Supplementary Online Content

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This supplementary material has been provided by the authors to give readers additional information about their work.

eMethods. Propensity Score Matching

Propensity score matching was performed to reduce the possibility of selection bias and confounding on the primary and secondary outcomes. The propensity scores were estimated using a multivariable logistic regression model in which the treatment status (administration of intravenous thrombolysis) was regressed on selected baseline covariates (listed below). We conducted a 1:1 pair matching using nearest-neighbor matching without replacement and with a caliper width of 0.2. Graphical comparison was used to assess the distributional similarity of propensity scores between the treatment and control group. After propensity score matching, balance across the treatment and control group was assumed for all selected baseline covariates given an absolute standardized mean difference (SMD) of <0.10. Propensity score matching was conducted using the Matchlt package in R statistical software (version 4.1.2, R Project for Statistical Computing) and RStudio statistical software (version 2021.09.1+372, Rstudio).

Selected baseline covariates for estimation of propensity scores:

- Age
- Sex
- National Institutes Health Stroke Scale (NIHSS) on admission
- Alberta Stroke Program Early CT Score (ASPECTS) on admission

	Number of Patients (n)				
	Control	Treatment			
All	513	390			
Matched	373	373			
Unmatched	140	17			
Discarded	0	0			

	Before Propensity Score Matching			After Propensity Score Matching		
	Control	Treatment	SMD*	Control	Treatment	SMD*
Age (years)	75.89	73.96	-0.14	75.27	74.95	-0.02
Male Sex (%)	43.1	41.0	-0.04	42.1	40.5	-0.03
NIHSS (points)	14.64	14.26	-0.06	14.23	14.33	0.02
ASPECTS (points)	7.84	8.37	0.29	8.31	8.35	0.02
*SMD_Standardized Mean Difference						



eFigure 1. Flowchart of Patient Inclusion and Exclusion Criteria

CT Score; IVT, Intravenous Thrombolysis; mRS, modified Rankin Scale; ACA, Anterior Cerebral Artery; ICA, Internal Carotid Artery, BA, Basilar Artery; VA, Vertebral Artery; PCA, Posterior Cerebral Artery; TICI, Thrombolysis in Cerebral Infarction

^a Multiple selection of patients possible.

eFigure 2. Propensity Score Matching

(A) Distributions of propensity scores before and after propensity score matching for the treatment and control group. (B) Scatter plots displaying the distributions of propensity scores before and after propensity score matching for the treatment and control group. (C) Standardized mean differences before propensity score matching (red) and after propensity score matching (blue) for all selected baseline covariates. Balance across the treatment and control group was assumed for all selected baseline covariates given an absolute standardized mean difference of <0.10 (dashed lines).



eFigure 3. Subgroup Analyses for Independent Ambulation

The forest plot demonstrates that odds of achieving independent ambulation at 90 days (defined as modified Rankin Scale sores of 0 to 3) favored the IVT+MT group across all predefined subgroups. Arrows indicate that the limits of the confidence interval are not fully shown.

Subgroup	No. of patients	MT Therapy	IVT+MT Therapy	Adjusted Odds Ratio fo	r 90-d mRS 0-3 (95% Cl)ª
		90-d m	RS 0-3 (%)		
Age					
≥70 years	533	14.4%	19.0%	1.59 (0.94 to 2.72)	
<70 years	213	34.0%	44.5%	1.96 (1.05 to 3.73)	
Sex					
male	308	27.4%	35.8%	1.88 (1.08 to 3.35)	}∎
female	438	14.4%	20.3%	1.66 (0.93 to 2.99)	∲ ∎ 1
Onset to admission					
≤270 minutes	369	21.6%	30.3%	2.15 (1.20 to 3.98)	i⊢∎i
>270 minutes or unknown	377	19.0%	19.4%	1.20 (0.65 to 2.21)	⊢:∎ 1
NIHSS score					
>10 points	522	8.2%	16.2%	2.11 (1.19 to 3.84)	⊢_∎ 1
≤10 points	224	45.7%	51.9%	1.56 (0.87 to 2.84)	
Occlusion site					
ICA	143	9.2%	17.4%	7.41 (1.70 to 44.33)	
MCA M1/M2	603	22.1%	28.8%	1.60 (1.05 to 2.45)	
ASPECTS					
6-10	693	20.5%	27.5%	1.74 (1.15 to 2.64)	⊢ ∎1
0-5	53	11.5%	14.8%	1.33 (0.27 to 7.41) ^b	⊢ !∎ I
TOAST classification					
Large-artery atherosclerosis	165	25.3%	31.1%	1.82 (0.81 to 4.28)	
Cardioembolism	373	16.9%	22.7%	1.55 (0.83 to 2.89)	1 −−−−1
Stroke of undetermined aetiology	163	17.8%	30.0%	2.57 (1.09 to 6.46)	• • •
Final mTICI grade					
0/1 (failed reperfusion)	481	15.2%	23.5%	1.83 (1.08 to 3.14)	
2a (partial reperfusion)	275	28.5%	31.9%	1.54 (0.84 to 2.87)	i .
^a Adjusted odds ratios of the p	redefined subg	groups. Result	ts were adjuste	d for age, sex, NIHSS,	1 2 3 4 5 6 7 8 9 10

ASPECTS and final mTICI grade. No adjustments were made for time from symptom onset to hospital admission and for the respective stratification variable.

Favors IVT+MT

^b Unadjusted odds ratio. Adjustment for baseline covariates was not performed due to limited number of patients in the subgroup (n=53).

Abbreviations:

MT, Mechanical Thrombectomy; IVT, Intravenous Thrombolysis; CI, Confidence Interval; NIHSS, National Institutes Health Stroke Scale; ICA, Internal Carotid Artery; MCA, Middle Cerebral Artery; ASPECTS, Alberta Stroke Program Early CT Score; TOAST, Trial of Org 10172 in Acute Stroke Treatment; mTICI, modified Treatment in Cerebral Infarction

eTable 1. Functional Outcomes and Safety Measures Before Propensity Score Matching

	Before Pro	opensity Sco	re Matching					
	Patients, I	Patients, No. (%)						
	All (n=903)	MT Therapy (n=513)	IVT+MT Therapy (n=390)	${oldsymbol{P}}^{(a)}$	Adjusted Risk Difference (95% Cl) ^b	Unadjusted Odds Ratio (95% Cl)	Adjusted Odds Ratio (95% Cl)⁵	Ρ
Primary Outcome								
90-day mRS score 0-2	124/903 (13.7)	47/513 (9.2)	77/390 (19.7)	<.001 ²	6.7% (4.9 to 8.5)	2.44 (1.66-3.62)	2.65 (1.47-4.78)	.001
Secondary Outcomes								
90-day mRS score 0-3	199/903 (22.0)	87/513 (17.0)	112/390 (28.7)	<.001 ²	7.4% (3.0 to 11.8)	1.97 (1.44-2.72)	1.88 (1.14-3.12)	.01
90-day mRS score, median (IQR)	6 (4-6)	6 (4-6)	5 (3-6)	<.001 ¹	NA	1.91 (1.49-2.45)⁰	2.13 (1.49-3.05)⁰	<.001
Safety Measures								
Symptomatic ICH within 24 hours	49/889 (5.5)	27/505 (5.3)	22/384 (5.7)	.80 ²	0.4% (-2.3 to 3.0)	1.08 (0.60-1.92)	0.99 (0.42-2.42)	0.98
Death within 90 days	460/903 (50.9)	294/513 (57.3)	166/390 (42.6)	<.001 ²	-12.5% (-19.5 to -5.5)	0.55 (0.42-0.72)	0.49 (0.32-0.76)	.001

^a Characteristics were compared between MT and IVT+MT patients with the use of either Mann-Whitney U test (1) for continuous variables or a chi-square test (2) for categorical variables.

^b Results were adjusted for age, sex, interval from symptom onset to hospital admission, NIHSS, ASPECTS and final mTICI grade.

^c Common odds ratios derived from ordinal logistic regression analysis. Values greater than 1 indicate a shift in the distribution of 90-day mRS scores towards lower values (better functional outcomes) favoring IVT+MT compared with MT alone.

Abbreviations:

MT, Mechanical Thrombectomy; IVT, Intravenous Thrombolysis, IQR, Interquartile Range; mRS, modified Rankin Scale; ICH, Intracranial Hemorrhage; NA, not applicable

eTable 2. Multivariable Logistic Regression Models

Used to identify independent determinants of functional independence at 90 days (modified Rankin Scale scores of 0 to 2) and independent ambulation at 90 days (modified Rankin Scale scores of 0 to 3) in stroke patients with unsuccessful mechanical reperfusion.

	90-day mRS score 0-2			90-day mRS score 0-3			
	β Coefficient	Adjusted Odds Ratio (95% Cl)	Р	β Coefficient	Adjusted Odds Ratio (95% Cl)	Ρ	
Independent Variables							
Age (per 10 years)	-0.432	0.65 (0.51-0.81)	<.001	-0.530	0.59 (0.47-0.73)	<.001	
Male Sex (yes)	-0.436	0.65 (0.36-1.17)	.149	-0.389	0.68 (0.40-1.14)	.14	
Stroke onset to admission (per minute)	0.002	1.00 (1.00-1.00)	.142	0.001	1.00 (1.00-1.00)	.19	
Admission NIHSS score (per 1 point)	-0.176	0.84 (0.79-0.88)	<.001	-0.174	0.84 (0.80-0.88)	<.001	
Administration of IVT (yes)	0.967	2.63 (1.41-5.11)	.003	0.563	1.76 (1.03-3.04)	.04	
ASPECTS 6-10 (yes)	-0.524	0.59 (0.09-2.70)	.541	-0.239	0.79 (0.19-2.76)	.72	
Final mTICI 0 (Reference)							
Final mTICI 1	0.232	1.26 (0.48-3.11)	.622	0.247	1.28 (0.57-2.80)	.54	
Final mTICI 2a	1.453	4.27 (2.28-8.25)	<.001	1.022	2.78 (1.60-4.89)	<.001	

n = 423 patients included. A P value of less than 0.05 was considered significant.

Abbreviations:

mRS, modified Rankin Scale; NIHSS, National Institutes of Health Stroke Scale; IVT, Intravenous Thrombolysis; ASPECTS, Alberta Stroke Program Early CT Score; mTICI, modified Thrombolysis in Cerebral Infarction.

eTable 3. Multivariable Ordinal Logistic Regression Model

Used to assess the relationship between the distribution of modified Rankin Scale scores at 90 days and selected independent variables (shift analysis).

	90-day mRS score				
	β Coefficient	Adjusted Common Odds Ratio (95% CI)	Р		
Independent Variables					
Age (per 10 years)	-0.530	0.59 (0.50-0.68)	<.001		
Male sex (yes)	0.008	1.01 (0.69-1.47)	.97		
Stroke onset to admission (per minute)	0.001	1.00 (1.00-1.00)	.42		
Admission NIHSS score (per 1 point)	-0.139	0.87 (0.84-0.90)	<.001		
Administration of IVT (yes)	0.682	1.98 (1.35-2.92)	<.001		
ASPECTS 6-10 (yes)	-0.647	0.52 (0.20-1.25)	.16		
Final mTICI 0 (Reference)					
Final mTICI 1	0.437	1.55 (0.88-2.70)	.13		
Final mTICI 2a	0.967	2.63 (1.75-3.97)	<.001		

 ${\sf n}$ = 423 patients included. A P value of less than 0.05 was considered significant.

Abbreviations:

mRS, modified Rankin Scale; NIHSS, National Institutes of Health Stroke Scale; IVT, Intravenous Thrombolysis; ASPECTS, Alberta Stroke Program Early CT Score; mTICI, modified Thrombolysis in Cerebral Infarction.