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Predictors of Functional Dependence Despite Successful Revascularization in Large-Vessel Occlusion Strokes

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Disclosures

Drs Liebeskind, Smith, and Duckwiler were employed by the University of California, which holds a patent on endovascular devices for stroke. Dr Ge is an employee of Stryker. Dr Liebeskind: consultant—Stryker and Covidien. Dr Albers: consultant—Covidien; equity interest—iSchemaView. Dr Budzik: consultant—Stryker and Covidien. Dr Gupta: consultant—Stryker, Covidien and Rapid Medical; royalties—UpToDate. Dr Jansen: consultant—Stryker and Covidien. Dr Jovin: consultant—Silk Road Medical. Dr Lutsep: consultant—Stryker. Dr Nogueira: consultant—Stryker (Trevo-2 Trial PI, DAWN Trial PI), Covidien (SWIFT and SWIFT-PRIME Trials Steering Committee, STAR Trial Core Laboratory) and Penumbra (3-D Separator Trial Executive Committee). Dr Rymer: honoraria and speaking engagements—Stryker and Covidien. Dr Smith: consultant—Stryker and Covidien.

Abstract

Background and Purpose—High revascularization rates in large-vessel occlusion strokes treated by mechanical thrombectomy are not always associated with good clinical outcomes. We evaluated predictors of functional dependence despite successful revascularization among patients with acute ischemic stroke treated with thrombectomy.

Methods—We analyzed the pooled data from the Multi Mechanical Embolus Removal in Cerebral Ischemia (MERCI), Thrombectomy Revascularization of Large Vessel Occlusions in Acute Ischemic Stroke (TREVO), and TREVO 2 trials. Successful revascularization was defined as thrombolysis in cerebral infarction score 2b or 3. Functional dependence was defined as a score of 3 to 6 on the modified Rankin Scale at 3 months. We assessed relationship of demographic, clinical, angiographic characteristics, and hemorrhage with functional dependence despite successful revascularization.

Results—Two hundred and twenty-eight patients with successful revascularization had clinical outcome follow-up. The rates of functional dependence with endovascular success were 48.6% for Trevo thrombectomy and 58.0% for Merci thrombectomy. Age (odds ratio, 1.04; 95% confidence interval, 1.02–1.06 per 1-year increase), National Institutes of Health Stroke Scale score (odds ratio, 1.08; 95% confidence interval, 1.02–1.15 per 1-point increase), and symptom onset to endovascular treatment time (odds ratio, 1.11; 95% confidence interval, 1.01–1.22 per 30-minute delay) were predictors of functional dependence despite successful revascularization. Symptom onset to reperfusion time beyond 5 hours was associated with functional dependence. All subjects with symptomatic intracranial hemorrhage had functional dependence.

Conclusions—One half of patients with successful mechanical thrombectomy do not have good outcomes. Age, severe neurological deficits, and delayed endovascular treatment were associated with functional dependence despite successful revascularization. Our data support efforts to minimize delays to endovascular therapy in patients with acute ischemic stroke to improve outcomes.

Keywords

stroke

Although intravenous (IV) tissue-type plasminogen activator (tPA) is the recommended treatment for eligible patients with acute ischemic stroke within 4.5 hours after symptom onset, the recanalization rate is low for patients with large intracranial vessel occlusions. Nonresponse to IV tPA is associated with poor clinical outcomes in patients with moderate-to-severe stroke.^{1,2} Endovascular intervention with intra-arterial (IA) thrombolysis or mechanical thrombectomy offers an alternative treatment for patients with large-vessel occlusion strokes who are ineligible for or refractory to IV tPA 8 hours after symptom onset. Greater than 80% revascularization rates can be achieved with mechanical thrombectomy, particularly with stent retrievers.^{3–8} Although studies have reported a strong association between better outcomes and successful revascularization, especially when tissue-level reperfusion is considered,^{4–6,9} and randomized trials between devices support better outcomes in the cohorts treated with more efficacious devices,^{6,7} the high revascularization rates with mechanical thrombectomy have not translated into better

outcome in randomized trials against IV tPA or in combination with IV tPA.^{10,11} The lack of clinical benefit with endovascular treatment in 2 recent trials may be related to a substantially delayed time to endovascular therapy and less use of contemporary technologies such as stent retriever.^{10,11} Even in the stent retriever studies, no more than 60% of the patients had an independent neurological outcome.^{6–8,12}

The factors leading to functional dependence despite successful revascularization remain unknown. Ideally, focusing the endovascular efforts to those patients who are not already destined to poor outcomes would improve the efficacy of treatment. We pooled data from the 3 prospective mechanical thrombectomy trials for acute ischemic stroke, namely the Multi Mechanical Embolus Removal in Cerebral Ischemia (MERCI), Thrombectomy Revascularization of Large Vessel Occlusions in Acute Ischemic Stroke (TREVO), and TREVO 2 trials, to identify the clinical features associated with functional dependence despite successful revascularization.

Methods

Trial Inclusion and Patients

We analyzed data from the Multi MERCI, TREVO, and TREVO 2 trials, in which the Merci Retriever (Stryker Neurovascular, Mountain View, CA) and the Trevo Retriever (Stryker Neurovascular) were used.^{4,6,8} The databases of all 3 trials were maintained at Stryker Neurovascular. Appropriate institutional review board or ethics committees approvals were obtained by participating centers in all 3 trials. All 3 trials enrolled patients with acute ischemic stroke who were either ineligible for or refractory to IV tPA. Endovascular thrombectomy therapy was initiated within 8 hours of symptom onset.

All 3 trials included patients with angiographically confirmed intracranial vessel occlusion in both anterior and posterior circulations. All patients were older than 18 years with National Institutes of Health Stroke Scale (NIHSS) score of greater than 8. The TREVO and TREVO 2 trials had age limit of 85 years and NIHSS score upper limit of 30 and 29, respectively. In the Multi MERCI trial, IV tPA was allowed within 3 hours after symptom onset. A 4.5-hour time window was used for IV tPA in the TREVO and TREVO 2 trials.

Procedures

Details of the trial protocol and primary results of the 3 trials have been reported previously.^{4,6,8} Different revascularization scores were used for these trials. For our pooled analysis, digital subtraction angiography of subjects in the Multi MERCI trial was reevaluated by the imaging core laboratory to derive thrombolysis in cerebral infarction (TICI) score.⁶ Successful revascularization was redefined as TICI 2b (major partial reperfusion of two thirds or more of the vascular distribution of the occluded artery) or 3 reperfusion flow in the target territory documented on the final angiogram after endovascular treatment.

Functional dependence was defined as a score of 3 to 6 on the modified Rankin Scale at 90 days. Functional independence was defined as a score of 0 to 2 on the modified Rankin Scale at 90 days.

Statistical Analysis

We compared patient demographic, angiographic characteristics, intracranial hemorrhage, and clinical outcome among the 3 trials. We analyzed continuous variables with the Wilcoxon rank-sum test or 2-sample *t* test and categorical variables with the Fisher's exact test. The calculation of odds ratio (OR) and 95% confidence intervals (CI) was assessed in all tests.

The key management times were documented in all 3 trials. We defined time from stroke onset to procedure termination as onset to reperfusion time in subjects with successful revascularization. We tested time interval variables as continuous and categorical data for the analysis, including symptom onset to reperfusion time, onset to groin puncture, groin puncture to first device pass, onset to first device pass, and groin puncture to reperfusion time. We determined the cutoff points at different values and dichotomizations for variables based on clinical judgment and previous literature. Full details about variables are provided in the Methods in the online-only Data Supplement.

The association of patient characteristics with functional dependence despite successful revascularization was analyzed with univariate and multivariate logistic regression models. The prespecified variables as potential predictors of functional dependence despite successful revascularization included age, baseline NIHSS score, sex, internal carotid artery occlusion, onset to reperfusion time, onset to groin puncture time, onset to first device pass time, groin puncture to reperfusion time, and number of pass with study device. A separate multivariate logistic regression analysis was performed, using the stratified continuous variables of age, NIHSS score, and onset to reperfusion time. We analyzed the relationship of hemorrhage and device-related serious adverse events with functional dependence despite successful revascularization. We further assessed the association of age, stroke severity, and time intervals with functional dependence despite successful revascularization by categorization into quartiles or tertiles. We used SAS statistical software, version 9.2.

Results

Trial participants were enrolled from 43 centers in 6 countries between January 2004 and December 2011. Among 402 subjects enrolled in the 3 trials, 235 subjects achieved successful revascularization with a TICI score of 2b or 3. The successful revascularization rate was higher in the primary Trevo thrombectomy than the primary Merci thrombectomy (75.0% versus 48.8%; $P < 0.0001$). A total of 228 subjects were included in outcome analysis. The pooled analysis population consisted of 73 subjects from Multi MERCI trial, 47 from TREVO trial, and 108 from TREVO 2 trial. The study profile is shown in Figure 1.

Baseline and angiographic characteristics of 109 subjects treated primarily with Trevo devices and 119 subjects with Merci devices are shown in Table I in the online-only Data Supplement. Intracranial hemorrhage, device-related serious adverse events, and 90-day mortality were similar between treatment groups.

Functional dependence despite successful revascularization was observed in 122 of 228 (53.5%) subjects. The rates of functional dependence with endovascular success were 48.6%

for Trevo thrombectomy and 58.0% for Merci thrombectomy. Baseline, angiographic characteristics, and hemorrhage between subjects with functional dependence and those with functional independence are shown in Table 1. In the entire cohort and Trevo thrombectomy group, subjects with functional dependence were older, had higher baseline NIHSS score, had higher systolic blood pressures and blood glucose, and more often had a cardioembolic stroke source and comorbidities (most common with hypertension and diabetes mellitus) than those with functional independence. In both Trevo and Merci thrombectomy groups, delay in revascularization contributes to functional dependence (Table 1 and Table II in the online-only Data Supplement). Symptom onset to groin puncture time, onset to first device pass, and onset to reperfusion time were all longer in subjects with functional dependence than functional independence. In the entire cohort, symptom onset to reperfusion time beyond 5 hours is associated with functional dependence (OR, 1.83; 95% CI, 1.07–3.12; $P=0.0306$). The same association is found in the Trevo group as well (OR, 2.48; 95% CI, 1.13–5.45; $P=0.0319$). In the entire cohort, functional dependence is related to intracranial hemorrhage and device-related serious adverse events, but not associated with number of device pass, rescue therapy, and location of vessel occlusion.

Factors independently associated with functional dependence despite successful revascularization are shown in Table 2. The risks of functional dependence increases with each year of age (OR, 1.04; 95% CI, 1.02–1.06), each point of NIHSS score (OR, 1.08; 95% CI, 1.02–1.15), and each 30-minute delayed symptom onset to groin puncture for endovascular treatment (OR, 1.11; 95% CI, 1.01–1.22) in the entire cohort. When dichotomized variables of age, NIHSS score, and symptom onset to reperfusion time were selected into the multivariate model, predictors of functional dependence in the entire cohort included NIHSS score ≥ 20 (OR, 2.31; 95% CI, 1.33–4.02; $P=0.003$) and onset to reperfusion >5 hours (OR, 1.83; 95% CI, 1.06–3.17; $P=0.03$). Predictors of functional dependence in the Trevo cohort included age >80 years (OR, 7.35; 95% CI, 1.31–41.12; $P=0.02$), NIHSS score ≥ 20 (OR, 2.66; 95% CI, 1.14–6.24; $P=0.02$), onset to reperfusion >5 hours (OR, 2.70; 95% CI, 1.14–6.39; $P=0.02$), and men (OR, 0.38; 95% CI, 0.16–0.90; $P=0.03$). There was no predictor of functional dependence in the Merci group. We found similar predictors of functional dependence when subjects with prestroke modified Rankin Scale ≥ 2 were excluded from the analysis.

The association of age, NIHSS score, and time intervals with functional dependence despite successful revascularization are shown in Tables 3 and 4. In the entire cohort, the ORs of functional dependence for every 30-minute delay in symptom onset to groin puncture time, onset to first device pass time, and onset to reperfusion time were 1.12 (95% CI, 1.02–1.24), 1.10 (95% CI, 1.01–1.21), 1.11 (95% CI, 1.01–1.21), respectively (Table 3). In the Trevo thrombectomy, every 10-year age increase is associated with a 92% relative increase in the odds of functional dependence. Every 5-point baseline NIHSS score increase is associated with a 78% relative increase in the odds of functional dependence after Trevo thrombectomy (Table 4). All subjects with symptomatic intracranial hemorrhage or severe parenchymal hematoma had functional dependence in both Trevo thrombectomy and Merci thrombectomy groups (Table III in the online-only Data Supplement).

Discussion

Our study shows that half of the patients with large intracranial vessel occlusion had functional dependence despite successful revascularization in the 3 endovascular thrombectomy trials. Older age, higher NIHSS score, delayed endovascular treatment, and procedural complications are associated with increased frequency of functional dependence despite endovascular success.

Our analysis confirms the superiority of the stent retriever over the Merci device for successful revascularization using TICI 2b or greater as the threshold. TICI and modified TICI scales are superior to TIMI scale for evaluating the extent of tissue reperfusion and predicting clinical outcome after IA therapy.^{9,10,13} The 75% reperfusion rate (TICI 2b) with Trevo devices in our study is higher than the 40% rate (modified TICI 2b) from IMS III trial, in which almost half of subjects in the endovascular group were treated only with IA tPA and seldom received the stent retriever.¹⁰ The variant thresholds for major partial perfusion (ie, 2/3 versus 1/2) may result in different rates of successful reperfusion, with about 20% of subjects with modified TICI 2b (ie, partial perfusion with 50%-66% of the target downstream territory) were not graded as TICI 2b.¹³ The discrepancy of successful TICI 2b or greater reperfusion with good functional outcome may suggest that even 66% territorial reperfusion may be suboptimal.

Advanced age is associated with functional dependence despite endovascular success. The proportion of functional dependence increased from only 28% in people aged ≤ 60 years to 82% in those aged >80 years after Trevo thrombectomy. The association of advanced age with functional dependence is consistent with the results in 2 studies of endovascular intervention with or without stent retriever.^{14,15} However, our finding of less favorable outcomes with older age (>80 years) has not been shown in a meta-analysis of IV tPA trials.¹⁶

The association of severe stroke (NIHSS score ≥ 20) with functional dependence despite endovascular success in our study is supported by a multicenter study of endovascular treatment before stent retriever.¹⁵ In our study, stent retriever thrombectomy and Merci thrombectomy had similar rates of functional dependence in this subset of severe stroke. In contrast, results from the SYNTHESIS trial indicated that endovascular treatment was not superior to IV tPA for acute stroke with the median NIHSS score 13.¹¹ This finding differs from a recent study which showed that strokes with NIHSS score ≥ 14 benefit most from endovascular therapy.¹⁷ These conflicting results may be related to the infrequent use of mechanical thrombectomy (33.3% versus 88%) and more than one third stroke with NIHSS score ≥ 10 in the SYNTHESIS trial.¹¹ This subset of patients with minor to moderate stroke does not benefit from IA therapy, as shown in a previous randomized trial.³

Age and NIHSS score have been found as outcome predictors for endovascular interventions with IA thrombolysis and mechanical thrombectomy.^{18,19} Despite this, IMS III trial results suggest that endovascular interventions, including the Merci Retriever, trend toward benefit over IV tPA alone for severe stroke with NIHSS ≥ 20 ,¹⁰ probably because these patients do

even worse with IV tPA or medical therapy. Advanced age and severe stroke should not be used as exclusion criteria for stent retriever thrombectomy.

Our pooled analysis provides new information about time-dependent benefits of mechanical thrombectomy on acute ischemic stroke. In our study, a 30-minute delay from stroke onset to endovascular treatment was associated with a 11% increase in the odds of functional dependence. Time delay from symptom onset to first device pass also trends toward functional dependence. These findings are consistent with prior studies of IV tPA in randomized trials and clinical practice that showed shorten delay in initiation of thrombolytic treatment to be associated with both increased benefit and reduced mortality.^{16,20} In the IMS III and SYNTHESIS trials, delayed onset to endovascular treatment may have contributed to the futility of endovascular therapy.^{10,11}

In addition, our study shows that symptom onset to reperfusion time beyond 5 hours is associated with functional dependence despite endovascular success after mechanical thrombectomy, especially with Trevo Retriever. Moreover, this relationship occurs only in moderate-to-severe stroke with NIHSS 8 to 19 but not in critical stroke with NIHSS ≥ 20 . These findings are consistent with prior studies of IA therapies, which revealed the association of onset to reperfusion time with favorable clinical outcome and survival.^{21,22} Reducing delay from image to groin puncture time for IA therapy is likely associated with improved clinical outcome.²³ Accordingly, our data suggest that clinical effectiveness with mechanical thrombectomy for acute ischemic stroke is critically time dependent, similar to IV tPA and IA thrombolysis.

Although groin puncture to reperfusion time (ie, procedure time in our study) is not associated with functional dependence, faster procedure time and shorter treatment delays in a recent multicenter study with stent retriever than those observed in our study can lead to better clinical outcomes.¹² These findings from our study emphasize the importance of minimizing delays of onset to treatment and onset to reperfusion times with mechanical thrombectomy for achieving the best clinical outcomes. Reduction time from symptom onset to reperfusion within 5 hours with stent retriever may improve favorable outcome.

This study has several limitations. Data with a small cohort size were retrospectively collected from 2 single-arm prospective trials and a randomized controlled trial with a post hoc analysis. We cannot exclude the possibility that unmeasured confounding variables may influence some of our findings. Physiological determinants of outcome including blood pressure and glucose as well as medical history were not analyzed in the main multivariate model because these variables were not collected in the TREVO trial. However, the associations of diabetes mellitus, hypertension, and atrial fibrillation with functional dependence despite endovascular success after mechanical thrombectomy were also found in model excluding the TREVO trial data. A recent study suggests that good collateral flow may correlate with favorable outcomes after stent retriever thrombectomy.¹² Early ischemic change measured by the Alberta Stroke Program Early CT Score (ASPECTS) may predict the clinical outcome and reperfusion after endovascular treatment.⁵ However, favorable baseline ASPECTS in prediction for benefit from endovascular therapy was not shown in the IMS III trial, which had a significant delay between baseline computed tomography scan

and reperfusion.²⁴ The associations of collateral flow and ASPECTS with functional dependence despite endovascular success after mechanical thrombectomy have not been investigated in this analysis. The impact of imaging selection with modalities such as multimodal computed tomography or MRI on functional dependence despite successful revascularization also remains unknown.

Conclusions

Our findings show functional dependence despite successful revascularization is relatively frequent in subjects with large-vessel occlusion strokes after endovascular thrombectomy treatment, particularly among old patients with severe neurological deficits and delayed endovascular treatment. A 30-minute delay from stroke onset to endovascular treatment is associated with a 11% increase in the odds of functional dependence. Symptom onset to reperfusion time beyond 5 hours is associated with functional dependence despite endovascular success after mechanical thrombectomy. Our data support minimizing delays to reperfusion in randomized controlled trials of mechanical thrombectomy with stent retriever alone or as an adjunctive therapy against IV tPA alone.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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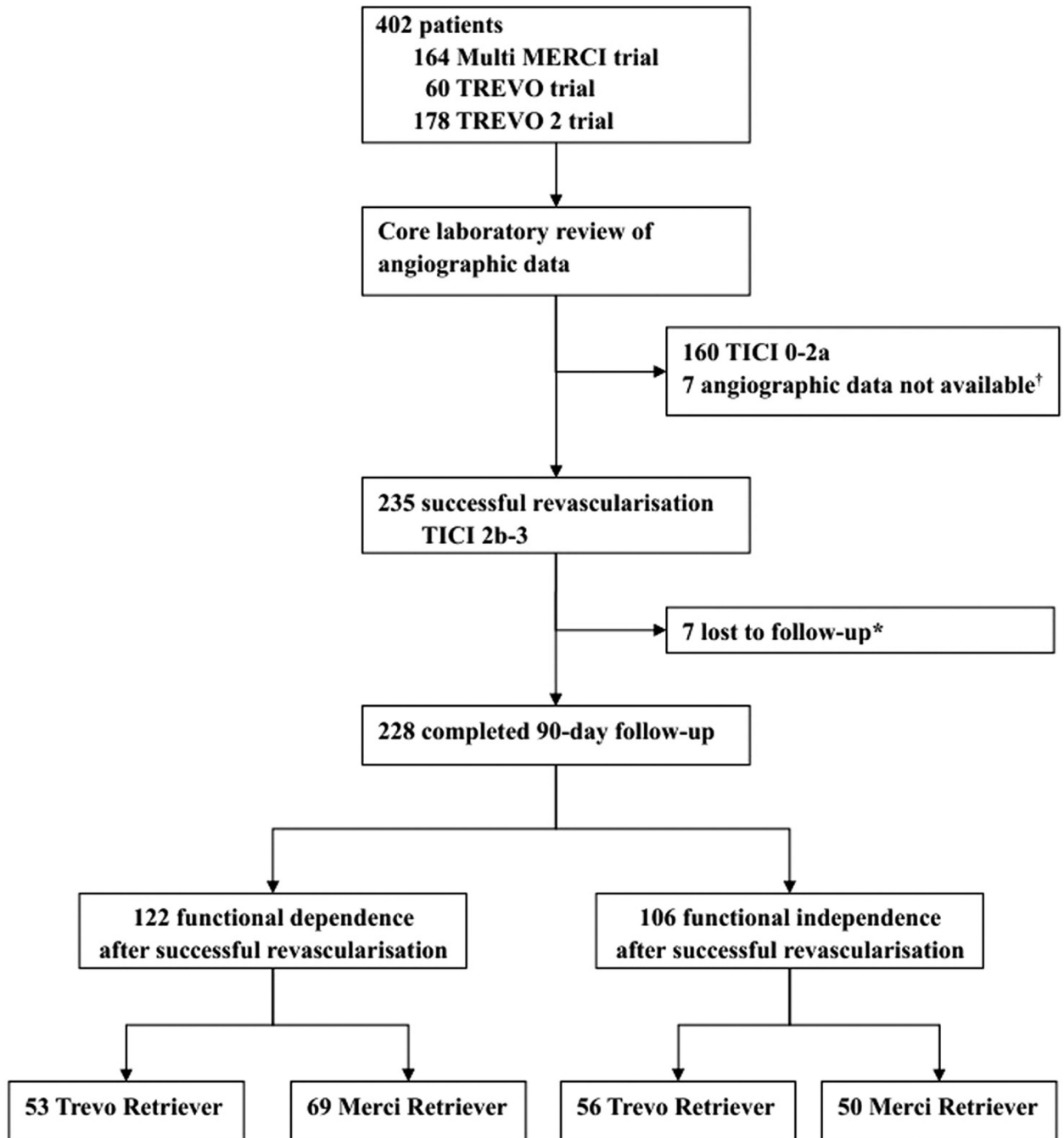


Figure. Study profile

TICI indicates thrombolysis in cerebral infarction. *Modified Rankin scores were not obtained for 7 patients at 90 days. †Angio-graphic images forwarded to the core laboratory were insufficient to allow TICI outcome assessment in 5 patients in Multi Mechanical Embolus Removal in Cerebral Ischemia (MERCI) trial. Technical reasons precluded the transfer of the angiographic images to the core laboratory in 2 patients in Thrombectomy Revascularization of Large Vessel Occlusions in Acute Ischemic Stroke (TREVO) trial.

Table 1
 Functional Dependence Despite Successful Revascularization With Trevo Retriever and Merci Retriever

Characteristics	All Patients (n=228)			Trevo Group (n=109)			Merci Group (n=119)		
	mRS 0-2 (n=106)	mRS 3-6 (n=122)	P Value	mRS 0-2 (n=56)	mRS 3-6 (n=53)	P Value	mRS 0-2 (n=50)	mRS 3-6 (n=69)	P Value
Age, y, mean (SD)	62 (16)	69 (13)	0.0003	62 (15)	71 (10)	0.0007	62 (18)	68 (15)	0.0661
Male sex	42/106 (40%)	57/122 (47%)	0.2878	20/56 (36%)	29/53 (55%)	0.0555	22/50 (44%)	28/69 (41%)	0.7117
White race	65/79 (82%)	85/102 (83%)	0.8455	23/29 (79%)	29/33 (88%)	0.4932	42/50 (84%)	56/69 (81%)	0.8092
NIHSS score, median	17 (14-20)	20 (16-22)	0.0002	16 (12-20)	20 (16-22)	0.0048	17 (15-21)	19 (17-22)	0.0268
Cardioembolic stroke source	37/70 (53%)	63/85 (74%)	0.0071	29/56 (52%)	41/53 (77%)	0.0089	8/14 (57%)	22/32 (69%)	0.5115
Systolic blood pressure, mmHg, median	138 (120-156)	153 (128-170)	0.0033	142 (130-160)	160 (140-177)	0.0266	131 (119-155)	150 (122-168)	0.0284
Diastolic blood pressure, mm Hg, median	77 (64-85)	76 (66-90)	0.5298	84 (72-91)	80 (74-92)	0.9888	72 (62-80)	72 (63-84)	0.2317
Blood glucose, mg/dL, mean (SD)	118 (30)	156 (70)	<0.0001	113 (28)	159 (68)	0.0010	121 (32)	154 (72)	0.0010
Medical history									
Hypertension	45/79 (57%)	90/102 (88%)	<0.0001	18/29 (62%)	29/33 (88%)	0.0354	27/50 (54%)	61/69 (88%)	<0.0001
Diabetes mellitus	8/79 (10%)	42/100 (42%)	<0.0001	5/29 (17%)	20/33 (61%)	0.0007	3/50 (6%)	22/67 (33%)	0.0005
Dyslipidemia	32/79 (41%)	56/99 (57%)	0.0360	14/29 (48%)	23/32 (72%)	0.0716	18/50 (36%)	33/67 (49%)	0.1883
Congestive heart failure	4/79 (5%)	27/96 (28%)	<0.0001	3/29 (10%)	12/31 (39%)	0.0164	1/50 (2%)	15/65 (23%)	0.0009
Atrial fibrillation	24/79 (30%)	47/100 (47%)	0.0311	12/29 (41%)	15/32 (47%)	0.7974	12/50 (24%)	32/68 (47%)	0.0126
Coronary artery disease	22/79 (28%)	44/100 (44%)	0.0296	5/29 (17%)	15/33 (45%)	0.0286	17/50 (34%)	29/67 (43%)	0.3433
Most proximal occlusion site									
Internal carotid artery	24/106 (23%)	30/122 (25%)	0.7570	9/56 (16%)	10/53 (19%)	0.8025	15/50 (30%)	20/69 (29%)	1.0000
Middle cerebral artery M1	58/106 (55%)	63/122 (52%)	0.6906	35/56 (63%)	31/53 (58%)	0.6988	23/50 (46%)	32/69 (46%)	1.0000
Middle cerebral artery M2	15/106 (14%)	14/122 (11%)	0.5571	6/56 (11%)	8/53 (15%)	0.5736	9/50 (18%)	6/69 (9%)	0.1652
Vertebrobasilar artery	9/106 (8%)	15/122 (12%)	0.3930	6/56 (11%)	4/53 (8%)	0.7429	3/50 (6%)	11/69 (16%)	0.1487
Intravenous tPA failure	53/106 (50%)	60/122 (49%)	1.0000	36/56 (64%)	32/53 (60%)	0.6969	17/50 (34%)	28/69 (41%)	0.5662
Intubation	49/80 (61%)	70/93 (75%)	0.0506	37/56 (66%)	40/53 (75%)	0.3011	12/24 (50%)	30/40 (75%)	0.0581
Time intervals, min, median									
Onset to groin puncture	221 (180-300)	260 (200-334)	0.1105	219 (175-302)	270 (176-345)	0.1870	245 (190-292)	252 (206-316)	0.4187
Groin puncture to first device pass	33 (23-43)	32 (24-44)	0.6371	33 (20-43)	29 (22-40)	0.7850	34 (27-42)	35 (26-45)	0.7900
Onset to first device pass	264 (206-347)	292 (233-374)	0.0962	257 (199-339)	293 (220-380)	0.1325	275 (222-353)	292 (235-370)	0.3833
Groin puncture to reperfusion	77 (57-101)	82 (53-105)	0.6702	80 (58-100)	82 (60-115)	0.7394	75 (56-104)	85 (53-104)	0.7692

Characteristics	All Patients (n=228)			Trevo Group (n=109)			Merci Group (n=119)		
	mRS 0-2 (n=106)	mRS 3-6 (n=122)	P Value	mRS 0-2 (n=56)	mRS 3-6 (n=53)	P Value	mRS 0-2 (n=50)	mRS 3-6 (n=69)	P Value
Onset to reperfusion	315 (256-410)	345 (281-422)	0.0642	285 (244-401)	374 (285-422)	0.0717	326 (268-416)	345 (280-420)	0.4148
Number of passes with	2 (1-3)	2 (1-3)	0.5221	2 (1-3)	2 (1-3)	0.4785	2 (1-3)	2 (1-3)	0.7960
Intra-arterial lytic use	14/106 (13%)	14/122 (11%)	0.6922	3/56 (5%)	2/53 (4%)	1.0000	11/50 (22%)	12/69 (17%)	0.6393
Use of rescue treatment	22/106 (21%)	32/122 (26%)	0.3527	8/56 (14%)	9/53 (17%)	0.7943	14/50 (28%)	23/69 (33%)	0.5549
Symptomatic ICH*	0/106 (0%)	17/122 (14%)	<0.0001	0/56 (0%)	8/53 (15%)	0.0023	0/50 (0%)	9/69 (13%)	0.0099
Device-related serious adverse events	1/106 (1%)	8/122 (7%)	0.0395	0/56 (0%)	4/53 (8%)	0.0526	1/50 (2%)	4/69 (6%)	0.3968

Functional dependence was defined as a score of 3 to 6 on the mRS at 90 days. The baseline data of race, medical history, blood pressure, and blood glucose were not available in the Thrombectomy Revascularization of Large Vessel Occlusions in Acute Ischemic Stroke (TREVO) trial. The data of suspected stroke cause and intubation were not available in the Multi Mechanical Embolus Removal in Cerebral Ischemia (MERC) trial. ICH indicates intracranial hemorrhage; mRS, modified Rankin Scale; NIHSS, National Institutes of Health Stroke Scale; and tPA, tissue-type plasminogen activator.

* Symptomatic ICH was defined as a point increase of 4 in the NIHSS within 24 hours with evidence of any blood in the brain or within the cranium identified on 24-hour head computed tomography/MRI scan, or any intracranial hemorrhage in which no further NIHSS scores were available beyond baseline and the patient died.

Table 2
Multivariate Analysis of Predictors of Functional Dependence Despite Successful Revascularization With Trevo Retriever and Merci Retriever

Variables	All Patients (n=228)		Trevo Group (n=109)		Merci Group (n=119)	
	Odds Ratio (95% CI)	P Value	Odds Ratio (95% CI)	P Value	Odds Ratio (95% CI)	P Value
Age, per year increase	1.04 (1.02–1.06)	0.0005	1.06 (1.02–1.10)	0.0026	1.02 (1.00–1.05)	0.0405
NIHSS score, per point increase	1.08 (1.02–1.15)	0.0055	1.13 (1.03–1.23)	0.0074	...	NS
Onset to groin puncture, per 30-min increase	1.11 (1.01–1.22)	0.0306	...	NS	...	NS

Variables with $P < 0.15$ in the univariate analysis were considered in multivariate logistic regression model-building process. Models were built using stepwise regression with variables entered into the model at the 0.15 significance level and removed at the 0.05 significance level. CI indicates confidence interval; NIHSS, National Institutes of Health Stroke Scale; and NS, not significant.

Table 3
Functional Dependence Despite Successful Revascularization by Time Intervals

Time Tertiles	All Patients (n=228)			Trevo Group (n=109)		
	Patient/Total (%)	Adjusted OR (95% CI)	P Value	Patient/Total (%)	Adjusted OR (95% CI)	P Value
Onset to groin puncture, min						
75–210	40/85 (47)	Reference	...	19/45 (42)	Reference	...
211–345	56/103 (54)	1.53 (0.83–2.83)	0.1770	21/42 (50)	1.45 (0.56–3.74)	0.4427
346–480	25/39 (64)	2.81 (1.20–6.55)	0.0168	13/22 (59)	2.76 (0.83–9.12)	0.0963
OR per 30 min or <i>P</i> for trend	...	1.12 (1.02–1.24)	0.0223	...	1.14 (0.99–1.32)	0.0623
Onset to first device pass, min						
105–240	36/79 (46)	Reference	...	17/42 (41)	Reference	...
241–375	55/100 (55)	1.64 (0.87–3.07)	0.1234	21/44 (48)	1.49 (0.58–3.85)	0.4069
376–510	28/44 (64)	2.57 (1.14–5.79)	0.0230	15/23 (65)	3.23 (0.95–10.98)	0.0597
OR per 30 min or <i>P</i> for trend	...	1.10 (1.01–1.21)	0.0379	...	1.12 (0.97–1.28)	0.1145
Onset to reperfusion, min						
120–270	26/61 (43)	Reference	...	10/32 (31)	Reference	...
271–420	65/115 (57)	1.83 (0.94–3.55)	0.0745	29/53 (55)	2.70 (0.98–7.42)	0.0549
421–570	31/52 (60)	2.79 (1.23–6.33)	0.0141	14/24 (58)	4.10 (1.17–14.42)	0.0279
OR per 30 min or <i>P</i> for trend	...	1.11 (1.01–1.21)	0.0228	...	1.13 (0.99–1.29)	0.0676

Data were adjusted for age, sex, and baseline National Institutes of Health Stroke Scale score. CI indicates confidence interval; and OR, odds ratio.

Table 4
 Functional Dependence Despite Successful Revascularization by Age and Baseline NIHSS Score

Variables	All Patients (n=228)			Trevo Group (n=109)		
	Patient/Total (%)	Adjusted OR (95% CI)	P Value	Patient/Total (%)	Adjusted OR (95% CI)	P Value
Age*						
60	28/69 (41)	Reference	...	8/29 (28)	Reference	...
60-70	27/50 (54)	1.85 (0.86-3.99)	0.1158	16/32 (50)	2.92 (0.92-9.30)	0.0698
70-80	42/73 (58)	2.08 (1.03-4.20)	0.0414	20/37 (54)	3.72 (1.20-11.51)	0.0225
>80	25/36 (69)	3.32 (1.35-8.16)	0.0089	9/11 (82)	14.29 (2.14-95.69)	0.0061
OR per 10 y or P for trend	...	1.41 (1.15-1.73)	0.0008	...	1.92 (1.27-2.89)	0.0019
NIHSS score [†]						
8-10	7/16 (44)	Reference	...	5/11 (46)	Reference	...
11-19	54/119 (45)	0.69 (0.22-2.17)	0.5298	20/54 (37)	0.31 (0.06-1.50)	0.1455
20	61/93 (66)	1.48 (0.46-4.70)	0.5092	28/44 (64)	1.09 (0.23-5.20)	0.9128
OR per 5 points or P for trend	...	1.47 (1.11-1.96)	0.0083	...	1.78 (1.12-2.81)	0.0144

CI indicates confidence interval; NIHSS, National Institutes of Health Stroke Scale; and OR, odds ratio.

* Data were adjusted for baseline NIHSS score, sex, and onset to reperfusion time.

[†] Data were adjusted for age, sex, and onset to reperfusion time.