

# Endovascular treatment for acute ischemic stroke in the Netherlands

|                                        |                                                   |                                                                                                   |
|----------------------------------------|---------------------------------------------------|---------------------------------------------------------------------------------------------------|
| <b>Submission date</b><br>03/06/2012   | <b>Recruitment status</b><br>No longer recruiting | <input type="checkbox"/> Prospectively registered<br><input checked="" type="checkbox"/> Protocol |
| <b>Registration date</b><br>24/07/2012 | <b>Overall study status</b><br>Completed          | <input type="checkbox"/> Statistical analysis plan<br><input checked="" type="checkbox"/> Results |
| <b>Last Edited</b><br>18/12/2023       | <b>Condition category</b><br>Circulatory System   | <input type="checkbox"/> Individual participant data                                              |

## Plain English summary of protocol

### Background and study aims

Ischaemic strokes occur when a blood clot blocks the flow of blood to the brain. Intra-arterial thrombolysis and mechanical thrombectomy is a new and promising treatment for patients with acute ischemic stroke. This treatment involves going into the brain arteries with a very small tube (catheter) to deliver treatment at the site of the blocked blood vessel that caused the stroke, reopening the blocked vessels and removing the clot. This study will investigate the effectiveness and safety of this treatment.

### Who can participate?

Patients with acute ischemic stroke who admitted to the ER of one of the participating medical centers within 6 hours from onset of symptoms.

### What does the study involve?

Participants are randomly allocated to receive either standard treatment for acute ischemic stroke or intra-arterial treatment as described above. Participants then undergo additional MRI or CT scans to check whether the blocked vessel has been reopened. After 1 week, a CT scan is carried out to assess the extent of the stroke. Blood is drawn at this time to look for blood clotting abnormalities. After 3 months, participants are approached by telephone to check on their general condition.

### What are the possible benefits and risks of participating?

Those who participate in the study will benefit from high level stroke assessment by experienced stroke neurologists. As the effectiveness of the treatment is not yet proven, the opportunity to receive this treatment cannot be called a benefit. The treatment is probably associated with an increased risk of bleeding inside the skull and at the site of the catheter insertion in the groin. Early studies so far suggest that the risks of the treatment are in balance with the expected benefit.

### Where is the study run from?

In 14 large hospitals in the Netherlands, listed on the trial website: <http://www.mrclean-trial.org/centers.htm>

When is the study starting and how long is it expected to run for?

December 2010 to January 2015

Who is funding the study?

The project is funded by the Dutch Heart Foundation, and by several companies, all of whom are listed on the trial website: <http://www.mrclean-trial.org/sponsors.htm>

Who is the main contact?

Prof. Diederik Dippel and Prof. Charles Majoie

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## Contact information

### Type(s)

Scientific

### Contact name

Prof Diederik Dippel

### Contact details

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## Additional identifiers

### Protocol serial number

NTR1804, NL 30557.078.10 v02, NL695

## Study information

### Scientific Title

Multicenter Randomized CLinical trial of Endovascular treatment for Acute ischemic stroke in the Netherlands

### Acronym

MR CLEAN

### Study objectives

The primary objective of this study is to estimate the effect of endovascular treatment on overall functional outcome after acute ischemic stroke of less than six hour duration, in patients with a symptomatic intracranial anterior circulation occlusion.

### Ethics approval required

Old ethics approval format

**Ethics approval(s)**

Medical Ethics Committee and Research Board of Erasmus MC University Medical Center

**Study design**

Multicenter randomized open-label treatment and blinded endpoint evaluation (PROBE design)

**Primary study design**

Interventional

**Study type(s)**

Treatment

**Health condition(s) or problem(s) studied**

Acute ischemic stroke, caused by intracranial proximal arterial occlusion (anterior circulation)

**Interventions**

Intervention arms consists of endovascular treatment by means of microcatheter guided local application of recombinant tissue plasminogen activator (rt-PA) or urokinase, and/or mechanical thrombectomy, by means of a retraction device, aspiration device, or retrievable stent.

The control arm consists of regular treatment according to current national clinical guidelines.

**Intervention Type**

Procedure/Surgery

**Primary outcome(s)**

Score on the modified Rankin scale at 3 months

**Key secondary outcome(s)**

Imaging parameters:

1. Vessel recanalization at 24 hours after treatment, assessed by CTA or MRA. The criteria for recanalization on CTA or MRA are based on a simplified TICI score (Table 5b),<sup>45</sup> and the clot burden score, proposed by Puetz et al (Table 5c)<sup>46</sup>.
2. Infarct size assessed by CT on day 5-7, using standard methods, including manual tracing of the infarct perimeter and semiautomated pixel thresholding.<sup>47, 48</sup> Infarct size at day 5-7 will be compared with plain CT and perfusion CT results (if available) at baseline.
3. CTA or MRA at 24 hours will be compared with baseline vessel imaging data, to estimate the recanalization rate. Perfusion CT at baseline is optional, but available at most centers.

Clinical parameters:

1. NIHSS, including NIH supplemental motor score, 50 at 24 hours
2. NIHSS at 1 week or at discharge.

Functional outcome:

1. Score on the EQ5D at 90 days
2. Barthel index at 90 days

**Completion date**

01/01/2015

# Eligibility

## Key inclusion criteria

1. A clinical diagnosis of acute stroke, with a deficit on the National Institutes of Health (NIH) stroke scale of 2 points or more
2. Computerised tomography (CT) or magnetic resonance imaging (MRI) scan ruling out intracranial hemorrhage
3. Intracranial arterial occlusion of the distal intracranial carotid artery or middle (M1/M2) or anterior (A1/A2) cerebral artery, demonstrated with Computed Tomography Angiography (CTA), magnetic resonance angiography (MRA), Digital subtraction angiography (DSA) or transcranial Doppler/duplex (TCD)
4. The possibility to start treatment within 6 hours from onset
5. Informed consent given
6. Age 18 or over

## Participant type(s)

Patient

## Healthy volunteers allowed

No

## Age group

Adult

## Lower age limit

18 years

## Sex

All

## Key exclusion criteria

General:

1. Arterial blood pressure > 185/110 mmHg
2. Blood glucose < 2.7 or > 22.2 mmol/L
3. Intravenous treatment with thrombolytic therapy in a dose exceeding 0.9 mg/kg alteplase or 90 mg
4. Intravenous treatment with thrombolytic therapy despite contra-indications, i.e. major surgery, gastrointestinal bleeding or urinary tract bleeding within the previous 2 weeks, or arterial puncture at a non-compressible site within the previous 7 days

For Intended mechanical thrombectomy:

Laboratory evidence of coagulation abnormalities, i.e. platelet count <40 x 10<sup>9</sup>/L, activated partial thromboplastin time (APTT)>50 sec or International Normalised Ratio (INR) >3.0.

For intended intra-arterial thrombolysis:

1. Cerebral infarction in the distribution of the relevant occluded artery in the previous 6 weeks
2. History of intracerebral hemorrhage
3. Severe head injury (contusion) in the previous 4 weeks
4. Clinical or laboratory evidence of coagulation abnormalities, i.e. platelet count <90 x 10<sup>9</sup>/L, APTT>50 sec or INR >1.7

**Date of first enrolment**

01/12/2010

**Date of final enrolment**

01/01/2015

## Locations

**Countries of recruitment**

Netherlands

**Study participating centre**

Erasmus MC University Medical Center Rotterdam

Rotterdam

Netherlands

3000CA

## Sponsor information

**Organisation**

Dutch Heart Foundation (Netherlands)

**ROR**

<https://ror.org/05nxhgm70>

## Funder(s)

**Funder type**

Charity

**Funder Name**

Dutch Heart Foundation (Nederlandse Hartstichting) (Netherlands) ref: 2008T030

**Alternative Name(s)**

Heart Foundation

**Funding Body Type**

Private sector organisation

**Funding Body Subtype**

Trusts, charities, foundations (both public and private)

**Location**  
Netherlands

## Results and Publications

**Individual participant data (IPD) sharing plan**  
Not provided at time of registration

**IPD sharing plan summary**  
Not provided at time of registration

### Study outputs

| Output type                     | Details | Date created | Date added | Peer reviewed? | Patient-facing? |
|---------------------------------|---------|--------------|------------|----------------|-----------------|
| <a href="#">Results article</a> | results | 01/01/2015   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/03/2016   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/06/2016   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 16/08/2016   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/09/2016   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/10/2016   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/12/2016   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 06/04/2017   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/05/2017   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/05/2017   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 20/06/2017   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/07/2017   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/10/2017   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/02/2018   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/02/2018   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/02/2018   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/03/2018   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/04/2018   |            | Yes            | No              |
| <a href="#">Results article</a> | results | 01/12/2018   |            | Yes            | No              |
|                                 | results | 01/05        |            |                |                 |

|                                               |                                                               |                |                |     |     |
|-----------------------------------------------|---------------------------------------------------------------|----------------|----------------|-----|-----|
| <a href="#">Results article</a>               |                                                               | /2019          |                | Yes | No  |
| <a href="#">Results article</a>               | Multivessel occlusions (MVO)                                  | 15/12<br>/2023 | 18/12<br>/2023 | Yes | No  |
| <a href="#">Protocol article</a>              | protocol                                                      | 01/09<br>/2014 |                | Yes | No  |
| <a href="#">Abstract results</a>              | Economic evaluation                                           | 14/10<br>/2021 | 15/10<br>/2021 | No  | No  |
| <a href="#">Abstract results</a>              | Results abstract European Stroke Organisation Conference 2021 | 03/09<br>/2021 | 29/03<br>/2023 | No  | No  |
| <a href="#">Other publications</a>            | post hoc analysis                                             | 01/12<br>/2019 | 12/12<br>/2019 | Yes | No  |
| <a href="#">Other publications</a>            | retrospective analysis                                        | 08/06<br>/2022 | 09/06<br>/2022 | Yes | No  |
| <a href="#">Participant information sheet</a> | Participant information sheet                                 | 11/11<br>/2025 | 11/11<br>/2025 | No  | Yes |
| <a href="#">Study website</a>                 | Study website                                                 | 11/11<br>/2025 | 11/11<br>/2025 | No  | Yes |