiopulmonary comorbidity leads to slight neurological disability before EVLOS onset, instead of aggravating impairment at stroke onset. Besides, our results are comparable to the rate of recanalization achieved in large randomized trials with similar inclusion criteria [2, 29-31]. There were more intraoperative embolization and less rescue therapy including both balloon angioplasty and stent implantation in the CC group, indicating that treatment was more difficult. According to the recent studies, the composition of the thrombus may impact mechanical thrombectomy outcome [9]. Fibrin-rich thrombi have a less favorable outcome, mostly due to their increased stiffness and resistance to mechanical thrombectomy [8]. It has been acknowledged that one of the major hidden cause of occlusions refractory to modern mechanical thrombectomy procedures is underlying severe intracranial atherosclerotic stenosis (ICAS) [32, 33]. According to the Chinese Intracranial Atherosclerosis study, ICAS was the most common vascular lesion of stroke patients in China [34]. We could infer that chronic cardiopulmonary comorbidity may be accompanied by more ICAS before stroke onset and cause in situ thrombotic occlusion.

The poor efficacy outcomes of patients with cardiopulmonary dysfunction could be indicated at earliest at 24 h after EVT by the NIHSS and last until at day 90 by mRS, which was consistent with a study of MR CLEAN and IMS III trial [35]. Additionally, results of safety outcomes were no difference. The overall mortality rate was numerically lower in the NCC group, but the difference was not significant. Patients with chronic cardiopulmonary comorbidities were more likely to death from other system failure. In agreement of the previous findings that

Table 4 Univariate analysis of 90-days outcomes

Characteristics	All patients (n=191)	Good outcome (n=81)	outcomePoor outcome1)(n = 110)		Good outcomePoor outcomeTest value(n = 81)(n = 110)	Test value	<i>P</i> -value	
 Male, n(%)	119 (62.3%)	54 (66.7%)	65 (59.1%)	1.140 ^c	0.286			
Medical history, n(%)								
Ischemic stroke	51 (26.7%)	16 (19.8%)	35 (31.8%)	3.470 ^c	0.063			
Diabetes mellitus	75 (39.3%)	22 (27.2%)	53 (48.2%)	8.644 ^c	0.003**			
Hypertension	129 (67.5%)	52 (64.2%)	77 (70.0%)	0.716 ^c	0.397			
Hyperlipidemia	74 (38.7%)	25 (30.9%)	49 (44.5%)	3.679 ^c	0.055			
Atrial fibrillation	92 (48.2%)	34 (42.0%)	58 (52.7%)	2.160 ^c	0.142			
Coronary heart disease	52 (27.2%)	13(16.0%)	39 (35.5%)	8.866 ^c	0.003**			
Smoking	37 (19.4%)	19 (23.5%)	18 (16.3%)	1.503 ^c	0.220			
Drinking	19 (9.9%)	10 (12.3%)	9 (8.1%)	0.903 ^c	0.342			
Cardiopulmonary comorbidities, n(%)	67 (35.1%)	17 (21.0%)	50 (45.5%)	12.262 ^c	0.000**			
Intravenous thrombolysis, n(%)	75 (39.3%)	38 (46.9%)	37 (33.6%)	3.448 ^c	0.063*			
General anesthesia, n(%)	109 (57.05%)	43 (53.1%)	66 (60.0%)	0.910 ^c	0.340			
Tandem lesion, n(%)	44 (23.0%)	17 (21.0%)	27 (24.5%)	0.333 ^c	0.564			
Balloon angioplasty, n(%)	49 (25.6%)	23 (28.4%)	26 (23.6%)	0.554 ^c	0.457			
Stent implantation, n(%)	27 (14.1%)	12 (14.8%)	14 (12.7%)	0.173 ^c	0.678			
Recanalization (mTICI 2b/3), n(%)	167 (87.4%)	79 (97.5%)	88 (80.0%)	13.049 ^c	0.000**			
Intraoperative embolization, n(%)	24 (12.6%)	7 (8.6%)	17 (15.5%)	1.971 ^c	0.160			
TOAST classification, n(%)				3.135 ^c	0.209			
Cardioembolism	70 (36.6%)	32 (39.5%)	38 (34.5%)					
Large-artery atherosclerosis	108 (56.5%)	41 (50.6%)	67 (60.9%)					
Other etiologies	13 (6.8%)	8 (9.9%)	5 (4.5%)					
Location of occlusion, n(%)				5.186 ^c	0.075*			
Anterior circulation	162 (84.8%)	74 (91.4%)	88 (80.0%)					
Posterior circulation	27 (14.1%)	7 (8.6%)	20 (18.2%)					
Both anterior and posterior	2 (1%)	0 (0%)	2 (1.8%)					
Age, mean ± SD	68.40 ± 13.68	65.70±13.21	70.38±13.74	-2.363 ^a	0.019**			
Baseline NIHSS, median(IQR)	16 (13, 21)	15 (12,17)	18.5 (14,23)	2715.00 ^b	0.000**			
Pre-stroke mRS, n(%)				9.4996 ^c	0.087*			
0	147 (77.0%)	70(86.4%)	77(70.0%)					
1	26 (13.6%)	9(11.1%)	17(15.5%)					
2	18 (9.4%)	2(2.5%)	16(14.5%)					
ODT (min), mean±SD	260.47 ± 225.07	235.23 ± 203.35	279.08±239.00	-1.335 ^a	0.184			
DPT (min), mean ± SD	91.04 ± 57.93	88.54 ± 63.60	92.88 ± 53.59	-0.504 ^a	0.615			
PRT (min), median(IQR)	67(36,109)	50(35,86)	78(39.5,130.75)	3353.00 ^b	0.002**			
ASPECTS, mean ± SD	8.52±1.67	8.74±1.53	8.35 ± 1.75	1.586 ^a	0.114			
Times of thrombectomy, median(IQR)	2 (1,3)	1 (1,2)	2 (1,4)	3046.00 ^b	0.000**			
NIHSS after EVT, median(IQR)	12(7,20)	7 (3,11)	17.5(12,22)	1310.50 ^b	0.000**			
NIHSS after EVT 24 h, median(IQR)	10 (4,17.5)	4 (2,6)	15(10.25,21.75)	637.50 ^b	0.000**			
Hemorrhage transformation	30(15.7%)	4(4.9%)	26(23.6%)	12.319 ^c	0.000**			
Brain herniation	19(9.9%)	0(0%)	19(17.3%)	15.536 ^c	0.000**			
Decompressive hemicraniectomy	10(5.2%)	0(0%)	10(9.1%)	7.770 ^c	0.005**			

IQR Interquartile range, SD Standard deviation, mTICI Modified thrombolysis in cerebral infarction, NIHSS The National Institutes of Health Stroke Scale, ODT Onset to door time, DPT Door to puncture time, PRT Puncture to recanalization time

* for *P*<0.1, ** for *P*<0.05

 $^a\,$ for t-value, $^b\,$ for Z-value, $^c\,$ for $\chi^2\,$ value

 Table 5
 Multivariate analysis of 90-days outcomes

Characteristics	Coefficient	Р	OR	95%Cl
Cardiopulmonary comor- bidities	-0.7860	0.048*	0.456	(0.209, 0.992)
Diabetes mellitus	-0.7796	0.039*	0.459	(0.219, 0.960)
Coronary heart disease	-0.4699	0.284	0.625	(0.265, 1.477)
Hyperlipidemia	-0.4017	0.310	0.670	(0.308, 1.453)
Ischemic stroke	-0.2793	0.491	0.756	(0.342, 1.674)
Baseline NIHSS	-0.1331	0.000*	0.880	(0.820, 0.935)
Age	-0.0075	0.567	0.993	(0.967, 1.018)
Intravenous thrombolysis	0.2958	0.394	1.344	(0.680, 2.655)

* for *P* < 0.05

comorbidity is associated with higher mortality [11], we demonstrate that heart and respiratory comorbidities were associated with poor neurological recovery and could cause more death related to multisystem failure.

In terms of the relationship between cardiopulmonary comorbidity and outcomes at day 90, our study find that cardiopulmonary comorbidities can also be a predictor of poor outcomes. Newly developed clinical decision models no longer based on single patient characteristics [36]. Instead, multivariable regression models tend to join multiple baseline clinical and radiological characteristics, and conclude that large variations in treatment can benefit patients. However, there is still no consensus in



Fig. 2 Forrest plot of multivariate analysis of 90-days outcomes. The red lines show the factor is significant in the logistic regression. The black lines show that there is no significance. * for P < 0.05, no_sig = no significant



Fig. 3 The receiver operating characteristic (ROC) curve of the logistic regression model for predicting outcomes. The x-axis meant the false-positive rate of the risk prediction

prognosis model on predictor factors selection, for it is uncertain about treatment benefit in specific subgroups. In our multivariable regression analysis, history of diabetes mellitus, higher baseline NIHSS and existence of cardiopulmonary comorbidity are associated with a poor prognosis. Thus, cardiopulmonary comorbidity is an independent risk factors to predict outcomes before EVT and assist clinical decision making.

Compared to the previous studies [31], our result firstly reports that cardiopulmonary comorbidity could lead to a worse ending. We have focused on cardiac and pulmonary dysfunction, which are not traditional factors of the vascular diseases, and those patients are generally underrepresented in randomized trial [7, 37]. Previously, the effect of cardiac and pulmonary dysfunction on cerebral arteries were studied separately, but few noticed the combined effect of them on patients with ELVOS and undergoing emergency thrombectomy. Patients with chronic heart failure and respiratory failure commonly have the diagnostic symptom of exertional dyspneoea varies in intensity [10, 38] and lead to more intensive condition after stroke. Early identification and timely treatment can reduce the mortality in neurological intensive care units to a certain extent.

Our study had limitations. First, there were the general limitations of the single-center and small-sample retrospective study design, which selection and information bias cannot be ruled out. Second, the revascularization techniques were not uniform and scattered. Third, process that comorbidity affected AIS prognosis were complicated, especially when the subjects were elderly people and our baseline characteristics were unbalanced in our study. Therefore, multi-center randomized controlled studies are needed for further validation.

Conclusion

Based on our results, we conclude that cardiopulmonary comorbidities are important risk factors that have been overlooked for a long time. Endovascular thrombectomy was safe in ALVOS patients with chronic cardiopulmonary diseases. However, the unfavorable outcomes were achieved in such patients. The effect of cardiopulmonary morbidity on ALVOS is mixed. Moreover, cardiopulmonary comorbidity was a powerful predictor of worse clinical outcomes and should be considered when neurologists make individualising selection of patients for EVT.

Supplementary Information

The online version contains supplementary material available at https://doi. org/10.1186/s12883-024-03660-w.

Supplementary Material 1: S1 Text. STROBE checklist. STROBE, Strengthening the Reporting of Observational Studies in Epidemiology.

Authors' contributions

J.D. was responsible for the conceptualization of the study. J.D., J.R.W. and Y.Q.C. were responsible for the methodology. J.D., Y.Q.C., X.K.K, B.D., T.L. and X.Y.Z. were involved in data curation. J.R.W. and X.Y.Z. were responsible for statistical analysis. J.R.W., T.L., D.X.L. and L.L. were responsible for investigation. Y.Q.C., L. L., X.K.K and J.D. validated the results. J.R.W. prepared the original draft. J.D. reviewed and edited the manuscript. J.D. and X.Y.Z. were responsible for funding acquisition. All authors reviewed the manuscript.

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Availability of data and materials

Restrictions apply to the availability of these data and they are not publicly available. However, data are available from the corresponding author upon reasonable request and with the permission of the institution.

Declarations

Ethics approval and consent to participate

The research protocol was reviewed and approved by the Ethics Committee of Strategic Support Force Medical Center with K2023-021-01. Written informed consent was obtained from all patients or their legal representatives. Written informed consent was obtained from all patients or their legal representatives.

Consent for publication

Not applicable.

Competing interests

The authors declare no competing interests.

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References

- Krishnamurthi RV, Ikeda T, Feigin VL. Global, regional and country-specific burden of ischaemic stroke, intracerebral haemorrhage and subarachnoid haemorrhage: a systematic analysis of the global burden of disease study 2017. Neuroepidemiology. 2020;54(2):171–9. https://doi.org/10. 1159/000506396.
- Bendszus M, Fiehler J, Subtil F, et al. Endovascular thrombectomy for acute ischaemic stroke with established large infarct: multicentre, openlabel, randomised trial. Lancet. Published online October 10, 2023:S0140-6736(23)02032-9. https://doi.org/10.1016/S0140-6736(23)02032-9.
- Alexandre AM, Valente I, Consoli A, et al. Posterior circulation endovascular thrombectomy for large vessels occlusion in patients presenting with NIHSS score ≤ 10. Life (Basel). 2021;11(12):1423. https://doi.org/10.3390/ life11121423.
- Alexandre AM, Valente I, Pedicelli A, et al. Mechanical thrombectomy in acute ischemic stroke due to large vessel occlusion in the anterior circulation and low baseline National Institute of Health Stroke Scale score: a multicenter retrospective matched analysis. Neurol Sci. 2022;43(5):3105– 12. https://doi.org/10.1007/s10072-021-05771-5.
- Goyal M, Hill MD, Saver JL, Fisher M. Challenges and opportunities of endovascular stroke therapy. Ann Neurol. 2016;79(1):11–7. https://doi. org/10.1002/ana.24528.
- Castro HHG, Alencar AP, Benseñor IM, Lotufo PA, Goulart AC. Multimorbidities are associated to lower survival in ischaemic stroke: results from a

Brazilian stroke cohort (EMMA Study). Cerebrovasc Dis. 2017;44(3–4):232–9. https://doi.org/10.1159/000479827.

- Downer MB, Li L, Carter S, Beebe S, Rothwell PM. Associations of multimorbidity with stroke severity, subtype, premorbid disability, and early mortality: Oxford vascular study. Neurology. 2023;101(6):e645-652. https://doi.org/10.1212/WNL.000000000207479.
- Pilato F, Valente I, Calandrelli R, et al. Clot evaluation and distal embolization risk during mechanical thrombectomy in anterior circulation stroke. J Neurol Sci. 2022;432: 120087. https://doi.org/10.1016/j.jns.2021.120087.
- Pilato F, Valente I, Alexandre AM, et al. Correlation between thrombus perviousness and distal embolization during mechanical thrombectomy in acute stroke. Diagnostics (Basel). 2023;13(3):431. https://doi.org/10. 3390/diagnostics13030431.
- Palmer T, Obst SJ, Aitken CR, et al. Fixed-intensity exercise tests to measure exertional dyspnoea in chronic heart and lung populations: a systematic review. Eur Respir Rev. 2023;32(169):230016. https://doi.org/ 10.1183/16000617.0016-2023.
- Beuker C, Köppe J, Feld J, et al. Association of age with 1-year outcome in patients with acute ischaemic stroke treated with thrombectomy: real-world analysis in 18 506 patients. J Neurol Neurosurg Psychiatry. Published online March 31, 2023;jnnp-2022-330506. https://doi.org/10. 1136/jnnp-2022-330506.
- Escobar C, Palacios B, Gonzalez V, et al. Burden of illness beyond mortality and heart failure hospitalizations in patients newly diagnosed with heart failure in Spain according to ejection fraction. J Clin Med. 2023;12(6): 2410. https://doi.org/10.3390/jcm12062410.
- Chinese Society of Neurology, Chinese Stroke Society. Chinese guidelines for diagnosis and treatment of acute ischemic stroke 2018. Chin J Neurol. 2018;51(9):666–82. https://doi.org/10.3760/cma.j.issn.1006-7876.2018.09. 004.
- Kernan WN, Ovbiagele B, Black HR, et al. Guidelines for the prevention of stroke in patients with stroke and transient ischemic attack: a guideline for healthcare professionals from the American Heart Association/American Stroke Association. Stroke. 2014;45(7):2160–236. https://doi.org/10. 1161/STR.00000000000024.
- Du J, Lin T, Kong X, et al. Characteristics and outcomes of endovascular treatment in the elderly with acute large vessel occlusion stroke. Chin J Cerebrovasc Dis. 2022;19(02):101–8.
- Lyden P, Raman R, Liu L, et al. NIHSS training and certification using a new digital video disk is reliable. Stroke. 2005;36(11):2446–9. https://doi.org/ 10.1161/01.STR.0000185725.42768.92.
- Adams HPJ, Bendixen BH, Kappelle LJ, et al. Classification of subtype of acute ischemic stroke. Definitions for use in a multicenter clinical trial. TOAST. Trial of Org 10172 in acute stroke treatment. Stroke. 1993;24(1):35– 41. https://doi.org/10.1161/01.str.24.1.35.
- Barber PA, Demchuk AM, Zhang J, Buchan AM. Validity and reliability of a quantitative computed tomography score in predicting outcome of hyperacute stroke before thrombolytic therapy. Lancet. 2000;355(9216):1670–4. https://doi.org/10.1016/S0140-6736(00)02237-6.
- Puetz V, Khomenko A, Hill MD, et al. Extent of hypoattenuation on CT angiography source images in basilar artery occlusion: prognostic value in the basilar artery international cooperation Study. Stroke. 2011;42(12):3454–9. https://doi.org/10.1161/STROKEAHA.111.622175.
- Yoo AJ, Simonsen CZ, Prabhakaran S, et al. Refining angiographic biomarkers of revascularization: improving outcome prediction after intra-arterial therapy. Stroke. 2013;44(9). https://doi.org/10.1161/STROK EAHA.113.001990.
- Xiao T, Wijnant SRA, van der Velpen I, et al. Lung function impairment in relation to cognition and vascular brain lesions: the Rotterdam study. J Neurol. 2022;269(8):4141–53. https://doi.org/10.1007/ s00415-022-11027-9.
- Zhang L, Liu W, Sun W, et al. Heart failure and ischemic stroke: a bidirectional and multivariable mendelian randomization study. Front Genet. 2021;12: 771044. https://doi.org/10.3389/fgene.2021.771044.
- Haeusler KG, Laufs U, Endres M. Chronic heart failure and ischemic stroke. Stroke. 2011;42(10):2977–82. https://doi.org/10.1161/STROKEAHA.111. 628479.
- Simmons CA, Poupore N, Nathaniel TI. Age stratification and stroke severity in the telestroke network. J Clin Med. 2023;12(4): 1519. https://doi.org/ 10.3390/jcm12041519.

- Ogoh S, Sugawara J, Shibata S. Does cardiac function affect cerebral blood flow regulation? J Clin Med. 2022;11(20): 6043. https://doi.org/10. 3390/jcm11206043.
- Ohyama Y, Ambale-Venkatesh B, Noda C, et al. Association of aortic stiffness with left ventricular remodeling and reduced left ventricular function measured by magnetic resonance imaging. Circ Cardiovasc Imaging. 2016;9(7):e004426. https://doi.org/10.1161/CIRCIMAGING.115.004426.
- Kim Y, Lee H, Son TO, et al. Reduced forced vital capacity is associated with cerebral small vessel disease burden in cognitively normal individuals. Neuroimage Clin. 2020;25:102140. https://doi.org/10.1016/j.nicl.2019. 102140.
- Weatherald J, Sattler C, Garcia G, Laveneziana P. Ventilatory response to exercise in cardiopulmonary disease: the role of chemosensitivity and dead space. Eur Respir J. 2018;51(2):1700860. https://doi.org/10.1183/ 13993003.00860-2017.
- Rg N, Ap J, Dc H, et al. Thrombectomy 6 to 24 hours after stroke with a mismatch between deficit and infarct. N Engl J Med. 2018;378(1). https:// doi.org/10.1056/NEJMoa1706442.
- Uchida K, Shindo S, Yoshimura S, et al. Association between alberta stroke program early computed tomography score and efficacy and safety outcomes with endovascular therapy in patients with stroke from largevessel occlusion: a secondary analysis of the recovery by endovascular salvage for cerebral ultra-acute embolism-Japan large ischemic core trial (RESCUE-Japan LIMIT). JAMA Neurol. 2022;79(12):1260–6. https://doi.org/ 10.1001/jamaneurol.2022.3285.
- Pilato F, Silva S, Valente I, et al. Predicting factors of functional outcome in patients with acute ischemic stroke admitted to neuro-intensive care unit-a prospective cohort study. Brain Sci. 2020;10(12): 911. https://doi. org/10.3390/brainsci10120911.
- Kang DH, Yoon W, Kim SK, et al. Endovascular treatment for emergent large vessel occlusion due to severe intracranial atherosclerotic stenosis. J Neurosurg. Published online June 1, 2018;1:1–8. https://doi.org/10.3171/ 2018.1JNS172350.
- Li W, Sui X, Li C, et al. Emergency angioplasty or stenting for stroke patients with intracranial atherosclerotic large vessel occlusion. J Atheroscler Thromb. 2023;30(2):160–9. https://doi.org/10.5551/jat.63381.
- Wang Y, Zhao X, Liu L, et al. Prevalence and outcomes of symptomatic intracranial large artery stenoses and occlusions in China: the Chinese intracranial atherosclerosis (CICAS) study. Stroke. 2014;45(3):663–9. https://doi.org/10.1161/STROKEAHA.113.003508.
- 35. Chalos V, van der Ende NAM, Lingsma HF, et al. National Institutes of Health Stroke Scale: an alternative primary outcome measure for trials of acute treatment for ischemic stroke. Stroke. 2020;51(1):282–90. https:// doi.org/10.1161/STROKEAHA.119.026791.
- Venema E, Mulder MJHL, Roozenbeek B, et al. Selection of patients for intra-arterial treatment for acute ischaemic stroke: development and validation of a clinical decision tool in two randomised trials. BMJ. 2017;357: j1710. https://doi.org/10.1136/bmj.j1710.
- Hanlon P, Hannigan L, Rodriguez-Perez J, et al. Representation of people with comorbidity and multimorbidity in clinical trials of novel drug therapies: an individual-level participant data analysis. BMC Med. 2019;17(1):201. https://doi.org/10.1186/s12916-019-1427-1.
- Poulos LM, Ampon RD, Currow DC, Marks GB, Toelle BG, Reddel HK. Prevalence and burden of breathlessness in Australian adults: the National Breathlessness Survey-a cross-sectional web-based population survey. Respirology. 2021;26(8):768–75. https://doi.org/10.1111/resp.14070.

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