

Endovascular thrombectomy for M2 occlusions: comparison between forced arterial suction thrombectomy and stent retriever thrombectomy

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ABSTRACT

Background To date there has been no direct comparison of two frequently used endovascular thrombectomy (EVT) methods (forced arterial suction thrombectomy (FAST) and stent retriever thrombectomy) in M2 occlusions. We review our experiences with EVT performed using FAST and stent retriever thrombectomy in such cases.

Methods The subjects comprised 41 patients with an M2 occlusion who underwent EVT (25 with FAST, 16 with stent retriever thrombectomy). The patients' data were retrospectively analyzed to evaluate the technical characteristics and angiographic outcome of the two EVT techniques.

Results Thrombolysis In Cerebral Infarction (TICI) grades 2b–3 using the first chosen technique did not differ significantly between the two techniques (FAST 64.0% vs stent retriever thrombectomy 81.2%, $p=0.305$). Time from groin puncture to reperfusion was significantly shorter for stent retriever thrombectomy (53.0 vs 38.5 min; $p=0.045$). Distal embolization occurred in three cases (12.0%) in the FAST group and in four (26.7%) in the stent retriever group ($p=0.362$). However, the two techniques did not differ significantly in the final TICI 2b–3 rate (72.0% vs 87.5%; $p=0.441$). A frequent angiographic finding regarding the failure of FAST was that the M2 occlusion was located immediately after severe acute angulation between M1 and M2.

Conclusions Stent retriever thrombectomy may provide faster reperfusion than FAST, while the FAST technique might be associated with lower distal embolization and a higher reperfusion rate for the first thrombectomy attempt, but without any significant difference in clinical outcome. When choosing the EVT method for M2 occlusions, consideration of the location of the occlusion and tortuosity between M1 and M2 might be helpful to achieve a better angiographic outcome.

INTRODUCTION

Recent trials regarding endovascular thrombectomy (EVT) in patients with acute ischemic stroke (AIS) due to proximal artery occlusion in the anterior circulation have demonstrated its benefits in terms of efficacy and safety when used with or without intravenous thrombolytic therapy.^{1–5} Based on data from these trials, updated guidelines on EVT for patients with AIS recommend using the stent retriever for occlusions of a proximal intracranial artery such as the distal internal carotid artery

(ICA) and proximal middle cerebral artery (MCA).^{6,7}

However, the risks and benefits of EVT for distal intracranial arterial occlusions (ie, M2 or M3 segments of the MCA) still remain debatable. Theoretically, the size of the ischemic core and ischemic penumbra for a distal intracranial occlusion is less than that of a proximal intracranial arterial occlusion; this difference can lessen the beneficial effects of EVT for M2 occlusions, while the thin vessel wall and narrow lumen of distal intracranial arteries can be associated with a higher risk of EVT complications.

Despite the lack of clarity regarding its benefits, the use of EVT with stent retrievers is recommended by the updated American Heart Association/American Stroke Association Korean Clinical Practice guidelines in carefully selected patients with AIS due to M2 segment occlusion.^{6,7} A few studies have retrospectively investigated the safety and efficacy of EVT in patients with M2 occlusions.^{8–11} These studies have examined the clinical outcome following EVT of M2 occlusions and have mostly used stent retriever thrombectomy as the EVT method.

To date there has been no direct comparison of two frequently used EVT methods (stent retriever and aspiration thrombectomy using a large-bore catheter) in patients with M2 occlusions. We therefore retrospectively analyzed the data of 41 patients with AIS due to occlusion of the M2 segment of the MCA who underwent EVT using a stent retriever or forced arterial suction thrombectomy (FAST) with the Penumbra reperfusion catheter. We compared the safety and efficacy of the two EVT methods in terms of technical characteristics. In addition, we aimed to identify predictors of a favorable clinical outcome after EVT in M2 occlusions.

MATERIALS AND METHODS

Patients

The subjects for this study were chosen from patients with acute stroke treated at our institute between January 2011 and June 2015 as per the prospectively recorded acute stroke endovascular treatment registry maintained at our center. The criteria for inclusion in the study were as follows: (1) Patients had AIS with angiographically confirmed M2 segment occlusion and National Institute of Health Stroke Scale (NIHSS) score of ≥ 6 at the time of arrival. The M2 segment was defined as part of the MCA extending from the



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genu to the top of the Sylvian fissure and the circular sulcus.¹²
 (2) Time from symptom onset to hospital arrival was <8 hours.
 (3) EVT was performed using the FAST technique with the Penumbra reperfusion catheter or the stent retriever technique. The study protocol and study design of retrospective analysis was approved by the local institutional board.

Acute stroke management and endovascular procedure

The initial neurological examination, including the assessment of the NIHSS score, was performed in the emergency room. Non-enhanced brain CT scans were performed immediately to rule out intracerebral hemorrhage (ICH). If no ICH was evident on CT scans, intravenous recombinant tissue plasminogen activator (rtPA; Actilyse; Boehringer Ingelheim, Basel, Switzerland) was administered in accordance with the guidelines for early management (within 3 hours of symptom onset up to December 2012 and within 4.5 hours since January 2013) of adults with ischemic stroke.¹³ Vascular imaging (CT angiography or MR angiography) and diffusion-weighted MRI were conducted subsequently. The institutional criteria for EVT were as follows: (1) AIS corresponding to arterial occlusion on CT angiography or MR angiography; (2) infarction volume on CT or diffusion-weighted MRI less than half of the corresponding vascular territory; and (3) no evidence of well-developed parenchymal hyperintensity on FLAIR or T2-weighted imaging in patients >6 hours from symptom onset.

Informed consent was obtained before initiating EVT. All interventional procedures were performed by three experienced neurointerventionalists (Y-WK, D-HK, and Y-SK) using a biplane angiography system. Transfemoral cerebral angiography was performed under conscious sedation. EVT was initiated after the assessment of the angioarchitecture of the occluded artery and collateral status. A 9 Fr balloon-tipped guide catheter (BGC; Optimo, Tokai Medical, Aichi, Japan) was advanced to the most distal portion of the accessible intracranial ICA.

The choice of EVT technique in each case was made according to the neurointerventionists' preferences, except that the stent retriever was used in cases when the diameter of M2 was smaller than the Penumbra reperfusion catheter.

For the FAST technique, the Penumbra reperfusion catheter (041 or 4Max; Penumbra, Alameda, California, USA) coaxially with a 0.014 inch microguidewire (Synchro-14; Stryker, Kalamazoo, Michigan, USA or Traxcess 14; Microvention, Columbia, USA) and a 0.021 inch microcatheter (Prowler Select Plus; Cordis, Miami, Florida, USA) were advanced to the proximal end of the occlusion and wedged to the clot under manual aspiration using a 50 mL syringe. After the balloon of the BGC was inflated, the Penumbra reperfusion catheter was withdrawn while continuing to maintain manual aspiration.¹⁴

For the stent retriever technique (Solitaire AB/FR; Covidien-Medtronic, Irvine, California, USA or TrevoProVue/XP; Stryker, Kalamazoo, Michigan, USA), the target occluded segment was navigated with a 0.014 inch microguidewire and a 0.021 inch microcatheter was advanced beyond the occlusion. The proximal and distal ends of the occlusion were then confirmed, and the stent retriever was deployed for several minutes. After the balloon of the BGC was inflated with remote aspiration through the guide catheter, the microcatheter and stent retriever were withdrawn into the guide catheter. If reperfusion was not achieved after three attempts of mechanical thrombectomy, we used rescue measures such as switching to another mechanical thrombectomy device¹⁵ and intra-arterial tirofiban infusion.¹⁶

Clinical and angiographic data collection

Data for the following parameters were collected from the patient records: demographics, clinical characteristics, complete medical history, and angiography findings. The NIHSS score was assessed at baseline and at day 7. The Alberta Stroke Program Early CT Score (ASPECTS) was calculated from initial diffusion-weighted image (DWI). The clinical outcome was assessed using the modified Rankin Scale (mRS) at 3 months, and a favorable clinical outcome was defined as a mRS score of ≤ 2 . The reperfusion status was assessed using the Thrombolysis In Cerebral Infarction (TICI) score, which was determined during the final angiography after thrombectomy. The TICI grade was independently evaluated by an experienced neuroradiologist (Y-SK) and neurologist (Y-HH) who were blinded to the patients' clinical information. If there was a discordance, the final assessment was determined by consensus. Successful reperfusion was defined as a TICI score of 2b or 3.

A control CT scan was acquired immediately after the EVT and 24 hours after the procedure to check for hemorrhagic complications. The type of intracranial hemorrhage was determined according to the European Cooperative Acute Stroke Study II classification system.¹⁷ Symptomatic intracranial hemorrhage was defined as any type of hemorrhage with an increase of ≥ 4 in NIHSS score within 24 hours.¹⁸

Statistical analysis

Statistical analyses were performed using SPSS V22.0 (IBM, Armonk, New York, USA). A χ^2 test or Fisher exact test was used for categorical data. The Student t-test or Mann-Whitney U test was used for continuous variables. Multiple linear regression analysis was performed to identify any independent predictor(s) of favorable clinical outcome at 3 months. Age, DWI-ASPECTS, initial NIHSS score, use of intravenous rtPA, EVT device, time from symptom onset to reperfusion, and reperfusion of TICI 2b–3 were included as potential confounders. A p value of <0.05 was considered statistically significant.

RESULTS

Overall results

The median age of the 41 enrolled patients was 72 years (IQR 64–79) and the median NIHSS score was 13 (IQR 8–17). Twenty-six patients had left MCA M2 occlusions and 20 patients (48.8%) received intravenous rtPA prior to EVT. The median time from symptom onset to door was 142 min and the median time from door to groin puncture was 99 min. Baseline characteristics are summarized in [table 1](#).

The first attempt at EVT was made with the Penumbra reperfusion catheter via the FAST technique in 25 patients (61.0%) and with the stent retriever technique in the remaining 16 patients (39.0%). For 70.7% (29/41) of the patients, successful reperfusion (TICI 2b–3) was achieved using the first choice device (primary reperfusion), while 17.1% (7/41) of the patients required rescue measures. The median time from groin puncture to reperfusion was 45 min. The successful reperfusion rate, as per the final angiography, was 78.0% and one patient (2.4%) developed symptomatic ICH. The median NIHSS score at 7 days after stroke onset was 2 (IQR 1–6), and a favorable clinical outcome at 3 months was achieved in 33 patients (80.5%).

Technical comparison between the FAST and stent retriever techniques

No statistically significant differences were noted between the FAST technique and the stent retriever technique in terms of

Table 1 Baseline characteristics

Variable	FAST	Stent retriever	p Value
Number	25	16	
Age, median (IQR)	71 (60.5–75)	76.5 (65.5–82)	0.121*
Female	8 (32%)	9 (56.2%)	0.124
NIHSS, median (IQR)	15 (8–17)	10.5 (6–17)	0.511*
NIHSS at 7 days, median (IQR)	2 (1–6)	1.5 (0.25–5.75)	0.465*
DWI-ASPECTS score	8 (7.5–9)	7 (6.75–9.0)	0.230*
Lesion side (left)	14 (56%)	12 (75%)	0.218
Intravenous tPA	13 (52%)	7 (43.8%)	0.606
Hypertension	15 (60%)	11 (68.8%)	0.570
Diabetes	5 (20%)	3 (18.8%)	>0.999†
Dyslipidemia	6 (24%)	7 (43.8%)	0.185
Atrial fibrillation	15 (60%)	12 (75%)	0.323
Coronary disease	2 (8%)	2 (12.5%)	0.637†
Previous stroke	3 (12.0%)	2 (12.5%)	>0.999†
Smoking	7 (28.0%)	3 (18.8%)	0.712†
Onset to door time, median (IQR)	137 (47–307)	143.5 (77.5–299.25)	0.897*

*Mann-Whitney U test.

†Fisher's exact t-test.

ASPECTS, Alberta Stroke Program Early CT Score; DWI, diffusion-weighted image; FAST, forced arterial suction thrombectomy; NIHSS, National Institute of Health Stroke Scale; tPA, tissue plasminogen activator.

the baseline characteristics. EVT device delivery time from guide catheter to target occlusion did not differ significantly between the two methods (4 min for FAST vs 3 min for the stent retriever, $p=0.120$). Both EVT techniques had a similar rate of successful reperfusion at the first thrombectomy attempt (56.0% vs 56.2%, $p=0.987$) and with the primary chosen method (64.0% vs 81.2%, $p=0.305$). The rate of successful reperfusion in less than 30 min from carotid catheterization to reperfusion was not significantly different (64.0% vs 75.0%, $p=0.460$). Five (19.2%) of the 26 patients in the FAST group required rescue treatment, including device switching to the stent retriever with intra-arterial tirofiban infusion, and two patients (12.5%) required conversion from the stent retriever technique to the FAST technique. The time from groin puncture to reperfusion was significantly shorter using the stent retriever technique (median 53 vs 38.5 min; $p=0.045$). However, the two techniques did not show a statistically significant difference in the final reperfusion rate (72.0% vs 87.5%; $p=0.441$) and a favorable clinical outcome at 3 months (84.0% vs 75.0%; $p=0.689$). There was no embolization of uninvolved territory in either group, but distal embolization occurred in three cases (12.0%) in the FAST group and in four cases (26.7%) in the stent retriever group ($p=0.362$). Symptomatic ICH occurred in one patient (4.0%) treated with the FAST technique. Angiographic data and clinical outcomes are summarized in [table 2](#). Multiple linear regression analysis showed that age

Table 2 Details of endovascular treatment and clinical outcomes

Variable	FAST	Stent retriever	p Value
Onset to door time, median (IQR)	137 (47–307)	143.5 (77.5–299.25)	0.897*
Door to puncture time, median (IQR)	99 (82.5–120.5)	98 (66.5–123.25)	0.626*
Puncture to recanalization time, median (IQR)	53 (41–74)	38.5 (31.5–57.25)	0.045*
Time intervals, median (IQR)			
Puncture to carotid catheterization	24 (16–35)	20.5 (15.25–23)	0.080*
Guide catheter to access to the target occlusion	4 (2.5–7.5)	3 (2–4)	0.120*
Start of thrombectomy to recanalization	16 (7–42.5)	15.5 (12.25–33.5)	0.678*
TICI 2b–3 by first thrombectomy attempt	14 (56.0%)	9 (56.2%)	0.987
TICI 2b–3 by primary chosen device	16 (64.0%)	13 (81.2%)	0.305†
TICI 2b–3 <60 min (puncture to recanalization)	15 (60.0%)	12 (75.0%)	0.323
TICI 2b–3 <30 min (carotid catheterization to recanalization)	16 (64.0%)	12 (75.0%)	0.460
Rescue treatments used	5 (20.0%)	2 (12.5%)	0.685†
Stent retriever	5	0	
FAST	0	2	
Intra-arterial tirofiban	5	0	
Final TICI 2b–3	18 (72.0%)	14 (87.5%)	0.441†
Distal embolization	3 (12.0%)	4 (26.7%)	0.392†
Uninvolved territory embolization	0	0	
Intracranial hemorrhage			
Hemorrhagic infarction 1	3 (12.0%)	2 (12.5%)	1.000†
Hemorrhagic infarction 2	0	1 (6.2%)	0.390†
Parenchymal hematoma 1	0	0	
Parenchymal hematoma 2	1 (4.0%)	0	1.000†
Symptomatic ICH	1 (4.0%)	0	1.000†
SAH	0	1 (6.2%)	0.390†
mRS 0–2 at 3 months	21 (84.0%)	12 (75.0%)	0.689†
Mortality	0	1 (6.2%)	0.390†

*Mann-Whitney U test.

†Fisher's exact t-test.

FAST, forced arterial suction thrombectomy; ICH, intracerebral hemorrhage; mRS, modified Rankin Scale; SAH, subarachnoid hemorrhage; TICI, Thrombolysis in Cerebral Infarction.

Table 3 Multiple linear regression analysis for favorable clinical outcome at 3 months

Variables	Unstandardized coefficients		Standardized coefficients	p Value	95% CI	
	B	S.E.	β		Lower	Upper
Age (/1 year)	-0.016	0.006	-0.463	0.009	-0.028	-0.004
DWI-ASPECTS	0.062	0.046	0.233	0.187	-0.032	0.157
Initial NIHSS	-0.002	0.012	-0.028	0.875	-0.027	0.023
Intravenous tPA	-0.083	0.145	-0.105	0.572	-0.380	0.214
EVT strategy	-0.043	0.148	-0.050	0.772	-0.346	0.260
Onset to reperfusion time(/10 min)	-0.005	0.002	-0.414	0.027	-0.010	-0.001
Successful reperfusion	0.092	0.143	0.096	0.528	-0.202	0.385

ASPECTS, Alberta Stroke Program Early CT Score; DWI, diffusion-weighted image; EVT, endovascular thrombectomy; NIHSS, National Institute of Health Stroke Scale; tPA, tissue plasminogen activator.

($\beta = -0.463$, $p = 0.009$) and time from symptom onset to reperfusion ($\beta = -0.414$, $p = 0.027$) were associated with a favorable clinical outcome at 3 months (table 3).

DISCUSSION

In this study the rates of successful reperfusion and favorable outcome following EVT for patients with an M2 occlusion were 78.6% and 81.0%, respectively, with a mortality rate of 2.4%. These results are comparable with those reported in recent retrospective studies on EVT performed using only the stent retriever technique for M2 segment occlusions.^{8–10} In this study we did not find any significant difference between the two evaluated techniques in terms of reperfusion rate or rate of favorable clinical outcome. However, time from groin puncture to reperfusion was significantly shorter using the stent retriever technique than with FAST. The FAST technique had a lower rate of distal embolization than the stent retriever technique, although the difference was not statistically significant.

Our study design did not allow for the assessment of the superiority of either EVT technique because there was no control group receiving medical treatment alone or treatment using other EVT techniques. To overcome this deficiency, we reviewed historical data involving various EVT strategies to treat M2 occlusions. One study on the natural history of acute stroke indicated a poor clinical outcome (mRS score ≥ 3) at 3 months in up to 47% of patients with a distal MCA occlusion not administered reperfusion therapy.¹⁹ Furthermore, studies on patients with distal MCA occlusions receiving intravenous thrombolytic agents found reperfusion rates of 30.8–68.4% and favorable clinical outcomes ranging from 48% to 81%.^{20–22} Finally, a recent study on EVT using the stent retriever technique reported

reperfusion rates of 78.5–93.3% and a favorable clinical outcome in 60–66.3% of M2 occlusion cases (table 4).^{8–10} These results are comparable with the overall result of the present study, which yielded a final successful reperfusion rate of 78.6% and a favorable clinical outcome in 81.0%.

For comparison of the technical characteristics of the two EVT techniques, we analyzed key time intervals of EVT in terms of puncture to carotid catheterization time, guide catheter to target access time, and target access to reperfusion time. With regard to the delivery of the thrombectomy device, it is known that the Penumbra reperfusion catheter is more difficult than the stent retriever because of its larger size compared with the microcatheter in stent retriever thrombectomy.²³ The tortuosity of the carotid siphon may impede the delivery of the Penumbra catheter to the occlusion site using the FAST technique, as was the case with the origin of the ophthalmic artery. However, our results showed no significant difference in time from guide catheter to target access. The Penumbra reperfusion catheter itself has a highly navigable flexible tip, and it could help to overcome such impedance with the use of a coaxial system (0.021 inch microcatheter and 0.014 inch microwire) and steam shaping of the catheter tip. In terms of the reperfusion process, time from the start of thrombectomy to final reperfusion was similar for both EVT techniques. Among the cases with successful reperfusion, the FAST technique achieved a higher reperfusion rate with first thrombectomy attempts than the stent retriever (77.8% vs 64.3%), although it had a lower final successful reperfusion rate. This finding suggests that the FAST technique might be less effective for repeated thrombectomy attempts. Therefore, early switching to other techniques might provide better angiographic outcomes in cases with unsuccessful reperfusion after a few attempts with the FAST technique.

Based on the review of angiographic data, we found that successful or unsuccessful reperfusion using the FAST technique was associated with the location of the thrombus and the tortuosity between M1 and M2. If the thrombus was located immediately after severe acute angulation, especially interposition from M1 to M2, it would disturb the optimal contact of the Penumbra catheter tip and the thrombus. Eventually, the FAST technique might be ineffective due to imprecise facing with the thrombus and Penumbra catheter tip. In these cases, we assume that it might be reasonable to use the stent retriever instead of the Penumbra reperfusion catheter.

We used a balloon guide catheter in both groups. There was no embolization of uninvolved territory in either group, but distal embolization occurred in 12% of cases in the FAST group and in 26.7% of cases in the stent retriever group. For our FAST technique, we did not cross the clot using a microcatheter and microguidewire so fragmentation or disruption of the clot may rarely occur (figure 1). We also believe that the small clot size in an M2 occlusion, which could be easily removed by

Table 4 Characteristics of reperfusion therapy studies for M2 occlusions

Studies	Age	Female (%)	Baseline NIHSS	Reperfusion method	Reperfusion status	mRS ≤ 2 at 3 months (%)
Flores <i>et al</i> ¹⁰ (n=65)	66 (mean)	47.7	16 (median)	Stent retriever \pm IA-tPA \pm IV-tPA	78.5% (TICI 2b–3)	60
Dorn <i>et al</i> ⁹ (n=16)	68.1 (mean)	N/A	13.7 (mean)	Stent retriever \pm IV-tPA	93.3% (TICI 2b–3)	60
Coutinho <i>et al</i> ⁸ (n=50)	71 (mean)	48	13 (median)	MERCI, Solitaire \pm IV-tPA	85% (mTICI 2b–3)	60
Current study (n=41)	72 (median)	41.5	13 (median)	FAST, Stent retriever \pm IV-tPA	78% (TICI 2b–3)	80.5

FAST, forced arterial suction thrombectomy; IA, intra-arterial; IV, intravenous; mRS, modified Rankin Scale; mTICI, modified thrombolysis in cerebral ischemia; NIHSS, National Institute of Health Stroke Scale; tPA, tissue plasminogen activator; TICI, Thrombolysis In Cerebral Infarction.

suction power via the Penumbra catheter, could also be a reason for less distal embolization in the FAST group.

In this study we noted a discrepancy in the results regarding the reperfusion rate and rate of favorable outcome between the two EVT techniques. Generally, a higher reperfusion rate is associated with better clinical outcomes.^{1–5} However, the rate of favorable clinical outcome was lower for the stent retriever technique than the FAST technique, despite a higher reperfusion rate. Various factors including arterial occlusion site, infarct volume, collateral status, use of rtPA, reperfusion status, time from symptom onset to reperfusion, and age have been established as predictors of clinical outcome in other studies.^{21–24–27} In this study, multivariate analysis showed that age and time from symptom onset to reperfusion were the independent predictors of favorable outcomes in EVT for M2 segment occlusion. Additionally, the absolute value of the β coefficient of age was greater than that of the time from symptom onset to reperfusion. We therefore speculate that age had the greatest impact on favorable clinical outcomes in our study, so the worse clinical outcomes in the stent retriever group—despite better reperfusion rates—can be explained by the fact that the patients in the stent retriever group had an older median age.

This study has some limitations. First, this was a retrospective single-center study although the data were collected prospectively, so there is a possibility of selection bias. Second, the sample size was small so the statistical power of the study might be weak. Third, there was no control group that did not undergo EVT, so we were not in a position to assess whether EVT should be done using FAST or the stent retriever technique.

CONCLUSIONS

Our findings suggest that EVT for M2 segment occlusion may be safe and feasible. EVT using stent retrievers was associated with more rapid reperfusion, while the FAST technique had a higher reperfusion rate for first thrombectomy attempts. There was no significant difference in terms of the clinical outcome between the two groups. In EVT for M2 occlusions, choosing the EVT technique based on the location of occlusion and the tortuosity between the M1 and M2 segments might result in better angiographic outcomes. Future prospective multicenter studies are necessary to further confirm our findings.

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