

Definition of protocol violations in the MR CLEAN LATE trial:

1. Cross-over:

- Subjects that were allocated to the control group, but eventually received groin puncture in order to undergo EVT
- Subjects that were allocated to the intervention group, but eventually did not receive groin puncture in order to undergo EVT.

2. Violations of the in- and exclusion criteria:

- Subjects < 18 years of age
- Subjects with a NIH Stroke Scale score < 2*
- Time of randomization <6h since last seen well or onset of symptoms
- Time of randomization >24h since last seen well or onset of symptoms
- Intracranial hemorrhage on baseline CT or MRI*
- No proximal intracranial anterior circulation occlusion (i.e. distal intracranial carotid artery or middle (M1/M2) cerebral artery) confirmed by neuro-imaging (CTA or MRA).*
- Insufficient collateral flow shown on baseline neuro-imaging (CTA).*
 - Collateral grade 0 (or absent collaterals)
 - And since 09-10-2020 also patients with collateral grade 1 (“poor collaterals” or >0% and ≤50% collateral flow)
- Pre-stroke mRS >2*
- Cerebral infarction in the previous 6 weeks with residual neurological deficits or signs of a large recent infarction on neuroimaging in the territory of the middle cerebral artery (MCA).*
- Clearly demarcated hypodensity in >1/3 of the MCA territory, consistent with current symptoms.*
- Participation in another medical or surgical intervention trial (excluding the MR ASAP and ARTEMIS).
- Eligibility for direct treatment (outside of this trial) with EVT beyond the 6 hour time window. This is the case when the subject adheres to all following criteria:*^
 - Distal intracranial carotid artery occlusion or M1 occlusion

- NIHSS ≥ 10
- Infarct core ≤ 25 ml
- Mismatch ratio total ischemic volume / infarct core ≥ 2
- Clinical evidence of hemorrhagic diathesis*
 - INR > 3
 - Platelet count $< 40 \times 10^9/L$
 - APTT > 50 sec

* For these criteria, the assessment of the local physicians at the moment of randomization will be leading in establishing whether or not the protocol has been violated.

^Eligibility for direct treatment was only after implementation of these criteria in the Dutch guidelines, where after these criteria were added as exclusion criteria for inclusion in the MR CLEAN LATE trial (15/03/2019)