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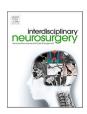
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# The association between diffusion-weighted imaging-Alberta Stroke Program Early Computed Tomography Score and the outcome following mechanical thrombectomy of anterior circulation occlusion

Hiroaki Hashimoto <sup>a,b,\*</sup>, Tomoyuki Maruo <sup>a</sup>, Yuki Kimoto <sup>a</sup>, Masami Nakamura <sup>a</sup>, Takahiro Fujinaga <sup>a</sup>, Hajime Nakamura <sup>c</sup>, Yukitaka Ushio <sup>a</sup>

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#### ABSTRACT

Background: Although preoperative diffusion-weighted imaging-Alberta Stroke Program Early Computed Tomography Score (DWI-ASPECTS) is well known as a predictor of outcomes after mechanical thrombectomy (MT) for large-vessel occlusion (LVO), assessment of changes in DWI-ASPECT from before to after MT is rare. Therefore, we clarified the relationship between the change in DWI-ASPECTS and clinical outcomes.

Methods: In this retrospective single-center study, we enrolled 63 cases of anterior LVOs treated with MT between April 2015 and March 2022. Preoperative and postoperative DWI-ASPECTSs were calculated. DWI-ASPECTSs were categorized into cortical-ASPECTSs (c-ASPECTSs) and subcortical ASPECTSs and assessed. Additionally, medical variables related to patients, such as sex, age, National Institutes of Health Stroke Scale (NIHSS) score, and premorbid modified Rankin Scale (mRS) score, were evaluated. A good outcome was defined as an mRS score of 0–2 at 3 months.

Results: Forty-five patients met the inclusion criteria. Nine (20 %) had a good outcome. The good outcome group showed significantly higher postoperative DWI-ASPECTs (median 8 vs. 5, p=0.012) and c-ASPECTSs (median 4 vs. 3, p=0.020) than the poor outcome group. No difference in DWI-ASPECTSs and c-ASPECTSs from before to after MT were significantly associated with the good outcome (p=0.017, p=0.016, respectively). The cut-off values for the good outcome on receiver operating characteristic curve analysis for differences between DWI-ASPECTSs and c-ASPECTSs were 0 [area under the curve (AUC) 0.77] and 0 [AUC 0.74]. Logistic regression analyses showed that baseline NIHSS score (odds ratio, 0.69; 95 % confidence interval 0.48–1.00; p=0.046) and postoperative DWI-ASPECTS (odds ratio, 2.27; 95 % confidence interval 1.02–5.04; p=0.039) were independent factors for the good outcome.

Conclusions: The good outcome of patients with anterior LVO was associated with no difference in DWI-ASPECTSs and c-ASPECTSs from before to after MT.

# 1. Introduction

Diffusion-weighted imaging (DWI)-Alberta Stroke Program Early

Computed Tomography Score (ASPECTS) and DWI lesion volume ( $Vol_{DWI}$ ) are negatively correlated [1,2], and in acute ischemic stroke (AIS) with large-vessel occlusion (LVO), DWI-ASPECTS 4 and  $Vol_{DWI}$  71

Abbreviations: AHA, American Heart Association; ASA, American Stroke Association; ASPECTS, Alberta Stroke Program Early Computed Tomography Score; AIS, acute ischemic stroke; c-ASPECTS, cortical ASPECTS; CCA, common carotid artery; CI, confidence interval; COV, cut-off value; DICOM, Digital Imaging and Communications in Medicine; DWI, diffusion-weighted imaging; ICA, internal carotid artery; IQR, interquartile range; LVO, large-vessel occlusion; MCA, middle cerebral artery; MRI, magnetic resonance imaging; mRS, modified Rankin Scale; MT, mechanical thrombectomy; mTICI, modified Thrombolysis in Cerebral Infarction; NIHSS, National Institutes of Health Stroke Scale; OR, odds ratios; ROC, receiver operating characteristic; SAH, subarachnoid hemorrhage; sc-ASPECTS, subcortical ASPECTS; tPA, tissue plasminogen activator; Vol<sub>DWI</sub>, DWI lesion volume.

<sup>&</sup>lt;sup>a</sup> Department of Neurosurgery, Otemae Hospital, Osaka, Osaka 540-0008, Japan

b Department of Neurological Diagnosis and Restoration, Graduate School of Medicine, Osaka University, Suita, Osaka 565-0871, Japan

<sup>&</sup>lt;sup>c</sup> Department of Neurosurgery, Graduate School of Medicine, Osaka University, Suita, Osaka 565-0871, Japan

<sup>\*</sup> Corresponding author at: Neurosurgeon/Medical Staff, Department of Neurosurgery, Otemae Hospital, Osaka, 540-0008, Japan. E-mail address: h-hashimoto1986@otemae.gr.jp (H. Hashimoto).

mL were suggested as cut-off values for the prediction of a poor outcome [3]. Previous studies reported that mechanical thrombectomy (MT) was effective for LVO patients who were within 6-12 h from the onset and showed ASPECTS  $\geq$  6 or 7/DWI-ASPECTS  $\geq$  6 [4–8], and recently, DAWN [9] and DEFUSE 3 [10] trials suggested that even in LVO over 6 h from the onset, MT was effective for patients whose ischemic core volume was <51 mL or 70 mL, respectively. Therefore, tissue-based evaluation of AIS is indispensable for the optimal indication of MT, and the correlation between ASPECTS and ischemic core volume was fair [11]. Therefore, high ASPECTS/DWI-ASPECTS is reported to be a predictor of good outcomes among patients undergoing MT [5,7,12]. In the American Heart Association (AHA)/American Stroke Association (ASA) guideline, MT is recommended for AIS-LVO based hours of symptom onset and ASPECTS, and MT is not recommended in AIS-LVO with AS-PECTS < 6 [13]. At our institution, MT has been chosen according to the guidelines, expecting good results. Unfortunately, although we adhered to these guidelines, some AIS patients undergoing MT showed poor outcomes. Additionally, we have frequently experienced DWI-ASPECTS worsening from before to after MT. Therefore, many studies focused on DWI-ASPECTS before MT to reveal factors to predict good or poor outcomes and to help physicians select AIS-LVO patients who would benefit from MT [11,14-20]. However, studies investigating DWI-ASPECTS after MT are few, and it remains unclear how changing DWI-ASPECTS from before to after MT relates to the outcomes of AIS patients under-

In this study, we enrolled patients with anterior circulation occlusion-related AIS who had MT treatment to clarify factors related to good outcomes after MT by investigating changes in DWI-ASPECTSs from before to after MT. Prediction of outcomes after MT is crucial for physicians to plan the treatment and rehabilitation for AIS patients.

#### 2. Methods

# 2.1. Patients and study setting

In this retrospective study, we enrolled patients who underwent MT because of AIS at Otemae Hospital between April 2015 and March 2022. We referred to the AHA/ASA guideline [13], and the following criteria for patient enrollment were set: (1) underwent magnetic resonance imaging (MRI) both before and after MT and completion of postoperative MRI within 21 postoperative days; (2) occlusion of anterior circulation including common carotid artery (CCA), internal carotid artery (ICA), M1 and M2 segments of the middle cerebral artery (MCA); (3) MT within 24 h of AIS onset; (4) preoperative DWI-ASPECTS  $\geq$  6 or preoperative DWI-ASPECTS < 6 within 6 h of AIS onset; (5) patients with clinical outcomes obtained at 3 months after onset. If the symptom onset was unknown, the onset was defined as the time when patients were last known to be in wellness. The Ethics Committee of the Otemae Hospital (Osaka, Japan, approval no. CT210421002) gave the ethical approval for this work, and it was conducted following the Declaration of Helsinki guidelines for experiments involving humans. Informed consent was obtained using the opt-out method from our center's website because of the retrospective and noninvasive nature of the study.

# 2.2. Data collection

We retrospectively collected and evaluated medical variables related to patients [sex, age, National Institutes of Health Stroke Scale (NIHSS) score [21], and premorbid modified Rankin Scale (mRS) score [22]]. Imaging findings (laterality of AIS and arterial occlusion site), treatment details [retrieval attempts, onset to reperfusion time, modified Thrombolysis in Cerebral Infarction (mTICI) scale grade, and combination with tissue plasminogen activator (tPA)], and outcomes [mRS score three months after onset, hemorrhagic infarction, and subarachnoid hemorrhage (SAH)] were included. We defined a good outcome as an mRS

score 0–2 at 3 months [23] and successful reperfusion after MT as mTICI grade 2b or 3 [24]. Two stroke specialists certified by the Japan Stroke Society independently evaluated DWI-ASPECTS (H.H. and T.M.). The DWI-ASPECTS was categorized into subcortical ASPECTS (sc-ASPECTS) with five structures (the caudate, lentiform nucleus, internal capsule, insular ribbon, and white matter) and cortical ASPECTS (c-ASPECTS) with six structures in MCA cortical regions [23]. For quantitative assessment of Vol<sub>DWI</sub>, Digital Imaging and Communications in Medicine (DICOM) DWI-MRI date were imported to MATLAB R2020b (Math-Works, Natick, MA, USA), and lesions identified by DWI were segmented manually using the image segmenter app in MATLAB (https://www.mathworks.com/help/images/ref/imagesegmenter-app.html). These procedures enabled us to calculate Vol<sub>DWI</sub>.

# 2.3. Statistical analyses

Categorical data and continuous variables are presented as patients (percentages) or median (interquartile range (IQR)), respectively. Clinical differences between the good and poor outcome groups were assessed using the chi-squared test for categorical variables. The two-tailed Wilcoxon rank-sum test was used for continuous variables. The correlation coefficient was calculated using Spearman correlation analysis. The receiver operating characteristic (ROC) curve was used to perform the sensitivity and specificity. The cut-off value (COV) was defined as the maximal Youden index (sensitivity + specificity - 1). A logistic regression model was used to calculate the odds ratios (OR) with 95 % confidence intervals (CIs). The significance level was 0.05. All statistical analyses were performed using the Statistical and Machine Learning Toolbox of MATLAB R2020b (MathWorks, Natick, MA, USA).

# 2.4. Data availability

All data in this study are available from the corresponding authors upon reasonable request and after additional ethics approval.

#### 3. Results

# 3.1. Baseline characteristics

Forty-five patients fulfilled the inclusion criteria of the 63 patients treated within the study period. The median (IQR) age was 76 years (62-86), and 47 % were females. The frequency of preoperative DWI-ASPECTS < 6 was 7 (16 %). The rate of good clinical outcomes was 20 % (9/45 patients). Successful reperfusion, defined as mTICI grade 2b or 3, was achieved in 82 %. Table 1 indicates the baseline characteristics, procedural variables, and complications between the good and poor outcome groups. No significant differences were noted between the two groups regarding the sex distribution, premorbid mRS score, laterality, arterial occlusion site, retrieval attempts, onset to reperfusion time, mTICI grade, combination with tPA therapy, frequency of preoperative DWI-ASPECTS < 6, and incidence of SAH. However, in the good outcome group, the age and NIHSS scores were significantly lower than those in the poor outcome group (p = 0.00097 and 0.00016, respectively: Wilcoxon rank-sum test). Additionally, the percentage of good outcomes in hemorrhagic infarctions was statistically lower than that in no hemorrhagic infarctions (8 % vs. 33 %, p = 0.036: chi-squared test).

### 3.2. DWI-ASPECTS and Vol<sub>DWI</sub>

There were significant negative correlations between DWI-ASPECTS and  $\mathrm{Vol}_{\mathrm{DWI}}$  before (correlation coefficients; -0.88,  $p=1.8\times10^{\cdot15}$ ) and after MT (correlation coefficients; -0.86,  $p=2.6\times10^{\cdot14}$ ). Fig. 1 indicates the box-and-whisker plot between DWI-ASPECTS and  $\mathrm{Vol}_{\mathrm{DWI}}$  before and after MT. In the preoperation and postoperation, all  $\mathrm{Vol}_{\mathrm{DWI}}$  of DWI-ASPECTS  $\geq 6$  showed uneven distribution; however, in postoperation, DWI-ASPECTS < 6

**Table 1**Descriptive statistics regarding baseline, procedural, and outcome parameters.

Characteristics	Patients n (%)	Good outcome (% or IQR)	Poor outcome (% or IQR)	<i>p</i> -value
Sex				
Male	24 (53)	6 (25)	18 (75)	0.37
Female	21 (47)	3 (14)	18 (86)	
Age (years)		61 (49-71)	83 (76-89)	*0.00097
NIHSS		12 (6-14)	19 (17-22)	*0.00016
Premorbid mRS score		0 (0–1)	1 (0–3)	0.18
Laterality				
Left	25 (56)	4 (16)	21 (84)	0.45
Right	20 (44)	5 (25)	15 (75)	
Arterial occlusion sit	te			
CCA + ICA +	32 (71)	8 (25)	24 (75)	0.19
MCA-M1				
MCA-M2	13 (29)	1 (8)	12 (92)	
Retrieval attempts		1 (1-3)	2 (1-2)	0.98
Onset to reperfusion		8 (4–11)	5 (4–8)	0.25
time (hours)				
mTICI grade				
2b + 3	37 (82)	7 (19)	30 (81)	0.70
Others	8 (18)	2 (25)	6 (75)	
tPA therapy				
Yes	18 (40)	2 (11)	16 (89)	0.22
No	27 (60)	7 (26)	20 (74)	
Preoperative DWI-AS	SPECTS			
≥6	38 (84)	9 (24)	29 (76)	0.15
<6	7 (16)	0 (0)	7 (100)	
Hemorrhagic infarct	ion			
Yes	24 (53)	2 (8)	22 (92)	*0.036
No	21 (47)	7 (33)	14 (67)	
SAH				
Yes	13 (29)	1 (8)	12 (92)	0.19
No	32 (71)	8 (25)	24 (75)	

Patients (%) or median (IQR) are demonstrated for categorical data or continuous variables, respectively. P values were calculated using the chi-squared test for categorical variables and the Wilcoxon rank-sum test for continuous variables. Statistically significant p values are flagged with an asterisk.

ASPECTS indicates Alberta Stroke Program Early Computed Tomography Score; CCA, common carotid artery; DWI, diffusion-weighted imaging; ICA, internal carotid artery; IQR, interquartile range; MCA-M1, middle cerebral artery-M1 segment; MCA-M2, middle cerebral artery-M2 segment; mRS, modified Rankin Scale; mTICI, modified Thrombolysis in Cerebral Infarction; NIHSS, National Institutes of Health Stroke Scale; SAH, subarachnoid hemorrhage; and tPA, tissue plasminogen activator.

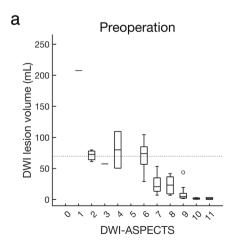
increased. In preoperation, DWI-ASPECTS, c-ASPECTS, and  $Vol_{DWI}$  values showed no significant differences between the good and poor outcome groups (Table 2). However, postoperative DWI-ASPECTS and c-ASPECTS were significantly higher in the good outcome group than in the poor outcome group (p=0.012 and 0.020, respectively: Wilcoxon rank-sum test), and postoperative  $Vol_{DWI}$  was significantly lower in the

**Table 2** Pre- and postoperative DWI-ASPECTS.

Characteristics	Patients n (%)	Good outcome (%)	Poor outcome (%)	<i>p</i> -value
Days of postoperative		1 (1–6)	1 (1–5)	0.95
MRI (days)				
Preoperation				
DWI-ASPECTS		8 (7–9)	7 (6–9)	0.47
sc-ASPECTS		4 (3-4)	3 (2-4)	0.64
c-ASPECTS		4 (4–6)	5 (2–6)	0.56
Vol <sub>DWI</sub> (mL)		9 (6.3-35.0)	27 (4.6-59.4)	0.31
Postoperation				
DWI-ASPECTS		8 (7–9)	5 (3-8)	*0.012
sc-ASPECTS		3 (3-4)	2 (1-4)	0.097
c-ASPECTS		4 (4–6)	3 (0-4)	*0.020
Vol <sub>DWI</sub> (mL)		13	57	*0.0057
		(6.8-23.4)	(21.8-123.2)	
Differences				
Pre - Postoperative		0 (0-0)	1 (0-3)	*0.011
DWI-ASPECTS				
No difference	16 (36)	6 (37.5)	10 (62.5)	*0.017
(differences = 0)				
Worsening	26 (58)	2 (8)	24 (92)	
(differences > 0)				
Pre - Postoperative		0 (0-0)	0 (0-1)	0.20
sc-ASPECTS				
Pre - Postoperative		0 (0-0)	0 (0-2)	*0.013
c-ASPECTS				
No difference	27 (60)	8 (30)	19 (70)	*0.016
(differences = 0)				
Worsening 16 (36)		0 (0)	16 (100)	
(differences > 0)				
Post - Preoperative		1 (-2.7-3.1)	18 (4.8-55.0)	*0.00086
Vol <sub>DWI</sub> (mL)				

Patients (%) or median (IQR) are demonstrated for categorical data or continuous variables, respectively. *P* values were calculated using the chi-squared test for categorical variables and the Wilcoxon rank-sum test for continuous variables. Statistically significant *p* values are flagged with an asterisk.

ASPECTS indicates Alberta Stroke Program Early Computed Tomography Score; c-ASPECTS, cortical ASPECTS; DWI, diffusion-weighted imaging; IQR, interquartile range; MRI, magnetic resonance imaging; sc-ASPECTS, subcortical ASPECTS; and  $Vol_{DWI}$ , DWI lesion volume.



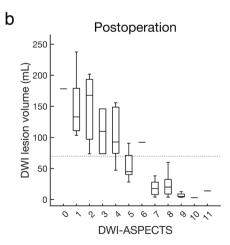


Fig. 1. Box-and-whisker plot of DWI-ASPECTS and DWI lesion volume (Vol<sub>DWI</sub>). (a) Values at preoperation. (b) Values at postoperation. Broken horizontal lines indicate 70 mL of Vol<sub>DWI</sub>. Boxes indicate interquartile range; whiskers, extreme values; horizontal lines in each box, median; and a circle, outlier. DWI indicates diffusion-weighted imaging; and ASPECTS, Alberta Stroke Program Early Computed Tomography Score.

good outcome group than in the poor outcome group (p=0.0057, Wilcoxon rank-sum test).

Next, we evaluated the differences between preoperative and postoperative values. In the good outcome group, the difference obtained from the preoperative minus the postoperative DWI-ASPECTS and c-ASPECTS was significantly lower than that in the poor outcome group (p = 0.011 and 0.013, respectively: Wilcoxon rank-sum test). Moreover, in no difference groups, both DWI-ASPECTS and c-ASPECTS showed a significantly higher rate of a good outcome than worsening groups (p =0.017 and 0.016, respectively: chi-squared test). The difference from the postoperative Vol<sub>DWI</sub> minus the preoperative Vol<sub>DWI</sub> in the good outcome group was significantly smaller than that in the poor outcome group (p = 0.00086: Wilcoxon rank-sum test). Fig. 2 indicates that the good outcome group showed no worsening of DWI-ASPECTS, c-AS-PECTS, and Vol<sub>DWI</sub> from preoperation to postoperation. Although the results of c-ASPECTS were similar to those of DWI-ASPECTS, results obtained from sc-ASPECTS showed no significance in preoperation, postoperation, and the difference. Additionally, we assessed the statistical difference between the no-difference group and the worsening group of c-ASPECTS (Supplemental Table S1). In the worsening group of c-ASPECTS, the baseline NIHSS and the retrieval attempts were significantly high (p = 0.032 and 0.045, respectively, Wilcoxon rank-sum test).

# 3.3. Factors for a good outcome

Table 3 indicates that NIHSS and postoperative DWI-ASPECTS were independent factors for a good outcome (OR = 0.69; 95 % CI, 0.48–1.00; p=0.046, OR = 2.27; 95 % CI, 1.02–5.04; p=0.039, respectively), as revealed by multivariate logistic regression analysis.

# 3.4. ROC analysis

The COV of age and NIHSS by ROC curve analysis for predicting good outcomes was 75 years old (area under the curve (AUC) 0.87; sensitivity 0.72, specificity 0.89) and 13 (AUC 0.92; sensitivity 0.92, specificity 0.78), respectively (Fig. 3a). The ROC curves obtained from DWI-ASPECTS and c-ASPECTS showed similarities. The COV obtained from the differences of DWI-ASPECTS and c-ASPECTS were 0 (AUC 0.77; sensitivity 0.67, specificity 0.78) and 0 (AUC 0.74; sensitivity 0.44, specificity 1.00), respectively (Fig. 3b and 3c). In Vol<sub>DWI</sub>, ROC curves showed high performance; the COV of postoperation and difference was 20 mL (AUC 0.83; sensitivity 0.78, specificity 0.78), and 8 mL (AUC 0.88; sensitivity 0.69, specificity 1.00), respectively (Fig. 3d).

**Table 3**Multivariate logistic regression analysis for predictors of good outcomes (mRS 0 to 2).

Characteristics	OR (95 % CI)	<i>p</i> -value
Age (years)	0.91 (0.81–1.01)	0.077
NIHSS	0.69 (0.48–1.00)	*0.046
Postoperative DWI-ASPECTS	2.27 (1.02–5.04)	*0.039

Statistically significant p values are flagged with an asterisk.

ASPECTS indicates Alberta Stroke Program Early Computed Tomography Score; CI, confidence interval; DWI, diffusion-weighted imaging; NIHSS, National Institutes of Health Stroke Scale; and OR, odds ratio.

# 4. Discussion

We hypothesized that an assessment of changing DWI-ASPECTS from preoperation to postoperation would enable us to reveal a novel insight for predicting the outcome of AIS after MT. We could show that the good outcome group showed statistically no-difference DWI-ASPECTS and c-ASPECTS from before to after MT. Postoperative DWI-ASPECTS and c-ASPECTS and their differences showed good performance for predicting good outcomes through ROC analyses. The cut-off values for predicting good outcomes were 75 years old for age, 13 for NIHSS, and 0 for differences of DWI-ASPECTS and c-ASPECTS. Logistic regression analyses showed that postoperative DWI-ASPECTS and NIHSS were independent predictive factors for a good outcome.

In our institute, the AHA/ASA Guideline [13] was referred for selecting AIS patients to provide proper treatment. The guideline recommends MT to AIS because of anterior circulation LVO patients who show (1) pre-stroke mRS score of 0 to 1; (2) age  $\geq$  18 years; (3) NIHSS score of  $\geq$  6; and (4) ASPECTS of  $\geq$  6. A low score of DWI-ASPECTS before MT was associated with poor outcome [14] and a high score of that associated with good outcome [20], and the ESCAPE and REVAS-CAT trial excluded AIS patients with ASPECTS < 6-7 or DWI-ASPECTS < 6 [5,8]. Thus, MT is not commonly recommended for large-volume AIS, and our results showed that DWI-ASPECTS < 6 indicated zero percent of a good outcome. In this study, we set the inclusion criteria using preoperative DWI-ASPECTS. Our results showed no significant difference in preoperative DWI-ASPECTS between the good and poor groups and that the median values of preoperative DWI-ASPECTS were over 6 scores. Our inclusion criteria that excluded the large volume AIS influenced these results, representing a selection bias.

Although preoperative DWI-ASPECTS has frequently been assessed as a factor for predicting outcomes after reperfusion therapy for AIS [1,3,12,16,18,19,25–27], studies investigating a relationship between the change in DWI-ASPECTS and reperfusion therapy for AIS are rare.

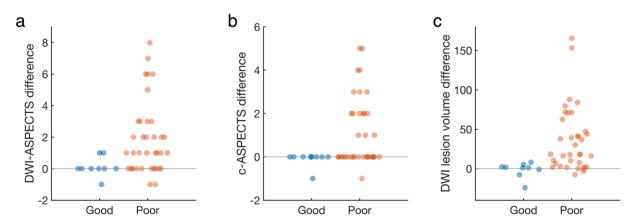


Fig. 2. Beeswarm plot between the good and poor groups The distribution patterns of differences between preoperation and postoperative values are shown. Differences obtained from preoperation minus postoperative values were used in DWI-ASPECTS (a), and c-ASPECTS (b). Differences obtained from postoperative minus preoperative values were used in DWI lesion volumes (c). ASPECTS indicates Alberta Stroke Program Early Computed Tomography Score; c-ASPECTS, cortical ASPECTS; and DWI, diffusion-weighted imaging.

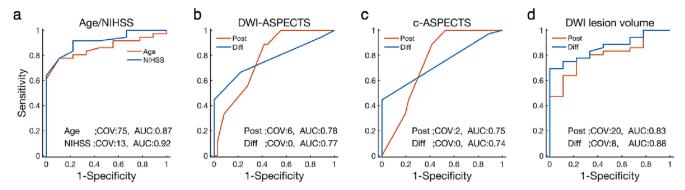


Fig. 3. Receiver operating characteristic (ROC) curves. ROC curves for predicting good outcomes (mRS: 0 or 2) are indicated and calculated from age and NIHSS (a), DWI-ASPECTS (b), c-ASPECTS (c), and Vol<sub>DWI</sub> (d). Cut-off values (COV) and area under the curve (AUC) are shown. ASPECTS indicates Alberta Stroke Program Early Computed Tomography Score; c-ASPECTS, cortical ASPECTS; Diff, differences; DWI, diffusion-weighted imaging; mRS, modified Rankin Scale; NIHSS, National Institutes of Health Stroke Scale; Post, postoperation; and Vol<sub>DWI</sub>, DWI lesion volume.

The ischemic brain lesion extension on 24 h was reported to be associated with the outcome of LVO-AIS [16]. Our ROC analyses could add a novel insight that no-difference of DWI-ASPECTS from before to after MT distinguished good outcomes from the poor outcomes. Moreover, we showed that the performance of the ROC curves in DWI-ASPECTS was similar to that in c-ASPECTS and that no difference in c-ASPECTS was associated with a good outcome (Table 2). A previous study showed a strong correlation between high c-ASPECTS and good outcomes in LVO-AIS, such as ASPECTS < 6 [23]. Therefore, while this study indicated that postoperative DWI-ASPECTS and c-ASPECTS were associated with outcomes, we inferred that a true factor influencing the outcome after MT was c-ASPECTS. We propose that if physicians experience a worsening of c-ASPECTS after MT, they should be careful of poor outcomes.

While accuracy for identifying AIS was higher in DWI than in computed tomography (CT) [28], initial CT alone was reported to be non-inferior to initial CT plus additional MRI concerning outcomes [29]. Moreover, since we showed that postoperative DWI-ASPECTS could distinguish good outcomes from poor outcomes, we think an MRI examination before MT could be skipped to shorten the time from examination to treatment.

Recently, several studies demonstrated that MT was effective for even large-volume ischemic strokes, such as preoperative ASPECTS  $\leq 5$  [23]/DWI-ASPECTS  $\leq 5$  [25,30]. However, in large ischemic core stroke, older age [25] and large  $Vol_{DWI}$  [25] was associated with poor outcomes, and successful recanalization [25,30–32] was associated with a good outcome. Moreover, even at ASPECTS < 6, high c-ASPECTS were reported to be a positive factor for good outcomes [23]. Therefore, in our institute, even though patients showed large ischemic volumes such as DWI-ASPECT < 6, we treated them by MT to realize successful recanalization. However, while we could achieve approximately 80 % successful recanalization in this study, no patients showed good outcomes in preoperative DWI-ASPECTS < 6.

According to our results, the age of the poor outcome group was significantly higher than that of the good outcome group. Patients aged  $\geq 80$  years after MT had worse functional outcomes and showed a lower percentage of successful recanalization [33]. However, studies between 2017 and 2019 indicated better outcomes even in older patients [33], and the efficacy of MT for patients aged  $\geq 80$  years over standard care was demonstrated [15,34]. A small infarct core and low NIHSS were associated with good outcomes in patients over 80 years of age [20]. Although the efficacy of MT for AIS patients with low NIHSS scores is controversial, NIHSS was reported as a factor associated with outcomes after MT [16]. A low NIHSS score was related to successful recanalization [35]. We also showed that NIHSS was an independent factor for predicting good outcomes by the multivariate logistic regression analysis. Another factor related to the outcome after MT was the first-pass effect of MT, which is related to a favorable outcome [36]. A previous

study indicated that the first three retrieval attempts were associated with improved clinical outcomes [37]. Our study could achieve less than three retrieval attempts in both the good and poor outcome groups.

We demonstrated a strong negative correlation between DWI-ASPECTS and  $Vol_{DWI}$ . All patients with DWI-ASPECTS  $\geq$  7, both preoperation and postoperation, had  $Vol_{DWI} < 70$  mL, in concordance with a previous study [1]. Additionally, ASPECTS correlated with ischemic core volume [11], and  $Vol_{DWI}$  correlated strongly with final infarct volume [28]. Therefore, DWI-ASPECTS and  $Vol_{DWI}$  are well-known factors related to the outcome of reperfusion therapy [3,19,27]. We added new results that both preoperative and postoperative DWI-ASPECTS  $\geq$  7 may be reliable surrogates of  $Vol_{DWI} < 70$  mL.

Regarding sites of occlusion arteries, the M2 segment of MCA was assessed by MR CLEAN [38], ESCAPE [5], EXTEND-IA [6], and SWIFT PRIME [7] and was not assessed by REVASCAT [8], DAWN [9], and DEFUSE3 [10]. Although the efficacy of MT for the M2 segment has been unclear [15], some studies showed the effectiveness of MT for LVO of the M2 segment [39–42]. AIS attributed to the M2 segment of the MCA is known as medium vessel occlusion (MeVO) [43]. While randomized studies have proven the safety and efficacy of MT for LVO [5–8,38], MT for MeVO has had no high-level evidence. However, since MeVO was observed in 25 %–40 % AIS [44] and showed substantial morbidity and the safety and efficacy of MT for MeVO have been presented [45,46], MT is increasingly performed for AIS-MeVO [47] in actual clinical situations. Therefore, our study included M2 occlusion, and there was no significant difference in arterial occlusion site, while a low percentage of the good outcome was observed in the M2 occlusion.

This study has some limitations. First, a small amount of our data was retrospectively collected from a single center. No control group treated with conventional medical care was enrolled, and the study was nonrandomized. Second, we focused on only AIS because of the anterior circulation occlusion, including CCA, ICA, M1, and M2 segments. Therefore, our findings could not be applied to AIS because of the posterior circulation occlusion. Third, we defined MRI obtained within 21 postoperation days as postoperative MRI. The number 21 was decided by our empirical experience, and it remains unclear how long after MT was useful for predicting outcomes. Finally, we did not measure the volume of the ischemic penumbra area before MT. While we revealed that the worsening of the c-ASPECTS group showed a statistically high score of NIHSS, it is possible that the worsening of c-ASPECTS is associated with the volume of the ischemic penumbra area because the volume of perfusion-weighted MRI correlated positively with NIHSS score [48].

In conclusion, an assessment of changes in DWI-ASPECTS from before to after MT revealed that the good outcome group showed no difference in DWI-ASPECTS and c-ASPECTS. The independent factors for predicting a good outcome after MT were low NIHSS and high postoperative DWI-ASPECTS. The cut-off values for a good outcome were 75 years old, NIHSS of 13, and a difference in DWI-ASPECTS of 0.

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#### CRediT authorship contribution statement

Hiroaki Hashimoto: Conceptualization, Methodology, Software, Validation, Formal analysis, Investigation, Data curation, Writing – original draft, Writing – review & editing, Visualization, Project administration, Funding acquisition. Tomoyuki Maruo: Conceptualization, Resources, Writing – review & editing, Supervision, Project administration. Yuki Kimoto: Investigation. Masami Nakamura: Investigation. Takahiro Fujinaga: Investigation. Hajime Nakamura: Writing – review & editing. Yukitaka Ushio: Supervision.

# **Declaration of Competing Interest**

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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# Appendix A. Supplementary material

Supplementary data to this article can be found online at https://doi.org/10.1016/j.inat.2023.101758.

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