

## **SUPPLEMENTAL MATERIAL**

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**Suppl. Table I.** Characteristics of analysed and patients in the MR CLEAN trial

	<b>Analysed patients (n=344)</b>	<b>MR CLEAN patients (n=500)</b>	<b>p</b>
<b>Allocation to intervention group (%)</b>	156 (45.3)	233 (46.6)	0.22
<b>Age (median [IQR])</b>	66 [56-76]	66 [56-76]	0.64
<b>Sex (male) (%)</b>	204 (59.3)	292 (58.4)	0.85
<b>NIHSS at baseline (median [IQR])</b>	18 [14-22]	18 [14-22]	0.83
<b>Previous stroke (%)</b>	41 (11.9)	54 (10.8)	0.69
<b>Atrial fibrillation (%)</b>	92 (26.7)	135 (27.0)	1
<b>Diabetes mellitus (%)</b>	48 (14.0)	68 (13.6)	0.96
<b>Smoking (%)</b>	101 (29.4)	143 (28.6)	0.87
<b>Myocardial infarction (%)</b>	56 (16.3)	76 (15.0)	0.74
<b>Hypertension (%)</b>	149 (43.3)	227 (45.4)	0.60
<b>Pre-stroke mRS ≤2 (%)</b>	312 (90.7)	479 (95.8)	1
<b>Systolic blood pressure at baseline (mmHg) (median [IQR])</b>	143 [130-160]	143 [130-160]	0.88
<b>Treatment with IV alteplase</b>	304 (88.4)	445 (89.0)	0.86
<b>ASPECTS ≥8 at baseline (%)</b>	269 (78.9)	373 (74.6)	0.25
<b>Location of intracranial occlusion (%)</b>			0.80
ICA	4 (1.2)	4 (0.8)	
ICA-T	90 (26.2)	134 (26.9)	
M1	213 (61.9)	319 (63.9)	
M2	34 (9.9)	39 (7.8)	
A1 or A2	3 (0.9)	3 (0.6)	
<b>Collateral status on baseline CTA (%)</b>			1
Absent collaterals	19 (5.6)	26 (5.2)	
Poor collaterals	94 (27.7)	136 (27.2)	
Moderate collaterals	134 (39.5)	198 (39.6)	
Good collaterals	92 (27.1)	133 (26.6)	
<b>Extracranial ICA&gt;50% stenosis at symptomatic side (%)</b>	40 (11.6)	57 (11.4)	1
<b>Extracranial ICA occlusion at symptomatic side (%)</b>	25 (7.3)	39 (7.8)	0.88
<b>Time from stroke onset to randomization (minutes) (median [IQR])</b>	189[145-248]	200 [150-261]	0.08

**Suppl. Table II.** Characteristics of patients according pattern of intracranial carotid artery calcification

	<b>Intimal dominance (n=118)</b>	<b>Medial dominance (n=152)</b>	<b>p</b>
<b>Allocated to intervention group</b>	57 (48.3)	65 (42.8)	0.433
<b>Age(median [IQR])</b>	63 [54-70]	76 [66-80]	<0.001
<b>Sex (male) (%)</b>	81 (68.6)	81 (53.3)	0.015
<b>NIHSS at baseline (median [IQR])</b>	18[15-21]	18[14-22]	0.330
<b>Previous stroke (%)</b>	11 (9.3)	22 (14.5)	0.274
<b>Atrial fibrillation (%)</b>	28 (23.7)	56 (36.8)	0.030
<b>Diabetes mellitus (%)</b>	12 (10.2)	31 (20.4)	0.035
<b>Smoking (%)</b>	42 (35.6)	35 (23.0)	0.033
<b>Myocardial infarction (%)</b>	11 (9.3)	40 (26.3)	0.001
<b>Hypertension (%)</b>	43 (36.4)	88 (57.9)	<0.001
<b>Pre-stroke mRS ≤2 (%)</b>	112 (94.9)	130 (85.5)	0.021
<b>Systolic blood pressure at baseline (mmHg) (median [IQR])</b>	140 [128-155]	152 [135-166]	<0.001
<b>Treatment with IV alteplase</b>	107 (90.7)	128 (84.2)	0.087
<b>ASPECTS ≥8 at baseline (%) *</b>	90 (76.9)	121 (79.6)	0.703
<b>Location of intracranial occlusion (%)</b>			0.366
ICA	1 (0.8)	3 (2.0)	
ICA-T	35 (29.7)	33 (21.7)	
M1	67 (56.8)	99 (65.1)	
M2	14 (11.9)	17 (11.2)	
A1 or A2	1 (0.8)	0	
<b>Collateral status on baseline CTA(%) †</b>			0.038
Absent collaterals	6 (5.2)	10 (6.6)	
Poor collaterals	24 (20.7)	53 (35.1)	
Moderate collaterals	56 (48.3)	51 (33.8)	
Good collaterals	30 (25.9)	37 (24.5)	
<b>Extracranial ICA&gt;50% stenosis at symptomatic side</b>	11 (9.3)	24 (15.8)	0.166
<b>Extracranial ICA occlusion at symptomatic side</b>	14 (11.9)	8 (5.3)	0.081
<b>Time from stroke onset to randomization ‡</b>	195 [148-268]	186 [146-239]	0.294
<b>Recanalization status on follow up CTA (%) §</b>			0.622
No recanalization	20 (20.2)	29 (27.1)	
Incomplete recanalization, no distal contrast	4 (4.0)	4 (3.7)	
Incomplete recanalization but any distal contrast	13 (13.1)	10 (9.3)	
Complete recanalization	62 (62.6)	64 (59.8)	
<b>Follow-up (5-7 days) infarct volume (mL) (median [IQR]   </b>	65 [26-115]	83 [30-161]	0.083

\* Alberta Stroke Program Early Computed Tomography Score (ASPECTS) was missing for three patients (1%).

† Data was missing for 5 (1%) patients.

‡ Data was missing for 3 (1%) patients.

§ Data was missing for 64 (19%) patients of which 28 patients died.

|| Data on follow-up infarct volume on NCCT 5-7 days was missing for 24 patients (9%) of which 8 patients died before assessment.

**Suppl. Table III.** Proportions of functional independence stratified by reperfusion grades in EVT-treated patients

mTICI-score *	EVT-patients with intimal ICAC (n=57)		EVT-patients with medial ICAC (n=65)	
	N (%)	mRS 0-2 (%)	N (%)	mRS 0-2 (%)
0-1	9 (19)	0/9	9 (17)	1/9 (11)
2a	13 (27)	2/13 (15)	11 (21)	3/11 (27)
2b-3	26 (54)	10/26 (38)	32 (62)	13/32 (40)
cOR (95%CI)	1.03 (0.89-1.18)		1.21 (1.07-1.38)	

\*Score of 0 indicates no perfusion or anterograde flow beyond occlusion site, 1: penetration of contrast but not perfusion, 2A: some perfusion <50% of vascular territory, 2B: substantial perfusion ≥50, 3: complete perfusion of vascular territory.

To assess impact of reperfusion on functional outcome with ordinal logistic regression, complete scale of dependent (mRS) and independent (mTICI-score) variables were used.

**Suppl. Table IV.** Serious adverse events according pattern of intracranial carotid artery calcification for both treatment groups.

	Patient with intimal calcification pattern		Patient with medial calcification pattern	
	Intervention group (n=57)	Control group (n=61)	Intervention group (n=65)	Control group (n=87)
<b>Any serious adverse event *</b>	26 (46%)	17 (28%)	33 (51%)	44 (51%)
Symptomatic intracerebral haemorrhage	0	4 (7%)	5 (8%)	7 (8%)
Recurrent ischemic stroke	2 (4%)	0	5 (8%)	0
Progression of ischemic stroke	14 (25%)	10 (16%)	13 (20%)	14 (16%)
Pneumonia	6 (11%)	4 (6%)	11 (17%)	23 (26%)
Cardiac ischemia	1 (2%)	1 (2%)	0	3 (3%)
Extracranial haemorrhage	0	0	0	1 (1%)
Allergic complication	1 (2%)	0	0	0
Other complication	9 (16%)	5 (8%)	3 (5%)	16 (18%)

\* Only first events of one type are listed. Patients experiencing multiple events of one type have been counted once.

## **Appendix**

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