

Supplemental material

Table S1: MINORS criteria for the selected studies included in the meta-analysis

| References | Q1 | Q2 | Q3 | Q4 | Q5 | Q6 | Q7 | Q8 | Q9 | Q10 | Q11 | Q12 | overall |
|---|----|----|----|----|----|----|----|----|----|-----|-----|-----|---------|
| Arregle <i>et al.</i> 2021 [12] | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 0 | 2 | 2 | 2 | 2 | 21 |
| Cooper <i>et al.</i> 2009 [5] | 2 | 2 | 2 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 20 |
| Diab, <i>et al.</i> 2016 [13] | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 0 | 2 | 2 | 2 | 2 | 18 |
| Fukuda, <i>et al.</i> 2014 [14] | 2 | 0 | 0 | 2 | 2 | 0 | 0 | 0 | 2 | 2 | 2 | 2 | 14 |
| Garcia-Cabrera, <i>et al.</i> 2013 [15] | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 0 | 0 | 2 | 2 | 2 | 19 |
| Grabowski, <i>et al.</i> 2010 [16] | 2 | 0 | 2 | 2 | 2 | 2 | 2 | 0 | 2 | 2 | 2 | 1 | 19 |
| Heiro, <i>et al.</i> 2007 [17] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 16 |
| Huang, <i>et al.</i> 2023 [18] | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 0 | 2 | 2 | 2 | 2 | 18 |
| Kim, <i>et al.</i> 2017 [19] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 16 |
| Lee Su Jin, <i>et al.</i> 2014 [20] | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 0 | 2 | 2 | 2 | 2 | 18 |
| Lee Seung-Jae <i>et al.</i> 2014 [21] | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 0 | 2 | 2 | 2 | 2 | 18 |
| Misfeld, <i>et al.</i> 2014 [22] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 16 |
| Okazaki, <i>et al.</i> 2013 [23] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 16 |
| Pinto, <i>et al.</i> 2024 [24] | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 0 | 2 | 2 | 2 | 2 | 21 |
| Roder, <i>et al.</i> 1997[25] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 16 |
| Ruttmann, <i>et al.</i> 2020 [26] | 2 | 2 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 18 |
| Santoshkumar <i>et al.</i> 1996 [27] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 1 | 1 | 14 |
| Scheggi, <i>et al</i> 2022[28] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 16 |
| Selton-Suty, <i>et al</i> 2016[29] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 16 |
| Sonneville, <i>et al</i> 2011[30] | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 0 | 2 | 2 | 2 | 2 | 22 |
| Suzuki, <i>et al</i> 2017[31] | 2 | 2 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 18 |
| Thuny, <i>et al</i> 2007[32] | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 0 | 2 | 2 | 2 | 2 | 22 |
| Tsai, <i>et al</i> 2024[33] | 2 | 0 | 0 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 16 |
| Wilbring <i>et al</i> 2014[34] | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 0 | 2 | 2 | 2 | 2 | 18 |
| Zaballos <i>et al</i> 2024[35] | 2 | 2 | 2 | 2 | 2 | 2 | 0 | 0 | 2 | 2 | 2 | 2 | 20 |

Q: question

Table S2: Subtypes of neurological complications and their relation to the primary outcome

| References | NC subtypes according to study design | NC subtypes used for analysis (M: major; m:minor) | OR of the primary outcome |
|--|--|--|----------------------------------|
| <i>Diab et al, 2016 (13)</i> | Major CVC (mRS >3) | M | 0.85 (0.55-1.30) |
| <i>Garcia-Cabrera et al, 2013 (15)</i> | Neurological complications together | | 1.58 (1.23-2.02) |
| | Encephalitis/meningitis | | 0.94 (0.61-1.45) |
| | Small ischemic complications | m | 1.41 (0.93-2.19) |
| | Moderate-severe ischemic complications | M | 1.63 (1.19-2.22) |
| | Cerebral hemorrhage | | 1.73 (1.10-2.71) |
| <i>Huang et al , 2023 (18)</i> | Symptomatic NC | M | 2.884 (2.574 – 3.213) |
| <i>Lee Seung-Jae et al, 2014 (20)</i> | Minor CVC | m | 0.856 (0.253-2.894) |
| | Major CVC (initial mRS >=3) | M | 2.865 (1.254-6.548) |
| <i>Ruttmann E. et al, 2020 (26)</i> | Uncomplicated stroke | m | 1.18 (0.79–1.77) |
| | Complicated stroke | M | 2.1 (1.24–3.54) |
| <i>Selton-Suty et al, 2016 (29)</i> | Asymptomatic NC | m | 0.393 (0.120-1.281) |
| | Symptomatic NC | M | 2.499 (1.623-3.894) |
| <i>Sonneville et al, 2011 (30)</i> | Symptomatic NC | | 1.21 (0.63–2.33) |
| | Neurological failure GCS <10 | M | 7.41 (2.89-18.96) |
| <i>Suzuki et al, 2017 (31)</i> | Complicated stroke | M | 10.763 (1.232-155.872) |
| <i>Thuny et al, 2007 (32)</i> | Silent cerebral embolism or TIA | m | 0.9 (0.46–1.61) |
| | Stroke | M | 1.6 (1.02–2.65) |

NC: neurological complications, CVC: cerebrovascular complications, mRS: modified Rankin Score, GCS: Glasgow coma scale, TIA: transient ischemic attack.

Table S3: Meta-regression analysis on the primary outcome

| Moderator | Beta | P value for interaction |
|--|-------------|--------------------------------|
| <i>Male sex</i> | -0.01 | 0.450 |
| <i>Mean age</i> | -0.02 | 0.003 |
| <i>Prosthesis involvement</i> | -0.00 | 0.854 |
| <i>Right heart valve involvement</i> | 0.02 | 0.645 |
| <i>Mitral valve involvement</i> | 0.01 | 0.124 |
| <i>Aortic valve involvement</i> | -0.01 | 0.073 |
| <i>S.A positive emoculture</i> | -0.00 | 0.675 |
| <i>Presence of valvular abscess or local complications</i> | 0.00 | 0.629 |
| <i>Valve surgery</i> | -0.00 | 0.330 |
| <i>Early surgery (< 7 days)</i> | -0.00 | 0.312 |
| <i>Heart failure at admission</i> | 0.00 | 0.937 |

SA: S.Aureus

Figure S1: Funnel plot

