



# Predictors of Functional Outcome and Mortality in Endovascular Treatment for Acute Basilar Artery Occlusion: A Single-Centre Experience

#### **OPEN ACCESS**

# Edited by:

Peter Sporns, University Hospital of Basel, Switzerland

#### Reviewed by:

Yingkun He, Henan Provincial People's Hospital, China Craig S. Anderson, University of New South Wales, Australia

#### \*Correspondence:

Ya Peng 289092792@qq.com Xucheng Zhu 853886168@qq.com

<sup>†</sup>These authors have contributed equally to this work and share first authorship

#### Specialty section:

This article was submitted to Stroke, a section of the journal Frontiers in Neurology

Received: 26 June 2021 Accepted: 16 September 2021 Published: 13 October 2021

#### Citation:

Cao J, Mo Y, Chen R, Shao H, Xuan J, Peng Y and Zhu X (2021) Predictors of Functional Outcome and Mortality in Endovascular Treatment for Acute Basilar Artery Occlusion: A Single-Centre Experience. Front. Neurol. 12:731300. doi: 10.3389/fneur.2021.731300 Jie Cao<sup>†</sup>, Yi Mo<sup>†</sup>, Ronghua Chen, Huaming Shao, Jinggang Xuan, Ya Peng<sup>\*</sup> and Xucheng Zhu<sup>\*</sup>

Department of Neurosurgery, The First People's Hospital of Changzhou/The Third Affiliated Hospital of Soochow University, Changzhou, China

**Background and Purpose:** The objective of this study was to identify prognostic factors of endovascular treatment in patients with acute basilar artery occlusion and add evidence about the efficacy and safety of endovascular treatment for acute basilar artery occlusion.

**Materials and Methods:** We reviewed the data of 101 patients with acute basilar artery occlusion receiving endovascular treatment from January 2013 to September 2019. Baseline characteristics and outcomes were evaluated. A favourable functional outcome was defined as a mRS of 0 to 2 assessed at the 3 month follow-up. The association of clinical and procedural characteristics with the functional outcome and mortality was assessed.

**Results:** The study population consisted of 101 patients: 83 males and 18 females. Successful recanalization was achieved in 99 patients (97.1%). A favourable clinical outcome was observed in 50 patients (49.5%), and the overall mortality rate was 26.7%. A favourable outcome was significantly associated with NIHSS score at admission and lung infection. Mortality was associated with NIHSS score at admission, the number of thrombectomy device passes, the postoperative pons-midbrain index, and diabetes mellitus.

**Conclusions:** This study suggested that NIHSS score at admission, the number of thrombectomy device passes, the postoperative pons-midbrain index, diabetes mellitus, and lung infection can predict the functional outcome and mortality. These initial results add evidence about the efficacy and safety of endovascular treatment for acute basilar artery occlusion and need to be confirmed by further prospective studies.

Keywords: acute basilar artery occlusion, endovascular treatment, functional outcome, mortality, thrombectomy

# INTRODUCTION

Acute basilar artery occlusion (ABAO) is an uncommon but potentially devastating neurological condition, accounting for ~20% of posterior circulation strokes (1) and approximately 1% of all ischaemic strokes (2), and is associated with a very poor outcome (3). With variable presentations and a broad differential diagnosis (4), BAO is associated with high rates of critical disability and mortality due to injury to the brainstem. Without active intervention or treatment, the patient mortality rate in BAO can exceed 90% (5, 6).

Compared with either intravenous or intraarterial thrombolysis, systematic meta-analyses of case series and registry data have indicated that mechanical thrombectomy provides the optimal potential for improved recanalization rates and more favourable clinical outcomes in patients with BAO, but well-conducted randomised controlled trials are needed (7–13). To date, the only large randomised controlled trial (the BASICS Study) has not shown obvious advantages of endovascular therapy over medical therapy, but too long time span, recruitment without achieving anticipated and partial presence of cross-group cases weaken the objectivity of the results (14). Further evidence in the use of endovascular treatment (EVT) in posterior-circulation strokes is required.

This study is based on data from a single-centre registry with 101 patients who received EVT. endovascular treatment in patients with acute basilar artery occlusion and. The purpose of the present study was to analyse prognostic factors of clinical outcomes of endovascular treatment for acute basilar artery occlusion and add evidence about the efficacy and safety of endovascular treatment for acute basilar artery occlusion.

# **METHODS**

### **Selection of Patients**

This was a retrospective study from one stroke centre in Changzhou, China. The time interval of this study ranged from January 2013 to December 2019. The clinical data of patients with ABAO confirmed by digital subtraction angiography treated with mechanical thrombectomy within 24 h from symptom onset were collected and statistically analysed.

Patients were enrolled if they met the following criteria: (a) age  $\geq 18$  years old; (b) diagnosis of BAO by CTA or MRA before EVT; (c) EVT performed within 24 h after symptom onset; and (d) mRS  $\leq 2$  before stroke. The exclusion criteria were as follows: (a) history of surgery or trauma within 2 months; (b) intracranial haemorrhage (ICH) or history of subarachnoid haemorrhage, tumour, ICH, or arteriovenous malformation; (c) large infarct core (exceeding two-thirds of the midbrain, pons, or either side of the cerebellum); (d) dysfunction of important organs; (e) definite bleeding tendency; and (f) voluntary abandonment of treatment or failure to follow doctor's instructions due to non-iatrogenic reasons.

All patients (or relatives) signed informed consent forms before treatment. This study was approved by the Ethics Committee of the Hospital.

# **Intervention Procedure**

During mechanical thrombectomy, patients were under local or general anaesthesia depending on the clinical circumstances. Each patient treated under general anaesthesia underwent tracheal intubation before anaesthesia. Oxygenation and strict blood pressure control were maintained during the procedure in all cases. All interventions were performed by senior neurointerventionalists under general or local anaesthesia. Successful recanalization after mechanical thrombectomy was defined by TICI 2b-3 (15). A 24-h follow-up non-enhanced CT was always performed to rule out complications of ICH.

In this study, the whole treatment process of mechanical thrombectomy was almost consistent with that in our previous study (16). Together with the extensive surgical experience and the progress of neurointerventional materials, the stent-retriever thrombectomy technique combined with an intermediate catheter or a direct aspiration first-pass technique (ADAPT) (17) was applied in our centre for appropriate patients.

Once atherosclerosis or dissection was considered the stroke aetiology, 10  $\mu$ g/kg tirofiban was immediately administered by an intravenous bolus and continued at 0.15  $\mu$ g/kg/min. Balloon angioplasty and/or stent placement was considered if there was obvious stenosis, while stent placement was used for the dissected artery.

One hundred milligrammes of aspirin and 75 mg of clopidogrel were administered at 24 h postprocedure. Tirofiban was stopped after overlapping medicine treatment for at least 6 h. Aspirin and clopidogrel were administered daily for 3 months.

### **Neuroimaging Assessment**

The time window was 24 h. Collected data included clinical data, such as age, sex, and vertebral artery lesions in the posterior circulation (occlusion or stenosis), cardiovascular risk factors, such as hypertension, diabetes mellitus, coronary artery disease, atrial fibrillation, history of ischaemic stroke or TIA, and smoking, stroke aetiology according to the Trial of Org 10172 in Acute Stroke Treatment (TOAST) criteria (18), and baseline NIHSS and Glasgow coma scale (GCS) scores at admission.

The baseline posterior circulation Acute Stroke Prognosis Early CT Score (Pc-ASPECTS) (19) and pons-midbrain index (PMI) (20) both at admission and within 24 h after surgery were evaluated based on preoperative and postoperative neuroimaging and CTA source images. The location of BA occlusion was also classified as proximal (from the vertebrobasilar junction to the origin of the anterior inferior cerebellar artery), middle (from the origin of the anterior inferior cerebellar artery), middle (from the superior cerebellar artery), or distal (distal to the origin of the superior cerebellar artery) according to previously published criteria (21). The OTT was defined as the time between symptom onset and the beginning of femoral artery puncture. Additionally, the puncture-to-recanalization time (PTR) was collected for reference.

The presence of hydrocephalus or evident ischaemic changes in the posterior circulation territory or ICH was reassessed via craniocerebral CT within 36 h after EVT. Symptomatic ICH was defined according to the definition of the safe thrombolysis in the stroke surveillance study (SITS-MOST): local or remote intraparenchymal haemorrhage type 2 in the 22- to 36-h follow-up imaging scans, combined with a neurological deterioration of  $\geq$ 4 points on the NIHSS from baseline, from the lowest NIHSS score between baseline and 24 h, or leading to death (22). Lung infection during the perioperative period and complications related to surgical instruments and operations (new vascular occlusion, reocclusion after successful opening, ICH, arterial dissection, vasospasm, haematoma or haemorrhage at the femoral artery puncture point, hypotension requiring medication, bradycardia requiring medication, etc.), considered a potential prognostic factors, were also documented.

#### **Clinical Outcome Measures**

The severity of stroke at the time of EVT was dichotomized as severe or mild to moderate, in conformity to the definition used in the Basilar Artery International Cooperation Study (BASICS) registry (6). Patients in a coma, with tetraplegia, or in a locked-in state were classified as having a severe stroke, whereas mild-tomoderate stroke was defined as any deficit that was less than severe. The functional outcome at 3 months was assessed with the mRS score obtained during a follow-up outpatient visit or via a semi-standardised telephone interview. A favourable outcome was defined as an mRS score of 0–2, in accordance with the definition used in the BASICS registry. Moreover, the patients were divided into two groups according to mortality to investigate the difference in the mortality rate between the two groups by risk factor stratification.

### **Statistical Analysis**

All statistical analyses were performed using the software package SPSS 22.0. The chi-square test or Fisher's exact test was used to compare categorical variables, and Student's *t*-test or the Mann–Whitney U test was used to compare continuous variables. A logistic regression model was used for multivariate analysis to determine whether the potential risk factors on univariate analysis remained independently associated with a favourable functional outcome and mortality at 90 days. All tests were 2-sided, and *p*-values of 0.05 or less were considered statistically significant.

# RESULTS

# **Baseline Characteristics**

A total of 101 patients who underwent EVT for ABAO between January 2013 and September 2019 were included in this analysis, including 20 patients in the mild-to-moderate group and 81 patients in the severe group. Baseline clinical and neuroimaging characteristics are presented in **Table 1**. There were 83 males and 18 females, with a mean age of  $62.2 \pm 12.91$  (mean  $\pm$  SD) years. Eight patients (7.8%) underwent IVT prior to EVT in this series. The median baseline NIHSS score was 30 (interquartile

**TABLE 1** | Baseline characteristics and functional outcome (mRS  $\leq$ 2 vs. mRS >2).

Characteristics	Overall ( <i>n</i> = 101)	Outcome			
		mRS ≤2 ( <i>n</i> = 50)	mRS >2 ( <i>n</i> = 51)	P-value	
Age (years), mean (SD)	62.2 (12.91)	62.5 (12.85)	62 (13.09)	0.853	
Male sex, No. (%)	83 (81.4)	42 (84)	41 (80.4)	0.636	
Hypertension, No. (%)	81 (79.4)	38 (76)	43 (84.3)	0.295	
Diabetes mellitus, No. (%)	20 (19.6)	7 (14)	13 (25.5)	0.147	
Baseline glycaemia (mmol/L), mean (SD)	8.3 (2.98)	7.84 (2.83)	8.76 (3.08)	0.065	
Atrial fibrillation, No. (%)	30 (29.4)	16 (32)	14 (27.5)	0.617	
Coronary heart disease, No. (%)	8 (7.8)	4 (8)	4 (7.8)	0.999	
Antithrombotic treatment, No. (%)	15 (14.7)	6 (12)	9 (17.6)	0.425	
History of ischaemic stroke or TIA, No. (%)	20 (19.6)	9 (18)	11 (21.6)	0.653	
Smoking, No. (%)	38 (37.3)	17 (34)	21 (41.2)	0.457	
GCS score at admission, median (IQR)	5 (4–7)	6 (4–9)	5 (4–6)	0.008	
NIHSS score at admission, median (IQR)	30 (22.5–36.5)	26 (20–32)	35 (28–38)	<0.001	
Preoperative Pc-ASPECTS, median (IQR)	9 (9–10)	9 (9–10)	9 (9–10)	0.234	
Preoperative PMI, median (IQR)	0 (0–0)	0 (0–0)	0 (0–0)	0.615	
IVT, No. (%)	8 (7.8)	4 (8)	4 (7.8)	0.999	
Stroke aetiology, No. (%)				0.435	
Atherosclerosis	49 (48.0)	21 (42)	28 (54.9)		
Cardioembolic	30 (29.4)	15 (30)	15 (29.4)		
Other	7 (29.4)	4 (8)	3 (5.9)		
Unknown	15 (14.7)	10 (20)	5 (9.8)		

SD, standard deviation; GCS, Glasgow Coma Scale; IQR, interquartile range; Pc-ASPECTS, posterior circulation Acute Stroke Prognosis Early CT Score; PMI, pons-midbrain index; IVT, intravenous thrombolysis.

#### TABLE 2 | Baseline characteristics and mortality.

Characteristics	Mortality				
	Yes ( <i>n</i> = 27)	No ( <i>n</i> = 74)	P-value		
Age (years), mean (SD)	63.9 (12.78)	61.6 (12.99)	0.451		
Male sex, No. (%)	20 (74.1)	63 (85.1)	0.321		
Hypertension, No. (%)	22 (81.5)	59 (79.7)	0.845		
Diabetes mellitus, No. (%)	9 (33.3)	11 (14.9)	0.039		
Baseline glycaemia (mmol/L), mean (SD)	9.2 (3.40)	7.97 (2.76)	0.049		
Atrial fibrillation, No. (%)	10 (37)	20 (27)	0.33		
Coronary heart disease, No. (%)	2 (7.4)	6 (8.1)	0.999		
Antithrombotic treatment, No. (%)	4 (14.8)	11 (14.9)	0.999		
History of ischaemic stroke or TIA, No. (%)	5 (18.5)	15 (20.3)	0.845		
Smoking, No. (%)	13 (48.1)	25 (33.8)	0.187		
GCS score at admission, median (IQR)	4 (3–6)	5.5 (4–8)	<0.001		
NIHSS score at admission, median (IQR)	37 (31–39)	28 (21–35)	<0.001		
Preoperative Pc-ASPECTS, median (IQR)	10 (9–10)	9 (9–9)	0.333		
Preoperative PMI, median (IQR)	0 (0–0)	0 (0–0)	0.209		
IVT, No. (%)	3 (11.1)	5 (6.8)	0.764		
Stroke aetiology, No. (%)			0.713		
Atherosclerosis	11 (40.7)	38 (51.4)			
Cardioembolic	10 (37)	20 (27)			
Other	2 (7.4)	5 (6.8)			
Unknown	4 (14.8)	11 (14.9)			

SD, standard deviation; GCS, Glasgow Coma Scale; IQR, interquartile range; Pc-ASPECTS, posterior circulation Acute Stroke Prognosis Early CT Score; PMI, pons-midbrain index; IVT, intravenous thrombolysis.

range [IQR], 22.5–36.5), and the median total GCS score was 5 (IQR, 4–7). Among the stroke risk factors, hypertension was most prevalent (81 of 101 patients [79.4%]). In this research, atherosclerosis, accounting for 48%, remained the most common cause of ABAO.

As shown in **Tables 1**, **2**, a favourable outcome was significantly associated with both the total GCS score at admission (p = 0.008) and the NIHSS score at admission (p < 0.001), which were also statistically associated with mortality (GCS score, p < 0.001; NIHSS score, p < 0.001), similar to diabetes mellitus (p = 0.039) and baseline glycaemia (mmol/L) (p = 0.049) (**Table 2**).

# Periprocedural Characteristics and Clinical Outcomes

**Tables 3**, **4** show the periprocedural characteristics and outcomes. The median OTT was 240 min (IQR, 180–340), and the median PTR was 65 min (IQR, 50–115). Seventy-six patients (74.5%) were treated under general anaesthesia. Compared with the pour outcome group, the PTR was significantly lower (p = 0.006) in the favourable outcome group, while a longer PTR seemed to be associated with a higher mortality rate (p = 0.002). The distal BA was the most common site of ABAO (49/101, 48%).

During the progression of mechanical thrombectomy, 17 patients (16.7%) were treated with mechanical thrombectomy in combination with IA thrombolysis, and 47 patients (46.1%)

received an intraarterial infusion of tirofiban. Direct balloon angioplasty combined with stenting without mechanical thrombectomy was performed in eight (11.8%) patients. Angioplasty with or without stenting after mechanical thrombectomy was performed in 46 (46.1%) patients: balloon angioplasty alone in 11 patients, stent placement alone in 11 patients, and balloon angioplasty combined with stenting in 24 patients. Among all 101 patients treated with angioplasty, TICI 2b-3 recanalization was achieved in 99 patients (97.1%). No significant effect of EVT on prognosis was found.

With regard to the postoperative neuroimaging evaluation based on the Pc-ASPECTS and PMI, the median Pc-ASPECTS was 6 (IQR, 5-8), and the median PMI was 2 (IQR, 1-3). It seemed that a higher postoperative Pc-ASPECTS was statistically related to a better functional outcome (p = 0.001), and a lower postoperative PMI was associated with both a better functional outcome (p = 0.001) and a lower mortality rate (p = 0.008). Overall, as shown in Table 1, a favourable functional outcome (mRS score 0-2) was reached by 50 (49.5%) patients, including 13 of 20 patients (65%) in the mild-moderate group and 37 of 81 patients (45.7%) in the severe group. Mortality occurred in 25 of 27 (92.6%) patients in the severe group (p = 0.048). Symptomatic ICH occurred in five patients (4.9%), all of whom experienced mortality (p = 0.001). Other complications included complications related to surgical instruments and operations in 11 patients, hydrocephalus in two patients, and perioperative TABLE 3 | Periprocedural characteristics and functional outcome (mRS ≤2 vs. mRS score >2).

Characteristics	Overall	Outcome			
	( <i>n</i> = 101)	mRS ≤ 2 ( <i>n</i> = 50)	mRS > 2 ( <i>n</i> = 51)	P-value	
Symptom-onset-to-treatment time (OTT), median (IQR), min	240 (180–340)	240.5 (158.75–322.5)	240 (180–380)	0.541	
$OTT \leq 360 \text{ min}, \text{ No. } (\%)$	79 (78.2)	44 (88)	35 (68.6)	0.018	
Puncture-to-recanalization time (PTR), median (IQR), min	65 (50–115)	60 (45-83.25)	90 (52–120)	0.006	
General anaesthesia, No. (%)	76 (74.5)	35 (70)	41 (80.4)	0.226	
Number of thrombectomy device passes, median (IQR), min	1 (1-2)	1 (1–2)	2 (10–2)	0.133	
BA occlusion site, No. (%)				0.807	
Proximal BA	30 (29.7)	15 (30)	15 (29.4)		
Mid BA	22 (21.7)	10 (20)	12 (23.5)		
Distal BA	49 (48.6)	26 (52)	23 (45.1)		
Endovascular treatment					
IA thrombolysis, No. (%)	17 (16.7)	9 (18)	8 (15.7)	0.756	
IA infusion of tirofiban, No. (%)	47 (46.1)	23 (46)	24 (47.1)	0.915	
Emergency angioplasty, No. (%)				0.41	
Simple stenting	11 (10.8)	5 (10)	6 (11.8)		
Simple balloon angioplasty	11 (10.8)	8 (16)	3 (5.9)		
Both stenting and balloon angioplasty	24 (23.5)	10 (20)	14 (27.5)		
None	55 (53.9)	27 (54)	28 (54.9)		
Postoperative Pc-ASPECTS, median (IQR)	6 (5–8)	7 (6–8)	6 (5–7)	0.001	
Postoperative PMI, median (IQR)	2 (1–3)	1 (1–2)	3 (1–4)	0.001	
Complications, No. (%)					
Complications related to surgical instruments and operations	11 (10.8)	4 (8)	7 (13.7)	0.356	
Symptomatic ICH	5 (4.9)	0 (0) 5 (9.8)		0.07	
Hydrocephalus	2 (2)	O (O)	2 (3.9)	0.484	
Lung infection	50 (49)	14 (28)	36 (70.6)	<0.001	
TICI 2b-3, No. (%)	99 (98.0)	50 (100)	49 (96.1)	0.157	

IQR, interquartile range; Pc-ASPECTS, posterior circulation Acute Stroke Prognosis Early CT Score; BA, basilar artery; IA, intraarterial; PMI, pons-midbrain index; ICH, intracranial haemorrhage.

lung infection in fifty patients. Lung infection was significantly associated with a poor outcome (p < 0.001).

### **Risk Factors for Clinical Outcomes**

On multivariate logistic analysis, three clinical factors were identified as predicting good clinical outcomes, including the NIHSS score at admission (OR 0.852; 95%CI, 0.748–0.970; p = 0.016) and lung infection (OR 0.135; 95%CI, 0.042–0.433; p = 0.001). Mortality was significantly associated with the NIHSS score at admission (OR 1.186; 95%CI, 1.005–1.399; p = 0.044), the number of thrombectomy device passes (OR 2.612; 95%CI, 1.190–5.731; p = 0.017), the postoperative PMI (OR 3.222; 95%CI, 1.544–6.726; p = 0.002), and diabetes mellitus (OR 25.037; 95% CI, 3.449–181.750; p = 0.001) (**Table 5**).

# DISCUSSION

In this study, we analysed our single-centre outcomes of EVT in patients with ABAO in the past 7 years and found that the rate of a favourable functional outcome, defined as an mRS score of 0-2 at 3 months, was 49.5%, with

an overall mortality rate of 26.7%. We obtained a high recanalization rate (97.1%) in patients with BAO treated with mechanical thrombectomy.

These results are comparable to those of other series published to date of patients with BAO treated with mechanical thrombectomy in terms of the rate of successful recanalization, a favourable outcome and mortality. According to previous studies, the recanalization rate of mechanical thrombectomy, including the last generation of mechanical devices in patients with ABAO, ranged from 75 to 94.8%, with higher rates in some studies performed with accepted EVT strategies with either stent retrievers, aspiration or a combination of both. Regarding the rate of a good functional outcome (mRS 0-2) and mortality, the results ranged from 29.4 to 46.1% and 21 to 40.9%, respectively (8, 10, 12, 13, 23). Based on two separate meta-analyses, the results from our study together with those of previous reports, showing higher recanalization rates and a better prognosis for patients with ABAO managed with endovascular thrombectomy when compared with drug therapy alone (either intravenous and/or intraarterial thrombolysis), suggest that our EVT strategy (mechanical thrombectomy combined with intracranial angioplasty or

#### TABLE 4 | Periprocedural characteristics and mortality.

Characteristics	Overall (n = 101)	Mortality		
		Yes (n = 27)	No (n = 74)	P-value
Severe group, No. (%)	81 (79.4)	25 (92.6)	56 (75.7)	0.048
Symptom-onset-to-treatment time (OTT), median (IQR), min	240 (180–340)	250 (200–386)	240 (153.75–330)	0.182
OTT ≤360 min, No. (%)	79 (78.2)	17 (63)	62 (83.8)	0.025
Puncture-to-recanalization time (PTR), median (IQR), min	65 (50–115)	93 (64–134)	60 (45–90)	0.002
General anaesthesia, No. (%)	76 (74.5)	20 (74.1)	56 (75.7)	0.869
Number of thrombectomy device passes, median (IQR), min	1 (1-2)	2 (1–3)	1 (1–2)	0.085
BA occlusion site, No. (%)				0.186
Proximal BA	30 (29.7)	7 (25.9)	23 (31.1)	
Mid BA	22 (21.8)	9 (33.3)	13 (17.6)	
Distal BA	49 (48)	11 (40.7)	38 (51.4)	
Endovascular treatment				
IA thrombolysis, No. (%)	17 (16.7)	4 (14.8)	13 (17.6)	0.979
IA infusion of tirofiban, No. (%)	47 (46.1)	11 (40.7)	36 (48.6)	0.481
Emergency angioplasty, No. (%)				0.558
Simple stenting	11 (10.8)	3 (11.1)	8 (10.8)	
Simple balloon angioplasty	11 (10.8)	1 (3.7)	10 (13.5)	
Both stenting and balloon angioplasty	24 (23.5)	8 (29.6)	16 (21.6)	
None	55 (53.9)	15 (55.6)	40 (54.1)	
Postoperative Pc-ASPECTS, median (IQR)	6 (5–8)	6 (3–8)	6 (5–8)	0.069
Postoperative PMI, median (IQR)	2 (1–3)	4 (0-6)	2 (1–3)	0.008
Complications, No. (%)				
Complications related to operations	11 (10.8)	5 (18.5) 6 (8.1)		0.26
Symptomatic ICH	5 (4.9)	5 (18.5)	5 (18.5) 0	
Hydrocephalus	2 (2)	1 (3.7)	1 (1.4)	0.465
Lung infection	50 (49)	16 (59.3)	34 (45.9)	0.236
TICI 2b-3, No. (%)	99 (97.1)	25 (92.6)	74 (100)	0.119

IQR, interquartile range; Pc-ASPECTS, posterior circulation Acute Stroke Prognosis Early CT Score; BA, basilar artery; IA, intraarterial; PMI, pons-midbrain index; ICH, intracranial haemorrhage.

TABLE 5 | Multifactor logistic regression for functional outcome and mortality.

Characteristics	Favourable outcome		Mortality	
	OR (95%CI)	P-value	OR (95%CI)	P-value
Diabetes mellitus			25.037 (3.449–181.750)	0.001
GCS score at admission	0.898 (0.620-1.299)	0.567	0.730 (0.431–1.237)	0.242
NIHSS score at admission	0.852 (0.748–0.970)	0.016	1.186 (1.005–1.399)	0.044
OTT ≤360 min	2.509 (0.589–10.689)	0.214	0.243 (0.049-1.195)	0.082
Puncture-to-recanalization time (PTR)	0.985 (0.969–1.001)	0.059	1.016 (0.998–1.035)	0.081
Number of thrombectomy device passes			2.612 (1.190-5.731)	0.017
Postoperative Pc-ASPECTS	1.427 (0.894–2.277)	0.136	1.545 (0.841–2.837)	0.161
Postoperative PMI	0.628 (0.382-1.034)	0.067	3.222 (1.544-6.726)	0.002
Lung infection	0.135 (0.042-0.433)	0.001		

GCS, Glasgow Coma Scale; Pc-ASPECTS, posterior circulation Acute Stroke Prognosis Early CT Score; PMI, pons-midbrain index; IVT, intravenous thrombolysis.

IA thrombolysis or IA infusion of tirofiban) has promise in patients with ABAO (24), despite the lack of good randomised controlled trials.

In fact, whether EVT is the best treatment for ABAO is controversial in some existing retrospective studies. In a multicentre clinical registry study (the EVT for Acute Basilar

6

Artery Occlusion Study, BASILAR), intravascular treatment within 24h of ABAO improved the functional outcome and reduced mortality, supporting the superiority of EVT compared to drug therapy alone (25). However, according to the first randomised controlled trial to assess the effect of contemporary EVT, including stent retriever-based mechanical thrombectomy, in the treatment of acute vertebrobasilar occlusion (Endovascular treatment vs. standard medical treatment for vertebrobasilar artery occlusion (BEST): an open-label, randomised controlled trial), the difference in the occurrence of a favourable outcome between patients receiving EVT and those receiving standard medical therapy alone statistically did not make sense (26). In addition, the BASICS Study also has not shown superiority of EVT over medical therapy. All of these studies have their own limitations. BASILAR study could not balance the systematic differences between the two treatment groups. BEST study had too many cross-group individuals, leading to an insufficient study quality. The limitation of BASICS Study has been mentioned in the previous part. In our study, factors associated with EVT, such as the high recanalization rate, the course of treatment and treatment-related complications, did not affect the prognosis of BAO patients; the number of thrombectomy device passes also did not affect the prognosis and seemed to be more related to the complex vascular structure and the pathophysiology of the embolism (27). A limitation of our work is the small sample size of patients with unsuccessful reperfusion. If we can obtain more samples of unsuccessful recanalization (TICI 1-2a), the results would be more convincing. Based on the fact that a substantial proportion ( $\sim$ 50.5%) of our patients had poor clinical outcomes (mRS score 3-6) despite TICI 2b-3 recanalization, we support that successful recanalization by EVT could be essential but not sufficient to obtain good clinical outcomes in patients with ABAO.

There is conflicting evidence on the importance of timing in treating ABAO. The current American Heart Association guidelines suggest applying mechanical thrombectomy in patients with causative occlusion of the BA by a groin puncture time within 6 h of symptom onset (class IIb; level of evidence C) (28), which is supported by several previous reports (23, 29, 30). A retrospective study of 215 patients from two endovascular centres by Bouslama et al. (31) suggests that the time to recanalization does not predict the outcome, which is in line with some other studies (13, 32). In our study, we found that a time to treatment of more than 6h was predictive of a poor outcome, and a shorter PTR could be associated with a favourable functional outcome and a lower mortality rate, based on a single-factor analysis. However, this correlation disappeared on multivariate analysis. In addition, we found that fewer thrombectomy device passes was associated with a lower mortality rate on multivariate logistic analysis. We tentatively propose that a delay from symptom onset, which is potentially related to a poorer functional outcome, should not discourage the application of reperfusion therapy in BAO patients; in addition, factors affecting the efficiency of mechanical thrombectomy, such as a complex cerebrovascular configuration or combination with other cerebrovascular lesions Several prior studies have noted the importance of clinical severity at admission and found an association between higher admission NIHSS scores and poorer clinical outcomes (23, 33). Our study confirms this finding. In addition, in contrast to a published single-centre registry (8) of 28 patients with BAO undergoing EVT, we did not find a statistically significant association between the clinical outcome and the GCS score at admission.

In the current literature, the Pc-ASPECT for the posterior circulation is the subject of discussion. Some existing studies prefer to use diffusion-weighted imaging (DWI) rather than CT for evaluation of the Pc-ASPECT (34, 35). Unfortunately, we failed to collect DWI data from most patients to test this view. As in our study, in the vast majority of patients who received both a pre-interventional and post-interventional Pc-ASPECT via CT, these scores were not predictive of the functional outcome. Although on the PMI requires CT for evaluation, the post-interventional PMI was highly predictive of mortality, as suggested by the multifactor analysis in our study. This trend is in line with other findings reported in the literature (20, 36).

In our study, diabetes mellitus was a predictor of mortality on multivariate logistic analysis, which is supported by a large single-centre report on 231 patients by Ravindren et al. (23). Most reports on this topic are not in line with this observation (11, 13, 30–33). Furthermore, we found that perioperative lung infection was predictive of an unfavourable functional outcome, despite the lack of supporting findings in the literature. In fact, lung infection is not uncommon, and most ABAO patients undergoing EVT still need prolonged bed rest, which accounts for the high probability of sputum excretion disorder and pulmonary infection. This finding could be helpful to enhance the importance of the prevention and control of perioperative lung infection.

Our study has some limitations that stem primarily from its retrospective design, small sample size, long duration, lack of randomisation and comparison with patients treated with medical management only, and heterogeneity of the centrespecific study population. Additionally, some data, including the ASITN/SIR grade, which could be important to the assessment of prognostic factors, were missing. In the future, more results from large randomised controlled trials may provide more evidence about the safety and efficacy of mechanical thrombectomy in ABAO.

# CONCLUSION

This study suggested that NIHSS score at admission, the number of thrombectomy device passes, the postoperative pons-midbrain index, diabetes mellitus, and lung infection can predict the functional outcome and mortality. These initial results add evidence about the efficacy and safety of endovascular treatment for acute basilar artery occlusion and need to be confirmed by further prospective studies.

# DATA AVAILABILITY STATEMENT

The original contributions presented in the study are included in the article/supplementary material, further inquiries can be directed to the corresponding author/s.

# **AUTHOR CONTRIBUTIONS**

YP and XZ conceived and designed the study, including quality assurance and control. JC and YM collected the data and wrote

# REFERENCES

- Mohlenbruch M, Stampfl S, Behrens L, Herweh C, Rohde S, Bendszus M, et al. Mechanical thrombectomy with stent retrievers in acute basilar artery occlusion. *AJNR Am J Neuroradiol*. (2014) 35:959–64. doi: 10.3174/ajnr.A3796
- Israeli-Korn SD, Schwammenthal Y, Yonash-Kimchi T, Bakon M, Tsabari R, Orion D, et al. Ischemic stroke due to acute basilar artery occlusion: proportion and outcomes. *Isr Med Assoc J.* (2010) 12:671–5.
- 3. Du S, Mao G, Li D, Qiu M, Nie Q, Zhu H, et al. Mechanical thrombectomy with the Solitaire AB stent for treatment of acute basilar artery occlusion: a single-center experience. *J Clin Neurosci.* (2016) 32:67–71. doi: 10.1016/j.jocn.2016.01.037
- Buchman SL, Merkler AE. Basilar artery occlusion: diagnosis and acute treatment. *Curr Treat Options Neurol.* (2019) 21:45. doi: 10.1007/s11940-019-0591-0
- Lindsberg PJ, Mattle HP. Therapy of basilar artery occlusion: a systematic analysis comparing intra-arterial and intravenous thrombolysis. *Stroke*. (2006) 37:922–8. doi: 10.1161/01.STR.0000202582.29510.6b
- Schonewille WJ, Wijman CA, Michel P, Rueckert CM, Weimar C, Mattle HP, et al. Treatment and outcomes of acute basilar artery occlusion in the Basilar Artery International Cooperation Study (BASICS): a prospective registry study. *Lancet Neurol.* (2009) 8:724–30. doi: 10.1016/S1474-4422(09)70173-5
- Gory B, Eldesouky I, Sivan-Hoffmann R, Rabilloud M, Ong E, Riva R, et al. Outcomes of stent retriever thrombectomy in basilar artery occlusion: an observational study and systematic review. *J Neurol Neurosurg Psychiatry*. (2016) 87:520–5. doi: 10.1136/jnnp-2014-310250
- Werner M, Lopez-Rueda A, Zarco F, Roman LS, Blasco J, Amaro S, et al. Mechanical thrombectomy in acute basilar artery occlusion: a safety and efficacy single centre study. *Interv Neuroradiol.* (2016) 22:310– 17. doi: 10.1177/1591019916631145
- Fahed R, Di Maria F, Rosso C, Sourour N, Degos V, Deltour S, et al. A leap forward in the endovascular management of acute basilar artery occlusion since the appearance of stent retrievers: a single-center comparative study. J Neurosurg. (2017) 126:1578–84. doi: 10.3171/2016.2.JNS151983
- Baik SH, Park HJ, Kim JH, Jang CK, Kim BM, Kim DJ. Mechanical thrombectomy in subtypes of basilar artery occlusion: relationship to recanalization rate and clinical outcome. *Radiology*. (2019) 291:730– 7. doi: 10.1148/radiol.2019181924
- Deb-Chatterji M, Flottmann F, Leischner H, Alegiani A, Brekenfeld C, Fiehler J, et al. Recanalization is the key for better outcome of thrombectomy in basilar artery occlusion. *Clin Neuroradiol.* (2019) 30:769– 75. doi: 10.1007/s00062-019-00850-9
- Meinel TR, Kaesmacher J, Chaloulos-Iakovidis P, Panos L, Mordasini P, Mosimann PJ, et al. Mechanical thrombectomy for basilar artery occlusion: efficacy, outcomes, and futile recanalization in comparison with the anterior circulation. J Neurointerv Surg. (2019) 11:1174–80. doi: 10.1136/neurintsurg-2018-014516
- Shore TH, Harrington TJ, Faulder K, Steinfort B. Endovascular therapy in acute basilar artery occlusion: a retrospective single-centre Australian analysis. *J Med Imaging Radiat Oncol.* (2019) 63:33–9. doi: 10.1111/1754-9485.12825
- Langezaal LCM, Van Der Hoeven E, Mont'alverne FJA, De Carvalho JJF, Lima FO, Dippel DWJ, et al. Endovascular therapy for stroke due to basilar-artery occlusion. N Engl J Med. (2021) 384:1910–20. doi: 10.1056/NEJMoa2030297

the paper. RC and JX designed the study's analytic strategy. HS reviewed and edited the manuscript. All authors read and approved the manuscript.

### FUNDING

This work was partly supported by Changzhou Science and Technology Support Program (No. CE20205025) and the Basic Research Project of the Changzhou Science and Technology Bureau (No. CJ20200111).

- Higashida RT, Furlan AJ, Roberts H, Tomsick T, Connors B, Barr J, et al. Trial design and reporting standards for intra-arterial cerebral thrombolysis for acute ischemic stroke. *Stroke.* (2003) 34:e109-37. doi: 10.1161/01.STR.0000082721.62796.09
- Cao J, Xuan J, Chen R, Zhu X, Shao H, Chao X, et al. Intra-arterial treatment for patients with severe acute vertebrobasilar occlusion: a singlecenter retrospective study. J Stroke Cerebrovasc Dis. (2017) 26:2376– 82. doi: 10.1016/j.jstrokecerebrovasdis.2017.05.028
- Boisseau W, Escalard S, Fahed R, Lapergue B, Smajda S, Maier B, et al. Direct aspiration stroke thrombectomy: a comprehensive review. *J Neurointerv Surg.* (2020) 12:1099–106. doi: 10.1136/neurintsurg-2019-015508
- Adams HP Jr., Bendixen BH, Kappelle LJ, Biller J, Love BB, 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) 24:35–41. doi: 10.1161/01.STR.24.1.35
- Puetz V, Sylaja PN, Coutts SB, Hill MD, Działowski I, Mueller P, et al. Extent of hypoattenuation on CT angiography source images predicts functional outcome in patients with basilar artery occlusion. *Stroke.* (2008) 39:2485– 90. doi: 10.1161/STROKEAHA.107.511162
- Garg R, Biller J. Neuroimaging predictors of clinical outcome in acute basilar artery occlusion. *Front Neurol.* (2017) 8:293. doi: 10.3389/fneur.2017.00293
- Archer CR, Horenstein S. Basilar artery occlusion: clinical and radiological correlation. *Stroke*. (1977) 8:383–90. doi: 10.1161/01.STR.8.3.383
- Wahlgren N, Ahmed N, Davalos A, Ford GA, Grond M, Hacke W, et al. Thrombolysis with alteplase for acute ischaemic stroke in the Safe Implementation of Thrombolysis in Stroke-Monitoring Study (SITS-MOST): an observational study. *Lancet.* (2007) 369:275–82. doi: 10.1016/S0140-6736(07)60149-4
- Ravindren J, Aguilar Perez M, Hellstern V, Bhogal P, Bazner H, Henkes H. Predictors of outcome after endovascular thrombectomy in acute basilar artery occlusion and the 6hr time window to recanalization. *Front Neurol.* (2019) 10:923. doi: 10.3389/fneur.2019.00923
- 24. Kumar G, Shahripour RB, Alexandrov AV. 荟萃分析:急性基底动脉 闭塞再通改善预后 Recanalization of acute basilar artery occlusion improves outcomes: a meta-analysis. J Neurointerv Surg. (2015) 7:868-74. doi: 10.1136/neurintsurg-2014-011418
- Writing Group for the Basilar Group, Zi W, Qiu Z, Wu D, Li F, Liu H, et al. Assessment of endovascular treatment for acute basilar artery occlusion via a nationwide prospective registry. *JAMA Neurol.* (2020) 77:561– 73. doi: 10.1001/jamaneurol.2020.0156
- 26. Liu X, Dai Q, Ye R, Zi W, Liu Y, Wang H, et al. Endovascular treatment versus standard medical treatment for vertebrobasilar artery occlusion (BEST): an open-label, randomised controlled trial. *Lancet Neurol.* (2020) 19:115–22. doi: 10.1016/S1474-4422(19)30395-3
- Bacigaluppi M, Semerano A, Gullotta GS, Strambo D. Insights from thrombi retrieved in stroke due to large vessel occlusion. J Cereb Blood Flow Metab. (2019) 39:1433–51. doi: 10.1177/0271678X198 56131
- 28. Powers WJ, Derdeyn CP, Biller J, Coffey CS, Hoh BL, Jauch EC, et al. 2015 American Heart Association/American Stroke Association Focused Update of the 2013 guidelines for the early management of patients with acute ischemic stroke regarding endovascular treatment: a guideline for healthcare professionals from the American Heart Association/American

Stroke Association. *Stroke.* (2015) 46:3020–35. doi: 10.1161/STR.0000000000 00074

- Vergouwen MD, Algra A, Pfefferkorn T, Weimar C, Rueckert CM, Thijs V, et al. Time is brain(stem) in basilar artery occlusion. *Stroke.* (2012) 43:3003– 6. doi: 10.1161/STROKEAHA.112.666867
- Mokin M, Sonig A, Sivakanthan S, Ren Z, Elijovich L, Arthur A, et al. Clinical and procedural predictors of outcomes from the endovascular treatment of posterior circulation strokes. *Stroke.* (2016) 47:782–8. doi: 10.1161/STROKEAHA.115.011598
- Bouslama M, Haussen DC, Aghaebrahim A, Grossberg JA, Walker G, Rangaraju S, et al. Predictors of good outcome after endovascular therapy for vertebrobasilar occlusion stroke. *Stroke.* (2017) 48:3252– 7. doi: 10.1161/STROKEAHA.117.018270
- Li C, Zhao W, Wu C, Shang S, Chen J, Ren M, et al. Outcome of endovascular treatment for acute basilar artery occlusion in the modern era: a single institution experience. *Neuroradiology*. (2018) 60:651– 9. doi: 10.1007/s00234-018-2011-7
- Rangaraju S, Jovin TG, Frankel M, Schonewille WJ, Algra A, Kappelle LJ, et al. Neurologic examination at 24 to 48 hours predicts functional outcomes in basilar artery occlusion stroke. *Stroke.* (2016) 47:2534–40. doi: 10.1161/STROKEAHA.116.0 14567
- 34. Lee WJ, Jung KH, Ryu YJ, Lee KJ, Lee ST, Chu K, et al. Acute symptomatic basilar artery stenosis: MR imaging predictors of early neurologic deterioration and long-term outcomes. *Radiology*. (2016) 280:193– 201. doi: 10.1148/radiol.2016151827

- 35. Manceau PF, Soize S, Gawlitza M, Fabre G, Bakchine S, Durot C, et al. Is there a benefit of mechanical thrombectomy in patients with large stroke (DWI-ASPECTS ≤ 5)? Eur J Neurol. (2018) 25:105–10. doi: 10.1111/ene.13460
- Pallesen LP, Khomenko A, Działowski I, Barlinn J, Barlinn K, Zerna C, et al. CT-angiography source images indicate less fatal outcome despite coma of patients in the Basilar Artery International Cooperation Study. *Int J Stroke*. (2017) 12:145–51. doi: 10.1177/1747493016669886

**Conflict of Interest:** The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

**Publisher's Note:** All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

Copyright © 2021 Cao, Mo, Chen, Shao, Xuan, Peng and Zhu. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.