

Supplemental Online Content

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eTable 1. Neuroimaging triage protocols of sites in the CLEAR Study

eTable 2. Selection criteria for late window (6-24h) thrombectomy between 2014 to 2020

eTable 3. Modified Rankin Score Assessment

eTable 4. Analysis of imaging modality with good outcome: Local patients

eTable 5. mTICI reperfusion and 90-day mRS by imaging modality

This supplemental material has been provided by the authors to give readers additional information about their work.

eTable 1. Neuroimaging triage protocols of sites in the CLEAR Study

| Site | CT ASPECTS | CTA routine | CTA Collateral for selection | CT selection beyond ASPECTS | CT vs advanced imaging | CTP criteria | MRI |
|--------------------------------|------------------------|---|------------------------------|----------------------------------|---------------------------------------|-------------------------------------|--|
| Bern | Not used | Yes | No | No | Advanced imaging routine | Tmax>6, rCBF<35%, ADC<620* | Same as CTP |
| Boston Medical Center | ≥5 | Yes | No | No | Advanced imaging rarely used | Rarely used | Rarely used |
| CHU Lille | No specified threshold | If contraindication to MRI | No | No | MRI Routine | No | Core <70 ml, Mismatch ratio 1.8, Mismatch volume > 15 ml |
| CHU Montreal | ≥5 | Yes | No | No | Advanced imaging rarely used | Rarely used, no guideline | Rarely used, no guideline |
| Cooper | No specified threshold | Yes | Sometimes | No | Attending dependent | No specified threshold; most <50 ml | No |
| Grady Memorial | No specified threshold | Yes | No | No | Advanced imaging (CTP) routine | No specified threshold | N/A |
| Heidelberg University Hospital | ≥6 | Yes | No | No | Attending dependent** | Core < 100 ml | Core < 100 ml |
| Lausanne University Hospital | ≥7 (NIHSS≥10) | Yes, until May 2018 (MRI initial imaging) | No | Core < 2/3 of affected territory | NCCT if contraindications to advanced | Core < 50 ml (NIHSS≥10); core | Same as CTP |

| | | | | | | | |
|-----------------------------|----------------------------|-----|-----|-------------------------------------|---|---|-----------------------------|
| | ≥ 8 (NIHSS <10) | | | | imaging or technical problems with CTP (movement artifact, injection failure, poor cardiac ejection fraction) | < 30 ml (NIHSS <10) | |
| Mercy Hospital | ≥ 6 | Yes | No | Yes, greater than 1/3 MCA territory | Availability of the lab | >1.2 mismatch ratio | No |
| SUNY Upstate Medical | ≥ 6 | Yes | Yes | No | CT only if clinical-imaging mismatch | Core < 70 ml | Core < 70 ml |
| University of Iowa | ≥ 6 | Yes | No | No | Advanced imaging only | Core < 70 ml | No |
| University of Massachusetts | ≥ 6 | Yes | Yes | No | CT only if clinical-imaging mismatch | Core < 70 ml | No |
| UT Health McGovern | ≥ 6 | Yes | No | No | CTP Routine | Core < 70 ml, significant mismatch | Rarely used |
| University of Toledo | ≥ 6 | Yes | Yes | No | CT only if ASPECTS 9 or 10 | No pre-specified parameters | No pre-specified parameters |
| Vall D'Hebron, Barcelona | No specific threshold | Yes | No | No | CTP if NCCT not favorable | Routine; CTP volumes not used for selection | Rarely used |

| | | | | | | | |
|--|--|--|--|--|--|-------------------|--|
| | | | | | | if NCCT favorable | |
|--|--|--|--|--|--|-------------------|--|

CHU: Centre Hospitalier de l'Université; ASPECTS: Alberta Early CT Program CT Score.

* Olea, Rapid and Syngo were used for perfusion processing. No thresholds for volumes were used.

** Refer to Nagel S, Herweh C, et al. JNIS supplement for additional details of their imaging selection protocol.

eTable 2. Selection criteria for late window (6-24h) thrombectomy between 2014 to 2020

| Site | Change in patient selection for mechanical thrombectomy between 2014 to 2020 |
|--------------------------------|---|
| Bern | No change, other than more distal occlusions being treated over time |
| Boston Medical Center | Selected more patients in extended window after 2018 |
| CHU Lille | Selected more patients in extended window after 2018 |
| CHU Montreal | No change |
| Cooper | Selected more patients in extended window after 2018; no change in imaging protocol before and after 2018 |
| Grady Memorial Hospital | No change |
| Heidelberg University Hospital | No change** |
| Lausanne University Hospital | <p>Until 2014, EVT if treatment initiated within 6 hours, NIHSS ≥ 6, CTA proximal LVO, and CTP showed $> 50\%$ penumbra and informed consent was available.</p> <p>Since October 2014, CTP criteria were replaced by ASPECTS ≥ 5 and lower NIHSS limit replaced by the presence of disabling deficit.</p> <p>Since May 2017, patients were treated with the same criteria up to 8 hours. After 8 hours, treatment was offered with modified DAWN criteria: i.e. in the presence of NIHSS ≥ 10 and ASPECTS ≥ 7, or if stroke was disabling, NIHSS was 1-10, and ASPECTS was ≥ 8.</p> <p>Since January 2018, late treatment was alternatively based on any NIHSS, core < 70 ml and mismatch ratio ((penumbra + core)/core) > 1.8, according to DEFUSE-3 criteria, and in accordance with European and American guideline recommendations.</p> |
| Mercy Hospital, Toledo | No change |

| | |
|---------------------------------|--|
| SUNY Upstate Medical University | After 2018, selection criteria resembled DAWN/DEFUSE-3 trial protocols |
| University of Iowa | After 2015, selection with RAPID software. EVT in extended window not widely implemented until after 2018 |
| University of Massachusetts | No change |
| UT Health McGovern | After 2018, CTP criteria were added to the stroke imaging selection protocol for extended window patients similar to DEFUSE-3, anterior circulation patients with NIHSS \geq 6, mRS 0-2. |
| University of Toledo | No change |
| Vall D'Hebron, Barcelona | After 2018, indications for EVT expanded in the extended window |

CHU: Centre Hospitalier de l'Université; EVT: endovascular therapy

** Refer to Nagel S, Herweh C, et al. JNIS supplement for additional details of their imaging selection protocol.

eTable3. Modified Rankin Score Assessment

| Site | Modified Rankin Score assessment by site | mRS blinding to mode of imaging selection |
|---------------------------------|---|---|
| Bern | Standard phone interview by study nurse | Unblinded |
| Boston Medical Center | Standard questionnaire used by clinician (in clinic or telehealth) or stroke quality coordinator (via telephone). | Provider could have been aware of the imaging modality used |
| CHU Lille | Standard questionnaire by trained neurologists | Unblinded |
| CHU Montreal | Standard approach by neurologists (trained by NINDS criteria) | Unblinded |
| Cooper University | Half of mRS obtained by neurology provider (MD, NP) during follow-up visit; half of mRS obtained with semi-structured telephone interview | Provider could have been aware of imaging modality of selection |
| Grady Memorial Hospital | Standard approach. If phone, used Bruno et al. questionnaire. * | Provider could have been aware of imaging modality of selection |
| Lausanne University Hospital | Standard approach at outpatient clinic or standard telephone interview, all with mRS-certified medical personnel | Blind to patient treatment in the acute phase |
| Mercy Hospital, Toledo | Standard approach by certified stroke unit nurse, conducted follow-up call | Not aware of imaging modality for treatment |
| University of Iowa | Standard approach | Unblinded |
| University of Massachusetts | Neurologist, stroke coordinator and dedicated conducted mRS with standard approach | The assessors were not aware of the mode of patient selection for treatment |
| UT Health McGovern | mRS was determined by a standard questionnaire, performed by certified stroke coordinators who conducted phone-based surveys | Provider could have been aware of imaging modality of selection |
| University of Toledo | mRS was conducted with a standard approach, conducted by stroke nurse | Provider could have been aware of imaging modality of selection |
| SUNY Upstate Medical University | mRS was conducted with a standard approach, conducted by stroke attending and NP in clinic | Provider could have been aware of imaging modality of selection |

| | | |
|-----------------------------|---|-----------|
| Vall D'Hebron, Barcelona | mRS was performed by the treating physician | Unblinded |
|-----------------------------|---|-----------|

*Reference: Bruno A, Akinwuntan AE, Lin C, Close B, Davis K, Baute V, Aryal T, Brooks D, Hess DC, Switzer JA, Nichols FT. Simplified modified rankin scale questionnaire: reproducibility over the telephone and validation with quality of life. Stroke. 2011 Aug;42(8):2276-9.

| eTable 4. Univariate and multivariate analysis of imaging modality with good outcome (mRS score 0–2) and ordinal mRS shift: Local patients | | | | |
|---|-------------------------------|-------|------------------|-------|
| | Univariate | | Multivariate | |
| | Odds ratio (95% CI), <i>P</i> | | | |
| Outcome mRS 0-2 | | | | |
| CT | Ref | | Ref | |
| CTP | 0.86 (0.56-1.30) | 0.461 | 0.71 (0.42-1.21) | 0.209 |
| MRI | 0.75 (0.37-1.51) | 0.416 | 0.69 (0.36-1.33) | 0.272 |
| Outcome mRS shift | | | | |
| CT | Ref | | | |
| CTP | 0.92 (0.65-1.31) | 0.652 | 0.74 (0.46-1.18) | 0.205 |
| MRI | 0.84 (0.53-1.35) | 0.481 | 0.78 (0.54-1.12) | 0.175 |

| eTable 5. mTICI Reperfusion and mRS at 90 days. | | | |
|---|----------------------------|----------------------------|----------|
| | mRS 90 days 0-2 (n=676) | mRS 90 days 3-6 (n=923) | <i>P</i> |
| Reperfusion mTICI | | | |
| | n (row %) | | |
| 0-2a | 25 (12.2) | 180 (87.8) | <0.0001 |
| 2b-3 | 651 (46.7) | 743 (53.3) | |
| Odds of good outcome (mRS score 0-2) for mTICI reperfusion 2b-3 | | | |
| | Odds ratio (95% CI) | | |
| Overall | 6.3 (4.4-8.9) | | <0.0001 |
| CT | 6.1 (2.2-16.5) | | 0.0004 |
| CTP | 5.1 (2.9-9.2) | | <0.0001 |
| MRI | 8.9 (6.7-11.9) | | <0.0001 |

TICI: thrombolysis in cerebral infarction