

Intravenous Thrombolysis for Acute Mild Ischemic Stroke Patients: Higher ABCD2 Score Associated with Better Outcome

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Abstract

Background: Up to 30% of patients with mild ischemic stroke suffer neurologic deterioration. However, optimal medical approaches of such patients remain controversial given the efficacy and safety of intravenous thrombolysis (IVT). The purpose of this study was to evaluate whether patients with acute mild stroke stratified with ABCD2 score (the risk of stroke on basis of age, blood pressure, clinical features, duration of symptoms, and presence of diabetes mellitus) could benefit from IVT.

Methods: Among 3321 patients with a final diagnosis of acute ischemic stroke or transient ischemic attack, we retrospectively included 227 patients identified with acute mild neurologic deficits (National Institution of Health Stroke Scale, NIHSS ≤ 5) treated with or without IVT. Odds ratios (OR) with their confidence intervals (CI) for outcomes between groups were assessed by using multivariable binary logistic regression analyses. And the heterogeneity of treatment effect magnitude for excellent outcome at 90 days (modified Rankin Scale [mRS] 0-1) was estimated in different subgroups.

Results: A total of 227 cases were enrolled, 108 receiving IVT and 119 treated with secondary stroke prevention strategies alone. Patients receiving IVT had more history of atrial fibrillation. At 7 days, 32 (29.6%) patients with IVT treatment versus 17 (14.3%) patients not receiving IVT achieved significant improvement (≥ 4 -point NIHSS score decrease; OR, 2.57; 95%CI, 1.29-5.12; $P=0.007$). At 90 days, excellent outcome was achieved in 85 (78.7%) patients treated with IVT versus 78 (65.5%) patients without IVT treatment (OR: 2.95; 95% CI, 1.44-6.03; $P=0.003$), especially in those with ABCD2 score ≥ 4 (78.4% versus 64.6%; OR, 2.81; 95%CI, 1.31-6.03; $P=0.008$). Besides, 7(6.5%) IVT-treated patients versus 2 (1.7%) non-IVT-treated patients developed intracranial hemorrhage (ICH; OR, 4.05; 95%CI, 0.82-19.96; $P=0.085$), among these only 1 (0.9%) was symptomatic ICH with IVT treatment.

Conclusions: For acute mild ischemic stroke patients, we reassured the safety and especially the efficacy of IVT at 7- and 90-days. Patients with 4 or more of ABCD2 score might benefit more from IVT.

Keywords: Prognosis, risk scores, stroke, intravenous thrombolysis.

Background

Mild stroke accounts for approximately two thirds of acute ischemic stroke patients in population-based studies[1, 2]. Nevertheless, minor neurological deficiency or rapidly improving symptoms are the most common reasons for withdraw from IVT in otherwise guideline-based eligible patients[3]. Previous data indicated that 30% of such patients had persistent disability at 90 days[4], and the risk of recurrent stroke at 90 days, 1 year and 5 years were 3.7%, 5.1% and 9.5%, respectively[5, 6].

Although the proportion of thrombolytic therapy for mild stroke patients has increased over the past decade, the efficacy and safety of recombinant tissue plasminogen activator (rt-PA) remains controversial. A series of studies suggested patients with mild deficiency could benefit from IVT[7-11]. Currently, the American Stroke Association gives a level I (strong) and IIb (weak) evidence-based score

respectively for IVT in patients with mild disabling and nondisabling ischemic stroke symptoms within 3h[12]. But there were also studies indicating no significance in the efficacy between IVT and antiplatelet therapy in patients with mild stroke[13, 14], especially in those with non-disabling deficiency[14]. And the patient treated with IVT was not recommended given antithrombotic therapy within 24 hours, even though an aggravation of the condition. In consideration of the controversial therapeutic decision-making, it is necessary to screen high-risk mild stroke patients for IVT.

The purpose of this observational study was to investigate whether patients with acute mild stroke symptoms (NIHSS score \leq 5) could achieve 7- and 90-day good outcome from intravenous rt-PA therapy by comparing with patients treated with timely secondary stroke prevention strategies not receiving rt-PA. We also hypothesized that ABCD2 score might be related to the prognosis of IVT patients.

Methods

Study Design

This study was designed as a retrospective study evaluating the efficacy and safety of IVT administered within 4.5 hours of symptom onset.

This study was approved by the First Affiliated Hospital of Soochow University Institutional Review Board. And the need for patient informed consent was waived by the same ethics committee. Consecutive acute ischemic stroke (AIS) patients with minor-to-mild stroke symptoms (NIHSS score \leq 5) were enrolled from our hospital between August 2016 and May 2018.

Patients Selection

Eligible cases were collected using the following inclusion criteria: (1) Clinical diagnosis of AIS or transient ischemic stroke (TIA) with NIHSS score 0 to 5; (2) Age 18 years or older; (3) Time from symptom onset within 4.5h for rt-PA group, however, it could be administered up to 24 hours after symptom onset in non-rt-PA group; (4) Available for a telephone interview at 90 days. The exclusion criteria were as follows: (1) Pre-stroke mRS score of \geq 2; (2) ICH on baseline computed tomography, and other contraindications to IVT and antithrombotic therapy.

Patient Management

Participants were divided into two groups according to different approaches about the management of mild stroke patients: (1) rt-PA group: Intravenous rt-PA within 4.5 hours of AIS onset or last known well time (0.9 mg/kg, maximum dose 90 mg with initial 10% of total dose given as bolus during 1 minute), followed by appropriate secondary stroke prevention strategies; (2) Non-rt-PA group: Optimal secondary stroke prevention strategies within 24h from AIS onset: including initiation of dual antiplatelets among patients with NIHSS \leq 3 (clopidogrel 75 mg per day for 90 days + aspirin 100 mg per day for the first 3 weeks); clopidogrel 75mg/ aspirin 100mg per day; or anticoagulation agents in the event of cardioembolism, and other measures such as statins and antihypertensive medication, etc.

Patient Data Collection

Baseline demographic and clinical information were collected by experienced stroke neurologists: including age, gender, previous history (hypertension, diabetes mellitus, atrial fibrillation, coronary heart disease, previous stroke or TIA and current smoking), medication history, stroke severity (measured by NIHSS score at admission and 7d), time from symptom onset, blood pressure at baseline, ABCD₂ score, stroke subtype (assessed using TOAST classification [Trial of Org 10172 in Acute Stroke Treatment])[15], and neurological imaging (collected at baseline, 24-36h, 7d or discharge if sooner in stroke unit). The 90d-mRS was assessed by telephone follow-up.

Definition of Clinical Information

All participants were hospitalized in the stroke unit assessed by neurologic specialists. (1) Efficacy outcomes: The achievement of an mRS of 0 to 1 at 90 days was defined as excellent outcome, and an mRS of 0 as perfect outcome. Significant improvement was referred as complete resolution of the neurologic deficit or an improvement of at least 4 points over baseline NIHSS score[16]. (2) Adverse outcomes: Early neurological deterioration (END) was defined as a NIHSS score increase of 2 or more within 7d after symptom onset excluding any CT- or MRI-documented ICH[17]. sICH was defined as CT-based ICH within 7 days with a NIHSS score increase of at least 4 or death[18]. The recurrent ischemic stroke within 3 months was assessed by stroke specialists via face-to-face interviews or telephone follow-up.

Statistical Analysis

Categorical variables are expressed as numbers (%). Continuous variables are expressed as mean (standard deviation [SD]) in the case of normal distribution, otherwise as median (interquartile range [IQR]), and the normality of distributions was evaluated by histograms and the Shapiro–Wilk test. The between-group differences in baseline characteristics were assessed as follows: continuous variables were compared with the Student t test or Mann-Whitney test as appropriate, while categorical variables were compared with chi-square test (Pearson's χ^2 or continuity correction). ORs with their CIs for efficacy and adverse outcomes were estimated respectively using multivariable binary logistic regression analyses in the whole study group. Moreover, we intended to estimate the heterogeneity of therapeutic effect magnitude for excellent outcome stratified by baseline NIHSS score (≤ 3 versus ≥ 3), stroke subtype (large artery atherosclerosis, cardioembolism, small vessel occlusion and other/undetermined etiology), disabling neurologic deficits (yes versus no), and baseline ABCD₂ score (≤ 4 versus ≥ 4).

The α -level of significance was $P < 0.05$ two-tailed. All analyses were performed using the SPSS software version 25.0.

Results

Population and Baseline Characteristics

Between August 2016 and May 2018, a total of 327 patients with AIS were treated with intravenous rt-PA in stroke unit within 4.5h, among which 111 (33.9%) with mild neurologic deficit (baseline NIHSS score ≤ 5). And 3 were excluded for lost during follow-up, resulting in 108 patients in the rt-PA group. Meanwhile, 2994 patients were admitted to the unit with AIS not receiving intravenous rt-PA. Among these 119 participants responding to inclusion criteria were enrolled in the non-rt-PA group (Figure 1).

The median age of rt-PA treated patients was 65.5 (IQR, 59-74), and 66 (IQR, 59-74) in rt-PA untreated patients. There were 30 patients (27.8%) in rt-PA group and 45 patients (37.8%) in non-rt-PA group were female. The median baseline NIHSS score in different medical approaches was both 3 (IQR, 2-4; $P=0.09$). And the median baseline ABCD₂ score among two groups was also matched (5 versus 5; IQR, 4-6 versus 5-6; $P=0.58$). Study groups were generally balanced as shown in Table 1, with the most common medical risk factors being hypertension (71.8%) and current smoking (34.8%). However, patients in rt-PA group had a significantly higher proportion of history of atrial fibrillation (18.5% versus 7.6%, $P=0.01$). Among patients with mild stroke, the most common stroke subtype was small vessel occlusion (37.9%) followed by large artery atherosclerosis (32.6%), with other/undetermined etiology (15.4%) and cardioembolism (14.1%) less frequent.

Efficacy outcomes

Figure 2 showed the comparisons in efficacy and adverse outcomes between different medical approaches. And multivariable-adjusted associations between covariates and efficacy/adverse outcomes were showed in Table 2. Excellent outcome at 90 days was achieved in 85 (78.7%) patients treated with rt-PA compared with 78 (65.5%) patients without rt-PA treatment (Figure 2), with an adjusted OR of 2.95 (95% CI, 1.44-6.03; $P=0.003$) after controlling for the effects of male (OR, 3.2; 95%CI, 1.60-6.28; $P=0.001$), history of diabetes mellitus (OR, 0.34; 95%CI, 0.16-0.70; $P=0.004$) and worse baseline NIHSS score (OR, 0.51; 95%CI, 0.38-0.69; $P<0.001$) (Table 2). Meanwhile, excellent outcome at 90 days achieved by intravenous rt-PA therapy was both observed in different time window from symptom onset (0-3h versus 3-4.5h, 83.9% versus 73.1%, $P=0.17$) (data not shown). Similar result was found considering perfect outcome (OR, 2.08; 95%CI, 1.17-3.70; $P=0.013$). Besides, worse stroke severity (OR, 0.66; 95%CI, 0.52-0.83; $P=0.001$) and history of diabetes mellitus (OR, 0.48; 95%CI, 0.24-0.97; $P=0.039$) were associated with lower rate of perfect outcome. Moreover, we detected significant improvement in 32 (29.6%) patients receiving rt-PA treatment (OR, 2.57; 95%CI, 1.29-5.12; $P=0.007$), with control for the effects of male (OR, 4.52; 95%CI, 1.78-11.49; $P=0.002$) and worse baseline NIHSS score (OR, 0.69; 95%CI, 0.53-0.91; $P=0.009$).

Adverse outcomes

The rate of END within 7 days was 13.0% in patients treated with rt-PA, whereas 10.1% in patients without rt-PA treatment, with no statistically significant difference between two groups (OR, 1.33; 95%CI, 0.59-3.01; $P=0.497$). History of diabetes mellitus was strongly associated with END at 7 days (OR, 3.37; 95%CI, 1.46-7.78; $P=0.004$). Any ICH within 7 days, occurred more frequently in patients treated with rt-PA compared with patients without rt-PA treatment (6.5% versus 1.7%; OR, 4.05; 95%CI, 0.82-19.96; $P=0.085$).

However, only 1 (0.9%) was sICH in rt-PA group. Older age, high level of systolic blood pressure at admission and worse stroke severity were associated with any ICH. While the recurrent ischemic stroke in 90 days was less frequent in patients treated with rt-PA compared with the other group (2.8% versus 4.2%; OR, 0.65; 95%CI, 0.15-2.79; $P=0.564$). Only male was strongly associated with lower incidence of recurrent ischemic stroke (OR, 0.064; 95%CI, 0.01-0.53; $P=0.011$).

Association Between Subgroups and Excellent Outcome

In the subgroup analysis stratified by ABCD₂ score (Figure 3), excellent outcome was achieved in 80 patients (78.4%) receiving intravenous rt-PA with ABCD₂ score ≥ 4 , compared with 73 (64.6%) patients not receiving intravenous rt-PA with ABCD₂ score ≥ 4 (OR, 2.81; 95%CI, 1.31-6.03; $P=0.008$). For patients with ABCD₂ score ≤ 4 , no significant of treatment effect was observed between two groups. Excellent outcome was achieved in both subgroups of baseline NIHSS score with significant difference. However, we detected no significant heterogeneity of therapeutic effect magnitude for excellent outcome stratified by stroke subtype and disabling neurologic deficits.

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Discussion

Our study implied a high proportion of 78.7% acute mild ischemic stroke (NIHSS score ≤ 5) patients receiving intravenous rt-PA within 4.5 hours achieved excellent outcome (mRS 0-1) at 90 days, and 29.7% with significant improvement at 7 days, indicating IVT being effective for patients with mild deficits. And

the encouraging results were both detected in the 0-3h (83.9%) and 3-4.5h (73.1%) time windows. Notably, we found that patients with mild stroke whose ABCD₂ score ≥4 could benefit more from IVT.

A series of randomized studies[7, 14], systematic review[8-10] and observational cohorts[11, 13] indicated controversial results regarding the efficacy of IVT among patients with acute mild ischemic stroke. The post hoc analysis in a rigorously selected sample (restricted to 106 participants with a baseline NIHSS score ≤5 within 3 hours from symptom onset) of the third International Stroke Trial (IST-3) suggested encouraging results of intravenous rt-PA efficacy in mild ischemic stroke (84% rt-PA versus 65% control; adjusted odds ratio, 3.31; 95% CI, 1.24–8.79; *P*=0.03)[7], which is consistent with our study and most post previous data[8-11]. Nevertheless, a large observational cohort from China suggested that intravenous rt-PA might potentially benefit patients with NIHSS score ≤5 within 4.5 hours from symptom onset, with no statistical significance (76% rt-PA versus 69.5% control; odds ratio, 1.48; 95% CI, 0.91–2.43; *P*=0.12)[13]. The difference in efficacy of IVT might be associated with the different time window selection, various definition of mild stroke and better secondary stroke prevention measurements. Currently, the preliminary results of Potential of rtPA for Ischemic Strokes with Mild Symptoms (PRISMS)[14], a double-blind, multicenter, randomized controlled trial, revealed that likelihood of excellent outcome at 90 days didn't increase among mild nondisabling AIS patients receiving IVT treatment, yet a higher risk of complication of sICH. However, the early termination with only 313 participants enrolled might preclude the definitive conclusions. Our study didn't detect significant difference for excellent outcome stratified by disabling deficiency in two medical approaches either. The disabling deficit is commonly defined as follows: complete hemianopia (NIHSS-3 ≥2), severe aphasia (NIHSS-9 ≥2), neglect NIHSS-11 ≥1), limb weakness that cannot resist gravity (NIHSS-5/6 ≥2), functional impairment with NIHSS >5, or any potential disabling deficit judged by experienced physician[14]. Nevertheless, the NIHSS scale may not be adequate to evaluate severity of mild stroke, especially in posterior circulation AIS patients[19], hence considerable patients are regarded as nondisabling deficiency, resulting in unfavorable outcome. So accurate description of disabling neurological deficits is warranted urgently to screen high-risk mild AIS patients.

As for the safety of intravenous rt-PA in mild stroke, we detected 7 (6.5%) receiving intravenous rt-PA patients occurred ICH within 7 days compared with the lower rate of 1.7% in non-rt-PA group, yet among these only 1 (0.9%) developed sICH from rt-PA group, lower than the proportion of 1.8%-4.1% previous studies had reported[14, 20, 21]. The significant association between post-IVT ICH with older age, worse stroke severity and higher admission glucose level was reported in a systematic review from 55 studies[22], similar with our study data. The incidence of sICH in a large single-center cohort[23] stratified by baseline NIHSS score (≤6 versus >6) was 2.0% and 8.1% (*P* = 0.001), respectively, while the difference was not detected with increasing NIHSS score within the range of 0 to 5 (*P* = 0.51)[20], indicating that intravenous rt-PA is relatively safe but not risk-free in mild stroke patients.

Our study demonstrated that acute mild stroke with ABCD₂ score ≥4, a fresh perspective to screen appropriate patients, might potentially benefit more from intravenous rt-PA treatment. ABCD₂ score was intended to aid clinical management and estimate the risk of stroke recurrence in patients with TIA and minor stroke[24]. In addition, ABCD₃-I score (range 0-13, addition of dual TIA within 7 days, ipsilateral

carotid artery stenosis \geq 50% and positive brain imaging) showed better validation for prediction of early and 90-day stroke recurrent risk, with clinical presentation (C), symptom duration (D) and cerebral/carotid imaging (I) being the most essential components[25-27]. Recent trial revealed the ABCD₂ score of 4 and more in TIA or minor stroke was significantly associated with longer-term risk of another stroke[6]. Also, these scores were used to select patients for intensive therapy[28, 29]. However, it remains unclear regarding the validation of the risk scores to screen high-risk candidates for intravenous rt-PA among mild stroke patients, so that further investigation in randomized trials or larger observational studies is needed. Besides, in clinical practice, physicians can as well refer to the stroke etiology of large artery atherosclerosis[13] and presentation of disability[12] to assist thrombolytic decision-making.

Thus, further investigations are warranted. Mild and Rapidly Improving Stroke Study (MaRISS; observational trial; NCT02072681) and Antiplatelet vs Rt-PA for Acute Mild Ischemic Stroke (ARAMIS; randomized, placebo-controlled trial; NCT03661411) are two ongoing trials investigating the safety and particularly the efficacy of intravenous rt-PA in patients with acute mild stroke.

The non-randomized design due to retrospective study is the most notable limitation of our research. Although, the baseline characteristics between two groups were matched generally, with significant difference only in history of atrial fibrillation, the well-known limitations of observational study still not be avoided. And the small sample size of 227 enrolled subjects in this study limited the power to assess the efficacy and safety of IVT among mild stroke patients. In addition, majorities of our enrolled participants had 4 points or more on baseline ABCD₂ score, which might cause bias and undermine the results. Considering the inadequacy to generalize the results of our research to nation-wide population with mild stroke, well-designed multicenter randomized clinical trials are needed.

Conclusions

A substantial proportion of patients deemed mild stroke (NIHSS \leq 5) with intravenous rt-PA therapy has excellent outcome both at 7- and 90-days, especially among patients with 4 or more of baseline ABCD₂ score. While, accurate identification of high-risk stroke is so essential to screen patients with mild deficiency for IVT, that further researches are warranted. In conclusion, given the significant percentage of excellent outcome and low risk of sICH, our results reassure the efficacy and safety of IVT therapy for acute mild ischemic stroke patients.

Abbreviations

ABCD₂ score: The risk of stroke on basis of age, blood pressure, clinical features, duration of symptoms, and presence of diabetes mellitus; ABCD₃-I score: Addition of dual symptom onsets within 7 days, ipsilateral carotid artery stenosis \geq 50% and positive brain imaging on the basis of ABCD₂ score; AIS: Acute ischemic stroke; ARAMIS: Antiplatelet vs rt-PA for acute mild ischemic stroke; CI: Confidence interval; END: Early neurological deterioration ICH: Intracranial hemorrhage; IQR: Interquartile range; IVT: Intravenous thrombolysis; MaRISS: Mild and rapidly improving stroke study; mRS: Modified Rankin scale;

NIHSS: National Institutes of Health Stroke Scale; OR: Odds ratio; PRISMS: Potential of rtPA for ischemic strokes with mild symptoms; rt-PA: Recombinant tissue plasminogen activator; SD: Standard deviation; sICH: Symptomatic intracranial hemorrhage; TIA: Transient ischemic stroke; TOAST: Trial of org 10172 in acute stroke treatment.

Declarations

Ethics approval and consent to participate

The use of data analysis was legally approved by the at the First Affiliated Hospital of Soochow University Institutional Review Board. This research was performed in accordance with the tenets of the Declaration of Helsinki as amended in 2008. The need for patient informed consent was waived by the same ethics committee.

Consent for publication

Not applicable.

Availability of data and material

The datasets used and analyzed during the current study are available from the corresponding author on reasonable request.

Competing interests

The authors declare that they have no competing interests.

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Authors' contributions

QF, XC and YK designed this study. RL and CH collected and analyzed the patient data, and were the major contributor in writing the manuscript. JZ, LZ, RL, CH, ZL and XL conducted the clinical assessments and follow-up of participants. All authors contributed to creating this manuscript and improved the final version.

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Tables

Table 1 Baseline Characteristics of Patients with Mild Stroke According to therapeutic approach.

Characteristics	rt-PA (n= 108)	Non-rt-PA (n= 119)	<i>P</i> value
Age, y, median (IQR)	65.5(59-74)	66(59-74)	0.83
Female, n (%)	30(27.8)	45(37.8)	0.11
Medical history, n (%)			
Hypertension	78(72.2)	85(71.4)	0.89
Diabetes mellitus	31(28.7)	28(23.5)	0.38
Atrial Fibrillation	20(18.5)	9(7.6)	0.01
Coronary heart disease	8(7.4)	10(8.4)	0.78
Previous stroke/TIA	16(14.8)	17(14.3)	0.91
Current smoking	42(38.9)	37(31.1)	0.22
Medications prior to onset, n (%)			
Antihypertension agents	62(57.4)	69(58.0)	0.93
Antidiabetic agents	24(22.2)	21(17.6)	0.39
Antiplatelet agents	15(13.9)	12(10.1)	0.38
Anticoagulant agents	3(2.8)	2(1.7)	0.67
Baseline NIHSS score, median (IQR)	3(2-4)	3(2-4)	0.09
Baseline ABCD2 score, median (IQR)	5(4-6)	5(5-6)	0.58
Stroke subtype, n (%)			0.25
Large artery atherosclerosis	36(33.3)	38(51.4)	
Cardioembolism	20(18.5)	12(10.1)	
Small vessel occlusion	38(35.2)	48(40.3)	
Other/Undetermined etiology	14(13.0)	21(17.6)	
Systolic blood pressure, mmHg, (mean ± SD)	153.8±21.2	149.3±18.9	0.09
Diastolic blood pressure, mmHg, (mean ± SD)	85.5±13.3	83.8±12.1	0.329

Abbreviations: ABCD2 score (age, blood pressure, clinical features, duration of symptoms, and presence of diabetes mellitus; range, 0 to 7, with higher scores indicating a higher risk of stroke); IQR, interquartile range; mRS, modified Rankin Scale; NIHSS, National Institutes of Health Stroke Scale; rt-PA, recombinant tissue plasminogen activator; SD, Standard Deviation; ang TIA, transient ischemic stroke.

Table 2. Multivariable-adjusted Associations Between Covariates and Clinical Outcomes.

Outcomes	variables	OR (95%CI)
Excellent outcome	Intravenous rt-PA	2.95 (1.44-6.03)
	Male	3.17 (1.60-6.28)
	Diabetes mellitus	0.34 (0.16-0.70)
	Baseline NIHSS score, per 1 point	0.51 (0.38-0.69)
Perfect outcome	Intravenous rt-PA	2.08 (1.17-3.70)
	Baseline NIHSS score, per 1 point	0.66 (0.52-0.83)
	Diabetes mellitus	0.48 (0.24-0.97)
Significant improvement	Intravenous rt-PA	2.57 (1.29-5.12)
	Male	4.52 (1.78-11.49)
	Baseline NIHSS score, per 1 point	0.69 (0.53-0.91)
	Diabetes mellitus	3.37 (1.46-7.78)
END	Diabetes mellitus	3.37 (1.46-7.78)
Any ICH	Baseline NIHSS score, per 1 point	3.81 (1.37-10.59)
	Age, per 1 year	1.26 (1.06-1.49)
	Systolic blood pressure, per 1 mmHg	1.10 (1.02-1.19)
	Large artery atherosclerosis	0.01 (1.06×10 ⁻⁴ -0.27)
Recurrent ischemic stroke	Male	0.06 (0.01-0.53)

Abbreviations: CI, confidence interval; END, early neurological deterioration; ICH, intracranial hemorrhage; NIHSS, National Institutes of Health Stroke Scale; OR, odds ratio; and rt-PA, recombinant tissue plasminogen activator.

Figures

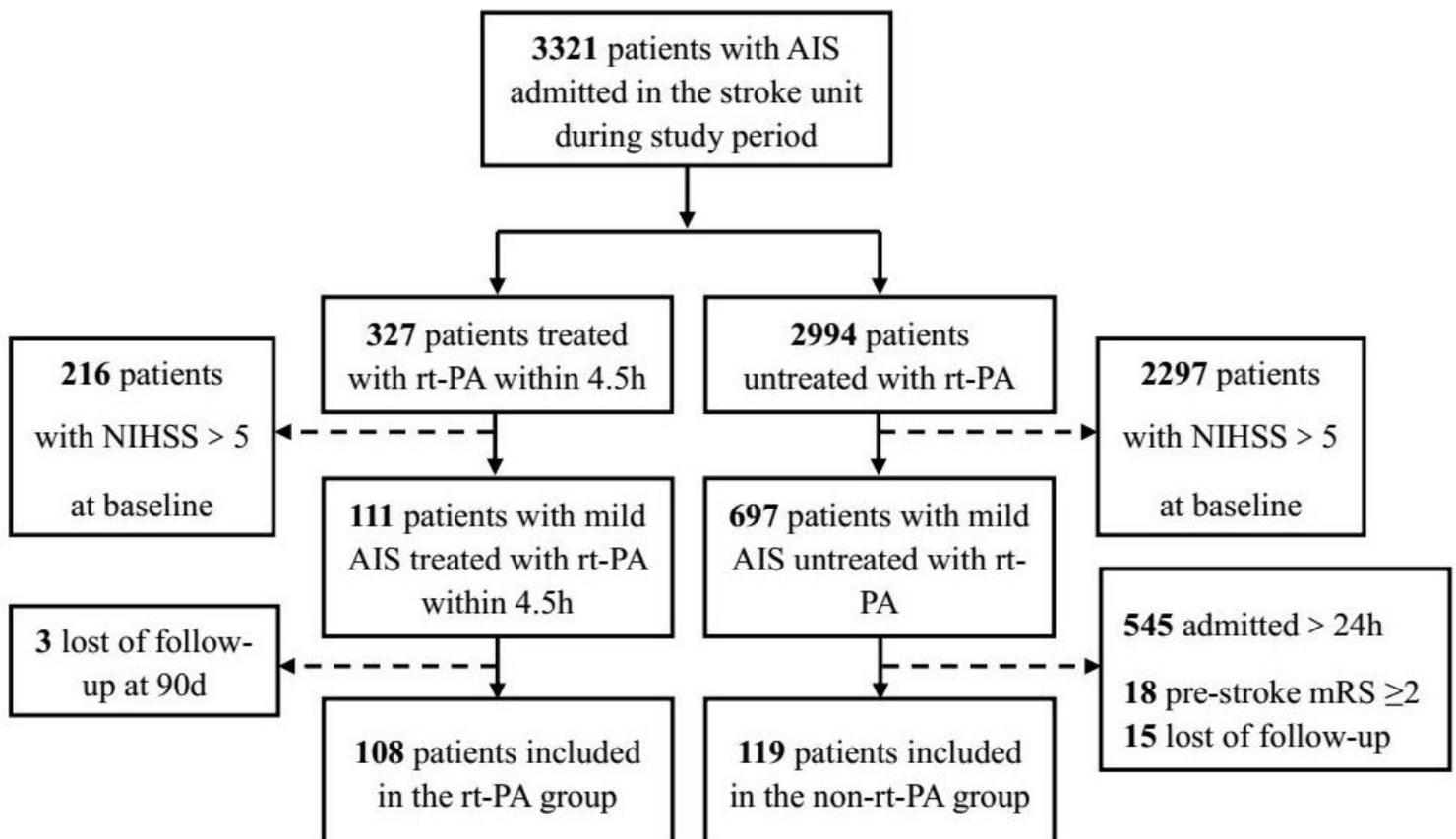


Figure 1

The Flowchart of The Study. Abbreviations: AIS, acute ischemic stroke; mRS, modified Rankin Scale; NIHSS, National Institutes of Health Stroke Scale; rt-PA, recombinant tissue plasminogen activator.

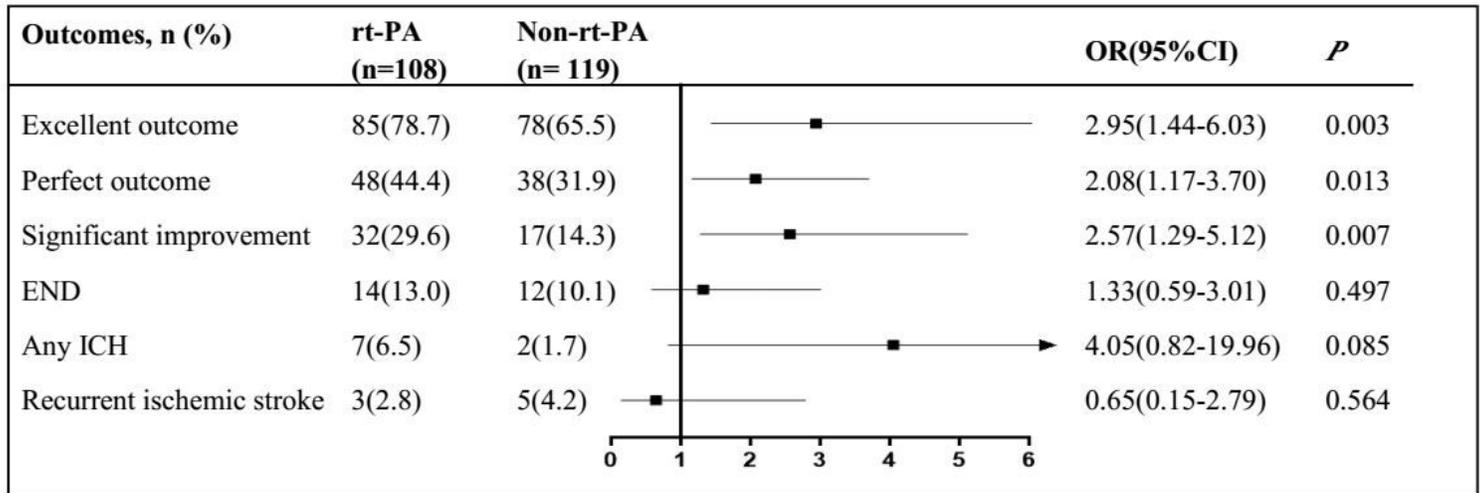


Figure 2

Comparisons of Clinical Outcomes with Mild Stroke Between Different Medical Approaches.

Abbreviations: CI, confidence interval; END, early neurological deterioration; ICH, intracranial hemorrhage; and OR, odds ratio.

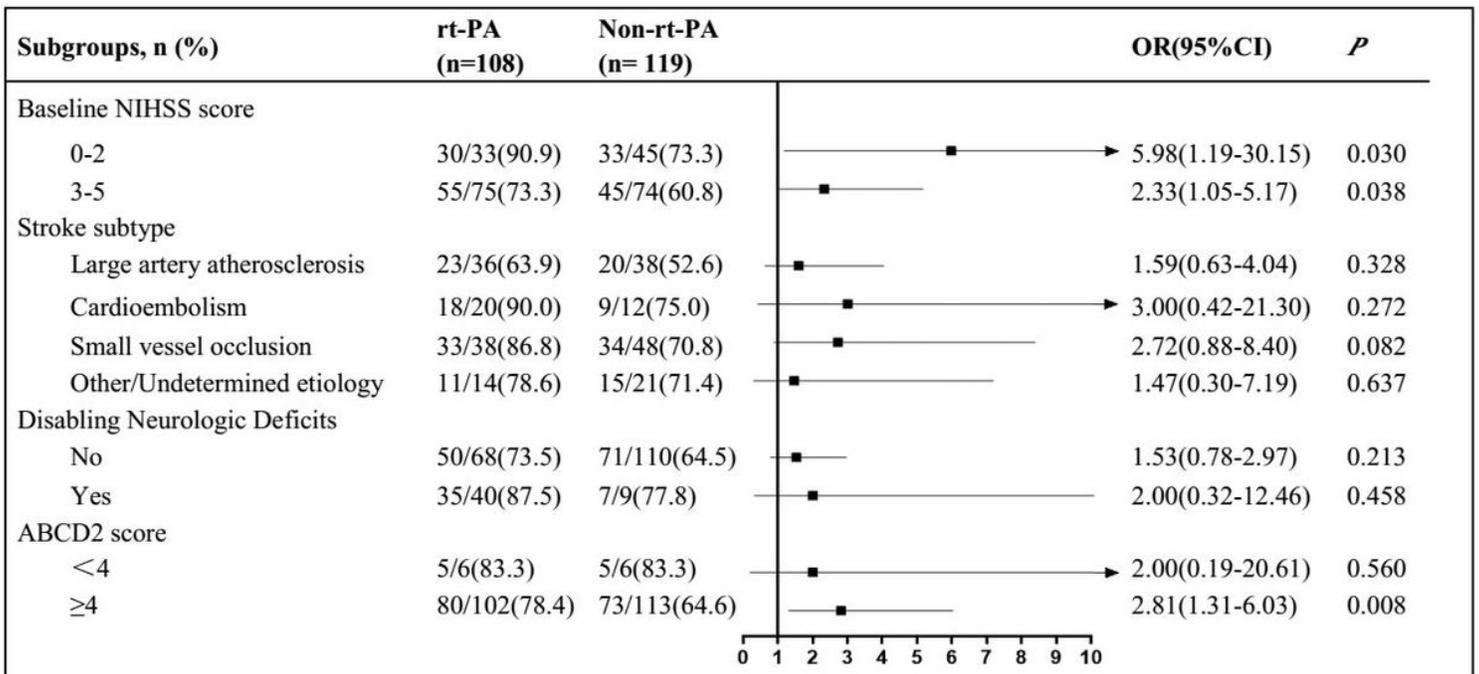


Figure 3

Subgroup Analysis with Mild Stroke for Excellent Outcome. Abbreviations: ABCD2 score (age, blood pressure, clinical features, duration of symptoms, and presence of diabetes mellitus; range, 0 to 7, with higher scores indicating a higher risk of stroke); CI, confidence interval; and OR, odds ratio.