

A comparison between acute large vessel occlusion in the posterior circulation and anterior circulation after endovascular treatment: the ANGEL-ACT registry experience

Xuan Sun, Jingyu Zhang (), Xu Tong (), Baixue Jia (), Dapeng Mo, Ning Ma, Feng Gao, Lian Liu, Xiaochuan Huo (), Ligang Song, Yiming Deng, Zhongrong Miao ()

ABSTRACT

Background and purpose Endovascular treatment (EVT) has shown an overwhelming benefit for acute anterior circulation artery occlusion (ACO). Whether it can achieve the same outcomes in posterior circulation artery occlusion (PCO) has not been well explained. We aimed to evaluate the characteristics and prognosis of ACO and PCO after EVT in a nationwide registry.

Method The present analysis was based on the prospective ANGEL-ACT Registry in China between November 2017 and March 2019. Demographic data, periprocedural times, recanalisation rate, intracranial haemorrhage (ICH) and 90-day functional outcomes were compared between the ACO and PCO groups. Results A total of 1793 patients were analysed including 397 (22.1%) consecutive patients with PCO and 1396 (77.9%) patients with ACO treated with EVT. A larger proportion of patients with PCO had intracranial atherosclerotic disease and received extra angioplasty during EVT. Successful recanalisation and 90-day favourable functional outcomes did not differ significantly between the two groups. Patients with PCO showed lower 24-hour ICH and symptomatic ICH rates. There was a trend towards higher mortality rate in the PCO group (22.09% vs 14.44%; adjusted OR 1.286 (95% CI 0.820 to 2.017), p=0.2731), especially when the onset to puncture time was over 6 hours (30.77% vs 11.13%; adjusted OR 2.673 (95% CI 1.280 to 5.583), p=0.0089, interactive p=0.0002). **Conclusions** In this large prospective multicentre registry, there was a significant difference in the characteristics and periprocedural features between patients with PCO and ACO. However, successful recanalisation and 90-day favourable functional outcomes in PCO were equivalent to those in ACO.

INTRODUCTION

Endovascular treatment (EVT) has shown an overwhelming benefit for acute anterior circulation artery occlusion (ACO).^{1–7} Evidence on the efficacy and safety of EVT for posterior circulation artery occlusion (PCO) is still lacking.⁸ With PCO, however, acute ischaemic

strokes (AIS) rarely occur with varied presentation and remain devastating with a high mortality and morbidity rate, especially without early reperfusion.⁹ With updates to EVT materials and the development of multimodal imaging evaluations, a larger number of clinical practices focus on EVT in PCO populations.¹⁰ ¹¹ Whether EVT can achieve the same outcomes in PCO as in ACO has not been well explained. We therefore analysed the data collected from a prospective multicentre registry¹² to investigate differences in the characteristics of ACO and PCO and to ascertain the treatment prognosis between the two groups.

METHOD

Patients' selection

Written informed consent was obtained from all patients to participate in the study, and the privacy of the research subjects was strictly protected. The data of the patients included in this study were retrieved from the ANGEL-ACT Registry. The ANGEL-ACT Registry is a multicentric, nationwide, prospective registry study launched in December 2017 and terminated in March 2019. The characteristics and operating protocols of ANGEL-ACT Registry have already been published.¹² The ANGEL-ACT Registry enrolled 2004 patients with AIS who had undergone EVT. Patients were enrolled according to the following inclusion criteria: (1) Age ≥ 18 years; (2) Diagnosis of AIS caused by imaging-confirmed intracranial large vessel occlusion (LVO); and (3) Initiation of any type of EVT, including intra-arterial thrombolysis, mechanical thrombectomy (MT), angioplasty and permanent stenting. The exclusion criteria were (1) Isolated cervical internal carotid artery or vertebral

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XS and JZ contributed equally.

XS and JZ are joint first authors.

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Interventional Neuroradiology Center, Beijing Tiantan Hospital, Capital Medical University, Beijing, China

Correspondence to

Professor Zhongrong Miao; miaozhongrong123@126.com





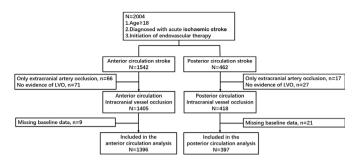


Figure 1 Flow chart of patient screening for the ANGEL-ACT registry. LVO, large vessel occlusion.

artery (VA) occlusion and (2) No evidence of LVO angiogram. Among the enrolled patients, 83 patients with isolated extracranial artery occlusion, 98 patients with no evidence of LVO on digital subtraction angiography, and 30 patients with missing baseline data were excluded from the data analysis, leaving 397 (22.1%) consecutive patients with PCO and 1396 (77.9%) patients with ACO. A flow chart of patient screening is presented in figure 1.

Imaging assessment

All enrolled patients were required to complete brain plain CT scan before the procedure. The examination method of plain CT scan combined with CT angiography (CTA) was recommended for patients with ACO, and MRI and angiography (MRI/MRA) was recommended for patients with PCO. If priority examination cannot be performed due to the limitation of central conditions, the imaging evaluation of brain tissue and blood vessels can be completed according to the actual situation. The head plain CT scan must be completed 24±3 hours after the procedure. If conditions permit, MRI/MRA can be performed to evaluate the final infarction volume, and CTA is preferred to evaluate whether there is vascular recanalisation or not.

Data collection

Data reporting preprocedural and procedural variables were extracted by reviewing the patient charts, procedure notes, angiograms, and progression and follow-up

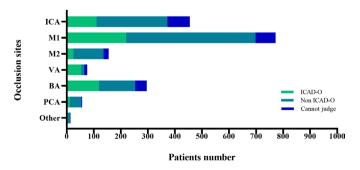


Figure 2 The distribution of artery occlusion site of the ANGEL-ACT registry. ICA, intracranial artery; ICAD-O,intracranial atherosclerotic disease related occlusion; M1, middle cerebral artery M1 segment; M2, middle cerebral artery M2 segment; BA, basilar artery; PCA, posterior cerebral artery; VA, vertebral artery.

notes. Preprocedural variables included in this analysis were age, sex, administration of intravenous thrombolysis (bridging IVT), National Institute of Health Stroke Scale (NIHSS) Score on admission, preantiplatelet treatment, vascular risk factors (such as previous ischaemic stroke, hypertension, diabetes mellitus, atrial fibrillation, alcohol consumption, and previous or current smoking habit), and pathogenesis of stroke (large artery atherosclerosis, cardioembolism and others). Procedural variables included the time from symptom onset to puncture and recanalisation (OPT, ORT), the types of endovascular procedures (stent thrombectomy, intra-arterial thrombolysis, catheter aspiration, balloon angioplasty and permanent stenting) and administration of heparin IIb/ IIIa receptor inhibitors during thrombectomy. Baseline and postprocedural modified Thrombolysis in Cerebral Infarction Scale Score, and occurrence of intracranial haemorrhage (ICH) on post-treatment imaging were reviewed by core laboratory image interpretation. All patients' images were selected separately and reviewed by two interventionalists blinded to the patient details.

Outcome measures

The primary outcome was functional independence, defined as a modified Rankin Scale (mRS) Score of 0–2 at 90 days. The 90-day follow-up was ascertained using a standardised telephone interview performed by trained investigators blinded to the baseline and procedural data. Secondary outcomes included (1) Technical efficacy according to successful reperfusion rate, defined as a postprocedural modified Thrombolysis in Cerebral Infarction Scale Score of 2b or 3; (2) Any type of ICH at 24hours; (3) Symptomatic ICH (sICH) according to the Heidelberg Bleeding Classification criteria at 24hours; and (4) Mortality within 90 days.

Statistical analysis

All data were presented as the median (IQR) for continuous and ordinal variables and a number (percentage) for categorical variables. A comparison of the baseline and procedural characteristics between both groups was performed using the Mann-Whitney U test for continuous and ordinal variables and Pearson's χ^2 test for categorical variables. A binary logistic regression model was used for multivariable analysis. Outcomes were further compared based on subgroups of OPT over or less than 6 hours. A multivariable model was constructed with variables with values of p<0.1 in univariate analysis using forward inclusion. The p value for inclusion in the model was <0.05. All analyses were performed in SAS, V.9.4, software (SAS Institute, Cary, North Carolina, USA).

RESULTS

A total of 1793 patients were included in the analysis: 1396 (77.9%) patients with ACO and 397 (22.1%) patients with PCO. The distribution of occlusion sites in both ACO and PCO groups is shown in figure 2. Baseline characteristics are listed in table 1. ...

Characteristic	ACO (n=1396)	PCO (n=397)	P value
Age (years)	66(56–73)	64(55–72)	0.0038
Male (n,%)	878 (62.89%)	301 (75.82%)	<0.0001
BMI	24.42±3.63	24.79±3.70	0.0479
Hypertension (n,%)	757 (54.23%)	272 (68.51%)	<0.0001
Hypercholesterolaemia (n,%)	109 (7.81%)	50 (12.59)	0.0031
Diabetes mellitus (n,%)	246 (17.62%)	86 (21.66%)	0.0674
CAD (n,%)	212 (15.19%)	63 (15.87%)	0.7390
Current smoker (n,%)	408 (29.23%)	140 (35.26%)	0.0212
Atrial fibrillation (n,%)	572 (40.97%)	78 (19.65%)	< 0.0001
History of stroke (n,%)	257 (18.41%)	89 (22.42%)	0.0741
History of TIA (n,%)	31 (2.22%)	22 (5.54%)	0.0006
Admission NIHSS	16(12–20)	22 (11–35)	<0.0001
ASPECTS/pc-ASPECTS	9 (7–10)	9 (7–10)	
SBP, mm Hg	145(130–160)	150(138–166)	< 0.0001
IV thrombolysis (n,%)	416 (29.80%)	108 (27.20%)	0.3157
Onset to thrombolysis, min	166 (120–180)	173 (120–180)	0.0410
General anaesthesia (n,%)	502 (35.96%)	221 (55.67%)	< 0.0001
ASTIN/SIR 3–4 (n,%)	522 (37.39%)	173 (43.58%)	0.0256
Underlying ICAD (n,%)*	359 (29.47%)	175 (51.47%)	< 0.0001
Tandem occlusion (n,%)	305 (21.85%)	74 (18.64%)	0.1671
IA thrombolysis(n,%)	99 (7.09%)	50 (12.59%)	0.0005
Mechanical thrombectomy (n,%)	1269 (90.9%)	338 (85.14%)	0.0009
Number of thrombectomy	2 (1–3)	1 (1–2)	< 0.0001
Heparin during MT (n,%)	669 (47.92%)	205 (51.64%)	0.1914
IIb/IIIa inhibitor (n,%)	677 (48.50%)	250 (62.97%)	< 0.0001
Extra angioplasty (n,%)	365 (26.15%)	160 (40.30%)	< 0.0001
Permanent stenting (n,%)	212 (15.19%)	113 (28.46%)	< 0.0001
Onset to puncture, min	300 (210–440)	298 (208–428)	0.2664
Onset to recanalisation, min	404 (300–555)	481.5 (360–765)	<0.0001
Puncture to recanalisation, min	83 (51–128)	93 (59–139)	0.0088
Stoke subtype by TOAST criteria†			
Large artery atherosclerosis (n,%)	634 (48.40%)	257 (67.99%)	< 0.0001
Cardioembolism (n,%)	517 (39.47%)	70 (18.52%)	
Other or unknown aetiology (n,%)	159 (12.14%)	51 (13.49%)	

*Missing 235 data.

†Missing 105 data.

ACO, anterior circulation artery occlusion; ASPECTS, Alberta Stroke Programme Early CT Score; ASTIN/SIR, American Society of Interventional and Therapeutic Neuroradiology/Society of Interventional Radiology Collateral Flow Grading System; BMI, body mass index; CAD, coronary artery disease; IA, intra-artery; ICAD, intracranial atherosclerotic disease; IV, intravenous; MT, mechanical thrombectomy; NIHSS, National Institute of Health Stroke Scale; pc-ASPECTS, posterior circulation Acute Stroke Prognosis Early CT Score; PCO, posterior circulation artery occlusion; SBP, systolic blood pressure; TIA, transient ischaemic attack; TOAST, Trial of Org 10172 in Acute Stroke Treatment.

Compared with patients with ACO, those with PCO were more likely to be male (75.8% vs 62.89%, p<0.0001) and younger (64 years, IQR (55–72) vs 66 years, IQR (56–73), p=0.0038) and had a history of hypertension (68.51% vs 54.23%, p<0.0001),

hypercholesterolaemia (12.59% vs 7.81%, p=0.0031), smoking (35.26% vs 29.23%, p=0.0212) and transient ischaemic attack (TIA) (5.54% vs 2.22%, p=0.0006); higher NIHSS Score at admission (22, IQR (11–35) vs 16, IQR (12–20), p<0.0001); and longer ORT (481.5,

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				Unadjusted		Adjusted 1*		Adjusted 2#	
	All patient	ACO (n=1396)	PCO (n=397)	OR (95% CI)	P value	aOR (95% CI)	P value	aOR (95% CI)	P value
Any ICH within 24 hours*	469 (26.16%)	402 (28.80%)	67 (16.88%)	0.502 (0.377 to 0.669) <0.0001 0.521 (0.390 to 0.695)	<0.0001		<0.0001	0.624 (0.416 to 0.935)	0.0224
Symptomatic ICH within 24 hours†	130 (7.66%)	118 (8.91%)	12 (3.22%)	0.340 (0.186 to 0.623)	0.0005	0.0005 0.363 (0.198 to 0.667)	0.0011	0.449 (0.202 to 0.998)	0.0494
Recanalisation‡	1577 (87.95%)	1577 (87.95%) 1227 (87.89%)	350 (88.16%)	1.026 (0.727 to 1.447)	0.8858	0.8858 1.016 (0.718 to 1.438)	0.9271	0.9271 0.9222 (0.510 to 1.668)	0.7880
mRS 0–2 at 90 days§	775 (45.16%)	607 (45.40%)	168 (44.33%)	0.958 (0.761 to 1.205)	0.7114	0.7114 0.895 (0.708 to 1.132)	0.3561	0.3561 1.304 (0.924 to 1.840)	0.1315
mRS 0–3 at 90 days ¶	948 (55.24%)	756 (56.54%) 192 (50.66%)	192 (50.66%)	0.789 (0.628 to 0.992)		0.042 0.735 (0.581 to 0.929)	0.0101	0.0101 0.988 (0.700 to 1.395)	0.9459
Mortality at 90 days**	273 (15.91%)	193 (14.44%) 379 (22.09%)		1.586 (1.187 to 2.120) <0.0001 1.704 (1.269 to 2.289)	<0.0001	1.704 (1.269 to 2.289)	0.0004	0.0004 1.286 (0.820 to 2.017)	0.2731
Adjusted 1* Adjust age and gender. Adjusted 2# Adjust all confounding factors in multivariable analysis. Adjust PRT, hypertension, extra angioplasty, ASTIN/SIR grade, general anaesthesia. *Adjust SBP, hypercholesterolaemia, general anaesthesia, ASTIN/SIR grade, intracra tAdjust SBP, hypercholesterolaemia, general anaesthesia, acTIN/SIR grade, intracra fadjust SBP, hypercholesterolaemia, general anaesthesia, activity general anaesthesia, and tadjust SBP, hypercholesterolaemia, general anaesthesia, mechanical thrombectomy. §Adjust general anaesthesia, admission NIHSS. ¶Adjust general anaesthesia, admission NIHSS. ACO, anterior circulation artery occlusion; aOR, adjusted OR; ASTIN/SIR, American i Flow Grading System; ICH, intracranial haemorrhage; mRS, modified Rankin Scale; blood pressure.	ge and gender. Il confounding factol nsion, extra angiople nolesterolaemia, gen ngioplasty, general a sisthesia, admission h isthesia, extra angiol riral anesthesia, adr tion artery occlusion r; ICH, intracranial h	rs in multivariable a asty, ASTIN/SIR gra naesthesia, <i>H</i> naesthesia, mecha NIHSS. plasty, admission h nission NIHSS. r; aOR, adjusted Ol aemorrhage; mRS,	analysis. ade, general anaes ASTIN/SIR grade, i inical thrombecton VIHSS. R; ASTIN/SIR, Ame R; ASTIN/SIR, Ame	Adjusted 1* Adjust age and gender. Adjust PRT, hypertension, extra angioplasty, ASTIN/SIR grade, general anaesthesia. *Adjust PRT, hypertension, extra angioplasty, general anaesthesia, ASTIN/SIR grade, intracranial atherosclerotic disease. †Adjust PRT, extra angioplasty, general anaesthesia, ASTIN/SIR grade, intracranial atherosclerotic disease. \$Adjust PRT, extra angioplasty, general anaesthesia, and therombectomy. \$Adjust general anaesthesia, admission NIHSS. ¶Adjust general anaesthesia, admission NIHSS. "Adjust general anaesthesia, admission NIHSS. "Adjust general anaesthesia, admission NIHSS. "Adjust general anaesthesia, admission NIHSS. "Of anterior circulation artery occlusion; aOR, adjusted OR; ASTIN/SIR, American Society of Interventional and Therapeutic Neuroradiology/Society of Interventional Radiology Collateral Flow Grading System; ICH, intracranial haemorrhage; mRS, modified Rankin Scale; NIHSS, National Institute of Health Stroke Scale; PCO, posterior circulation artery occlusion; SBP, systolic blood pressure.	lisease. and Thu	arapeutic Neuroradiology/6 Ith Stroke Scale; PCO, poc	Society of II	tterventional Radiology Coll lation artery occlusion; SBP	ateral systolic

IQR (360–765) vs 404, IQR (300–555), p<0.0001). Furthermore, they were more likely to have intracranial atherosclerotic disease (ICAD, 51.47% vs 29.47%, p<0.0001) and receive intra-artery thrombolysis (12.59% vs 7.09%, p=0.0005) and IIb/IIIa receptor inhibitors (62.97% vs 48.50%, p<0.0001) and permanent stenting (28.46% vs 15.19%, p<0.0001) during EVT.

Regarding the pathogenesis of stroke, large artery atherosclerosis (67.99%) was more often observed in the PCO group, while cardiac embolism (18.52%) and other types of pathogeneses (13.49%) were observed less often. Stent thrombectomy and catheter aspiration were performed more often in patients with ACO than in patients with PCO. Balloon angioplasty and permanent stenting were performed more often in patients with PCO than in those with ACO.

A comparison of recanalisation rates, postoperative haemorrhage and outcomes after a 90-day follow-up is shown in table 2. ICH, whether symptomatic or asymptomatic, occurred more frequently in ACO. PCO showed lower 24-hour ICH (16.88% vs 28.80%; OR 0.502 (95% CI 0.377 to 0.669), p<0.0001; adjusted OR 0.624 (95% CI 0.416 to 0.935), p=0. 0224) and sICH (3.22% vs 8.91%; OR 0.340 (95% CI 0.186 to 0.623), p=0.0005; adjusted OR 0.449 (95% CI 0.202 to 0.998), p=0.0494) rates than did ACO. Successful recanalisation and a favourable functional outcome at 90 days (mRS 0-2) did not differ significantly between PCO and ACO in either unadjusted or adjusted logistic regression (88.16% vs 87.89% and 44.33% vs 45.40%). The mRS distribution of the two groups is shown in figure 3. It can be clearly seen from the data and graphs that there was a trend towards higher mortality rate in the PCO group than in the ACO group (22.09% vs 14.44%; OR 1.586 (95% CI 1.187 to 2.120), p<0.0001; adjusted OR 1.286 (95% CI 0.820 to 2.017), p=0.2731).

In the subgroup analysis, we found that patients with PCO showed similar independent functional outcomes (57.87% vs 54.51%) and mortality (15.32% vs 16.46%) at 90 days and a lower sICH risk within 24 hours than patients with ACO when OPT was less than 6 hours.

However, when OPT was over 6 hours, PCO showed a devastating mortality rate compared with ACO (30.77% vs 11.13%; OR 3.548 (95% CI 2.260 to 5.569), p<0.0001; adjusted OR 2.673 (95% CI 1.280 to 5.583), p=0.0089, interactive p=0.0002) (table 3).

Furthermore, multivariate logistic regression analysis adjusting for baseline characteristics was performed in the two groups separately. In the ACO group, age <66 years (OR 1.681 (95%CI 1.255 to 2.252)), NIHSS Score <16 (OR 2.131 (95%CI 1.594 to 2.850)), Alberta Stroke Programme Early CT Score (ASPECTS) ≥9 (OR 1.489 (95% CI 1.090 to 2.036)) and OPT>6 hours (OR 1.359 (95%CI 1.006 to 1.837)) were significant predictors for 90-day functional independence consistent with the majority of reported studies. While in the PCO group, we just found NIHSS Score <16 (OR 2.890 (95%CI 1.773 to 4.694)) were a significant predictor for 90-day functional independence (figure 4). Finally, we compared the baseline characteristics and clinical outcomes of different occlusion sites, namely the VA-V4 segment, basilar artery (BA) and posterior cerebral artery (PCA) in the PCO group, and showed the results in the online supplemental table 1 and online supplemental table 2. Besides patients with PCA showed a lower successful recanalisation rate and no death at 90 days, other clinical outcomes did not relate to location site (online supplemental figure 1). A 90-day mRS distribution of VA, BA and PCA is shown in online supplemental figure 2.

DISCUSSION

Our present study aimed to compare the clinical characteristics and outcomes between patients with PCO and those with ACO treated by EVT in real situations. In this report, we found that the pathogenesis of stroke and treatment procedures were significantly different in PCO and ACO, which affected their clinical outcome.

Similar to many other studies,^{13 14} we found that large artery atherosclerosis and ICAD were more common in PCO than in ACO and that the larger percentage of men, higher BMI, and history of hypertension, hypercholesterolaemia, TIA and smoking related to ICAD were also

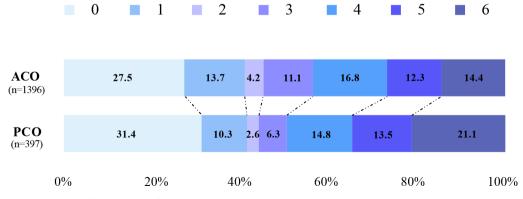


Figure 3 Outcome of the modified Rankin Scale at 90 days between the anterior circulation artery occlusion (ACO) and posterior circulation artery occlusion (PCO) groups.

			Multivariable logistic re	egression							
			Unadjusted			Adjusted 1*			Adjusted 2#		
	ACO	РСО	OR (95% CI)	P value	P value for interaction	aOR (95% Cl)	P value	P value for interaction	aOR (95% Cl)	P value	P value for interaction
Any ICH within 24 hours*	24 hours*										
OPT											
≤6hours	246 (28.34%)	40 (16.19%)	0.489 (0.338 to 0.707)	0.0001	0.8034	0.510 (0.352 to 0.739)	0.0004	0.8229	0.706 (0.412 to 1.209)	0.2041	0.1524
>6 hours	153 (29.77%)	27 (18.24%)	0.526 (0.333 to 0.832)	0.0060		0.539 (0.340 to 0.854)	0.0085		0.481 (0.250 to 0.926)	0.0285	
Symptomatic	Symptomatic ICH within 24 hours†	+									
OPT											
≤6hours	77 (9.45%)	3 (1.32%)	0.128 (0.040 to 0.409)	0.0005	0.0115	0.140 (0.044 to 0.449)	0.0009	0.0118	0.249 (0.066 to 0.943)	0.0406	0.2331
>6 hours	40 (8.06%)	9 (6.25%)	0.760 (0.360 to 1.606)	0.4724		0.763 (0.359 to 1.618)	0.4801		0.609 (0.200 to 1.858)	0.3839	
mRS 0-2 at 90 days‡) days‡										
OPT											
≤6hours	353 (43.05%)	118 (50.21%)	1.334 (0.998 to 1.784)	0.0518	0.0004	1.266 (0.941 to 1.703)	0.1189	0.0004	1.812 (1.145 to 2.868)	0.0112	0.0038
>6 hours	246 (48.91%)	50 (34.97%)	0.562 (0.382 to 0.826)	0.0034		0.514 (0.346 to 0.763)	0.0010		0.938 (0.532 to 1.655)	0.8262	
mRS 0-3 at 90 days§) days§										
ОРТ											
≤6hours	447 (54.51%)	136 (57.87%)	1.146 (0.855 to 1.537)	0.3613	<0.0001	1.077 (0.798 to 1.453)	0.6286	<0.0001	1.360 (0.857 to 2.158)	0.1920	0.0025
>6 hours	300 (77.86%)	56 (39.16%)	0.436 (0.298 to 0.637)	<0.0001		0.398 (0.269 to 0.590)	<0.0001		0.717 (0.408 to 1.259)	0.2468	
Mortality at 90 days¶) days¶										
OPT											
≤6hours	135 (16.46%)	36 (15.32%)	0.918 (0.615 to 1.370)	0.6748	<0.0001	0.979 (0.653 to 1.467)	0.9181	<0.0001	0.690 (0.363 to 1.314)	0.2590	0.0002
>6 hours	56 (11.13%)	44 (30.77%)	3.548 (2.260 to 5.569)	<0.0001		3.829 (2.409 to 6.085)	<0.0001		2.673 (1.280 to 5.583)	0.0089	
Adjusted 1* Adju Adjusted 2# Adj *Adjust PRT, hyr †Adjust BRP, hyr ‡Adjust general \$Adjust general \$Adjust general	Adjusted 1* Adjust age and gender. Adjusted 2# Adjust all confounding factors in multivariable analysis. Adjust PRT, hypertension, extra angioplasty, ASTIN/SIR grade, gen tAdjust SBP, hypercinolesterolaemia, general anaesthesia, ASTIN/SI &Adjust general aneschesia, admission NIHSS.	ctors in multivariable oplasty, ASTIN/SIR ç general anaesthesia on NIHSS. gioplasty, admissior admission NIHSS.	Adjusted 1* Adjust age and gender. Adjusted 2# Adjust all confounding factors in multivariable analysis. Adjust PRT, hypertension, extra angioplasty, ASTIN/SIR grade, general anaesthesia. Adjust SBP, hypercholestenclaemia, general anaesthesia, ASTIN/SIR grade, intracrania Adjust general anaesthesia, admission NIHSS. Adjust general anaesthesia, axtra angioplasty, admission NIHSS.	al atherosolerotic disease.	tic disease.				Adjusted 1* Adjust age and gender. Adjusted 2# Adjust all confounding factors in multivariable analysis. "Adjust PRT, hypercholesteroiaemia, general anaesthesia. "Adjusts SBP, hypercholesteroiaemia, general anaesthesia, addission anaesthesia, ASTIN/SIR grade, intracranial atherosclerotic disease. Adjust sgeneral anaesthesia, admission NIHSS. "Adjust sgeneral anaesthesia, admission NIHSS.		

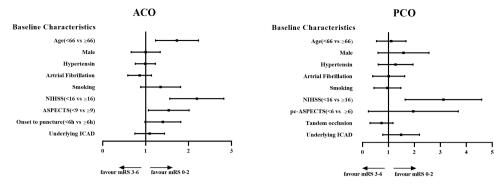


Figure 4 Predictors of a 90-day mRS Score 0–2 between the acute anterior circulation artery occlusion (ACO) and posterior circulation artery occlusion (PCO) groups. ASPECTS, Alberta Stroke Programme Early CT Score; ICAD, intracranial atherosclerotic disease; mRS, modified Rankin Scale; NIHSS, National Institute of Health Stroke Scale; pc-ASPECTS, posterior circulation Acute Stroke Prognosis Early CT Score.

observed in patients with PCO. Elderly patients and atrial fibrillation were more often observed in the ACO group.¹⁵

Due to the significant difference in pathogenesis between the two groups, EVT strategies were not the same. MT is more frequently used to treat ACO, while the combination of intra-artery thrombolysis and/or IIb/ IIIa receptor inhibitors is more prevalent in PCO for resolving ICAD lesions. In our study, rescue angioplasty was also an important means for maintaining recanalisation, as balloon angioplasty and permanent stents are more commonly used in patients with PCO. It is clear that the technique of successful recanalisation was satisfactory in both patients with PCO and patients with ACO, with success rates of 88.16% and 87.89%, respectively, slightly higher than those reported by a meta-analysis.¹⁶ The independent functional outcomes at 90 days seemed to be equivalent between the PCO and ACO groups (44.33% vs 45.40%). We were glad to see that the efficacy of properly performed EVT in patients with PCO could achieve similar outcomes to that in patients with ACO in real clinical practice.

Even with comparable successful recanalisation rates and independent functional outcomes, patients with PCO thrombectomy had relatively worse outcomes than those with ACO, as shown by a higher rate of mortality in our analysis (22.09% vs 14.44%). There was a high chance that futile recanalisation occurred in patients with PCO. Previous studies¹⁷⁻¹⁹ pointed out that age, stroke severity, baseline NIHSS Score and duration of the procedure were related to poor outcome in patients with PCO. Early symptoms of PCO are complex and not easy to recognise, and the time from onset to initiation of treatment is longer; these factors often result in ineligibility for intravenous thrombolysis and a delay of initiation of the EVT procedure.²⁰ Another reason for the high mortality rate of PCO might be attributable to disturbing brainstem symptoms, such as a comatose state, dysphagia, tracheostomy, hypostatic pneumonia and complications due to a long-term bedridden state. However, the PCO mortality in our study was slightly lower than that in a meta-analysis¹⁶ including 474 patients with PCO (26.6%) and the EVT arm group in the Basilar Artery Occlusion Endovascular

Intervention versus Standard Medical Treatment (BEST) Trial¹⁰ and The EVT for Acute Basilar Artery Occlusion Study (BASILAR)¹¹ (33% and 47.3%).

ICAD may be the main determining factor that results in different outcomes between patients with PCO and patients with ACO. ICAD was more frequently observed in patients with PCO and perhaps played an absolutely different role in the mechanisms and outcomes of stroke in PCO and ACO.²⁰ A study compared the EVT procedural and clinical outcomes among patients with ACO with and without ICAD. The results showed that the recanalisation rate and clinical outcomes did not differ between patients with or without ICAD in ACO.²¹ Another study compared the recanalisation rate and clinical outcomes of MT in different subtypes of basilar artery occlusion (BAO), and they found that BAO with in situ atherosclerotic thrombosis had a worse recanalisation rate and poorer clinical outcome than embolic BAO.²² In our study, we also found that more patients suffered TIAs before stroke in PCO. With stroke recurrence, the risk of cumulative disability and poor prognosis increases accordingly,²³ which may be a reason for a poor outcome in PCO. Therefore, we should strengthen the identification and intervention of patients with ICAD in the early stage, especially for those who present with symptomatic posterior circulation ICAD and suffer recurrent TIAs. Early angioplasty may be reasonable and should be considered from a more positive perspective.

Similar to other research findings,²⁴ patients with PCO also showed a lower rate of ICH than patients with ACO, including sICH and any type of ICH, even though intraartery thrombolysis and IIb/IIIa receptor inhibitors were more commonly used in our study. This might be explained by the larger volume of ischaemic tissue in ACO, which usually results from cardiac embolism, and oral preanticoagulant agents, which are also a risk factor.²⁵ Compared with ACO, PCO typically causes smaller ischaemic lesion volumes, which may reduce the rate of ICH after EVT. In our study, we found that there are more patients with good collateral circulation presented American Society of Interventional and Therapeutic Neuroradiology/Society of Interventional Radiology Score 3–4 grade in PCO than

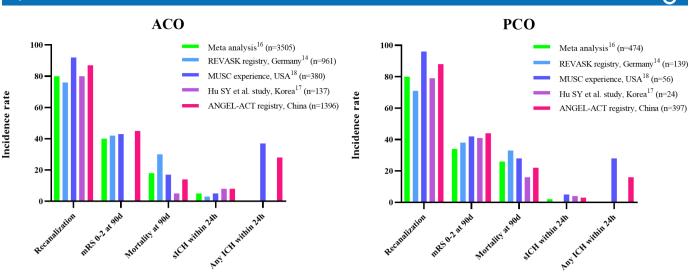


Figure 5 The clinical outcomes compared with other published studies. ACO, anterior circulation artery occlusion; ICH, intracranial haemorrhage; mRS, modified Rankin Scale; PCO, posterior circulation artery occlusion; sICH, symptomatic intracranial haemorrhage.

ACO (43.58% vs 37.39%, p=0.0256), which maybe result in a lower chance of ischaemia-reperfusion injury after recanalisation. A study reported that blood-brain barrier derangements were more frequently observed in ACO (22%) than in PCO (7%).²⁶ Another reason is that the number of MT device passes is higher in patients with ACO, which has been verified to be associated with ICH after EVT.²⁷ We present a brief comparison of our findings with other studies^{14 16-18} in figure 5.

Subgroup analysis according to the OPT time window gave rise to different functional outcomes. We found that successful recanalisation, 90-day favourable functional outcome and mortality in PCO were equivalent to ACO within a 6-hour OPT time window. However, when OPT was over 6 hours, the risk of 90-day death in PCO was 2.673 times higher than that in ACO. The test for interaction showed a significant difference in the clinical outcomes affected by ischaemic location at different OPT times.

Regardless of anterior and posterior circulation infarction, EVT in the early stage of onset is more to save the brain tissue in the ischaemic penumbra, while the collateral circulation in the PCO is better, which can quickly restore brain function and achieve the same clinical prognosis as the ACO. However, as the time of cerebral ischaemia is prolonged, the brainstem infarction caused by the progression of PCO often causes malignant cerebral infarction, brain herniation, death and other events that badly affect the clinical prognosis of patients, resulting in totally different clinical outcomes compared with ACO. This means that a selected OPT time window should be applied for assessment at admission for both patients with ACO and patients with PCO.

Finally, we tried to assess the potential predictive factors related to the 90-day independent outcome of ACO and PCO groups separately by multivariable analysis. The relevant data of the ACO group have been presented in our formerly published paper. And we found age <66 years, NIHSS Score <16, ASPECTS \geq 9 were significant predictors for 90-day functional independence consistent with the majority of reported studies. However, in the PCO group, we just found patients with lower NIHSS Score have a towards to a better clinical outcome. It is possible that in the PCO group, disease severity and brainstem infarction play a greater role in prognosis. Since we only adjusted baseline data in the multivariable analysis, there perhaps exist other possible predictors that need to be figured out in our later research.

Some limitations should also be mentioned before generalising the present findings. First, this was not a randomised study, and measured or unmeasured variables may represent potential confounders that cannot be ruled out despite adjustment. Second, this study was exclusively conducted in the Chinese population. Delays in the process of EVT in China and the variety and complexity of surgical procedures, as well as the high proportion of ICAD in Chinese patients, more rescue balloon angioplasty and permanent stenting, and differences in perioperative antithrombotic therapy will affect the extrapolation of the results of our study to other populations. The sample size of patients with PCO was relatively small compared with that of patients with ACO. We compared the different characteristics and prognoses between ACO and PCO, which could be analysed separately in later trials.

CONCLUSION

Although the pathogenesis, anatomy and EVT strategies are different, with timely recanalisation, patients with PCO could obtain a similar clinical benefit to that of patients with ACO.

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Contributors ZM designed and led the study, responsible for the overall content as the guarantor. XS and JZ analyzed the data and prepared the first draft of the report. XS and JZ contributed equally to the work. XT and BJ were responsible for acquisition of data. DM, NM, FG, LL, XH, LS and YD were involved in revising the article for important intellectual content. All authors critically reviewed the article and approved the final version.

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ORCID iDs

Jingyu Zhang http://orcid.org/0000-0001-5723-9272 Xu Tong http://orcid.org/0000-0002-3277-6941 Baixue Jia http://orcid.org/0000-0001-8782-5344 Xiaochuan Huo http://orcid.org/0000-0003-1264-5132 Zhongrong Miao http://orcid.org/0000-0002-0165-9814

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