

ESO Guideline on Mobile Stroke Units for Prehospital Stroke Management

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Disclosures of the MWG

(listed in Supplementary Table 1 of the publication)

1. Intellectual disclosures:

Karianne Larsen: MSU study investigator, PRESTO Board member

Simona Sacco: Co-chair of the Guideline Board of the European Stroke Organization

Thorsten Steiner: ATACH-2

Guillaume Turc: Chairman ESO Guideline board, Co-chairman 2019 ESO-ESMINT Guidelines on mechanical thrombectomy, Co-chairman 2021 ESO Guidelines on IVT

Georgios Tsivgoulis: Section Editor: “Stroke” journal, Associate Editor: “Therapeutics advances in Neurological Disorders” journal, Chair ESO Industry Roundtable

Silke Walter: Investigator of MSU studies, PRESTO Board member, Co-Chair of WISE, Co-Chair of ESOTA

2. Financial disclosures:

Heinrich Audebert: Speaker honoraria from Lundbeck, Boehringer Ingelheim (<€10,000 within last 3 years); Principal investigator INSPIRE-TMS project (co-funded by Pfizer); employed by the Charité Universitätsmedizin Berlin (receives institutional share from the STEMOMANIFOLD manufacturer (Meytec GmbH), no personal/team access to this money)

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Thorsten Steiner: Personal fees: Bayer, Boehringer, BMS-Pfizer, Daiichi Sankyo, Alexion

Guillaume Turc: Lecturing fees for Guerbet France, Travel support by Samsung Electronics France

Georgios Tsivgoulis: Participation in advisory meetings & satellite symposia for Boehringer-Ingelheim; Novartis, Sanofi, Biogen, Genesis Pharma, Teva, Merck-Serono, Bayer, Daiichi-Sankyo, Allergan, Specifar, Actavis, Shire, Medtronic, CSL Behring, Abbvie, Abbott, Takeda, Biomarin, Abbvie, Ipsen, Abbott, Takeda, Roche. Unrestricted research grants from Novartis, Genesis Pharma, Teva, Shire, Merck-Serono, Medtronic, Boehringer-Ingelheim, Allergan, Abbott, Amicus, Abbvie, Ipsen, Abbott, Sanofi, Bayer, Roche

Background

Mobile Stroke Units for prehospital acute stroke patient management

- Multiple publications indicate
 - earlier treatment
 - increased numbers of patients receiving treatment
 - optimised triage to the individually required level of care
 - 2 recent studies add data to clinical outcome

Walter et al., 2012; Ebinger et al., 2014; Bowry et al., 2015; Ebinger et al., 2015; Gomes et al., 2015; Wendt et al., 2015; Taqui et al., 2017; Shuaib et al., 2018; Helwig et al., 2019; Czap et al., 2020; Zhao et al., 2020; Cooley et al., 2021; Ebinger et al., 2021; Grotta et al., 2021; Larsen et al., 2021; Zhou et al., 2021



Methods

- ESO Standard operating Procedure for Guidelines with GRADE methodology
(Steiner et al., 2021)
- 3 PICO questions
- 14 outcomes
- RCTs and observational studies considered
- in case of overlapping populations: disentangling by personal communication with authors
(Turc et al., 2022)

Population	Intervention and Comparator	
	Mobile Stroke Unit	Conventional management
Suspected stroke patients (PICO1)	Outcomes	Clinical outcomes: <ul style="list-style-type: none">• mRS• Time to treatment• Triage• Proportion treated
Confirmed acute ischaemic stroke patients (PICO 2)		Safety outcomes <ul style="list-style-type: none">• All-cause mortality• Bleeding complications• Mimics treated• Haematoma growth
Confirmed acute ICH patients (PICO 3)		

Methods: Importance of outcomes (Delphi votes):

1. Suspected stroke patients:

mortality (90/7 days): 7.9 and 7.6,
sICH: 7.5,
mimics treated (IVT): 6.4,
major extracranial bleeding: 6.1

2. AIS:

functional outcomes: excellent: 8.9, any better: 8.8, good: 8.4,
proportion receiving treatment: 7.5, time to therapy: 7.3,
mortality (90/7 days): 8.0 and 7.0, “Golden hour” IVT: 7.1,
sICH: 7.3, LVO triaged to tertiary care: 6.8,
major extracranial bleeding: 6.0

3. ICH:

functional outcomes: good 8.5, any better 8.3, excellent 7.6,
mortality (90/7 days): 8.1 and 7.5
ICH triaged to tertiary care: 6.3,
size of haematoma expansion: 6.0

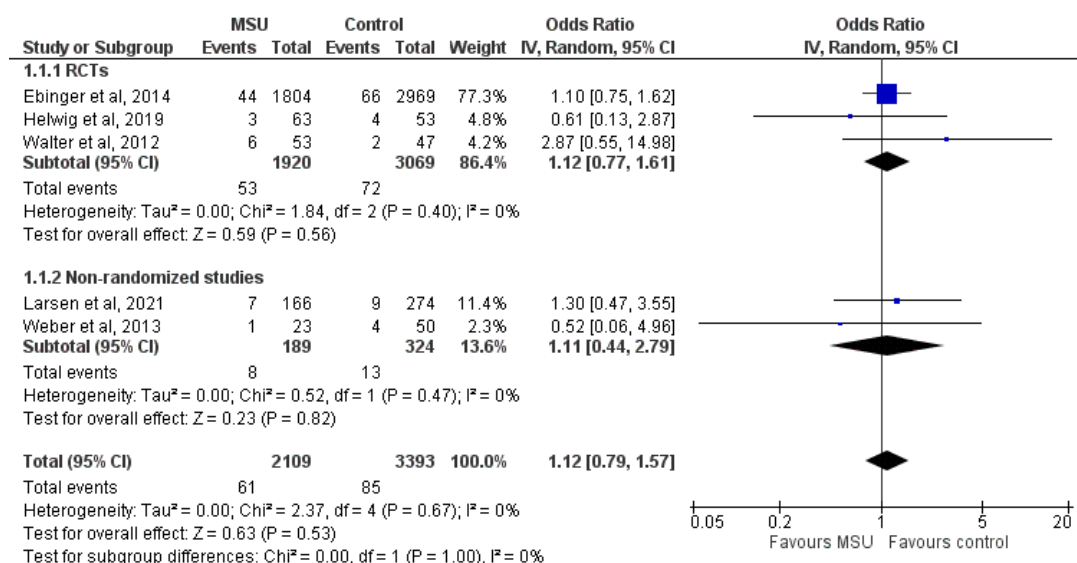
→ 23 sets of analysis and
additional sensitivity
analysis with further 3 sets
of analysis

→ **1 combined
recommendation**

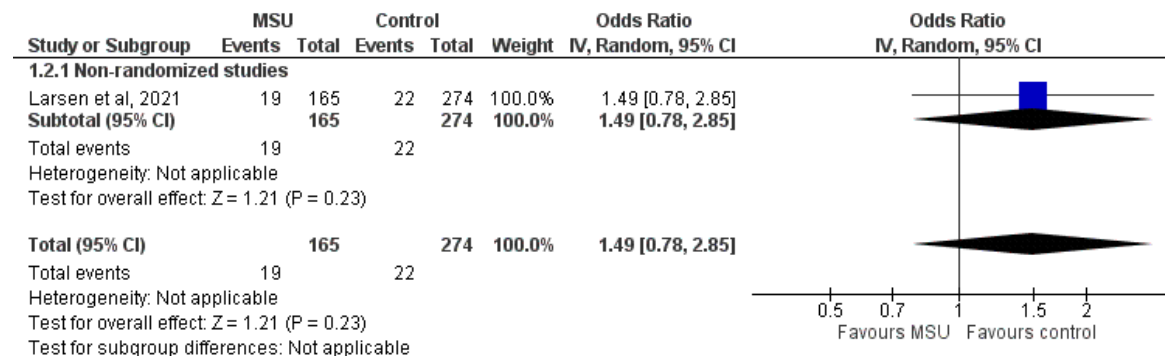
Suspected Stroke Patients

MSU vs conventional management

Safety outcome: all-cause mortality 7/90 days



Quality: **Low** ⊕⊕
(Bias)

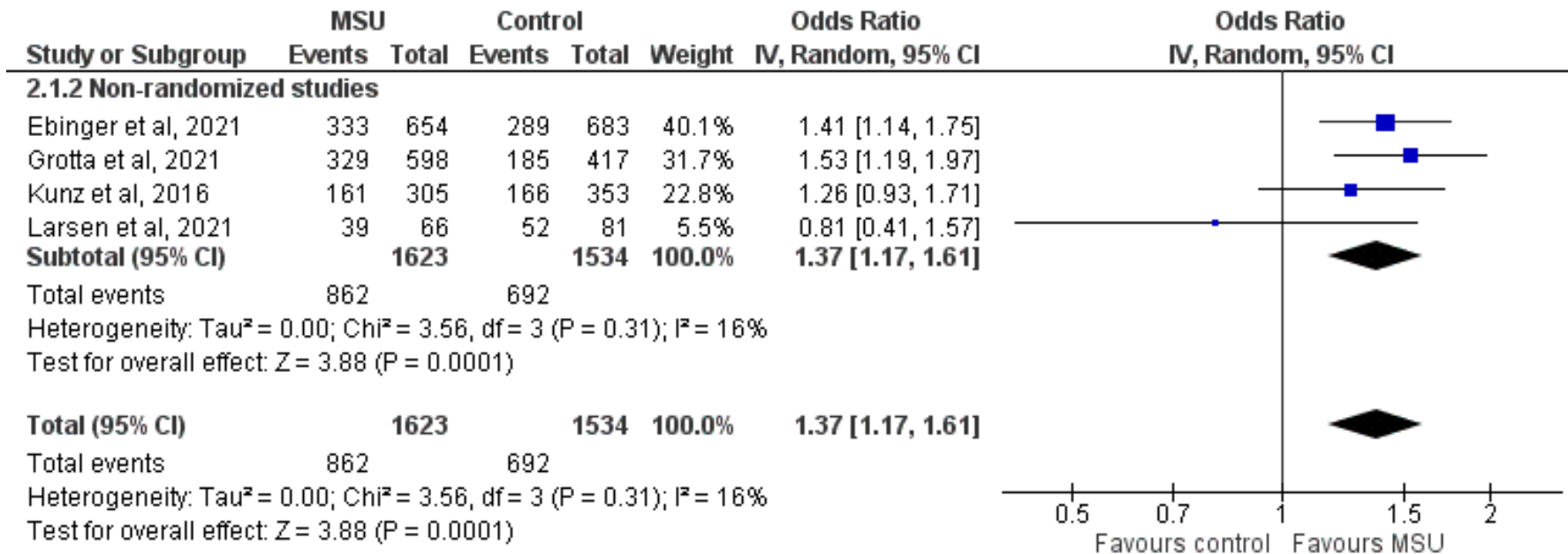


Quality: **Very low** ⊕
(Imprecision)

Acute Ischaemic Stroke Patients

MSU vs conventional management

Excellent functional outcome (mRS 0-1)

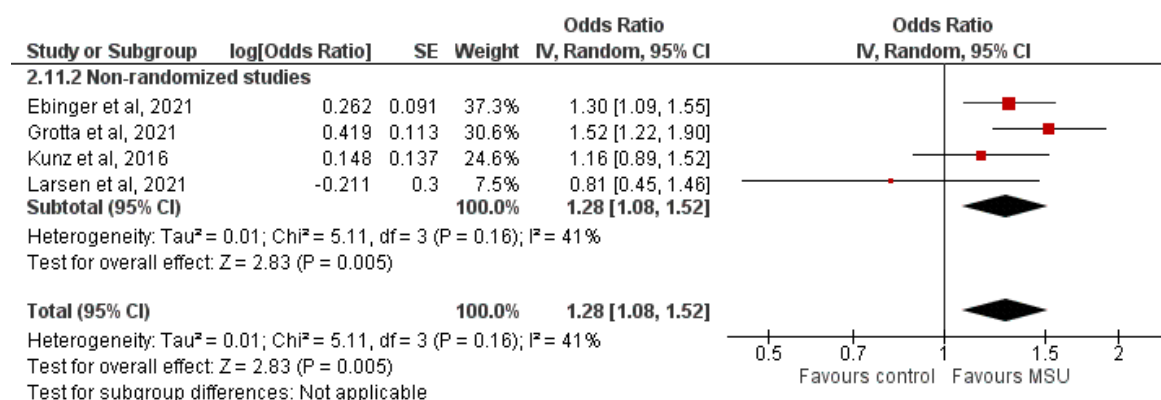
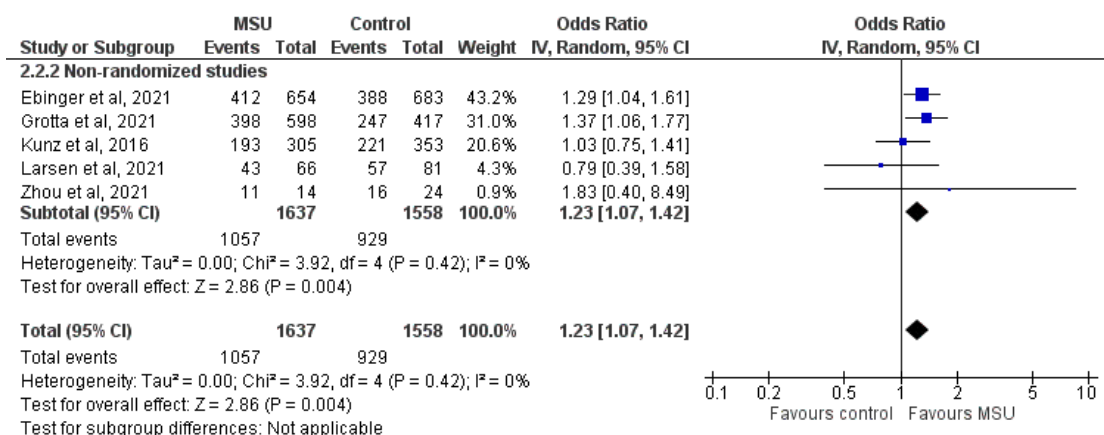


Quality: Moderate ⊕⊕⊕

Acute Ischaemic Stroke Patients

MSU vs conventional management

Good (mRS 0-2) and any better functional outcome



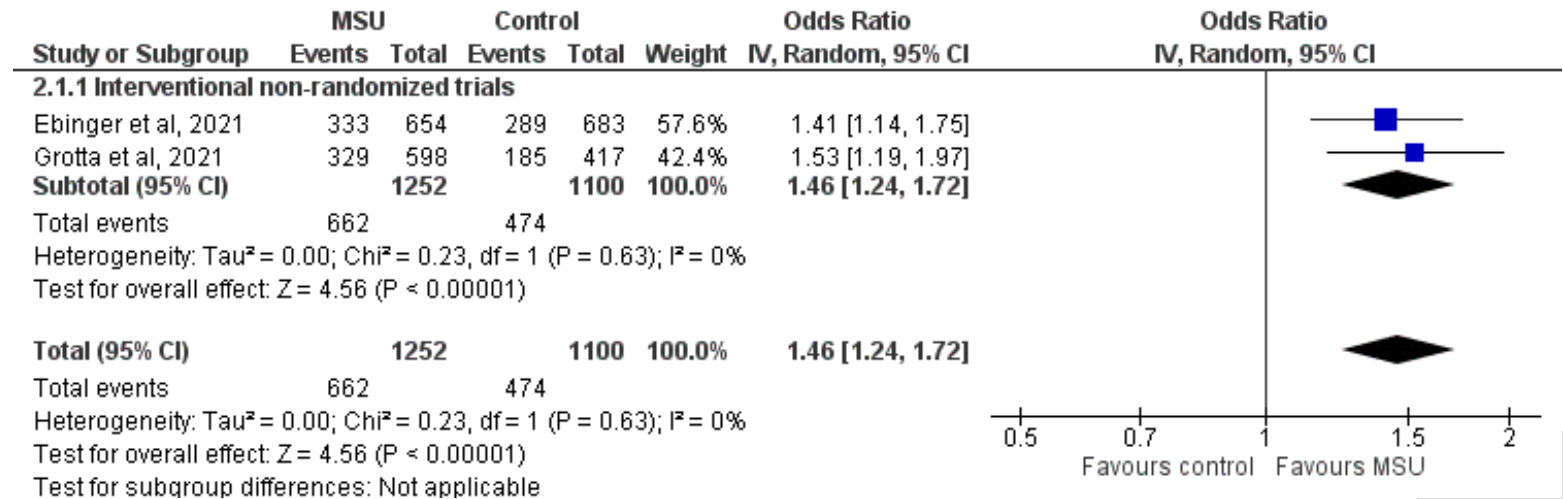
Quality: **Moderate** ⊕⊕⊕

Additional information for Acute Ischaemic Stroke Patients: Sensitivity analysis

MSU vs conventional management

Excellent functional outcome (mRS 0-1)

B_PROUD/BEST-MSU:
prospective interventional
studies with blinded
endpoint

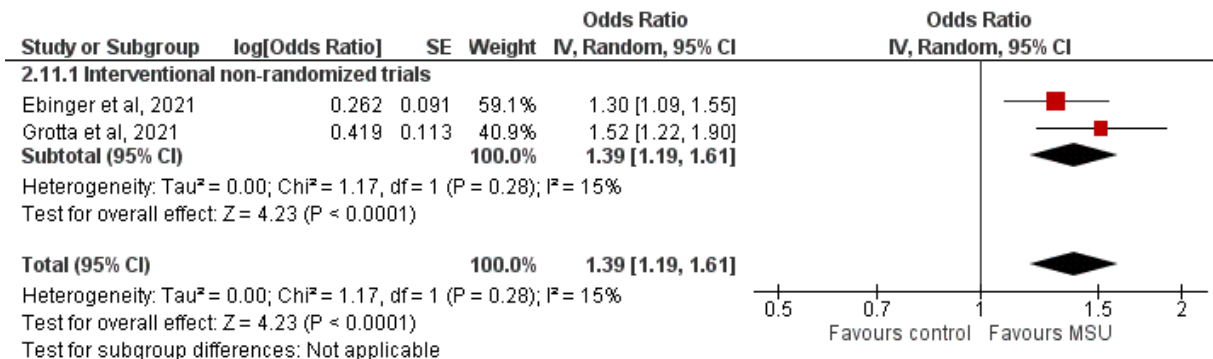
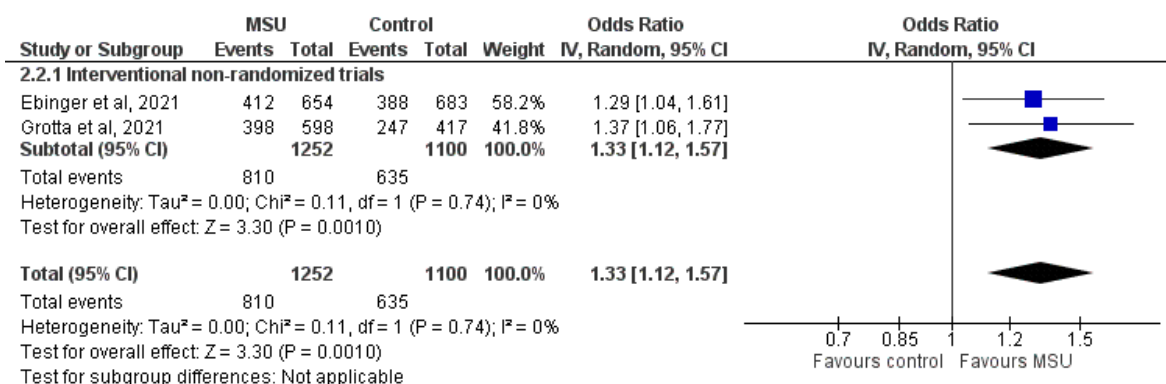


Quality: Moderate ⊕⊕⊕

Additional information for Acute Ischaemic Stroke Patients: Sensitivity analysis: B_PROUD/BEST-MSU

MSU vs conventional management

Good (mRS 0-2) and any better functional outcome

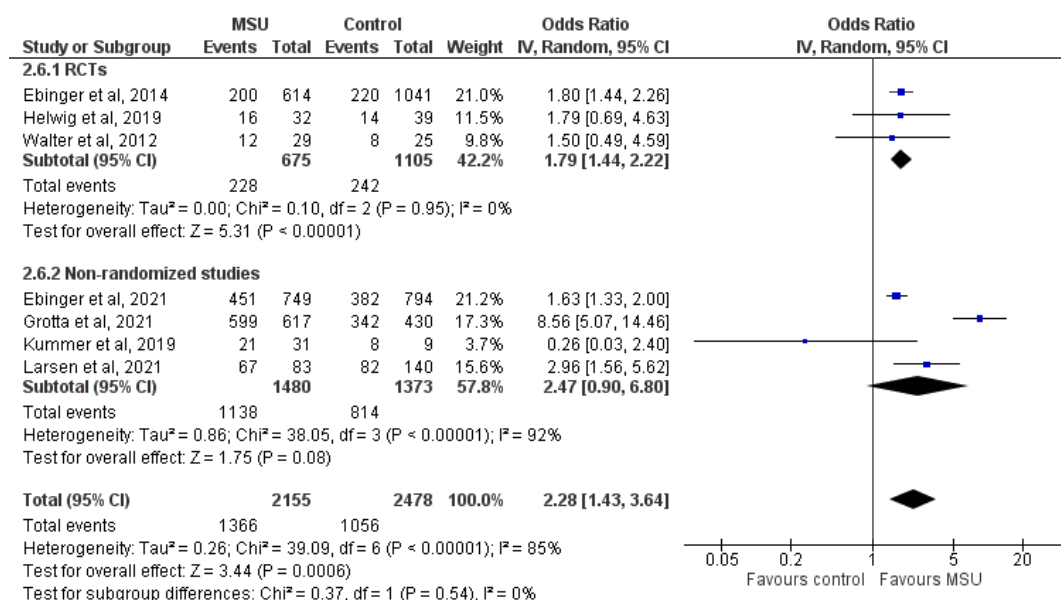


Quality: Moderate ⊕⊕⊕

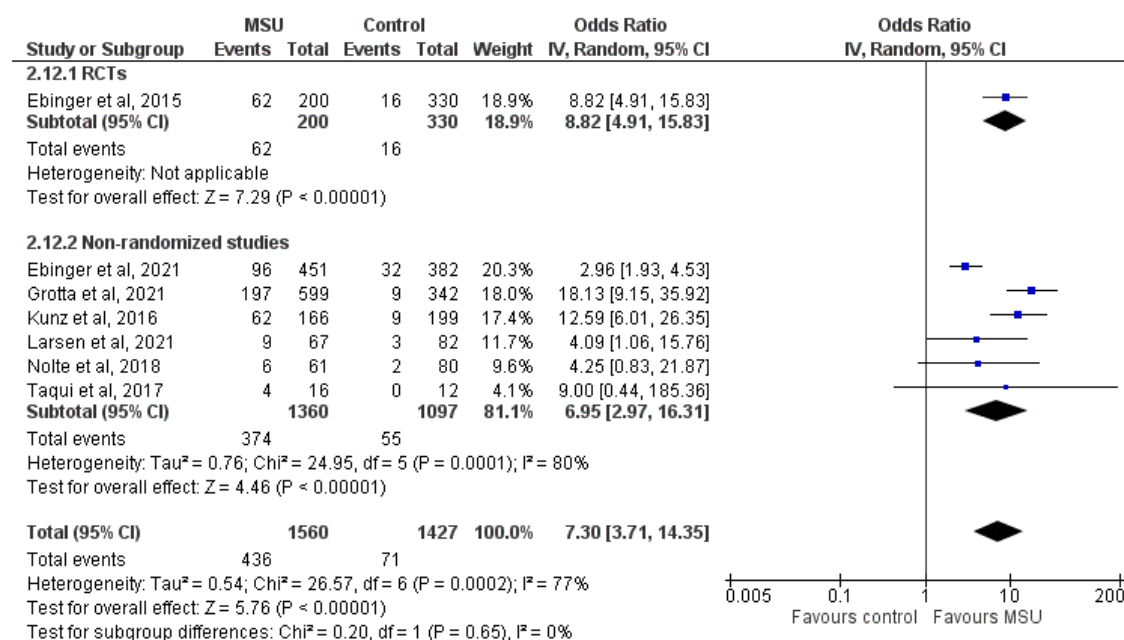
Acute Ischaemic Stroke Patients

MSU vs conventional management

Proportion of patients receiving IVT and of patients with “Golden hour” IVT



Quality: **Low** ⊕⊕ (Bias, Inconsistency)

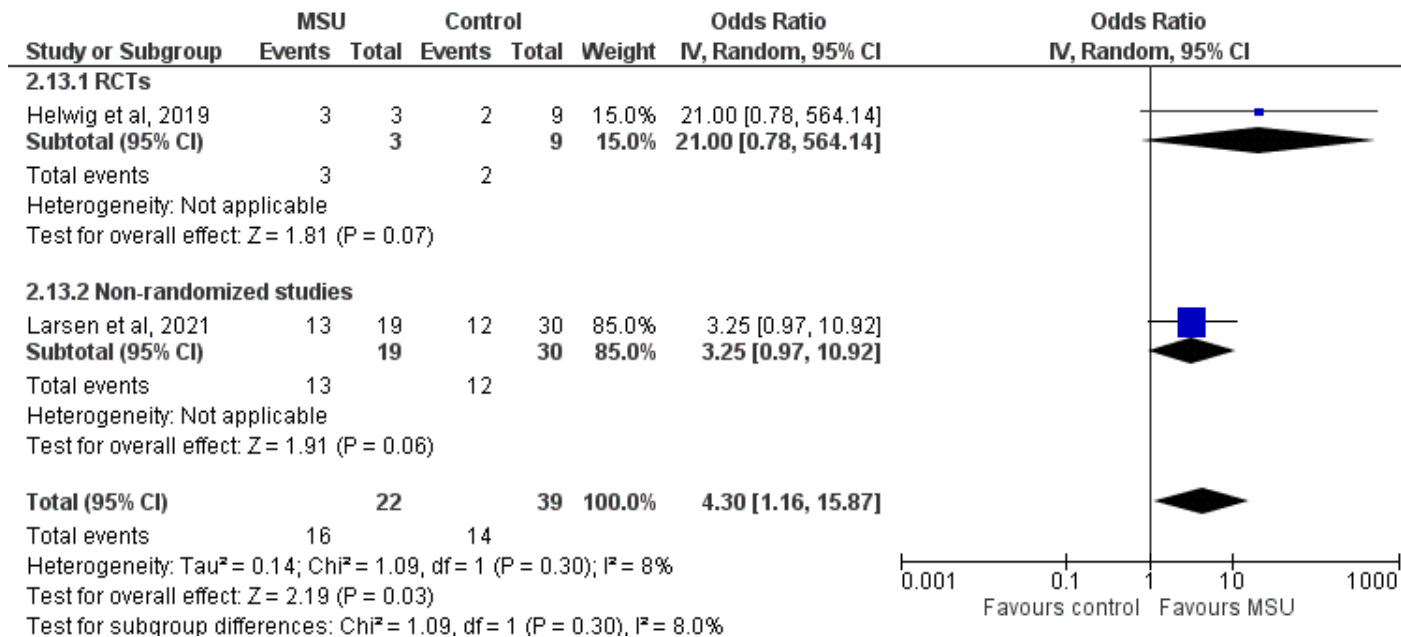


Quality: **Moderate** ⊕⊕⊕

Acute Ischaemic Stroke Patients – subgroup LVO patients

MSU vs conventional management

Proportion of patients primarily transferred to MT capable centres

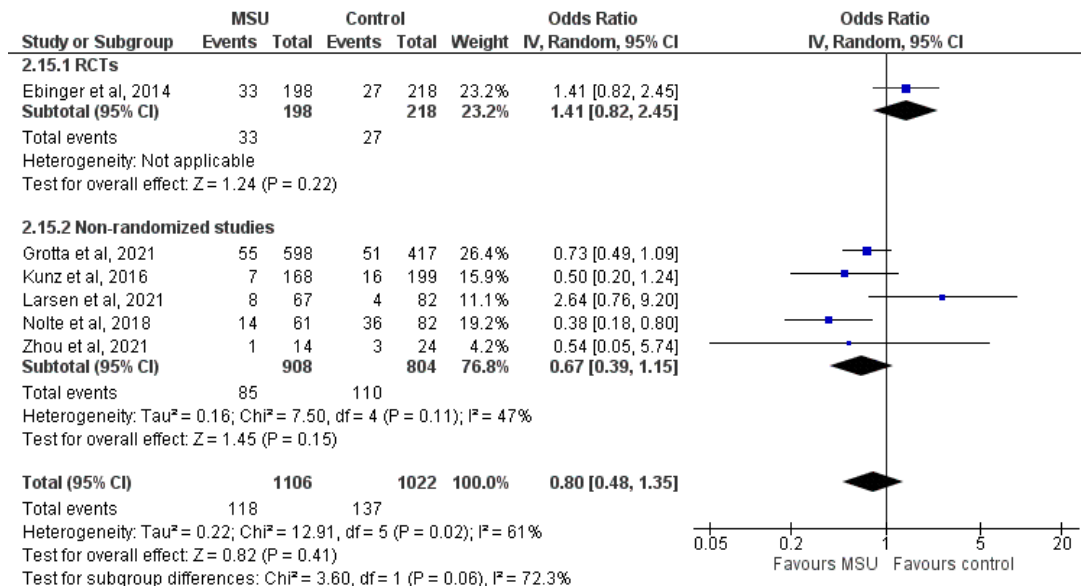


Quality: Low ⊕⊕ (Bias, Imprecision)

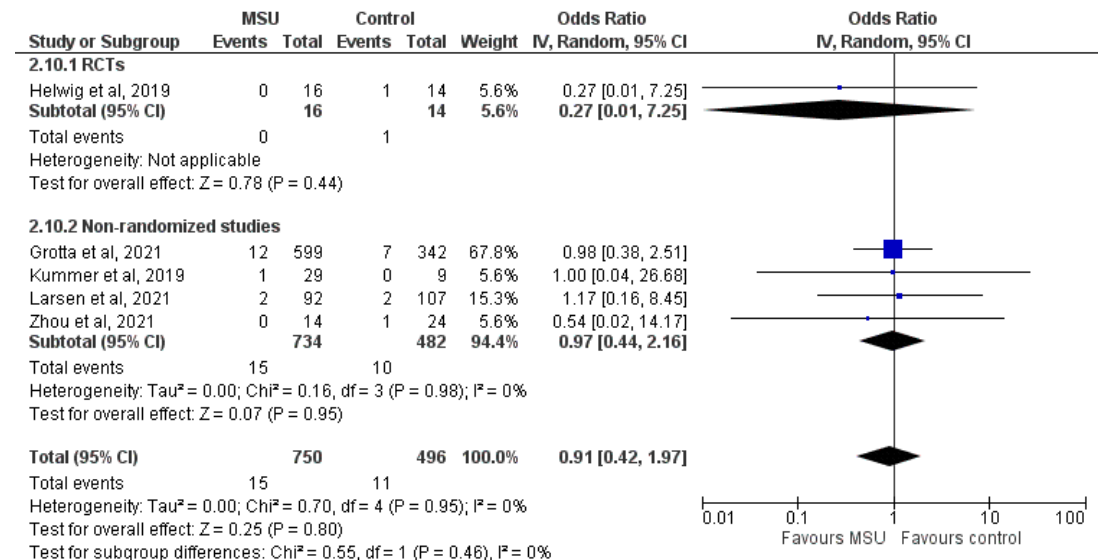
Acute Ischaemic Stroke Patients

MSU vs conventional management

All-cause mortality (90 days) and sICH among IVT-treated patients



Quality: Very low ⊕
(Imprecision)

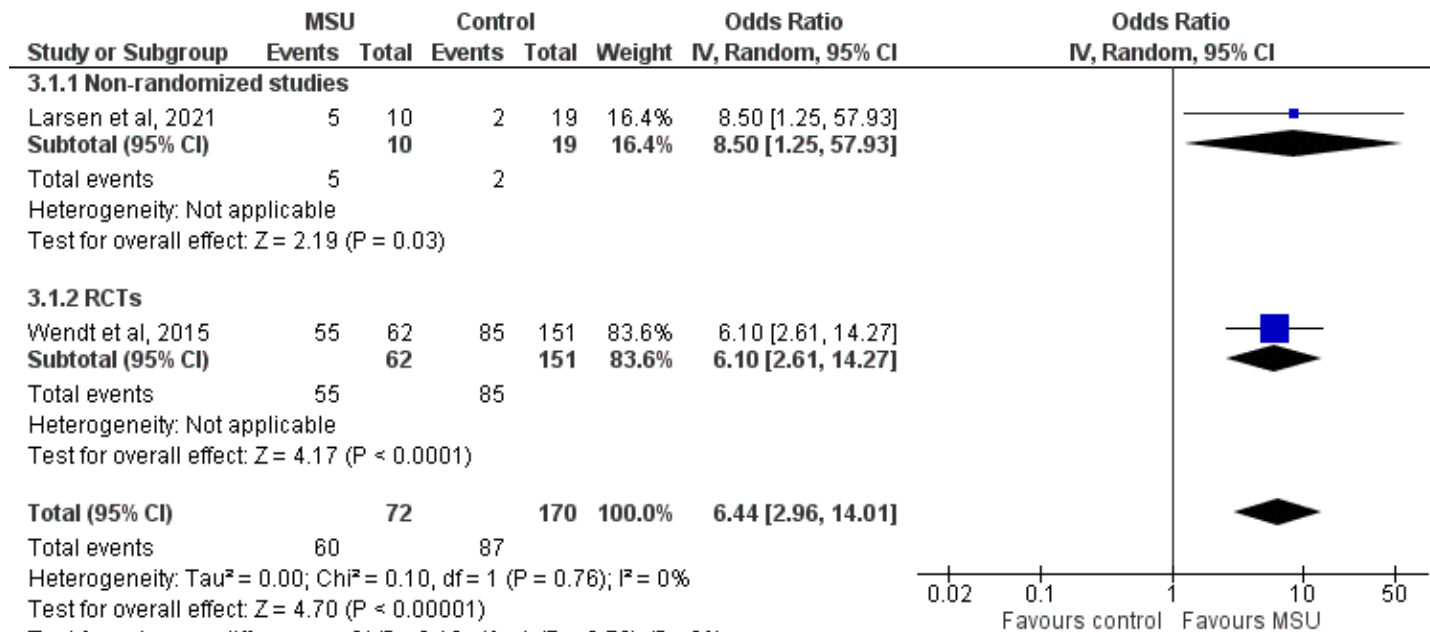


Quality: Very low ⊕ (Bias,
Indirectness, Imprecision)

Acute Intracerebral Haemorrhage Patients

MSU vs conventional management

Proportion primarily transported to tertiary stroke centres

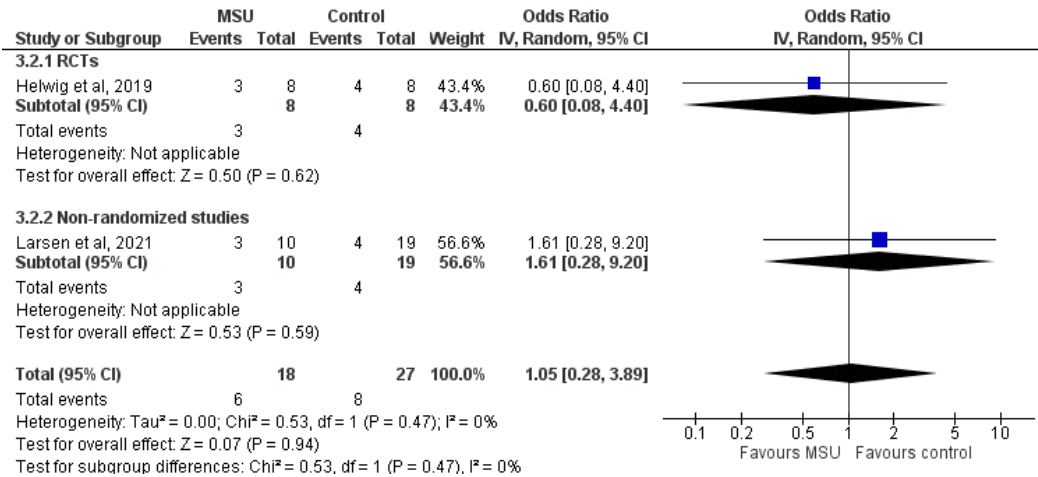


Quality: Low ⊕⊕
(Imprecision)

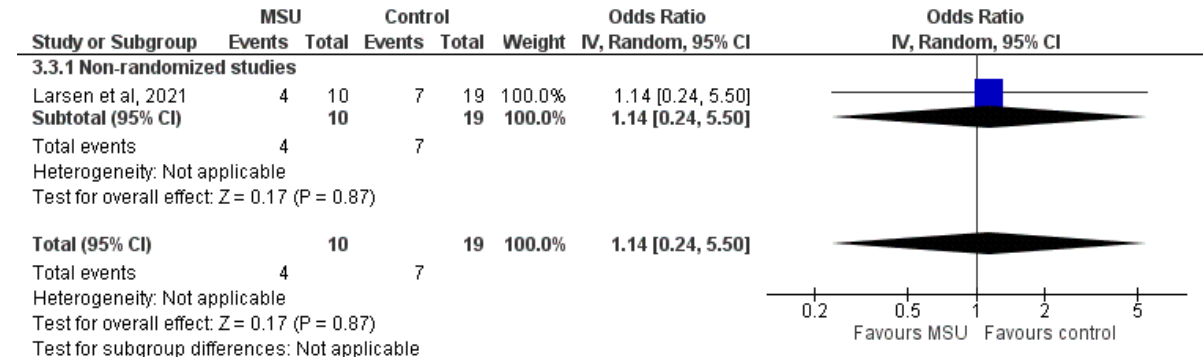
Acute Intracranial Haemorrhage Patients

MSU vs conventional management

Safety outcome: all-cause mortality (7/90 days)



Quality: Very low ⊕
(Bias, Imprecision)



Quality: Very low ⊕
(Imprecision)

Expert Consensus Statement

Acute Intracranial Haemorrhage Patients

In confirmed **acute intracerebral haemorrhage** patients, we **suggest** prehospital **management with Mobile Stroke Units** over conventional management because the **timely transport** of these patients **to tertiary stroke centres** is crucial for optimal therapeutic management.

Delphi voting result: 6/8 for and 2/8 against the statement

Evidence-based Recommendation

MSU care vs conventional management

We **suggest** the use of Mobile Stroke Units over conventional care for the prehospital management of patients with suspected stroke, for the following reasons:

- In patients with **acute ischaemic stroke**, prehospital management with a MSU **improves functional outcomes, increases the rates of treatment** with intravenous thrombolysis, **including** the rates of thrombolysis within the **golden hour** and **shortens onset to treatment** time without any safety concerns.

Quality of evidence: **Moderate** ⊕⊕⊕

Evidence-based Recommendation (*continued*)

MSU care vs conventional management

- In patients with **intracranial haemorrhage**, prehospital management with a MSU **increases** the proportion of **primary transport to tertiary care stroke** centres, without concerns on short-term mortality.

Quality of evidence: **Low** ⊕⊕

- In **other patients** (e.g. stroke mimics), **no signal of safety concerns** was identified.

Quality of evidence: **Very low** ⊕

Overall strength of recommendation: Weak ↑

Expert Consensus Statement

MSU staffing

When considering **MSU care**, to maintain the same benefits of clinical studies in routine practice and based on the current evidence, **including specialist neurological expertise** either by an in-person stroke expert or by remote consultation and a streamlined process of care **are essential**.

Delphi voting result: 8/8 for the statement

Conclusion

- Prehospital stroke management has evolved
- Evidence-based recommendation to manage suspected stroke patients in the prehospital setting
- Further research needed in:
 - Dispatch accuracy
 - Different geographical settings (rural)
 - Cost-effectiveness
 - Novel diagnostic equipment
- Ongoing studies will add more evidence (e.g. STOP-MSU, TASTEa, B_PROUD 2.0, ASPHALT)

Risk of bias in each study reporting data on excellent functional outcome (mRS 0-1 at 90 days)

	Confounding	Selection of participants	Classification of interventions	Deviation from intended interventions	Missing data	Measurement of outcomes	Selection of the reported result	Overall risk of bias
B_PROUD - Ebinger et al JAMA 2021	+	+	+	+	+	+	+	Low risk
BEST-MSU - Grotta et al NEJM 2021	+	+	+	+	+	+	+	Low risk
Kunz et al Lancet Neurol 2016	+	-	+	+	+	-	+	Moderate risk
Larsen et al Eur J Neurol 2021	+	-	+	?	+	+	+	Moderate risk

Legend:

- Low risk
- Moderate risk
- Serious risk
- Critical risk
- No information

according to the ROBINS-I tool