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Organised inpatient (stroke unit) care for stroke: network meta-analysis (Review)

Langhorne P, Ramachandra S, Stroke Unit Trialists' Collaboration

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[Intervention Review]

Organised inpatient (stroke unit) care for stroke: network meta-analysis

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ABSTRACT

Background

Organised inpatient (stroke unit) care is provided by multi-disciplinary teams that manage stroke patients. This can be provided in a ward dedicated to stroke patients (stroke ward), with a peripatetic stroke team (mobile stroke team), or within a generic disability service (mixed rehabilitation ward). Team members aim to provide co-ordinated multi-disciplinary care using standard approaches to manage common post-stroke problems.

Objectives

- To assess the effects of organised inpatient (stroke unit) care compared with an alternative service.
- To use a network meta-analysis (NMA) approach to assess different types of organised inpatient (stroke unit) care for people admitted to hospital after a stroke (the standard comparator was care in a general ward).

Originally, we conducted this systematic review to clarify:

- The characteristic features of organised inpatient (stroke unit) care?
- Whether organised inpatient (stroke unit) care provide better patient outcomes than alternative forms of care?
- If benefits are apparent across a range of patient groups and across different approaches to delivering organised stroke unit care?

Within the current version, we wished to establish whether previous conclusions were altered by the inclusion of new outcome data from recent trials and further analysis via NMA.

Search methods

We searched the Cochrane Stroke Group Trials Register (2 April 2019); the Cochrane Central Register of Controlled Trials (CENTRAL; 2019, Issue 4), in the Cochrane Library (searched 2 April 2019); MEDLINE Ovid (1946 to 1 April 2019); Embase Ovid (1974 to 1 April 2019); and the Cumulative Index to Nursing and Allied Health Literature (CINAHL; 1982 to 2 April 2019). In an effort to identify further published, unpublished, and ongoing trials, we searched seven trial registries (2 April 2019). We also performed citation tracking of included studies, checked reference lists of relevant articles, and contacted trialists.

Selection criteria

Randomised controlled clinical trials comparing organised inpatient stroke unit care with an alternative service (typically contemporary conventional care), including comparing different types of organised inpatient (stroke unit) care for people with stroke who are admitted to hospital.

Data collection and analysis

Two review authors assessed eligibility and trial quality. We checked descriptive details and trial data with co-ordinators of the original trials, assessed risk of bias, and applied GRADE. The primary outcome was poor outcome (death or dependency (Rankin score 3 to 5) or requiring institutional care) at the end of scheduled follow-up. Secondary outcomes included death, institutional care, dependency, subjective health status, satisfaction, and length of stay. We used direct (pairwise) comparisons to compare organised inpatient (stroke unit) care with an alternative service. We used an NMA to confirm the relative effects of different approaches.

Main results

We included 29 trials (5902 participants) that compared organised inpatient (stroke unit) care with an alternative service: 20 trials (4127 participants) compared organised (stroke unit) care with a general ward, six trials (982 participants) compared different forms of organised (stroke unit) care, and three trials (793 participants) incorporated more than one comparison.

Compared with the alternative service, organised inpatient (stroke unit) care was associated with improved outcomes at the end of scheduled follow-up (median one year): poor outcome (odds ratio (OR) 0.77, 95% confidence interval (CI) 0.69 to 0.87; moderate-quality evidence), death (OR 0.76, 95% CI 0.66 to 0.88; moderate-quality evidence), death or institutional care (OR 0.76, 95% CI 0.67 to 0.85; moderate-quality evidence), and death or dependency (OR 0.75, 95% CI 0.66 to 0.85; moderate-quality evidence). Evidence was of very low quality for subjective health status and was not available for patient satisfaction. Analysis of length of stay was complicated by variations in definition and measurement plus substantial statistical heterogeneity ($I^2 = 85\%$). There was no indication that organised stroke unit care resulted in a longer hospital stay. Sensitivity analyses indicated that observed benefits remained when the analysis was restricted to securely randomised trials that used unequivocally blinded outcome assessment with a fixed period of follow-up. Outcomes appeared to be independent of patient age, sex, initial stroke severity, stroke type, and duration of follow-up. When calculated as the absolute risk difference for every 100 participants receiving stroke unit care, this equates to two extra survivors, six more living at home, and six more living independently.

The analysis of different types of organised (stroke unit) care used both direct pairwise comparisons and NMA.

Direct comparison of stroke ward versus general ward: 15 trials (3523 participants) compared care in a stroke ward with care in general wards. Stroke ward care showed a reduction in the odds of a poor outcome at the end of follow-up (OR 0.78, 95% CI 0.68 to 0.91; moderate-quality evidence).

Direct comparison of mobile stroke team versus general ward: two trials (438 participants) compared care from a mobile stroke team with care in general wards. Stroke team care may result in little difference in the odds of a poor outcome at the end of follow-up (OR 0.80, 95% CI 0.52 to 1.22; low-quality evidence).

Direct comparison of mixed rehabilitation ward versus general ward: six trials (630 participants) compared care in a mixed rehabilitation ward with care in general wards. Mixed rehabilitation ward care showed a reduction in the odds of a poor outcome at the end of follow-up (OR 0.65, 95% CI 0.47 to 0.90; moderate-quality evidence).

In a NMA using care in a general ward as the comparator, the odds of a poor outcome were as follows: stroke ward - OR 0.74, 95% CI 0.62 to 0.89, moderate-quality evidence; mobile stroke team - OR 0.88, 95% CI 0.58 to 1.34, low-quality evidence; mixed rehabilitation ward - OR 0.70, 95% CI 0.52 to 0.95, low-quality evidence.

Authors' conclusions

We found moderate-quality evidence that stroke patients who receive organised inpatient (stroke unit) care are more likely to be alive, independent, and living at home one year after the stroke. The apparent benefits were independent of patient age, sex, initial stroke severity, or stroke type, and were most obvious in units based in a discrete stroke ward. We observed no systematic increase in the length of inpatient stay, but these findings had considerable uncertainty.

PLAIN LANGUAGE SUMMARY

Organised inpatient (stroke unit) care

Review question

Does organised inpatient (stroke unit) care improve the recovery of people with stroke in hospital compared with conventional care in general wards?

Background

Organised inpatient (stroke unit) care is a form of care provided in hospital by nurses, doctors, and therapists who specialise in looking after people with stroke. They aim to work as a co-ordinated team to provide the most appropriate care tailored to the needs of individual people with stroke.

Study characteristics

We identified 29 trials involving 5902 participants (search completed 2 April 2019). Participants who were recruited had had a recent stroke and required admission to hospital. Organised inpatient (stroke unit) care was provided in a variety of ways including stroke ward (care provided in a discrete stroke ward), mixed rehabilitation ward (setting seeking to improve care for people with stroke within a mixed rehabilitation ward), and mobile stroke team (peripatetic team looking after people with stroke across a range of wards).

Key results

At an average of 12 months after their stroke, people who received organised inpatient (stroke unit) care were more likely to be alive (an extra two people surviving for every 100 receiving stroke unit care; moderate-quality evidence) and living at home (an extra six patients for every 100 receiving stroke unit care; moderate-quality evidence). They also were more likely to be independent in daily activities (an extra six patients for every 100 receiving stroke unit care; moderate-quality evidence). The apparent benefits were seen in men and women, older and younger patients, and people with different types of stroke and different stroke severity. Benefits were most obvious when the stroke unit was based in a discrete stroke ward.

Quality of the evidence

We downgraded the quality of evidence to 'moderate' for the main outcomes because it was impossible to hide the treating service from participants or healthcare workers. These conclusions were not dependent on trials judged to be of lower quality because of poor design or missing data. More information was missing for some of the other outcome measures and analyses, which we have downgraded to low-quality evidence.

Conclusion

People with stroke who receive organised inpatient (stroke unit) care are more likely to be alive, living at home, and independent in looking after themselves one year after their stroke. Apparent benefits were seen across a broad range of people with stroke. Various types of stroke units have been developed. The best results appear to come from stroke units based in a dedicated stroke ward.

SUMMARY OF FINDINGS

Summary of findings 1. Organised inpatient (stroke unit) care versus alternative service

Organised inpatient (stroke unit) care compared with alternative service

Patient or population: adults with acute stroke

Settings: hospital

Intervention: organised inpatient (stroke unit) care

Comparison: alternative service (contemporary conventional care)

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	Number of participants (studies)	Quality of the evidence (GRADE)	Comments
	Assumed risk	Corresponding risk				
	Alternative service	Organised inpatient (stroke unit) care				
Poor outcome by the end of scheduled follow-up (modified Rankin score 3 to 6 or requiring institutional care; median 12-month follow-up) (Analysis 1.1)	577 per 1000	517 per 1000 (497 to 547)	OR 0.77 (0.69 to 0.87)	5336 (26)	⊕⊕⊕⊖ moderate ^a	Sensitivity analysis based on trial quality suggested no alteration of conclusions
Death by the end of scheduled follow-up (median 12-month follow-up) (Analysis 1.2)	219 per 1000	199 per 1000 (179 to 209)	OR 0.76 (0.66 to 0.88)	5902 (29)	⊕⊕⊕⊖ moderate ^a	Sensitivity analysis based on trial quality suggested no alteration of conclusions
Death or institutional care by the end of scheduled follow-up (median 12-month follow-up) (Analysis 1.3)	405 per 1000	345 per 1000 (315 to 375)	OR 0.76 (0.67 to 0.85)	4887 (25)	⊕⊕⊕⊖ moderate ^a	Sensitivity analysis based on trial quality suggested no alteration of conclusions
Death or dependency by the end of scheduled follow-up	609 per 1000	549 per 1000 (519 to 567)	OR 0.75 (0.66 to 0.85)	4854 (24)	⊕⊕⊕⊖ moderate ^a	Sensitivity analysis based on trial quality suggested no alteration of conclusions

(modified Rankin score 3 to 6; median 12-month follow-up) (Analysis 1.4)								
Subjective health status score	There was a pattern of improved results among stroke unit survivors, with results attaining statistical significance in 2 individual trials	N/A	843	⊕⊕⊕⊕		Data from 3 trials only		
Participant quality of life (Nottingham Health Profile; Quality of Life Scale)			(3)	very low ^{a,b,c}		High rate of missing data		
Patient satisfaction or preference	We could find no systematically gathered information on patient preferences	N/A	N/A	N/A		No data available		
Length of stay (days) in a hospital or institution (Analysis 1.5)	<table border="0"> <tr> <td>Mean length of stay across the control groups ranged from 12.1 to 123 days</td> <td>Mean length of stay for the intervention groups was, on average, 4.3 days less (7.9 days less to 0.7 days more)</td> </tr> </table>	Mean length of stay across the control groups ranged from 12.1 to 123 days	Mean length of stay for the intervention groups was, on average, 4.3 days less (7.9 days less to 0.7 days more)	SMD 0.16 lower (0.33 lower to 0.01 higher)	4162 (19)	⊕⊕⊕⊕ low ^{a,b}		Different definitions and imprecise measures of length of stay were reported
Mean length of stay across the control groups ranged from 12.1 to 123 days	Mean length of stay for the intervention groups was, on average, 4.3 days less (7.9 days less to 0.7 days more)							

*The basis for the **assumed risk** (e.g. median control group risk across studies) is provided in footnotes. The **corresponding risk** (and its 95% confidence interval) is based on the assumed risk in the comparison group and the **relative effect** of the intervention (and its 95% CI).
CI: confidence interval; N/A: not applicable; OR: odds ratio; SMD: standardised mean difference.

GRADE Working Group grades of evidence.

High quality: further research is very unlikely to change our confidence in the estimate of effect.

Moderate quality: further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.

Low quality: further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

Very low quality: we are very uncertain about the estimate.

^aDowngraded for potential risk of performance bias.

^bDowngraded for unexplained heterogeneity.

^cDowngraded for imprecision

Summary of findings 2. Stroke ward versus general medical ward

Organised inpatient (stroke unit) care compared with general medical ward care for stroke

Patient or population: adults with acute stroke

Settings: hospital

Intervention: stroke ward care

Comparison: general medical ward care

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	Number of participants (studies)	Quality of the evidence (GRADE)	Comments
	Assumed risk	Corresponding risk				
	General medical ward care	Stroke ward care				
Poor outcome by the end of scheduled follow-up (modified Rankin score 3 to 6 or requiring institutional care; median 12-month follow-up) (Analysis 2.1)	549 per 1000	499 per 1000 (459 to 529)	OR 0.78 (0.68 to 0.91)	3321 (14)	⊕⊕⊕⊖ moderate ^d	Sensitivity analysis based on trial quality suggested no alteration of conclusions
Death by the end of scheduled follow-up (median 12-month follow-up) (Analysis 2.2)	242 per 1000	202 per 1000 (172 to 222)	OR 0.75 (0.63 to 0.90)	3523 (15)	⊕⊕⊕⊖ moderate ^d	Sensitivity analysis based on trial quality suggested no alteration of conclusions
Death or institutional care by the end of scheduled follow-up (median 12-month follow-up) (Analysis 2.3)	383 per 1000	323 per 1000 (283 to 353)	OR 0.74 (0.63 to 0.87)	2924 (13)	⊕⊕⊕⊖ moderate ^d	Sensitivity analysis based on trial quality suggested no alteration of conclusions
Death or dependency by the end of scheduled follow-up (modified Rankin score 3 to 6; median 12-month follow-up) (Analysis 2.4)	602 per 1000	532 per 1000 (502 to 572)	OR 0.75 (0.64 to 0.88)	2839 (12)	⊕⊕⊕⊖ moderate ^d	Sensitivity analysis based on trial quality suggested no alteration of conclusions
Subjective health status score Participant quality of life (Nottingham Health Profile; Quality of Life Scale)	There was a pattern of improved results among stroke unit survivors, with results attaining statistical significance in 2 individual trials		N/A	535 (2)	⊕⊕⊕⊖ very low ^{a,b,c}	Data from 3 trials only High rate of missing data
Patient satisfaction or preference	We could find no systematically gathered information on patient preferences		N/A	N/A	N/A	No data available
Length of stay (days) in a hospital or institution (Analysis 2.5)	Mean length of stay across control groups	Mean length of stay for the intervention groups was, on average, 2.2 days less (5.2	SMD 0.13 lower (0.29 lower to 0.04 higher)	2547 (10)	⊕⊕⊕⊖ low ^{a,b}	Different definitions and imprecise measures of length of stay were reported

ranged from 12.8 days less to 0.8 days more)
to 123 days

*The basis for the **assumed risk** (e.g. median control group risk across studies) is provided in footnotes. The **corresponding risk** (and its 95% confidence interval) is based on the assumed risk in the comparison group and the **relative effect** of the intervention (and its 95% CI).
CI: confidence interval; N/A: not applicable; OR: odds ratio; SMD: standardised mean difference.

GRADE Working Group grades of evidence.

High quality: further research is very unlikely to change our confidence in the estimate of effect.

Moderate quality: further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.

Low quality: further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

Very low quality: we are very uncertain about the estimate.

^aDowngraded for potential risk of performance bias.

^bDowngraded for unexplained heterogeneity.

^cDowngraded for imprecision

Summary of findings 3. Mobile stroke team versus general medical ward

Mobile stroke team care compared with general medical ward care for stroke

Patient or population: adults with acute stroke

Settings: hospital

Intervention: mobile stroke team care

Comparison: general medical ward care

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	Number of participants (studies)	Quality of the evidence (GRADE)	Comments
	Assumed risk	Corresponding risk				
	General medical ward care	Mobile stroke team care				
Poor outcome by the end of scheduled follow-up (modified Rankin score 3 to 6 or requiring institutional care; median 12-month follow-up) (Analysis 3.1)	712 per 1000	672 per 1000 (582 to 752)	OR 0.80 (0.52 to 1.22)	438 (2)	⊕⊕⊕⊕ low ^{a,b}	As dependency data were complete, these are the same data as for death or dependency

Death by the end of scheduled follow-up (median 12-month follow-up) (Analysis 3.2)	259 per 1000 (189 to 359)	279 per 1000 (189 to 359)	OR 1.08 (0.71 to 1.65)	438 (2)	⊕⊕○○ lowa,b	
Death or institutional care by the end of scheduled follow-up (median 12-month follow-up) (Analysis 3.3)	481 per 1000	531 per 1000 (451 to 611)	OR 1.27 (0.84 to 1.93)	438 (2)	⊕⊕○○ lowa,b	
Death or dependency by the end of scheduled follow-up (modified Rankin score 3 to 6; median 12-month follow-up) (Analysis 3.4)	712 per 1000	672 per 1000 (582 to 752)	OR 0.80 (0.52 to 1.22)	438 (2)	⊕⊕○○ lowa,b	As dependency data were complete, these are the same data as for poor outcome
Subjective health status score Participant quality of life (EuroQol)	No apparent differences between groups		N/A	308 (1)	⊕○○○ very lowa,b	Data from 1 trial only
Patient satisfaction or preference	We could find no systematically gathered information on patient preferences		N/A	N/A	N/A	No data available
Length of stay (days) in a hospital or institution (Analysis 3.5)	No data available		N/A	N/A	N/A	No data available

*The basis for the **assumed risk** (e.g. median control group risk across studies) is provided in footnotes. The **corresponding risk** (and its 95% confidence interval) is based on the assumed risk in the comparison group and the **relative effect** of the intervention (and its 95% CI).
CI: confidence interval; N/A: not applicable; OR: odds ratio.

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Low quality: further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

Very low quality: we are very uncertain about the estimate.

^aDowngraded for potential risk of performance bias.

^bDowngraded for imprecision.

Summary of findings 4. Mixed rehabilitation ward versus general medical ward
Mixed rehabilitation ward care compared with general medical ward care for stroke
Patient or population: adults with acute stroke

Settings: hospital

Intervention: mixed rehabilitation ward care

Comparison: general medical ward care

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	Number of participants (studies)	Quality of the evidence (GRADE)	Comments
	Assumed risk	Corresponding risk				
	General medical ward care	Mixed rehabilitation wardcare				
Poor outcome by the end of scheduled follow-up (modified Rankin score 3 to 6 or requiring institutional care; median 12-month follow-up) (Analysis 4.1)	574 per 1000	474 per 1000 (404 to 554)	OR 0.65 (0.47 to 0.90)	630 (6)	⊕⊕⊕⊕ low ^{a,b}	As dependency data were complete, these are the same data as for death or dependency
Death by the end of scheduled follow-up (median 12-month follow-up) (Analysis 4.2)	171 per 1000	161 per 1000 (101 to 211)	OR 0.91 (0.58 to 1.42)	630 (6)	⊕⊕⊕⊕ low ^{a,b}	
Death or institutional care by the end of scheduled follow-up (median 12-month follow-up) (Analysis 4.3)	451 per 1000	371 per 1000 (291 to 451)	OR 0.71 (0.51 to 0.99)	578 (5)	⊕⊕⊕⊕ low ^{a,b}	
Death or dependency by the end of scheduled follow-up (modified Rankin score 3 to 6; median 12-month follow-up) (Analysis 4.4)	574 per 1000	474 per 1000 (404 to 554)	OR 0.65 (0.47 to 0.90)	630 (6)	⊕⊕⊕⊕ low ^{a,b}	As dependency data were complete, these are the same data as for poor outcome
Subjective health Status score	No data available		N/A	N/A	N/A	No data available

Patient satisfaction or preference	We could find no systematically gathered information on patient preferences		N/A	N/A	N/A	No data available
Length of stay (days) in a hospital or institution (Analysis 4.5)	Mean length of stay across control groups ranged from 30.5 to 129.5 days	Mean length of stay for the intervention groups was, on average, 3.9 days more (13.5 days less to 21.5 days more)	MD 0.08 more (0.21 lower to 0.37 higher)	387 (3)	⊕⊕⊕⊕ very low ^{a,b,c}	Different definitions and imprecise measures of length of stay were reported

*The basis for the **assumed risk** (e.g. median control group risk across studies) is provided in footnotes. The **corresponding risk** (and its 95% confidence interval) is based on the assumed risk in the comparison group and the **relative effect** of the intervention (and its 95% CI).

CI: confidence interval; MD: mean difference; N/A: not applicable; OR: odds ratio.

GRADE Working Group grades of evidence.

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^bDowngraded for imprecision.

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BACKGROUND

Description of the condition

Stroke is now the third leading cause of disability (Murray 2012), and it is the second leading cause of mortality worldwide (Lozano 2012). The global disease burden of stroke increased by 19% between 1990 and 2010 (Murray 2012), and current projections estimate the number of deaths worldwide will rise to 6.5 million in 2015 and to 7.8 million in 2030 (Strong 2007). Interventions that are applicable to a majority of people with stroke and that aim to reduce associated mortality and disability are essential.

During their initial illness, people with stroke are frequently admitted to hospital, where they can receive care in a variety of ways and in a range of settings. Traditionally, care for people with stroke was provided within departments of general (internal) medicine, neurology, or medicine for the elderly, where they would be managed alongside a range of other patient groups. A more focused approach to the treatment of people with stroke in hospital has been developed.

Description of the intervention

Organised inpatient (stroke unit) care is the term used to describe focused care for people with stroke in hospital under a multi-disciplinary team of individuals who specialise in stroke management (SUTC 1997a). This concept is not new, and its value has been debated for more than 30 years (Ebrahim 1990; Garraway 1985; Langhorne 1993; Langhorne 1998; Langhorne 2012). In essence, the debate has concerned whether the perceived effort and cost of focusing the care of people hospitalised with stroke within specially organised units would be matched by tangible benefits for the people receiving that care. In particular, would more people survive and make a good recovery as a result of organised inpatient (stroke unit) care?

Why it is important to do this review

A systematic review of all available trials previously described the range of characteristics of stroke unit care and addressed the question of whether improving the organisation of inpatient stroke care can bring about improvements in important patient outcomes (SUTC 1997a). This review continues to be extended and updated within the Cochrane Library (SUTC 2001; SUTC 2007; SUTC 2013). Since the last update, network meta-analysis (NMA) has become established as an approach for handling multiple comparisons. We have added NMA to our updated review.

OBJECTIVES

- To assess the effects of organised inpatient (stroke unit) care compared with an alternative service (usually contemporary conventional care)
- To use a network meta-analysis (NMA) approach to assess different types of organised inpatient (stroke unit) care for people admitted to hospital after a stroke (the standard comparator was care in a general ward)

Originally, this systematic review was conducted to address four broad questions.

- What are the characteristic features of organised inpatient (stroke unit) care?

- Can organised inpatient (stroke unit) care provide better patient outcomes than alternative forms of care?
- Are any benefits apparent across a range of patient groups?
- Are any benefits apparent across different approaches to delivering organised stroke unit care? In particular, we hypothesised that organised care would be more effective than care provided in general medical wards, but that different forms of organised care would achieve similar outcomes.

Within the current version of this review, we wished to establish whether previous conclusions were altered by the inclusion of new outcome data from recent trials and further analysis via NMA.

METHODS

Criteria for considering studies for this review

Types of studies

We included all randomised controlled clinical trials that compared an organised system of inpatient (stroke unit) care with an alternative form of inpatient care. This was usually contemporary conventional care but could include an alternative model of organised inpatient (stroke unit) care (see [Types of interventions](#)). Previous versions of this review included trials with quasi-random treatment allocation (such as bed availability or date of admission) (SUTC 1997a; SUTC 2001; SUTC 2007). However, in an effort to ensure that this ongoing systematic review focuses on data from trials with strict randomisation procedures, we excluded all quasi-randomised trials in the previous update (SUTC 2013). We would have included cluster-randomised trials, but we identified none. We excluded cross-over trials because of cross-over of effects.

Types of participants

Any person admitted to hospital who had suffered a stroke was eligible. We recorded the delay between stroke onset and hospital admission but did not use this as an exclusion criterion. We used a clinical definition of stroke: focal neurological deficit due to cerebrovascular disease, excluding subarachnoid haemorrhage and subdural haematoma.

Types of interventions

Organised inpatient (stroke unit) care can be considered a complex organisational intervention comprising multi-disciplinary staff providing a complex package of care to people with stroke in hospital. In the original version of this review, the first question was whether organised inpatient (stroke unit) care could improve outcomes compared with contemporary conventional care (usually in general wards) (SUTC 1997a). We then had to modify the analyses in a minor way to reflect the emerging pattern of service organisation and to allow the comparison of 'more organised' versus 'less organised' services (for which the latter was usually contemporary conventional care). We did this because some recent trials have addressed new questions and included comparisons of two services, both of which met the basic definition of organised (stroke unit) care. In the original service descriptions used in this review (SUTC 1997a), service organisation was considered as a hierarchy comprising the following.

- Stroke ward: where a multi-disciplinary team including specialist nursing staff based in a discrete ward cares exclusively

for people with stroke. This category included the following subdivisions.

- * Acute stroke units that accept patients acutely but discharge early (usually within seven days); these appear to fall into three broad subcategories.
 - 'Intensive' model of care with continuous monitoring, high nurse staffing levels, and the potential for life support.
 - 'Semi-intensive' model of care with continuous monitoring, high nurse staffing, but no life support facilities.
 - 'Non-intensive' model of care with none of the above.
- * Rehabilitation stroke units that accept patients after a delay, usually of seven days or longer, and focus on rehabilitation.
- * Comprehensive (i.e. combined acute and rehabilitation) stroke units that accept patients acutely but also provide rehabilitation for at least several weeks if necessary. Both the rehabilitation unit model and the comprehensive unit model offer prolonged periods of rehabilitation.
- Mixed rehabilitation ward: where a multi-disciplinary team including specialist nursing staff in a ward provides a generic rehabilitation service but not exclusively caring for people with stroke.
- Mobile stroke team: where a peripatetic multi-disciplinary team (usually excluding specialist nursing staff) provides care in a variety of settings.
- General medical ward: where care is provided in an acute medical or neurology ward without routine multi-disciplinary input.

For the NMA eligibility assessment, we considered the transitivity (similarity) of trials included in the network (see [Data synthesis](#)), which requires all interventions to be legitimate alternatives. All of the four main categories have been used to provide care for unselected acute stroke patients and can be considered broadly comparable for the purpose of an NMA. The general medical ward group was the reference group,

Types of outcome measures

Primary outcomes

In the previous version of this review, the primary analyses examined death, dependency, and the requirement for institutional care at the end of scheduled follow-up of the original trial ([SUTC 2013](#)). We categorised dependency into two groups, where we took 'independent' to mean that an individual did not require physical assistance for transfers, mobility, dressing, feeding, or toileting. We considered individuals who failed any of these criteria 'dependent'. The criteria for independence were approximately equivalent to a modified Rankin score of 0 to 2 or a Barthel Index greater than 18 out of 20 ([Wade 1992](#)). We took the requirement for long-term institutional care to mean care in a residential home, a nursing home, or a hospital at the end of scheduled follow-up.

In view of changes in reporting standards, for this update we have now provided a single composite primary outcome: poor outcome: death or dependency or requiring institutional care (if dependency data were not available). This allowed us to keep the primary focus of the review (i.e. the focus on independent survival as an outcome), while optimising the quantity of data available.

Secondary outcomes

Secondary outcome measures now include:

- death;
- death or institutional care;
- death or dependency;
- patient subjective health status (measured using tools such as the Nottingham Health Profile, EuroQoL, Short Form-36);
- patient and carer satisfaction (recorded on a Likert scale or as responses to statements); and
- duration of stay in hospital or institution or both.

Outcomes are reported at the end of scheduled follow-up. Some trials subsequently provided supplementary extended follow-up data, which are presented separately.

Search methods for identification of studies

See the methods for the Cochrane Stroke Group [Specialised register](#). We searched for trials in all languages and arranged the translation of relevant papers published in languages other than English.

Electronic searches

We searched the trials registers of the Cochrane Stroke Group (2 April 2019). In addition, in collaboration with the Cochrane Stroke Group Information Specialist, we searched the Cochrane Central Register of Controlled Trials (CENTRAL; 2019; Issue 4), in the Cochrane Library (searched 2 April 2019) ([Appendix 1](#)); MEDLINE Ovid (1946 to 1 April 2019) ([Appendix 2](#)); Embase Ovid (1974 to 1 April 2019) ([Appendix 3](#)); and the Cumulative Index to Nursing and Allied Health Literature (CINAHL) EBSCO (1982 to 2 April 2019) ([Appendix 4](#)).

We searched the following registers of ongoing trials using the keyword 'stroke'.

- US National Institutes of Health Ongoing Trials Register ClinicalTrials.gov (www.clinicaltrials.gov; searched 2 April 2019).
- World Health Organization International Clinical Trials Registry Platform (apps.who.int/trialsearch; searched 2 April 2019).
- CenterWatch Clinical Trials Listing Service (www.centerwatch.com; searched 13 August 2018).
- Community Research & Development Information Service (of the European Union) (cordis.europa.eu/en/home.html; searched 13 August 2018).
- South African National Clinical Trial Register (www.sanctr.gov.za; searched 13 August 2018).
- The Internet Stroke Center - Stroke Trials Registry (www.strokecenter.org/trials; searched 13 August 2018).
- Clinical Trials Results register (www.clinicaltrialresults.org; searched 2 April 2019).

Searching other resources

In an effort to identify further published, unpublished, and ongoing trials, we:

- performed citation tracking using Web of Science Cited Reference Search for all included studies;

- searched the reference lists of included trials and all relevant articles;
- obtained further information from individual trialists; and
- contacted other researchers in the field and publicised our preliminary findings at stroke conferences in UK, Scandinavia, Germany, Netherlands, Switzerland, Spain, Canada, Brazil, Argentina, Australia, Belgium, USA, India, Sri Lanka, Singapore, Italy, and Hong Kong.

Data collection and analysis

Selection of studies

For this updated review, one review author (PL) read the titles and abstracts of records obtained through the electronic searches and excluded obviously irrelevant studies. We obtained full copies of the remaining studies, and two review authors (PL and SR) independently selected studies for inclusion based on the following eligibility criteria.

- Randomised controlled trial.
- Service intervention providing a form of organised inpatient (stroke unit) care.
- Service aim to improve functional recovery and survival after stroke.
- Trial of stroke patients.

We tried to establish the characteristics of unpublished trials through discussion with the Cochrane Stroke Group Information Specialist before analysing the results.

Data extraction and management

If possible, the principal review author (PL) obtained descriptive information about the service characteristics of organised inpatient (stroke unit) care and conventional care settings through a structured interview or correspondence conducted with the trial co-ordinators ($n = 19$). We obtained additional information from published sources. We then allocated trials to service subgroups. We confirmed outcome data from published sources and supplemented them with unpublished information provided by the co-ordinator of each individual trial. We asked trialists to provide information on the number of participants who were dead or dependent and the number requiring institutional care or missing at the end of scheduled follow-up. For this updated review, two review authors (PL, SR) independently extracted information using a standard data extraction form.

We sought subgroup information on potential effect modifiers primarily for the combined outcome of death or requiring institutional care. We obtained unpublished aggregated data for a majority of trials, but insufficient quantities of individual patient data were available to allow a comprehensive individual patient data analysis.

We obtained subgroup data regarding the following participant groups (see [SUTC 1997a](#) for details).

- Age: up to 75 years or 75 years or older.
- Sex: men or women.

- Stroke severity: dependency at the time of randomisation (usually within one week of the index stroke).
 - * Mild stroke: equivalent to a Barthel Index of 10 to 20 out of 20 during the first week.
 - * Moderate stroke: equivalent to a Barthel Index of 3 to 9 out of 20 during the first week.
 - * Severe stroke: equivalent to a Barthel Index of 0 to 2 out of 20 during the first week.
- Stroke type: ischaemic or haemorrhagic based on neuroimaging.

Assessment of risk of bias in included studies

We assessed risk of bias using Cochrane's 'Risk of bias' tool, as described in Chapter 8 of the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2011). Two review authors identified the method of random sequence generation, the method of concealment of treatment allocation, blinding of participants and personnel, the presence of blinding of outcome assessment, completeness of follow-up, and evidence of selective reporting. We used these potentially important factors in sensitivity analyses, but we did not use them as exclusion criteria. The principal review author then used this information to inform the categorisations within the 'Summary of findings' tables and the GRADE allocations.

Measures of treatment effect

When our analyses of poor outcome, death, dependency, or institutionalisation at the end of scheduled follow-up were reported, we analysed these using the odds ratio (OR) and the 95% confidence interval (CI) for an adverse outcome.

We aimed to record length of stay in hospital or in an institution as the mean and standard deviation (SD). When only medians were available, we assumed these were approximate to the mean. When no other data were provided with the mean value, we inferred the SD as being at least as large as those in comparable trials using the same measure. Because length of stay was reported in a variety of ways, we checked the results obtained with the mean difference (MD) using the standardised mean difference (SMD) and the 95% CI.

We anticipated that measures of subjective health status would be analysed as mean differences, and measures of satisfaction would be analysed as odds ratios for particular responses.

Unit of analysis issues

We anticipated that most trials would have a simple parallel-group design in which each individual was randomised to one of two treatment groups. When a trial had three (or more) treatment groups, we planned to analyse each treatment arm as a separate study. We have not included cross-over trials because of the likelihood or carryover effects.

Dealing with missing data

When data were missing for the outcome of death, dependency, or institutionalisation, we assumed the participant to be alive, independent, and living at home. We aimed to explore the implications of these assumptions in sensitivity analyses.

Assessment of heterogeneity

We planned to determine heterogeneity by visually inspecting the forest plot and by using the I^2 statistic. We defined

significant heterogeneity as I^2 greater than 50%. When significant heterogeneity occurred, we explored potential sources using pre-planned sensitivity analyses. Assessments of transitivity and consistency are discussed in the NMA section under [Data synthesis](#).

Assessment of reporting biases

We employed a comprehensive search strategy in an effort to avoid reporting biases. To identify unpublished studies, we searched trial registers and contacted trialists and other experts in the field. We planned to inspect funnel plots if enough studies were available.

Data synthesis

Pairwise comparisons

When we did not have access to individual patient data, two review authors (PL, SR) extracted data from published reports. When individual patient data were available, we checked them for internal consistency and consistency with published reports. One review author entered data into the Review Manager software ([RevMan 2014](#)), and a second review author checked the entries. We analysed binary outcome data using OR and 95% CI. We analysed continuous outcome data using SMD and 95% CI. By default, we used a fixed-effect model first, but we corroborated results by using a random-effects model if heterogeneity was significant.

When data were available, we carried out subgroup analyses for age, sex, stroke severity, and stroke type. Through subgroup analyses, we considered the degree of interaction between subgroups ([Higgins 2011](#)).

Network meta-analysis

In this version of the review, we include a newer approach to meta-analysis in the form of a network meta-analysis (NMA) of trial data. The original aim of this review was to compare the effects of organised inpatient (stroke unit) care versus conventional care. We expected that within this broad definition, the included trials would comprise a range of treatment comparisons (which could include stroke wards, mobile stroke teams, mixed rehabilitation wards) with conventional care in general wards. In addition, later trials have addressed newer questions comparing different forms of organised inpatient (stroke unit) care (e.g. stroke ward versus mobile stroke team).

We have retained the previous analysis, which was organised in a hierarchical manner (organised stroke care versus alternative service; organised stroke care versus general ward; different systems of organised care). However, we now include an NMA to explore, when possible, the impact of different systems of stroke care. We used *MetaInsight* software, which uses a frequentist approach and is designed specifically for this role - to enable us to conduct our NMA ([Owen 2019](#)).

An NMA uses information from both direct and indirect estimates of treatment effect ([Tonin 2017](#)). Direct estimates are provided by a head-to-head comparison (e.g. treatment A versus treatment B). Indirect estimates are provided by two or more head-to-head comparisons that share a common comparator (e.g. when A versus B is the comparison of interest, then trials with A versus C and with B versus C are used). A trial network is then formed, using trials that allow, through direct and indirect comparisons, calculation of the relative effects of all treatments versus each other (or versus a single comparator). The results of such analyses are usually

presented as comparisons against a common comparator group (e.g. general ward). It is also possible to present a rank analysis, which ranks treatment groups on the likelihood of being most/least effective.

A key assumption in NMA is that of transitivity (or similarity). This concerns the validity of making indirect comparisons and assumes that treatment effects are 'exchangeable' across the included trials, and that all treatments are 'jointly randomisable'. In other words, all treatment categories could feasibly be randomised in the same trial, and those that are not treatment arms in any given trial are 'missing at random' ([Lu 2006](#)). As this assumption cannot be formally tested statistically, it was judged through consideration of trial settings and characteristics, patient characteristics, and treatment mechanisms, and to investigate if any differences would be expected to modify relative treatment effects. Two review authors independently extracted the relevant information, and the principal review author made the final judgement.

A second important assumption (known as the consistency assumption) assumes that it is feasible to make indirect comparisons between two treatments, and that the indirect evidence is consistent with the direct evidence ([Lu 2006](#)). The consistency assumption can be evaluated statistically by comparing the difference between the direct estimate and the indirect estimate for each loop of evidence. Therefore, we examined for any important differences in numerical results between direct, indirect, and network results.

GRADE and 'Summary of findings'

We constructed 'Summary of findings' tables and used GRADE criteria to assess the quality of evidence. 'Summary of findings' tables included the new primary outcome (poor outcome) and the three main clinical outcomes included in previous reviews (death, death or requiring institutional care, death or dependency) plus subjective health status, patient satisfaction, and length of stay in a hospital or institution. All were recorded at the end of scheduled follow-up.

This update included an NMA whereby different types of organised inpatient (stroke unit) care were compared with care provided in a general ward (see below). For the NMA, we used the approach of the GRADE group as outlined below ([Brignardello 2018](#); [Puhan 2014](#)).

- Present direct and indirect treatment estimates for each comparison of the evidence network.
- Rate the quality of each direct and indirect effect estimate (downgrading for risk of bias, inconsistency, indirectness, imprecision, and publication bias).
- Present the NMA estimate for each comparison of the evidence network.
- Rate the quality of each NMA effect estimate (as above).

Subgroup analysis and investigation of heterogeneity

Subgroup analyses involved a re-analysis stratified by participant or service subgroup using tabular subgroup data provided by trialists or obtained from published sources. We used a fixed-effect approach unless heterogeneity was statistically significant, and all subgroup analyses considered the degree of interaction between subgroups ([Higgins 2011](#)). We applied subgroup analyses only to

the main (first) comparison to minimise the risk of false-positive results.

Sensitivity analysis

We planned sensitivity analyses around key aspects of risk of bias that we identified during our assessment of risk of bias (i.e. concealment of treatment allocation, blinding of outcome assessment, completeness of follow-up, and a fixed period of follow-up). We applied sensitivity analyses only to the main (first) comparison.

RESULTS

Description of studies

This is the fifth update of this Cochrane Review. The key references are described in the following relevant tables: [Characteristics of included studies](#); [Characteristics of excluded studies](#);

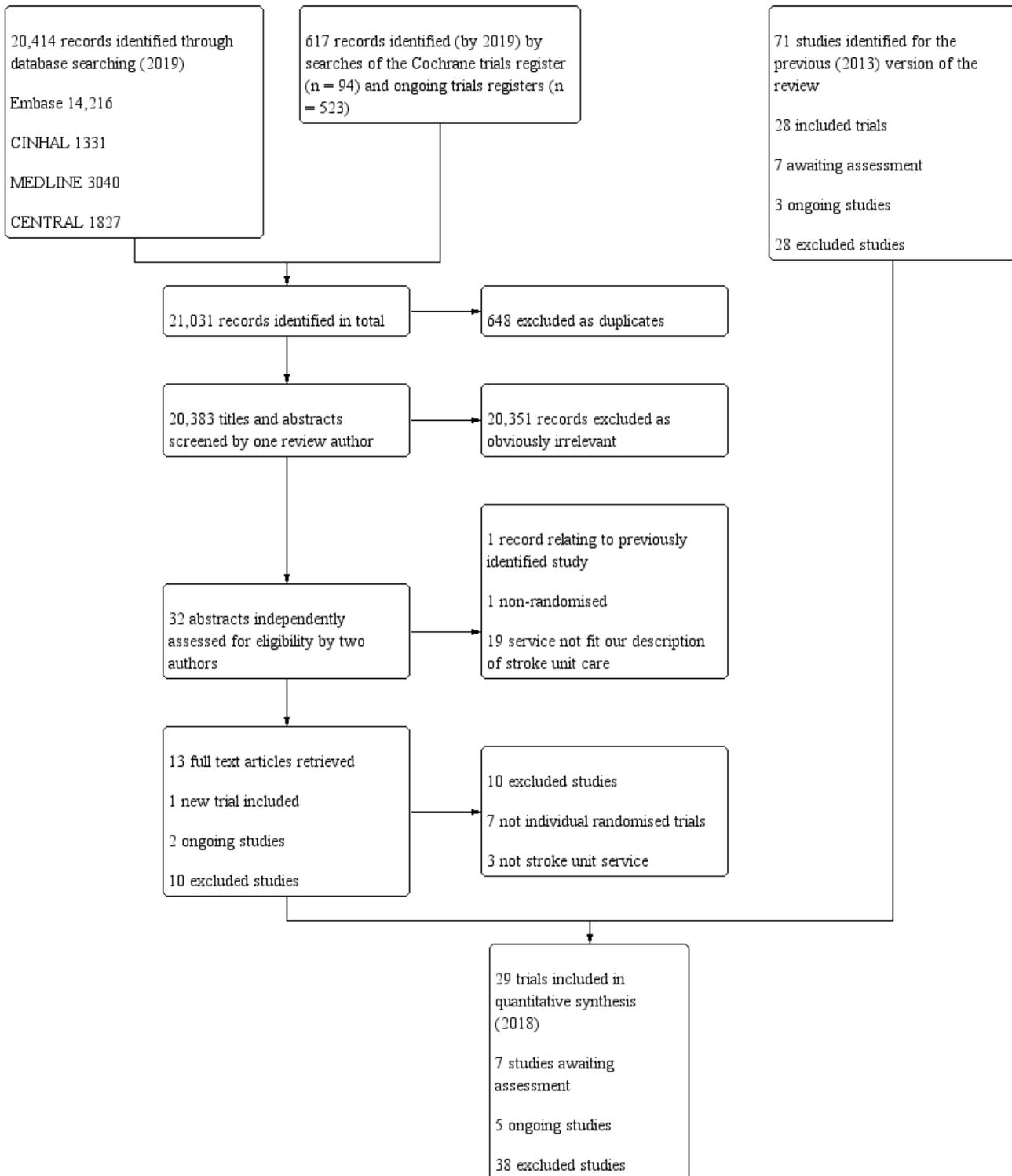
[Characteristics of studies awaiting classification](#); [Characteristics of ongoing studies](#).

Results of the search

The search strategy for previous versions of this review yielded 28 eligible trials ([Included studies](#)), seven awaiting classification ([Studies awaiting classification](#)), three ongoing studies ([Ongoing studies](#)), and 28 excluded studies ([Excluded studies](#)).

For this updated review, searches of Embase, CINHALL, MEDLINE, and CENTRAL revealed 16,562 records. Searches of Cochrane trials registers and other ongoing trials registers identified 432 new potentially eligible trials for consideration based on the four selection criteria ([Figure 1](#)). After exclusion of duplicate records and those that were obviously irrelevant, we were left with 32 abstracts for screening. Of these, one was a new record of a previously identified study ([Goteborg-Sahlgren 1994](#)), one was not randomised, and 19 described interventions that did not match organised inpatient (stroke unit) care.

Figure 1. Flow diagram illustrating the results of updated searches.



Of the remaining 13 articles retrieved, we excluded 10: seven were not randomised (Akhtar 2015; Al-Qahtany 2014; Fu 2006; He 2014; Inoue 2013; Rai 2016; Raiborirug 2017), and three did not meet the definition of stroke unit (Felix 2016; Janssen 2014; Middleton 2018). Two are ongoing studies (China (Wang) 2015; Russia 2017), and we included one new trial (New South Wales 2014).

Therefore, this updated review incorporates 29 randomised controlled trials with 5902 participants.

Included studies

Service characteristics within organised (stroke unit) care and conventional care settings

Descriptive information was available for all trials: for eight trials, we had access to published information ([Birmingham 1972](#); [Guangdong 2008](#); [Guangdong 2009](#); [Huaihua 2004](#); [Hunan 2007](#); [Illinois 1966](#); [New South Wales 2014](#); [New York 1962](#)); for two trials, we had detailed unpublished information ([Beijing 2004](#); [Joinville 2003](#)); and for the remaining 19 trials, we carried out a structured interview with the trial co-ordinator to determine the service characteristics.

Our original publication outlined the features of stroke unit trials ([SUTC 1997a](#)). In summary, organised inpatient (stroke unit) care was characterised by:

- co-ordinated multi-disciplinary rehabilitation;
- staff with a specialist interest in stroke or rehabilitation, or both;
- routine involvement of carers in the rehabilitation process; and
- regular programmes of education and training.

Several factors indicating more intensive or more comprehensive input of care were also associated with the stroke unit setting. Various service models of care exist ([Table 1](#)), but core characteristics that were invariably included in the stroke unit setting were (1) multi-disciplinary staffing, that is, medical, nursing, and therapy staff (usually including physiotherapy, occupational therapy, speech therapy, social work); and (2) co-ordinated multi-disciplinary team care incorporating meetings at least once per week ([SUTC 1997a](#)). When both of the compared services could satisfy the description of stroke unit care, the local conventional system of care was taken as the control service.

Service comparisons within the 29 trials with outcome data are detailed in [Table 2](#). The total number of comparisons is greater than the number of trials because in three trials, participants could be randomised to one of two alternatives to stroke unit care; two of these trials used a stratified randomisation procedure ([Nottingham 1996](#); [Orpington 1993](#)), and one did not ([Dover 1984](#)). In two small trials, the conventional care (general medical) group also received input from a specialist nurse ([Illinois 1966](#); [New York 1962](#)). Although this was not strictly general medical ward care, we have included this information because relatively little novel nursing

input appears to be available. Exclusion of these trials would not substantially alter the conclusions of the systematic review. For one trial, some participants appear to have been treated outside the rehabilitation wards (i.e. by peripatetic team care), but the number is unclear ([New York 1962](#)). This trial is currently classified as a mixed rehabilitation ward.

Four trials compared a model of stroke unit care using integrated traditional Chinese medicine (TCM) (e.g. acupuncture, herbal remedies) versus standard 'Western medicine' stroke unit care ([Guangdong 2008](#); [Hunan 2007](#)), or a general medical ward ([Guangdong 2009](#)). One trial compared a comprehensive stroke ward within a neurology unit with a general medical ward ([Huaihua 2004](#)). The duration of rehabilitation provided in all four trials was unclear, and only two trials reported the timing of randomisation ([Guangdong 2009](#); [Huaihua 2004](#)).

Of the 29 included trials, 23 incorporated rehabilitation lasting several weeks if required: 17 of these units admitted participants acutely, and eight after a delay of one or two weeks. Two trials compared an acute stroke unit with early transfer to conventional rehabilitation if required ([Groningen 2003](#); [New South Wales 2014](#)). One trial proved difficult to categorise as it contained elements of an acute unit but offered some rehabilitation ([Athens 1995](#)). It is classified here as a comprehensive stroke unit trial. The duration of rehabilitation was unclear for two Chinese trials ([Guangdong 2008](#); [Hunan 2007](#)). No trials evaluated an 'intensive care' model of a stroke unit.

The classification of trials is outlined in [Table 1](#), and the numbers in each comparison are shown in [Table 2](#).

Excluded studies

See [Characteristics of excluded studies](#).

Of the 38 excluded studies, 21 were not strictly randomised, six were evaluations of care pathways, four had no available outcome data, four evaluated an intervention that did not fit our description of organised inpatient (stroke unit) care, two treated intervention and control participants within the same unit, and one reported retrospective data from a previous study.

Risk of bias in included studies

See the 'Risk of bias' graph ([Figure 2](#)), the 'Risk of bias' summary ([Figure 3](#)), and the [Characteristics of included studies](#) table.

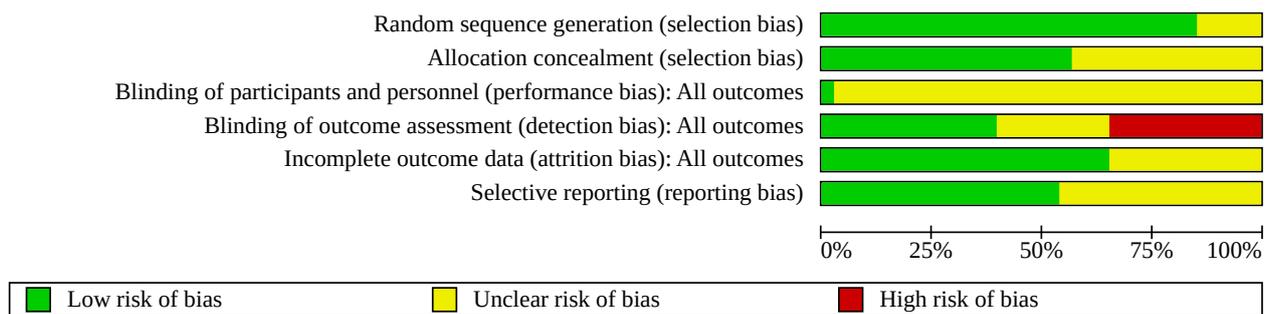
Figure 2. Risk of bias summary: review authors' judgements about each risk of bias item for each included study.

	Random sequence generation (selection bias)	Allocation concealment (selection bias)	Blinding of participants and personnel (performance bias): All outcomes	Blinding of outcome assessment (detection bias): All outcomes	Incomplete outcome data (attrition bias): All outcomes	Selective reporting (reporting bias)
Athens 1995	+	+	?	-	+	+
Beijing 2004	+	?	?	?	+	+
Birmingham 1972	?	?	?	-	?	+
Dover 1984	+	+	?	-	+	?
Dover 1984 (GMW)	+	+	?	-	+	?
Dover 1984 (MRW)	+	+	?	-	+	?
Edinburgh 1980	+	+	?	-	?	+
Goteborg-Ostra 1988	+	+	?	?	?	?
Goteborg-Sahlgren 1994	+	+	?	+	+	+
Groningen 2003	+	+	?	+	+	+
Guangdong 2008	+	?	?	?	?	?
Guangdong 2009	+	?	?	?	+	?
Helsinki 1995	+	+	?	+	+	?
Huaihua 2004	?	?	?	?	?	?
Hunan 2007	+	?	?	+	?	?
Illinois 1966	+	?	?	-	+	?
Joinville 2003	+	?	?	+	+	+
Kuopio 1985	+	+	?	+	+	?
Manchester 2003	+	+	?	+	+	+
Montreal 1985	+	+	?	+	+	?
Newcastle 1993	?	?	?	-	+	?
New South Wales 2014	+	+	+	+	+	?
New York 1962	+	?	?	?	?	?

Figure 2. (Continued)

New South Wales 2014	+	+	+	+	+	?
New York 1962	+	?	?	?	?	?
Nottingham 1996	+	?	?	+	?	+
Nottingham 1996 (GMW)	+	?	?	+	+	+
Nottingham 1996 (MRW)	+	?	?	+	+	+
Orpington 1993	+	+	?	-	?	+
Orpington 1993 (GMW)	+	+	?	-	?	+
Orpington 1993 (MRW)	+	+	?	-	?	+
Orpington 1995	?	?	?	?	+	+
Orpington 2000	+	+	?	+	?	+
Perth 1997	?	?	?	+	+	+
Svendborg 1995	+	+	?	?	+	?
Tampere 1993	+	+	?	-	+	+
Trondheim 1991	+	+	?	?	+	+

Figure 3. Risk of bias graph: review authors' judgements about each risk of bias item presented as percentages across all included studies.



Allocation

Sixteen trials used a secure and clearly concealed randomisation procedure (both random sequence generation and allocation concealment), and we judged these to be at low risk of bias (Athens 1995; Dover 1984; Edinburgh 1980; Goteborg-Ostra 1988; Goteborg-Sahlgren 1994; Groningen 2003; Helsinki 1995; Kuopio 1985; Manchester 2003; Montreal 1985; New South Wales 2014; Orpington 1993; Orpington 2000; Svendborg 1995; Tampere 1993; Trondheim 1991). The remaining trials were at unclear risk of bias.

Blinding

It is very challenging to blind participants or treating staff to treatment allocation, and only one trial reported any attempts to do so (New South Wales 2014). The remaining trials had unclear risk of bias. However, it is worth noting that most studies completed long-term follow-up at a time when participants and families often could not recall any details of their acute treatment.

Twelve trials used an unequivocally blinded final assessment for all participants (Goteborg-Sahlgren 1994; Groningen 2003; Helsinki 1995; Hunan 2007; Joinville 2003; Kuopio 1985; Manchester 2003; Montreal 1985; New South Wales 2014; Nottingham 1996;

Orpington 2000; Perth 1997). Eight trials had unclear risk of bias (Beijing 2004; Goteborg-Ostra 1988; Guangdong 2008; Guangdong 2009; Huaihua 2004; Orpington 1995; Svendborg 1995; Trondheim 1991), and we judged eight trials to be at high risk of bias (Athens 1995; Birmingham 1972; Dover 1984; Edinburgh 1980; Illinois 1966; Newcastle 1993; Orpington 1993; Tampere 1993).

Incomplete outcome data

We judged 19 trials to be at low risk of bias (Athens 1995; Beijing 2004; Dover 1984; Goteborg-Sahlgren 1994; Groningen 2003; Guangdong 2009; Helsinki 1995; Illinois 1966; Joinville 2003; Kuopio 1985; Manchester 2003; Montreal 1985; New South Wales 2014; Newcastle 1993; Orpington 1995; Perth 1997; Svendborg 1995; Tampere 1993; Trondheim 1991). The remaining nine trials were at unclear risk (Birmingham 1972; Edinburgh 1980; Goteborg-Ostra 1988; Guangdong 2008; Huaihua 2004; Hunan 2007; Nottingham 1996; Orpington 1993; Orpington 2000).

Ten trials had minor omissions of death and place of residence data (26 stroke unit participants and 35 controls in total) (Birmingham 1972; Dover 1984; Edinburgh 1980; Manchester 2003; Montreal 1985; New South Wales 2014; Nottingham 1996; Orpington 1993; Orpington 2000; Tampere 1993). For the purpose of our analysis, we

assumed these participants were alive and living at home, which may have introduced a minor bias in favour of the control group.

Selective reporting

We judged selective reporting bias to be low risk in 15 trials, largely because we obtained unpublished data from the trialists (Athens 1995; Beijing 2004; Birmingham 1972; Edinburgh 1980; Goteborg-Sahlgren 1994; Groningen 2003; Joinville 2003; Manchester 2003; Nottingham 1996; Orpington 1993; Orpington 1995; Orpington 2000; Perth 1997; Tampere 1993; Trondheim 1991). We classified the remaining 13 trials as having unclear risk of reporting bias (Dover 1984; Goteborg-Ostra 1988; Guangdong 2008; Guangdong 2009; Helsinki 1995; Huaihua 2004; Hunan 2007; Illinois 1966; Kuopio 1985; Montreal 1985; New South Wales 2014; Newcastle 1993; Svendborg 1995).

Other potential sources of bias

Most of the Stroke Unit Trialists Collaboration members carried out trials that are included in the review. However, trialists were not involved in selection or assessment of their own trials.

Effects of interventions

See: [Summary of findings 1 Organised inpatient \(stroke unit\) care versus alternative service](#); [Summary of findings 2 Stroke ward versus general medical ward](#); [Summary of findings 3 Mobile stroke team versus general medical ward](#); [Summary of findings 4 Mixed rehabilitation ward versus general medical ward](#)

Results of the systematic review are presented in five sections as pairwise comparisons followed by NMA.

Pairwise comparisons

These comparisons are listed in five sections as follows.

- Section 1. Organised inpatient (stroke unit) care versus alternative care (Comparison 1).
 - * First, we have outlined the main outcomes for the comparison of organised inpatient (stroke unit) care with an alternative service. Therefore, this section examines the impact of all types of organised inpatient (stroke unit) care on patient outcomes. For trials where both services compared could satisfy the definition of stroke unit care (Table 1), we have presented the system of care that we considered to be conventional care in the trial as the control service.
 - * This section includes analyses of different subgroups of participants and sensitivity analyses by trial quality.
 - * We have then described the results for the most common comparisons of different forms of organised stroke unit care versus a general medical ward: stroke ward, mobile stroke team, and mixed rehabilitation ward.
- Section 2. Stroke ward versus general medical ward (Comparison 2).
- Section 3. Mobile stroke team versus general medical ward (Comparison 3).
- Section 4. Mixed rehabilitation ward versus general medical ward (Comparison 4).

Finally, we have presented the results for any direct comparisons of organised care in a stroke ward versus a different form of organised stroke unit care.

- Section 5. Different systems of organised care: stroke ward versus alternative organised care (Comparison 5).

Section 1. Organised inpatient (stroke unit) care versus alternative service (Comparison 1)

Outcome 1.1. Poor outcome by the end of scheduled follow-up

Outcome data were available for 26 trials (5336 participants) (Analysis 1.1). The summary result indicated a significant reduction in the odds of a poor outcome (odds ratio (OR) 0.77, 95% confidence interval (CI) 0.69 to 0.87; moderate-quality evidence) recorded at the end of scheduled follow-up (median follow-up 12 months; range 6 weeks to 12 months) with no significant heterogeneity. The main methodological difficulties when dependency was used as an outcome were the degree of blinding at final assessment and the potential for bias if the assessor was aware of the treatment allocation. The results were unchanged when restricted to those trials in which an unequivocally blinded final assessment for all participants was undertaken (OR 0.75, 95% CI 0.62 to 0.91) (Goteborg-Sahlgren 1994; Groningen 2003; Helsinki 1995; Joinville 2003; Kuopio 1985; Manchester 2003; Montreal 1985; New South Wales 2014; Nottingham 1996; Orpington 2000).

Outcome 1.2. Death by the end of scheduled follow-up

Outcome data were available for all 29 trials (5902 participants) in which an organised inpatient (stroke unit) intervention was compared with an alternative service (Analysis 1.2). Case fatality recorded at the end of scheduled follow-up (median follow-up 12 months; range 6 weeks to 12 months) was lower in the organised (stroke unit) care group in 22 of 29 trials. The overall summary estimate included an OR of 0.76 (95% CI 0.66 to 0.88; moderate-quality evidence). A borderline significant subgroup interaction was reported ($P = 0.04$), with more positive effects seen in subgroups based on trials of stroke wards.

Outcome 1.3. Death or institutional care by the end of scheduled follow-up

Outcome data were available for 24 trials (4887 participants) (Analysis 1.3). The median duration of follow-up was 12 months (range 6 weeks to 12 months). The summary result indicated a significant reduction in the odds of a patient dying or requiring long-term institutional care (OR 0.76, 95% CI 0.67 to 0.85; moderate-quality evidence). A subgroup interaction was noted ($P = 0.01$), with more positive effects usually seen in subgroups based on trials of stroke wards. When we excluded trials that had a very short or variable period of follow-up, we found that the overall estimate of apparent benefit was unaffected (OR 0.75, 95% CI 0.65 to 0.86) (Beijing 2004; Goteborg-Ostra 1988; Groningen 2003; Illinois 1966; Montreal 1985; New York 1962; Orpington 1993; Orpington 1995).

Outcome 1.4. Death or dependency by the end of scheduled follow-up

Outcome data were available for 24 trials (4854 participants) (Analysis 1.4). The summary result indicated a significant reduction in the odds of the combined adverse outcomes of death or dependency (OR 0.75, 95% CI 0.66 to 0.85; moderate-quality evidence) with no significant heterogeneity. The main methodological difficulties when dependency was used as an outcome were the degree of blinding at final assessment and the potential for bias if the assessor was aware of the treatment allocation. The results were unchanged (OR 0.75, 95% CI 0.62 to 0.91) when restricted to those trials in which an unequivocally

blinded final assessment for all participants was undertaken (Goteborg-Sahlgren 1994; Groningen 2003; Helsinki 1995; Joinville 2003; Kuopio 1985; Manchester 2003; Montreal 1985; New South Wales 2014; Nottingham 1996; Orpington 2000).

Outcomes 1.5 and 1.6. Length of stay (days) in a hospital or institution or both

Length of stay data were available for 19 individual trials (4162 participants) (Analysis 1.5; Analysis 1.6). Mean (or median) length of stay ranged from 11 to 162 days in stroke unit groups and from 12 to 129 days in control groups. Thirteen trials reported a shorter length of stay in the organised inpatient (stroke unit) group, and six a more prolonged stay. The calculation of a summary result for length of stay was subject to major methodological limitations: length of stay was calculated in different ways (e.g. acute hospital stay, total stay in hospital or institution), two trials recorded median rather than mean length of stay, and in two trials the SD had to be inferred from the P value or from the results of similar trials. Overall, use of a random-effects model revealed no significant reduction in length of stay in the stroke unit group. The summary estimate was complicated by considerable heterogeneity that limits the extent to which more general conclusions can be inferred.

We re-analysed results according to whether length of stay was defined as stay in acute hospital only or total length of stay in a hospital or institution in the first year after stroke (Analysis 1.6). We found no significant difference between the two groups and no reduction in heterogeneity.

Participant satisfaction and subjective health status

Only three trials recorded outcome measures related to participant subjective health status (Nottingham Health Profile; EuroQol Quality of Life Scale) (Manchester 2003; Nottingham 1996; Trondheim 1991). In Nottingham 1996 and Trondheim 1991, there was a pattern of improved results among stroke unit survivors with results attaining statistical significance in the two trials. However, for Manchester 2003, there was no statistically significant difference between study groups. We could find no systematically gathered information on participant preferences.

Outcomes 1.7 to 1.10. Poor outcome, death, death or institutional care, and death or dependency at five-year follow-up

Three trials (1139 participants) carried out supplementary studies extending participant follow-up to five years post stroke for the outcome of death (Athens 1995; Nottingham 1996; Trondheim 1991), and two trials (535 participants) carried out supplementary studies extending participant follow-up to five years post stroke for the outcomes of death or institutionalisation and death or dependency (Nottingham 1996; Trondheim 1991). The OR for adverse outcomes continued to favour stroke unit care but with some heterogeneity: poor outcome 0.54 (95% CI 0.22 to 1.34), death 0.74 (95% CI 0.59 to 0.94), death or institutional care 0.59 (95% CI 0.33 to 1.05), and death or dependency 0.54 (95% CI 0.22 to 1.34).

Outcomes 1.11 to 1.14. Poor outcome, death, death or institutional care, and death or dependency at 10-year follow-up

Three trials (1139 participants) extended follow-up to 10 years post stroke for the outcome of death (Athens 1995; Nottingham 1996; Trondheim 1991), and two trials (535 participants) extended follow-up to 10 years post stroke for the outcomes of death or institutionalisation and death or dependency (Nottingham 1996; Trondheim 1991). Again, the summary results continued to favour stroke unit care but with increased heterogeneity and loss of statistical significance: poor outcome OR 0.70 (95% CI 0.27 to 1.80), death OR 0.66 (95% CI 0.43 to 1.03), death or institutional care OR 0.57 (95% CI 0.37 to 0.88), and death or dependency OR 0.70 (95% CI 0.27 to 1.80).

Sensitivity analyses by trial characteristics

Sensitivity analyses were applied only to the main (first) comparison to test confidence in the main hypothesis. In view of the variety of trial methods described, we carried out a sensitivity analysis based only on those trials with low risk of bias based on (1) secure concealment of allocation procedures, (2) unequivocally blinded outcome assessment, and (3) a fixed period of near complete follow-up.

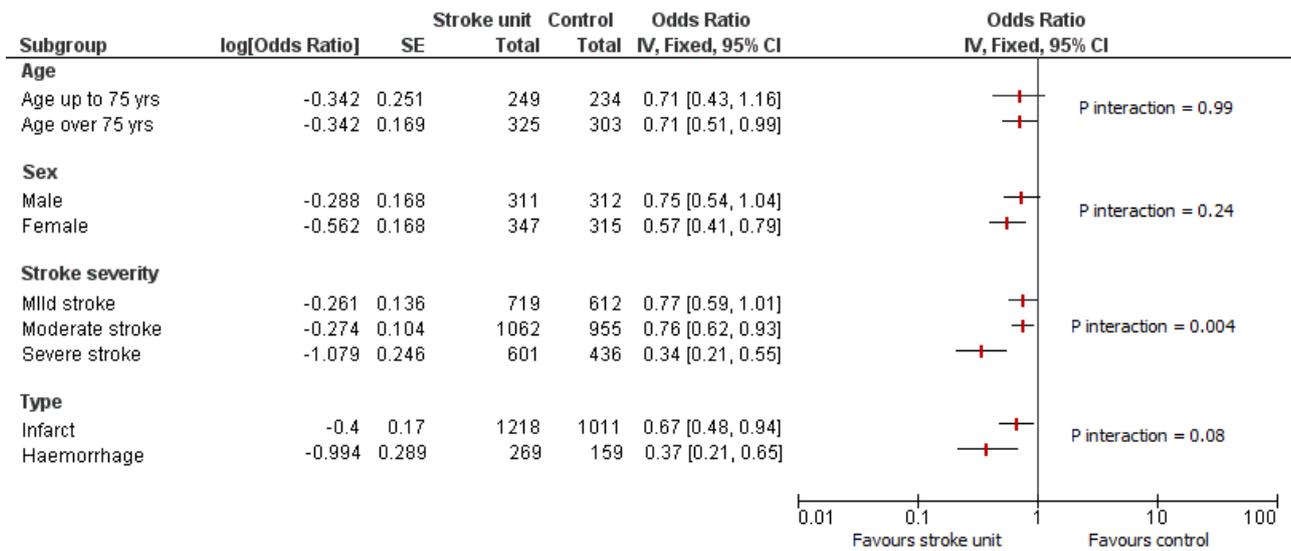
- Secure concealment of allocation procedures: restricting analyses to the 16 trials with clearly reported random sequence generation and concealment of allocation did not substantially alter the odds of a poor outcome (0.79, 95% CI 0.69 to 0.91) (Figure 2).
- Unequivocally blinded outcome assessment: restricting analyses to the 12 trials with clearly blinded outcome assessment did not substantially alter the odds of a poor outcome (OR 0.73, 95% CI 0.62 to 0.89) (Figure 2).
- A fixed period of near complete follow-up: restricting analyses to the 15 trials that clearly reported a fixed period of follow-up (with > 90% completeness of follow-up) did not substantially alter the odds of a poor outcome (OR 0.79, 95% CI 0.64 to 0.91) (Included studies).

Eight trials met all of these quality criteria (Goteborg-Sahlgren 1994; Groningen 2003; Helsinki 1995; Kuopio 1985; Manchester 2003; New South Wales 2014; Nottingham 1996; Orpington 2000). Within this group of trials, stroke unit care was associated with similar reductions in the odds of a poor outcome (OR 0.75, 95% CI 0.62 to 0.91).

Subgroup analyses by patient characteristics

Pre-defined subgroup analyses were based on previous versions of this review, and each subgroup analysis included data from at least nine trials (at least 1111 participants) (SUTC 1997a). These were based on participants' age, sex, and initial stroke severity. For this updated version of the review, we have incorporated additional data based on pathological stroke type (ischaemic or haemorrhagic stroke). See Figure 4.

Figure 4. Subgroup analysis by patient characteristics: poor outcome at the end of scheduled follow-up. Analyses used the generic inverse variance approach. P values relate to the subgroup interaction.



Caution is needed when interpreting these subgroup analyses, particularly as a relatively small number of outcome events were observed, which limits statistical power. Therefore, subgroup analyses were applied only to the main (first) comparison. Furthermore, the results may change depending on the outcome chosen. These results indicate that in general, the magnitude of benefit seemed greater for participants with more severe stroke (the only significant subgroup interactions were for stroke severity: $P = 0.004$). However, stroke unit benefits are apparent across a range of participant subgroups (i.e. age, sex, initial stroke severity, and stroke type).

Section 2. Stroke ward versus general medical ward (Comparison 2)

Analyses comparing a stroke ward with a general medical ward comprised two subgroups of stroke ward: comprehensive and rehabilitation (see Table 1).

Outcome 2.1. Poor outcome by the end of scheduled follow-up

Fourteen trials (3321 participants) compared care in a stroke ward with care in a general ward (Analysis 2.1). Stroke ward care showed a reduction in the odds of a poor outcome by the end of scheduled follow-up (OR 0.78, 95% CI 0.68 to 0.91; moderate-quality evidence) with no subgroup interaction between the different types of stroke ward. Some minor heterogeneity (59%) was noted. Re-analysis with a random-effects model did not alter the conclusions.

Outcome 2.2. Death by the end of scheduled follow-up

Fifteen trials (3523 participants) compared care in a stroke ward with care in a general ward (Analysis 2.2). Stroke ward care showed a reduction in the odds of death by the end of scheduled follow-up (OR 0.75, 95% CI 0.63 to 0.90; moderate-quality evidence) with no subgroup interaction between the different types of stroke ward and no significant heterogeneity.

Outcome 2.3. Death or institutional care by the end of scheduled follow-up

Thirteen trials (2924 participants) compared care in a stroke ward with care in a general ward (Analysis 2.3). Stroke ward care showed a reduction in the odds of death or institutional care by the end of scheduled follow-up (OR 0.74, 95% CI 0.63 to 0.87; moderate-quality evidence) with no subgroup interaction between the different types of stroke ward and no significant heterogeneity.

Outcome 2.4. Death or dependency by the end of scheduled follow-up

Twelve trials (2839 participants) compared care in a stroke ward with care in a general ward (Analysis 2.4). Stroke ward care showed a reduction in the odds of death or dependency by the end of scheduled follow-up (OR 0.75, 95% CI 0.64 to 0.88; moderate-quality evidence) with no subgroup interaction between the different types of stroke ward. Some heterogeneity (63%) was noted. Re-analysis with a random-effects model did not alter the conclusions.

Outcome 2.5. Length of stay (days) in a hospital or institution

Ten trials (2547 participants) compared care in a stroke ward with care in a general ward (Analysis 2.4). Overall, stroke ward care showed no reduction in length of stay (weighted mean difference (WMD) -2.19, 95% CI -5.19 to 0.82; low-quality evidence). We found substantial heterogeneity (78%) and a significant subgroup interaction between the different types of stroke ward, which limits confidence in the results.

Participant satisfaction and subjective health status

Only two trials recorded outcome measures related to participant subjective health status (Nottingham Health Profile; EuroQol Quality of Life Scale) (Nottingham 1996; Trondheim 1991). We found a pattern of improved results among stroke unit survivors with results attaining statistical significance in the two trials. We could find no systematically gathered information on participant preferences.

Section 3. Mobile stroke team versus general medical ward (Comparison 3)

Outcome 3.1. Poor outcome by the end of scheduled follow-up

Two trials (438 participants) compared care from a mobile stroke team with care in a general ward ([Analysis 3.1](#)). Stroke team care did not show a reduction in the odds of a poor outcome at the end of scheduled follow-up (OR 0.80, 95% CI 0.52 to 1.22; low-quality evidence) with no significant heterogeneity. Please note that as data were complete for dependency, this result is the same (as shown in [Analysis 3.4](#)).

Outcome 3.2. Death by the end of scheduled follow-up

Two trials (438 participants) compared care from a mobile stroke team with care in a general ward ([Analysis 3.2](#)). Stroke team care did not show a reduction in the odds of death by the end of scheduled follow-up (OR 1.08, 95% CI 0.71 to 1.63; low-quality evidence) with no significant heterogeneity.

Outcome 3.3. Death or institutional care by the end of scheduled follow-up

Two trials (438 participants) compared care from a mobile stroke team with care in a general ward ([Analysis 3.3](#)). Stroke team care did not show a reduction in the odds of death or institutional care by the end of scheduled follow-up (OR 1.27, 95% CI 0.84 to 1.93; low-quality evidence) with no significant heterogeneity.

Outcome 3.4. Death or dependency by the end of scheduled follow-up

Two trials (438 participants) compared care from a mobile stroke team with care in a general ward ([Analysis 3.4](#)). Stroke team care did not show a reduction in the odds of a poor outcome at the end of scheduled follow-up (OR 0.80, 95% CI 0.52 to 1.22; low-quality evidence) with no significant heterogeneity.

Outcome 3.5. Length of stay (days) in a hospital or institution

No data were available for this outcome.

Participant satisfaction and subjective health status

Only [Manchester 2003](#) recorded outcome measures related to participant subjective health status (EuroQol Quality of Life Scale). We could find no systematically gathered information on participant preferences.

Section 4. Mixed rehabilitation ward versus general medical ward (Comparison 4)

Outcome 4.1. Poor outcome by the end of scheduled follow-up

Six trials (630 participants) compared care in a mixed rehabilitation ward with care in a general ward ([Analysis 4.1](#)). Mixed rehabilitation ward care showed a reduction in the odds of a poor outcome at the end of scheduled follow-up (OR 0.65, 95% CI 0.47 to 0.90; moderate-quality evidence) with no significant heterogeneity. Please note that as data were complete for dependency, this result is the same (as shown in [Analysis 4.4](#)).

Outcome 4.2. Death by the end of scheduled follow-up

Six trials (630 participants) compared care in a mixed rehabilitation ward with care in a general ward ([Analysis 4.2](#)). Mixed rehabilitation ward care showed a reduction in the odds of death by the end of scheduled follow-up (OR 0.91, 95% CI 0.58 to 1.42; low-quality evidence) with no significant heterogeneity.

Outcome 4.3. Death or institutional care by the end of scheduled follow-up

Six trials (630 participants) compared care in a mixed rehabilitation ward with care in a general ward ([Analysis 4.3](#)). Mixed rehabilitation ward care showed a reduction in the odds of death or institutional care by the end of scheduled follow-up (OR 0.71, 95% CI 0.51 to 0.99; low-quality evidence) with no significant heterogeneity.

Outcome 4.4. Death or dependency by the end of scheduled follow-up

Six trials (630 participants) compared care in a mixed rehabilitation ward with care in a general ward ([Analysis 4.4](#)). Mixed rehabilitation ward care showed a reduction in the odds of death or dependency by the end of scheduled follow-up (OR 0.65, 95% CI 0.47 to 0.90; moderate-quality evidence) with no significant heterogeneity.

Outcome 4.5. Length of stay (days) in a hospital or institution

Three trials (387 participants) compared care in a stroke ward with care in a general ward ([Analysis 4.5](#)). Overall, stroke ward care showed no reduction in length of stay (WMD 3.85, 95% CI -13.49 to 21.18; very low-quality evidence) with no significant heterogeneity.

Participant satisfaction and subjective health status

No data were available for these outcomes.

Section 5. Different systems of organised care: stroke ward versus alternative organised care (Comparison 5)

The analyses in Section 2 above indicate that organised inpatient (stroke unit) care in a stroke ward is an effective model of care. Several recent trials have compared different ways of providing care in a stroke ward. We therefore analysed those trials that directly compared a stroke ward with another form of organised inpatient (stroke unit) care that met the basic inclusion criteria ([Table 1](#)).

Of the nine trials identified for which outcome data were available, one compared an acute stroke ward with a mixed rehabilitation ward ([Tampere 1993](#)). Two small trials compared care in an acute stroke ward with care in a comprehensive stroke ward ([Groningen 2003](#); [New South Wales 2014](#)). The Dutch trial compared care in an acute unit where there was an emphasis on close management of physiological variables ([Groningen 2003](#)). The Australian trial compared two systems of care: acute stroke ward (with transfer to a rehabilitation ward if required) with a stroke ward that combined acute care and rehabilitation (comprehensive stroke ward) ([New South Wales 2014](#)). One trial compared a stroke ward that combined acute care and rehabilitation (comprehensive stroke ward) with a general medical ward where care was co-ordinated by a multi-disciplinary team (mobile team care) ([Orpington 2000](#)). Two compared a stroke ward with integrated traditional Chinese medicine (TCM) with a 'Western medicine' stroke ward ([Guangdong 2008](#); [Hunan 2007](#)). Three trials incorporated designs in which participants could be randomised to a stroke rehabilitation ward or to conventional care in a general medical ward or a mixed rehabilitation ward within a Department of Geriatric Medicine ([Dover 1984](#); [Nottingham 1996](#); [Orpington 1993](#)). Data were available for both of these participant subgroups.

Acute stroke ward versus mixed rehabilitation ward

Outcomes 5.1, 5.2, 5.3, 5.4, and 5.5. Poor outcome, death, death or institutional care, death or dependency by the end of scheduled follow-up, and length of stay in hospital or institution

The trial comparing an acute unit with a mixed rehabilitation unit did not show any statistically significant difference in the odds of a poor outcome nor in death, death or requiring institutional care, death or dependency, or length of stay data ([Tampere 1993](#)).

Acute stroke ward versus comprehensive stroke ward

Outcomes 5.1, 5.2, 5.3, 5.4 and 5.5. Poor outcome, death, death or institutional care, death or dependency by the end of scheduled follow-up, and length of stay in hospital or institution

We found no consistent evidence that an acute stroke ward was superior to a comprehensive stroke ward. The Dutch trial suggested improved outcomes within the acute ward where there was an emphasis on close management of physiological variables ([Groningen 2003](#)). However, this pilot study was underpowered to demonstrate major differences. The recent Australian trial compared a conventional model of care (acute stroke ward with transfer to rehabilitation if required) with all care provided in one (comprehensive) ward ([New South Wales 2014](#)). Study authors suggested that care in the comprehensive ward may be more efficient (with more rapid recovery), but the key outcomes reported here were not statistically different.

Comprehensive stroke ward versus mobile stroke team

Outcomes 5.1, 5.2, 5.3, 5.4, and 5.5. Poor outcome, death, death or institutional care, death or dependency by the end of scheduled follow-up, and length of stay in hospital or institution

One trial compared a comprehensive stroke ward (providing acute care and rehabilitation) with admission to a general ward where care was provided by a mobile stroke team ([Orpington 2000](#)). Study authors found statistically significant ($P < 0.001$) reductions in death and the combined outcome of death or institutional care among the comprehensive stroke ward group. Fewer comprehensive stroke ward participants had a poor outcome (were dead or dependent) at the end of follow-up, but this result did not achieve statistical significance. However, [Orpington 2000](#) is the only trial in this analysis comparing comprehensive stroke wards with an alternative service, so these results require confirmation. Results show no significant difference in length of stay.

Rehabilitation stroke ward versus alternative service

Outcomes 5.1, 5.2, 5.3, 5.4 and 5.5. Poor outcome, death, death or institutional care, death or dependency by the end of scheduled follow-up, and length of stay in hospital or institution

We noted a pattern of improved outcomes in the stroke rehabilitation ward with fewer deaths (OR 0.50, 95% CI 0.28 to

0.90) but no statistically significant reduction in the composite end points of poor outcome, death or requiring institutional care, and death or dependency. However, the numbers were small and no definitive conclusions could be drawn. Interpretation of length of stay data was complicated by substantial heterogeneity. There was no evidence of a systematic increase in length of stay.

Stroke ward plus TCM versus alternative service

Outcome 5.2. Death at the end of scheduled follow-up

There was no significant difference in the odds of death in a stroke ward with integrated TCM when compared with a standard 'Western medicine' stroke ward ([Guangdong 2008](#); [Hunan 2007](#)). The type of care provided in a stroke unit with integrated TCM has not been well described. The overall estimate is based on the results of a single trial, and no definitive conclusions can be drawn.

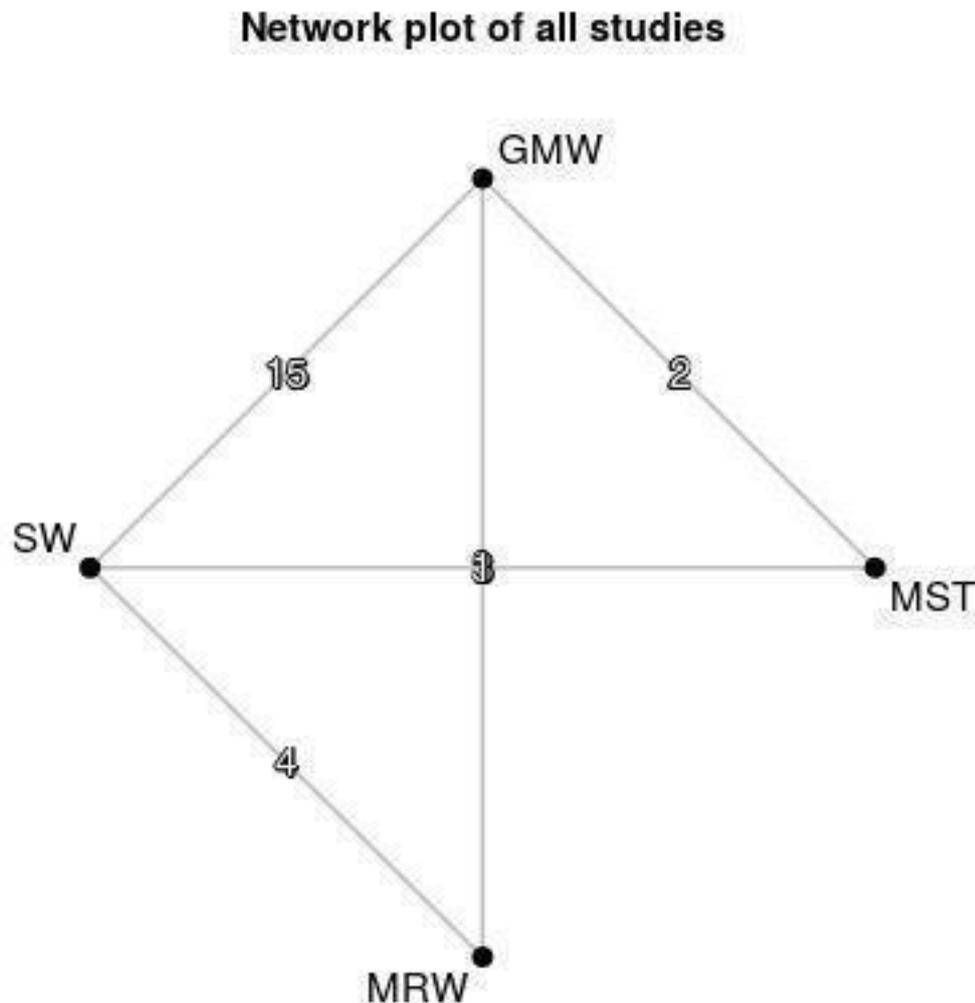
Analyses by service characteristics (including network meta-analysis (NMA))

In planning our analyses, we specified in advance that an important question for service planning would be whether the benefits of organised (stroke unit) care depended upon the establishment of a ward dedicated only to stroke care (stroke ward) or could be achieved through a mobile stroke team or a generic disability service (mixed rehabilitation unit) that specialises in the management of disabling illness including stroke. We explored these questions in Sections 2, 3, and 4 above, but we also performed an NMA to explore the impact of different systems of stroke care. We used *MetainSight* software, designed specifically for this role, to conduct our NMA (<https://crsu.shinyapps.io/metainsightb/>).

[Table 1](#) shows the categories of organised inpatient (stroke unit) care grouped according to our service classification. In view of the difficulty of defining precisely different forms of stroke wards and the small number of comparisons of different forms of stroke wards ([Table 2](#)), we have analysed the 'stroke ward' group as a single entity. Further comparisons within this group are shown in [Analysis 5.1](#), [Analysis 5.2](#), [Analysis 5.3](#), [Analysis 5.4](#), and [Analysis 5.5](#).

[Table 3](#) and [Figure 5](#) show trial comparisons within the NMA. We believe that the transitivity (or similarity) assumption was met, as all included trials recruited people with acute or subacute stroke for treatment within a hospital system of care ([Characteristics of included studies](#)). Subgroup analyses by patient characteristics indicate that there is no substantial treatment modification by patient age, sex, or type, but possibly by stroke severity ([Figure 4](#)). Although a range of stroke severities was evident in the recruited participants, they were usually well distributed within trials.

Figure 5. Network meta-analysis plot for different types of organised care. The nodes show the service groups (GMW: general medical ward; MRW: mixed rehabilitation ward; MST: mobile stroke team; SW: stroke ward), with care in a GMW as the reference.



We evaluated the consistency assumption statistically by comparing the difference between direct and indirect estimates for each loop of evidence. We examined for any inconsistency (i.e. important differences in numerical results between direct, indirect, and network results), and we presented OR estimates for each of the three comparisons. We showed the inconsistency tables from the NMA for analyses of poor outcome (Table 4), death (Table 5), death or institutional care (Table 6), and death or dependency (Table 7). These tables show the results of direct and indirect comparisons plus the NMA results. We found no statistically significant differences ($P > 0.05$) between any of the direct and indirect comparisons, but confidence intervals were wide.

The NMA used the general medical ward group as the comparator, as this was clinically relevant and was the most common comparator reported by trialists (Table 2). Figure 6 shows the NMA result for a poor outcome at the end of scheduled follow-up. The lowest odds of a poor outcome was seen with care in a stroke ward. A rank analysis, which orders treatments according to their relative effectiveness (the first ranked treatment is most likely to be the most effective treatment compared with the other treatments in the network), revealed that care in a stroke ward was the optimal option, although our confidence was limited by substantial imprecision. Finally, including a sensitivity analysis featuring only the six eligible trials that met all of the quality criteria listed in the

sensitivity analysis above did not alter the ranking but resulted in wider confidence intervals (Goteborg-Sahlgren 1994; Helsinki 1995; Kuopio 1985; Manchester 2003; Nottingham 1996; Orpington 2000).

Figure 6. Network meta-analysis plot for different types of organised care. The outcome is poor outcome at the end of scheduled follow-up. The treatment column shows the service groups (GMW: general medical ward; MRW: mixed rehabilitation ward; MST: mobile stroke team; SW: stroke ward). The results are the odds ratio (95% confidence interval) for the odds of a poor outcome, with care in a GMW as the reference (OR = 1.0).

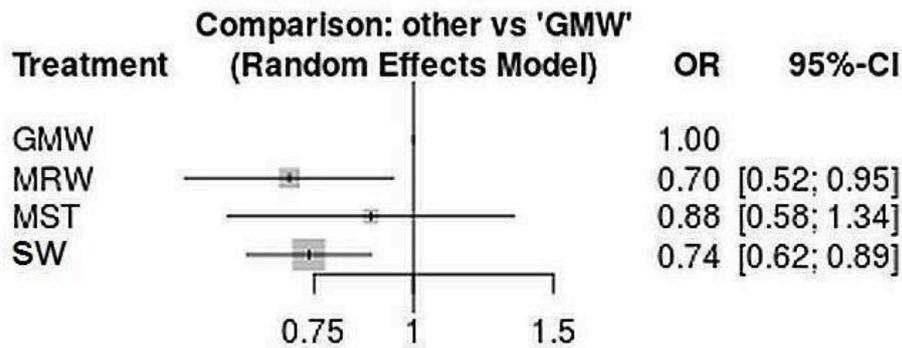


Figure 7 shows the NMA result for the outcome of death at the end of scheduled follow-up. The lowest odds of death was seen with care in a stroke ward. No statistically significant reductions were seen

for mobile stroke team or mixed rehabilitation ward care. A rank analysis revealed that care in a stroke ward was the optimal option. The sensitivity analysis did not alter the ranking.

Figure 7. Network meta-analysis plot for different types of organised care. The outcome is death at the end of scheduled follow-up. The treatment column shows the service groups (GMW: general medical ward; MRW: mixed rehabilitation ward; MST: mobile stroke team; SW: stroke ward). The results are the odds ratio (95% confidence interval) for the odds of a poor outcome, with care in a GMW as the reference (OR = 1.0).

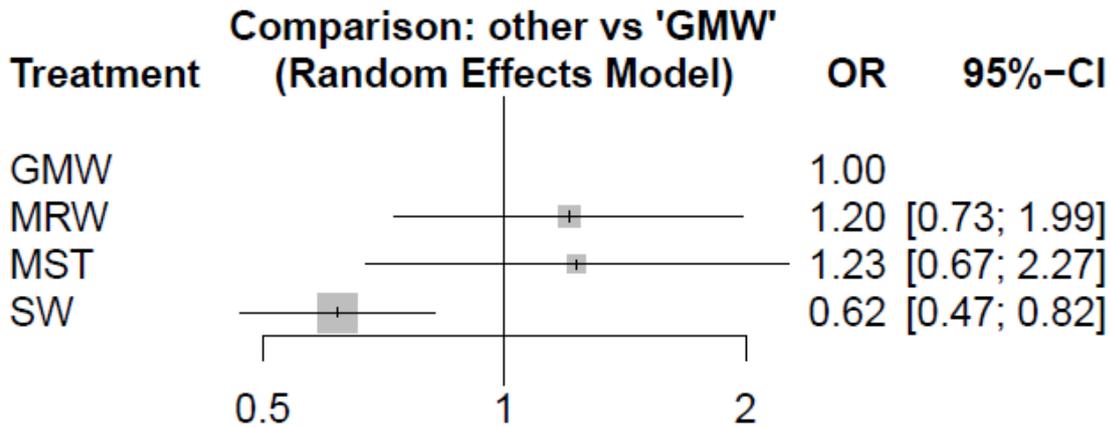


Figure 8 shows the NMA result for the outcome of death or institutional care at the end of scheduled follow-up. The lowest odds of an adverse outcome was seen with care in a stroke ward. Reductions in poor outcome were of borderline significance

for mixed rehabilitation ward, and no statistically significant reductions were seen for mobile stroke team. A rank analysis showed that care in a stroke ward was the optimal option. The sensitivity analysis did not alter the ranking.

Figure 8. Network meta-analysis plot for different types of organised care. The outcome is death or institutional care at the end of scheduled follow-up. The treatment column shows the service groups (GMW: general medical ward; MRW: mixed rehabilitation ward; MST: mobile stroke team; SW: stroke ward). The results are the odds ratio (95% confidence interval) for the odds of a poor outcome, with care in a GMW as the reference (OR = 1.0).

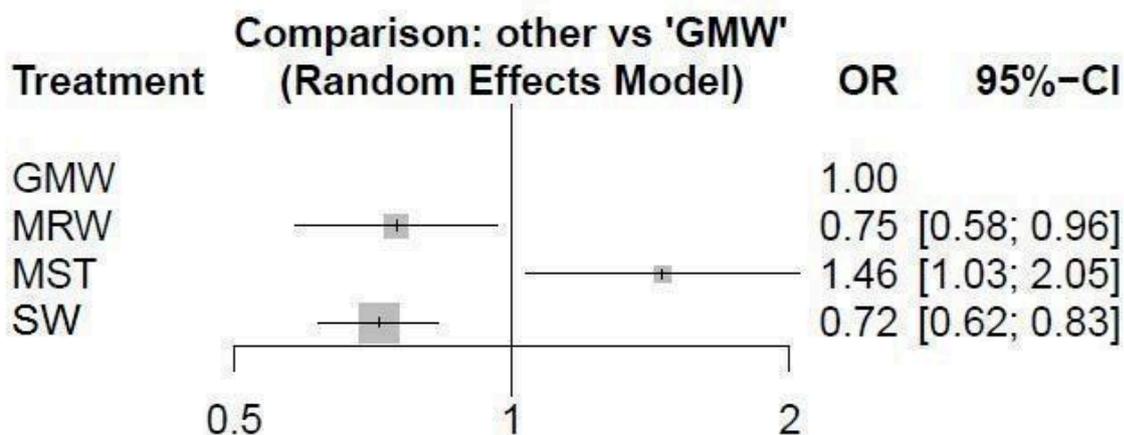
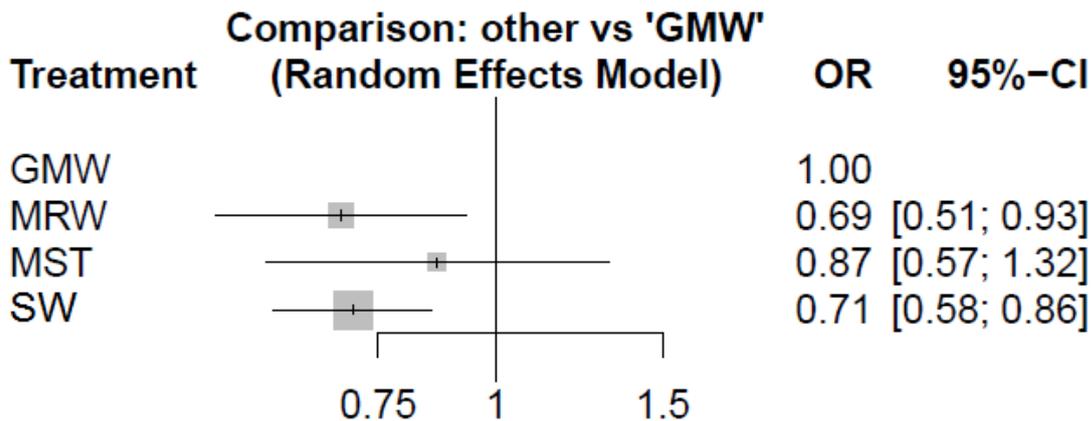


Figure 9 shows the NMA result for the poor outcome (death or dependency) at the end of scheduled follow-up. The lowest odds of poor outcome was seen with care in a stroke ward or in a mixed rehabilitation ward. No statistically significant reductions

were seen for mobile stroke team. A rank analysis found that care in a mixed rehabilitation ward was the optimal option. The sensitivity analysis did not alter the ranking.

Figure 9. Network meta-analysis plot for different types of organised care. The outcome is death or dependency at the end of scheduled follow-up. The treatment column shows the service groups (GMW: general medical ward; MRW: mixed rehabilitation ward; MST: mobile stroke team; SW: stroke ward). The results are the odds ratio (95% confidence interval) for the odds of a poor outcome, with care in a GMW as the reference (OR = 1.0).



Finally, we present a 'Summary of findings' table for the NMA (Table 8).

DISCUSSION

Summary of main results

Main analysis

The updated information in Section 1 confirms our previous observations that people receiving organised inpatient (stroke unit) care were more likely to survive, regain independence, and return home than those receiving an alternative service. These benefits were based on moderate-quality evidence. The observed reductions in combined adverse outcomes (poor outcome, death or institutionalisation, death or dependency) are relatively robust and would be negated only by several new neutral trials. The three trials that have extended follow-up for 5 or 10 years suggest sustained benefit among stroke unit patients.

The requirement for long-term care is a useful surrogate for disability and is likely to be less prone to observer bias than measurement of disability (Barer 1993). The absolute rates of institutionalisation, however, will be influenced by a variety of national and cultural factors. The combined adverse outcome of death or dependency is a more direct measure of patient outcome but is subject to potential observer bias where final assessments were not carried out in a blinded manner. The sensitivity analysis based on those trials that used an unequivocally blinded assessment suggested that such bias has not seriously influenced the results.

The analysis of length of stay is complicated by different methods of reporting results, widely varying control group lengths of stay, and statistically significant heterogeneity between trials. The most reasonable conclusion appears to be that no systematic increase in length of stay is associated with organised (stroke unit) care, and there may be a modest reduction.

Subgroup analyses by patient characteristics

In any discussion of the comparison of results from different subgroups, it is worth bearing in mind that the main issue is not whether a subgroup result is statistically different from the comparator result, but whether there is statistically significant heterogeneity between estimates of effect in each of the relevant subgroups. Our analyses are limited by relatively low statistical power and so must be interpreted with great caution. The subgroup analyses indicate that observed benefits of organised stroke unit care are not limited to any one subgroup of patients. Apparent benefits were seen in people of both sexes, younger and older than 75 years, with ischaemic or haemorrhagic stroke, and across a range of stroke severities. The apparent interaction between stroke severity and outcome must be interpreted with caution. People with more severe stroke symptoms are at greater risk of death or requiring institutional care and hence stand to gain more from treatment. People with a mild stroke appeared to benefit from stroke unit care when death or dependency was the chosen outcome (Figure 4), but this effect was less certain for the outcomes of death and death or institutional care (data not shown).

Analyses by different types of service

An original aim of this review was to compare effects of organised inpatient (stroke unit) care with effects of contemporary conventional care. We expected that within this broad definition, the included trials would comprise a range of treatment comparisons (which could include stroke wards, mobile stroke teams, and mixed rehabilitation wards) with conventional care in a general ward. However, later trials have addressed newer questions by comparing different forms of organised inpatient (stroke unit) care (e.g. stroke ward versus mobile stroke team). This means that although the analysis in Section 1 tells us about the likely impact of organising inpatient care, it is much more limited in providing advice about specific service models. We have therefore retained the previous analyses but now include a network meta-analysis (NMA) to explore, when possible, the impact of different models of organised inpatient (stroke unit) care.

Of the four approaches to organised inpatient (stroke unit) care tested in the NMA (stroke ward, mixed rehabilitation ward, mobile stroke team, general medical ward), care within a ward for stroke patients (stroke ward) was generally the most effective approach. Mixed rehabilitation units also showed some reduction in the composite outcomes (poor outcome, death or requiring institutional care, death or dependency).

In direct comparisons with care in a general medical ward, two different types of stroke wards (comprehensive stroke ward and mixed rehabilitation ward) tended to be more effective. There was a similar but less convincing pattern for rehabilitation stroke units. However, mobile stroke team care appeared to have a neutral effect. Apparent benefits were seen in units with acute admission policies, as well as in those with delayed admission policies and in units that could offer a period of rehabilitation lasting several weeks.

The final section (Section 5) focused on trials that directly compared two different forms of care, both of which met our basic definition of organised inpatient (stroke unit) care: multi-disciplinary team care co-ordinated through regular meetings. The results of this analysis indicate improved results from a comprehensive stroke ward over a mobile stroke team. Results also suggest better survival within the stroke rehabilitation ward setting as opposed to the mixed rehabilitation ward setting. Comparisons of a comprehensive ward with alternative pathways of care were limited by wide confidence intervals. No firm conclusions could be drawn for the comparisons of a stroke ward integrated with traditional Chinese medicine (TCM) versus a 'Western medicine' stroke ward.

In summary, subgroup analyses indicate that organised inpatient (stroke unit) care based in a dedicated stroke ward is likely to be most effective. Within the stroke ward approach, benefits were apparent for both comprehensive and rehabilitation stroke wards, indicating that organised stroke unit care is of benefit in both acute and rehabilitation phases of care. Mixed rehabilitation wards also showed some benefit in reducing dependency.

Overall completeness and applicability of evidence

This update includes one new trial (47 participants), and the overall conclusions remain unaltered from those provided in previous versions of the review. The review now summarises data from a total of 29 trials (5902 participants) from 12 countries

in Asia, Australia, Europe, North America, and South America. A majority of trials have been performed in high-income countries. Observational studies indicate that stroke units are likely to be effective in low- or middle-income countries (Langhorne 2018; Seenan 2007; Urimubenshi 2017), but there is a shortage of information from randomised clinical trials (Langhorne 2012).

As discussed above, our subgroup analyses suggest that the benefits of organised inpatient (stroke unit) care are seen across a wide range of stroke patients. The current analysis does not explain how stroke units may improve patient outcomes, but this could be the result of greater staff expertise, better diagnostic procedures, better nursing care, early mobilisation, prevention of complications, or more effective rehabilitation procedures (Langhorne 1998).

Since the original publication of this review, stroke services in many developed countries have undergone substantial reorganisation in line with national strategies and clinical practice guidelines to enable improvements in access to stroke unit care. More recently, stroke services in many countries have been further reorganised to reflect a two-tiered (or hub-and-spoke) model of care in which a central 'comprehensive stroke centre' (based around a 'hyper-acute stroke unit') is equipped with facilities for acute intravenous or intra-arterial treatments, intensive monitoring, advanced imaging, and neurosurgery. These then serve a number of 'primary stroke centres' or stroke units within a hospital network or geographical location. Although these developments appear almost intuitive to many stroke clinicians, they have never been formally tested in randomised controlled trials with important patient outcomes. Until such trials are available, stroke services should ensure that every stroke patient receives the core service characteristics identified in randomised trials of organised inpatient (stroke unit) care.

Costs and benefits

Stroke units appear to improve outcomes, but at what cost? In cost terms, length of stay is likely to dominate any individual component of acute patient care and rehabilitation. Longer-term costs are likely to be dominated by the need for nursing care. Studies from several developed countries have shown that fixed costs (particularly nursing staff salaries) account for over 90% of spending on people with acute stroke (Warlow 2008). Remedial therapy represents only a small proportion of the total cost of hospitalisation. In one analysis, stroke unit care was not clearly associated with an increase in total health and social care costs, but these conclusions were sensitive to some variation in cost estimates (Major 1998). More research is required to elucidate the cost implications of stroke units.

Quality of the evidence

The quality of evidence was made more uniform in the previous review update by the exclusion of several quasi-randomised controlled clinical trials that were originally included in the data synthesis (see [Description of studies](#)) (SUTC 2013). The main goal of this change was to simplify the inclusion criteria for this and future updates. However, it is worth noting that exclusion of these trials did not affect the overall estimate of treatment effect (SUTC 2013).

The most common potential source of bias is the difficulty of concealing treatment allocation from participants (patients) and treating staff (performance bias); this is very difficult to achieve

with this type of intervention. Although we have downgraded our recommendations for the risk of performance bias, it is worth noting that most studies completed long-term follow-up at a time when participants and families often could not recall any details of their treatment.

We judged some trials to be at high risk of bias due to poor allocation concealment and unblinded outcome assessment; in others, these important methodological aspects were not clearly reported, making a judgement of risk of bias difficult. We have not downgraded the GRADE recommendations because (1) the key outcomes (survival, return home) are unlikely to be sensitive to observer bias, and (2) sensitivity analyses restricted to the eight trials at low risk of bias did not alter effect sizes of the estimates. In particular, effect sizes for the composite adverse outcomes of death or institutionalisation and death or dependency remained largely unaltered.

We recognise that some of the included trials are relatively old, possibly applying entirely different standards of care from those used currently. Similarly, although a majority of included trials were conducted fairly recently, most would still have been undertaken in an era without routine access to re-perfusion therapies (intravenous thrombolysis or mechanical thrombectomy) for acute ischaemic stroke. Although essentially all stroke patients would be eligible for admission to a stroke unit, only a small proportion would be eligible for treatment with re-perfusion therapies even in the most established acute centres (Langhorne 2012). Moreover, all included trials were randomised; therefore any differences in the standard of care should not have had a confounding effect on the final conclusions.

Potential biases in the review process

Through a comprehensive search strategy and established connections with other researchers in the field, we have gone to considerable lengths to identify all potentially relevant studies. However, we did not search Chinese databases, and we were unable to classify or obtain useable outcome data for 7 of the 11 Chinese studies that we did identify (Anhui 2008; China (Hao) 2010; China (Pei) 2011; China (Wang) 2008; China (Wu) 2007; Haikou 2007; Shanghai 2006). We recognise that the absence of data from these studies in our meta-analysis could potentially introduce bias.

Methodological limitations may also have influenced the analysis of descriptive information about service organisation (SUTC 1997a). First, our system of classifying services is an attempt to bring some structure to a complex topic, and it will lack precision (Table 1). In addition, we collated service descriptions retrospectively through discussion with the trialists who ran the organised (stroke unit) care. Our findings may therefore be biased towards the expectations of trialists and by a tendency to discuss the results with trialists who ran the organised stroke unit care more than with those who ran the conventional care unit. At best, this represents a strictly factual account of service characteristics; at worst, it represents a consensus view of the trialists about which features of stroke unit care were effective.

Agreements and disagreements with other studies or reviews

This systematic review has been updated multiple times over the last 25 years with broadly similar conclusions (SUTC 1997a; SUTC

1997b; SUTC 2001; SUTC 2007; SUTC 2013). Other versions of the review have tended to focus on the method (Sun 2013), on a subgroup of trials (Chan 2013), or on studies of implementation in routine services (Seenan 2007; Urimubenshi 2017). Although the emphasis may have varied between reviews, the overall conclusion about the effectiveness of organised inpatient (stroke unit) care has not changed.

AUTHORS' CONCLUSIONS

Implications for practice

Moderate-quality evidence shows that people with acute stroke are more likely to survive, return home, and regain independence if they receive organised inpatient (stroke unit) care. This is typically provided by a co-ordinated multi-disciplinary team operating within a discrete stroke ward that can offer a substantial period of rehabilitation if required. There are no firm grounds for restricting access according to a person's age, sex, stroke severity, or pathological stroke type (i.e. ischaemic or haemorrhagic). Stroke unit care provided in a dedicated stroke ward seems to be most effective, with some evidence for effectiveness of a mixed rehabilitation ward model.

Since the original publication of this review, many stroke services have been re-organised to support newer treatments. Many have promoted a two-tiered model of care in which a central 'comprehensive stroke centre' is equipped with facilities ('hyperacute' stroke units) for re-perfusion therapies (intravenous thrombolysis and mechanical thrombectomy), advanced imaging, and neurosurgery. These in turn serve a number of 'primary stroke centres' within a hospital network or geographical location. These newer service models have not been formally tested in randomised controlled trials. Until such trials are available, clinicians and planners must ensure that every stroke patient receives the core service characteristics described in randomised trials of organised inpatient (stroke unit) care.

Implications for research

Future trials should focus on examining the potentially important components of stroke unit care and on performing direct comparisons of different models of organised stroke unit care, particularly with regard to the hyperacute stroke unit model. In low-income healthcare settings, appropriately powered clinical trials could help define how barriers to the establishment of stroke units could be overcome (Langhorne 2012). Outcome measures should not include only the outcomes of death, dependency, and institutionalisation; they should also include the domains of patient satisfaction, quality of life, and cost. Pre-planned collaboration between comparable trials could alleviate some of the problems of retrospective systematic reviews by, for example, ensuring that similar variables and outcomes are recorded in any new trial.

Anyone carrying out a relevant randomised trial of a stroke service component is invited to contact Peter Langhorne regarding a future collaborative review.

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* Indicates the major publication for the study

CHARACTERISTICS OF STUDIES

Characteristics of included studies [ordered by study ID]

Athens 1995

Study characteristics

Methods	RCT Sealed envelopes Unblinded follow-up
Participants	People with acute stroke admitted to emergency department within 24 hours of symptoms Excluded TIA or recurrent stroke
Interventions	Small (6-bed) ward within Internal Medicine department Used the American Heart Association protocol, management of physiological abnormalities, multi-disciplinary team approach Compared with conventional care in general medical ward
Outcomes	Death, cause of death, length of stay Recorded up to 6.5 years (12-month data used in primary analysis)
Notes	Unpublished at present

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomised ... using numbered opaque sealed envelopes"
Allocation concealment (selection bias)	Low risk	"Opaque sealed envelopes"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Beijing 2004

Study characteristics

Methods	RCT Divided randomly using SPSS software package
Participants	People with stroke admitted to hospital with first or recurrent stroke Subarachnoid haemorrhage and tumour were excluded
Interventions	New comprehensive stroke unit, early multi-disciplinary rehabilitation

Organised inpatient (stroke unit) care for stroke: network meta-analysis (Review)

Beijing 2004 (Continued)

Control participants were admitted to general medical or general neurology ward

Outcomes Death, NIHSS, Barthel Index, Oxford Handicap Scale, patient satisfaction at time of discharge

 Notes Some unpublished data included
 No institutional care available

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Divided randomly into two groups using SPSS software package"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment not described
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Unclear
Incomplete outcome data (attrition bias) All outcomes	Low risk	Similar numbers in treatment (n = 20) and control (n = 21) groups with missing data
Selective reporting (reporting bias)	Low risk	Data on all pre-specified outcomes reported

Birmingham 1972
Study characteristics

Methods	RCT
Participants	People with stroke within 2 weeks of stroke onset Able to tolerate active rehabilitation
Interventions	Intensive rehabilitation in rehabilitation centre (mixed rehabilitation unit) (n = 29) vs normal care in general medical ward (n = 23) Organised care provided for months if required
Outcomes	Death and functional status at end of follow-up (6 to 8 months)
Notes	Timing of outcomes not clearly stated Intervention not clearly defined 3 control participants lost to follow-up

Risk of bias

Bias	Authors' judgement	Support for judgement
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Birmingham 1972 (Continued)

Random sequence generation (selection bias)	Unclear risk	"Evenly divided on a random basis"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment not described
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	3 control participants (almost 10%) lost to follow-up
Selective reporting (reporting bias)	Low risk	Pre-specified outcomes reported

Dover 1984
Study characteristics

Methods	RCT
Participants	People with stroke up to 9 weeks after stroke onset (majority within 3 weeks) Fit for transfer to rehabilitation ward
Interventions	Stroke rehabilitation ward (dedicated stroke unit) (n = 116) vs general medical ward (n = 89) or geriatric medical ward (mixed rehabilitation unit) (n = 28) Organised care provided for months if required
Outcomes	Death, Rankin score, place of residence, length of stay in hospital up to 1 year after stroke
Notes	Randomisation resulted in marginally poorer prognosis in participants in the control group Numbers differ slightly from the published report after re-analysis of original data 2 control participants lost to follow-up

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Random allocation ... by the secretary opening the next in a stack of serially numbered sealed envelopes"
Allocation concealment (selection bias)	Low risk	Adequate allocation concealment
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal

Dover 1984 (Continued)

Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data explained; broadly similar numbers between intervention and control groups
Selective reporting (reporting bias)	Unclear risk	Rankin score pre-specified but not reported Disability reported with a different measure

Dover 1984 (GMW)
Study characteristics

Methods	RCT Subgroup of Dover 1984 (stroke unit vs general medical ward)
Participants	People with stroke up to 9 weeks after stroke onset (majority within 3 weeks) Fit for transfer to rehabilitation ward
Interventions	Stroke rehabilitation ward (dedicated stroke unit) (n = 98) vs general medical ward (n = 89) Organised care provided for months if required
Outcomes	Death, Rankin score, place of residence, length of stay in hospital up to 1 year after stroke
Notes	Stroke severity subgroup data inferred from distribution in the whole group

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Random allocation ... by the secretary opening the next in a stack of serially numbered sealed envelopes"
Allocation concealment (selection bias)	Low risk	Adequate allocation concealment
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data explained; broadly similar numbers between intervention and control groups
Selective reporting (reporting bias)	Unclear risk	Rankin score pre-specified but not reported Disability reported in an alternate way

Dover 1984 (MRW)

Study characteristics

Methods	RCT Subgroup of Dover 1984 (stroke unit vs mixed rehabilitation ward)
Participants	People with stroke up to 9 weeks after stroke onset (majority within 3 weeks) Fit for transfer to rehabilitation ward.
Interventions	Stroke rehabilitation ward (dedicated stroke unit) (n = 18) vs geriatric medical ward (mixed rehabilitation unit) (n = 28) Organised care provided for months if required
Outcomes	Death, Rankin score, place of residence, length of stay in hospital up to 1 year after stroke
Notes	Stroke severity subgroup data inferred from distribution in the whole group

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Random allocation ... by the secretary opening the next in a stack of serially numbered sealed envelopes"
Allocation concealment (selection bias)	Low risk	Adequate allocation concealment
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	Missing outcome data explained; broadly similar numbers between intervention and control groups
Selective reporting (reporting bias)	Unclear risk	Rankin score pre-specified but not reported Disability reported in an alternate way

Edinburgh 1980

Study characteristics

Methods	RCT
Participants	People with acute stroke within 7 days of stroke onset Stroke of moderate severity
Interventions	Comprehensive stroke ward (dedicated stroke unit) (n = 155) vs general medical ward (n = 156)

Edinburgh 1980 (Continued)

Organised care provided for a maximum of 16 weeks

Outcomes Death, dependency, place of residence, length of initial hospital admission up to 1 year after stroke

Notes 6 intervention and 10 control participants lost to follow-up

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation using numbered sealed envelopes
Allocation concealment (selection bias)	Low risk	Serially numbered sealed envelopes
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	4 participants in control group 'dropped out' after randomisation; no outcome data provided
Selective reporting (reporting bias)	Low risk	Outcomes were not clearly pre-specified but all expected outcomes are reported

Goteborg-Ostra 1988
Study characteristics

Methods	RCT
Participants	People with acute stroke within 7 days of stroke
Interventions	Comprehensive stroke ward (n = 215) within general medical service vs conventional care in general medical ward (n = 202)
Outcomes	Death, Barthel Index, place of residence, length of hospital stay recorded at discharge
Notes	Not yet published

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated randomisation in closed envelopes

Goteborg-Ostra 1988 (Continued)

Allocation concealment (selection bias)	Low risk	Adequate allocation concealment
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Unclear
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Unclear
Selective reporting (reporting bias)	Unclear risk	Unclear

Goteborg-Sahlgren 1994
Study characteristics

Methods	RCT
Participants	People with acute stroke within 7 days of onset
Interventions	Combined service continuum linking 2 acute and 2 rehabilitation stroke wards (n = 166) vs conventional care in general medical ward (n = 83)
Outcomes	Death, dependency (Barthel Index), place of residence, satisfaction, length of hospital stay up to 1 year
Notes	–

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Serially numbered sealed envelopes (randomization in blocks of 10)"
Allocation concealment (selection bias)	Low risk	Adequate allocation concealment
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias)	Low risk	All dichotomous outcomes reported, but proportionately more follow-up assessments missing in control group (7/83) than in intervention group (6/166)

Goteborg-Sahlgren 1994 (Continued)

All outcomes

Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported
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Groningen 2003
Study characteristics

Methods	RCT Blinded assessment of outcomes
Participants	People with acute ischaemic stroke admitted within 24 hours (conscious, hemiparetic, no prior dependency)
Interventions	Acute stroke unit with continuous physiological monitoring and intervention for 48 hours All other care as per conventional stroke unit Transfer to conventional stroke unit after 48 hours Conventional stroke unit: comprehensive stroke ward with intermittent physiological monitoring Both units had a multi-disciplinary team meeting once per week Both units had discharge for rehabilitation at about 2 weeks
Outcomes	Death or poor outcome (institutional care or Rankin score > 3 or Barthel Index < 12) recorded at 3 months Complications and interventions, length of stay

Notes

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomised ... using an envelope system on a one to one basis ..."
Allocation concealment (selection bias)	Low risk	Opaque sealed envelopes
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal but similar care routines
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Low risk	All pre-specified outcome data reported

Guangdong 2008

Study characteristics

Methods	RCT
Participants	People (56 men) with acute ischaemic stroke; timing of randomisation unclear Mean age intervention group: 61.4 years (SD 9.05); mean age control group: 60.9 years (SD 8.2)
Interventions	Stroke unit plus integrated traditional Chinese medicine (n = 58) vs 'Western medicine' stroke unit (n = 42)
Outcomes	Death, NIHSS at 30 days, Barthel Index Length of follow-up unclear
Notes	Limited translated data available

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Random number generator
Allocation concealment (selection bias)	Unclear risk	Unclear
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Unclear
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Unclear
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Unclear
Selective reporting (reporting bias)	Unclear risk	Unclear

Guangdong 2009

Study characteristics

Methods	RCT
Participants	Participants (137 men) with acute ischaemic stroke, randomised on admission Average age 61.9 years in intervention group vs 63.4 years in control group
Interventions	Stroke unit with integrated traditional Chinese medicine (n = 100) vs general medical ward (n = 100)

Helsinki 1995 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomisation was carried out in blocks of 10, with numbered sealed envelopes"
Allocation concealment (selection bias)	Low risk	Adequate allocation concealment
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data ITT analysis
Selective reporting (reporting bias)	Unclear risk	Not all pre-specified outcome data reported

Huaihua 2004
Study characteristics

Methods	RCT
Participants	People (292 men) with acute ischaemic stroke, randomised on admission Age 38 to 79 years (mean age 59.2 years)
Interventions	Comprehensive stroke unit within neurology department (n = 324) vs general medical ward (n = 73)
Outcomes	Death or poor outcome at 1 year Functional ability at 1 year, but scale used not clear
Notes	Limited translated data available

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Randomised" Numbers in intervention group much greater than in control group
Allocation concealment (selection bias)	Unclear risk	Unclear
Blinding of participants and personnel (performance bias)	Unclear risk	Unclear

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Huaihua 2004 (Continued)

All outcomes

Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Unclear
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Unclear
Selective reporting (reporting bias)	Unclear risk	Unclear

Hunan 2007
Study characteristics

Methods	RCT
Participants	People (163 men: 61.2%) with acute stroke; timing of randomisation unclear Mean age in intervention group: 62.3 years (SD 10.7); mean age in control group: 61.2 years (SD 11.8)
Interventions	Stroke unit with integrated traditional Chinese medicine (n = 139) vs Western medicine stroke unit (n = 127)
Outcomes	Death and NIHSS, Barthel Index and mRS at 90 days Length of stay
Notes	Limited translated data available

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Random number generator
Allocation concealment (selection bias)	Unclear risk	Unclear
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Unclear
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Unclear

Hunan 2007 (Continued)

Selective reporting (reporting bias)	Unclear risk	Unclear
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Illinois 1966
Study characteristics

Methods	RCT with 3:2 allocation to intervention:control
Participants	People with stroke up to 1 year after stroke onset Appropriate for rehabilitation service
Interventions	Rehabilitation service (mixed rehabilitation unit) (n = 56) vs general medical ward (which had some specialist nursing input) (n = 35) Organised care provided for months if required
Outcomes	Functional status and place of residence at end of follow-up
Notes	Intervention and control services not clearly defined No deaths reported

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Fisher's table of random numbers
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment not described
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Unclear
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data reported
Selective reporting (reporting bias)	Unclear risk	Outcomes were not clearly pre-specified

Joinville 2003
Study characteristics

Methods	RCT by means of randomised numbers in the emergency room Blinded follow-up
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Organised inpatient (stroke unit) care for stroke: network meta-analysis (Review)

Joinville 2003 (Continued)

Participants	Clinical stroke diagnosis (confirmed on CT scan) within 7 days of onset
Interventions	Comprehensive stroke unit within neurology department (n = 35) vs conventional care in general medical ward
Outcomes	Death, Rankin score, length of stay up to 6 months
Notes	-

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"List of randomized numbers available in the emergency room"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment not described
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Kuopio 1985
Study characteristics

Methods	RCT Blinded assessment of outcome
Participants	People with stroke within 7 days of stroke onset Able to tolerate intensive rehabilitation
Interventions	Intensive rehabilitation in neurological rehabilitation unit (mixed rehabilitation ward) (n = 50) vs general ward (n = 45) Organised care provided for months if required
Outcomes	Death, Lehman (disability) score, place of residence, total time in hospital up to 1 year after stroke
Notes	Majority of people screened failed to meet inclusion criteria for the trial

Risk of bias

Kuopio 1985 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomised using sealed envelopes"
Allocation concealment (selection bias)	Low risk	"Sealed envelopes"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Unclear risk	Outcomes were not clearly pre-specified

Manchester 2003
Study characteristics

Methods	RCT Telephone randomisation and blinded follow-up
Participants	People with acute stroke within 5 days of symptoms No recent myocardial infarction or fracture
Interventions	Mobile stroke team (stroke physician, therapist) in 2 acute hospitals provided early assessment and advice to staff, co-ordinated early therapy input, encouraged guideline adherence Controls received usual medical ward-based care
Outcomes	Death, institutional care, dependency, simple questions, Nottingham extended ADL score, Frenchay Aphasia Screening Test, EuroQol, Hospital Anxiety and Depression Scale Recorded up to 12 months
Notes	5 intervention and 4 control missing from final follow-up 23 people underwent secondary randomisation in trial of early supported discharge team

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Offsite office using a computer generated schedule"
Allocation concealment (selection bias)	Low risk	"Allocated using a simple computer generated procedure ... initially and then in the later stages a minimisation procedure"

Manchester 2003 *(Continued)*

Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	Proportionately small; similar numbers missing from intervention and control groups at 12 months
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Montreal 1985
Study characteristics

Methods	RCT Blinded assessment of outcome
Participants	Unselected people with stroke within 7 days of stroke onset
Interventions	Mobile stroke team (dedicated stroke unit) (n = 65) vs conventional care on general medical ward (n = 65) Study ended at 6 weeks post stroke
Outcomes	Death, Barthel Index, place of residence, length of initial hospital stay up to 6 weeks after stroke
Notes	Short follow-up period 1 intervention and 3 control patients lost to follow-up

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Patients were stratified ... "block randomization within each stratum"
Allocation concealment (selection bias)	Low risk	"Two series of numbered sealed envelopes"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	3 participants (1 intervention; 2 control) removed from study due to non-stroke diagnosis following randomisation

Montreal 1985 *(Continued)*

1 additional participant not admitted from the emergency room

Selective reporting (reporting bias)	Unclear risk	Not all pre-specified outcomes reported
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New South Wales 2014
Study characteristics

Methods	RCT: prospective, single-blind, randomised controlled trial across 4 hospitals
Participants	People with stroke (ischaemic and haemorrhagic) on day 1 of admission if vacant beds existed in both acute and rehabilitation hospitals and person had an acute stroke within previous 24 to 48 hours with sufficient neurological impairment and disability to require ongoing rehabilitation
Interventions	<p>The 'intervention' was the 'early' commencement of a rehabilitation process, with a significant portion of acute care to be spent in a rehabilitation setting</p> <p>Traditional stroke care (TSC): participants were admitted into an acute stroke unit (ASU) and were transferred to a rehabilitation unit at the end of their acute stroke phase (after completion of investigations and acute treatment, and when medically stable as per usual practice). Therefore participants were cared for in 2 different stages (acute and rehabilitation) by different nursing and allied health teams</p> <p>Comprehensive stroke care (CSC): participants were pressed to be transferred to a rehabilitation bed, aiming within 24 to 48 hours after arrival at ASU (or the next working day if weekend). This occurred when participants were still in acute stroke phase and might require attention to acute medical problems should they arise. Hence it was not equivalent to early transfer to a rehabilitation unit. Participants in the CSC arm were cared for by the same nursing and allied health team for a larger portion of their hospital stay</p> <p>All standard and best possible care was given to participants in both arms and the same treatment interventions were available to participants admitted to either arm (with the exception of rehabilitation process allowed to happen earlier after faster arrival in rehabilitation setting in the CSC arm)</p>
Outcomes	<p>Death</p> <p>FIM at discharge and at 3 months</p> <p>Modified Rankin score at discharge</p> <p>FIM efficiency (11) (change in FIM score ÷ total LOS) between participants who received CSC and those who received TSC. The FIM efficiency is an indicator of the rate of functional improvement per day of hospital stay</p> <p>Total hospital length of stay (acute and rehabilitation units combined)</p> <p>All FIM assessments were performed by the same research officer who had passed training sessions in performing the assessment tool (i.e. 1 research officer for each pair of acute rehabilitation units/hospitals)</p> <p>Length of stay for all participants was decided by the team caring for the participants, and information was extracted from medical records and counter-checked with the team by research officers upon discharge of participants</p>
Notes	Numbers independent at 3 months calculated as number FIM > 110 calculated from mean and SD

Risk of bias

New South Wales 2014 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomization was generated centrally by a biostatistician using a computer software program Stata11 (StataCorp LP, College Station, TX, USA)"
Allocation concealment (selection bias)	Low risk	"The generated results were concealed and stored locally. Clinicians at the rehabilitation services were not informed as to whether a patient was randomized into the study"
Blinding of participants and personnel (performance bias) All outcomes	Low risk	"Clinicians at the rehabilitation services were not informed as to whether a patient was randomized into the study" "All standard and best possible care was given to participants in both arms, and the same treatment interventions were available to patients admitted to either arm"
Blinding of outcome assessment (detection bias) All outcomes	Low risk	"The research officers who conducted baseline measures and subsequent FIM assessments at discharge and 90 days telephone follow-up interview were also blind to group allocation" "The rehabilitation teams were blind to the group allocation of the patients"
Incomplete outcome data (attrition bias) All outcomes	Low risk	A total of 4/47 (9%) participants withdrew
Selective reporting (reporting bias)	Unclear risk	Unclear

New York 1962
Study characteristics

Methods	RCT
Participants	People with stroke up to 2 months after stroke Appropriate for rehabilitation centre
Interventions	Mixed rehabilitation team working in rehabilitation centre or attending participants in other wards (n = 42) vs programme of care in general ward (n = 40) that had some specialist nursing input Organised care provided for months if required
Outcomes	Functional status and place of residence at end of follow-up (approximately 1 year)
Notes	No deaths reported Minor anomaly in published data table Not clear how many participants were managed in a peripatetic way

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomly drawn unmarked envelopes

New York 1962 *(Continued)*

Allocation concealment (selection bias)	Unclear risk	Allocation concealment not reported
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Unclear
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Unclear
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Unclear
Selective reporting (reporting bias)	Unclear risk	Unclear

Newcastle 1993
Study characteristics

Methods	RCT
Participants	Stroke patients within 3 days of stroke onset
Interventions	Mixed rehabilitation ward in geriatric medicine department (n = 34) vs general medical ward (n = 33) Organised care provided for months if required
Outcomes	Death, Barthel Index, Rankin score, place of residence, length of stay in hospital up to 6 months after stroke
Notes	Majority of patients screened failed to meet trial inclusion criteria

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Stratified based on continence and then randomly allocated"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment unclear
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias)	Low risk	No missing outcome data

Organised inpatient (stroke unit) care for stroke: network meta-analysis (Review)

Newcastle 1993 (Continued)

All outcomes

Selective reporting (reporting bias)	Unclear risk	Unclear
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Nottingham 1996
Study characteristics

Methods	RCT with 5:4 allocation of intervention:control Blinded assessment of outcome
Participants	Patients with stroke at 2 weeks after stroke onset Able to participate actively in rehabilitation
Interventions	Stroke rehabilitation ward in department of geriatric medicine (n = 176) vs conventional care in geriatric medical (mixed rehabilitation) ward (n = 63) or general medical ward (n = 76) Organised care provided for months if required
Outcomes	Death, Barthel Index, place of residence, Nottingham Health Profile, length of hospital stay up to 1 year after stroke
Notes	Some cross-over from general medical ward to geriatric medicine department 3 intervention and 4 control participants lost to follow-up

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Stratified based on admission ward ... then randomly allocated"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment unclear
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Small numbers (3 intervention; 4 control) lost to follow-up Some secondary outcome assessments not completed or partially completed; this varied between groups
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Nottingham 1996 (GMW)
Study characteristics

Methods	RCT Subgroup of Nottingham (stroke unit vs general medical ward)
Participants	People with stroke at 2 weeks after stroke onset Able to participate actively in rehabilitation
Interventions	Stroke rehabilitation ward in department of geriatric medicine (n = 78) vs conventional care in geriatric medical (mixed rehabilitation) ward (n = 63) Organised care provided for months if required
Outcomes	Death, Barthel Index, place of residence, Nottingham Health Profile, length of hospital stay up to 1 year after stroke
Notes	Some cross-over from general medical ward to geriatric medicine department

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Stratified based on admission ward ... then randomly allocated"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment unclear
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	Some secondary outcome assessments not completed or partially completed; this varied between groups
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Nottingham 1996 (MRW)
Study characteristics

Methods	RCT Subgroup of Nottingham (stroke unit vs mixed rehabilitation ward)
Participants	People with stroke at 2 weeks after stroke onset Able to participate actively in rehabilitation
Interventions	Stroke rehabilitation ward in department of geriatric medicine (n = 98) vs conventional care in general medical ward (n = 76)

Nottingham 1996 (MRW) *(Continued)*

Organised care provided for months if required

Outcomes	Death, Barthel Index, place of residence, Nottingham Health Profile, length of hospital stay up to 1 year after stroke
Notes	Some cross-over from general medical ward to geriatric medicine department

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Stratified based on admission ward... then randomly allocated"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment unclear
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	Some secondary outcome assessments not completed or partially completed; this varied between groups
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Orpington 1993
Study characteristics

Methods	RCT
Participants	People with stroke who had survived for 2 weeks Suitable for transfer to rehabilitation ward
Interventions	Stroke rehabilitation ward (n = 124) vs conventional care in geriatric (mixed rehabilitation unit) (n = 73) or general medical (n = 48) ward Organised care provided for months if required
Outcomes	Death, Barthel Index, place of residence, length of initial hospital stay at end of follow-up 2 intervention and 5 control patients lost to follow-up
Notes	Variable duration of follow-up (hospital discharge)

Risk of bias

Bias	Authors' judgement	Support for judgement
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Orpington 1993 (Continued)

Random sequence generation (selection bias)	Low risk	"Randomised with the use of Geigy table of random numbers"
Allocation concealment (selection bias)	Low risk	"Randomisation was computerized"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	2 intervention and 5 control participants lost to follow-up
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Orpington 1993 (GMW)
Study characteristics

Methods	RCT Subgroup of Orpington 1993 (stroke unit vs general medical ward)
Participants	People who survived stroke for 2 weeks Suitable for transfer to rehabilitation ward
Interventions	Stroke rehabilitation ward (n = 53) vs conventional care in general medical (n = 48) ward Organised care provided for months if required
Outcomes	Death, Barthel Index, place of residence, length of initial hospital stay at end of follow-up
Notes	Stroke severity subgroup data inferred from distribution in whole group

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomised with the use of Geigy table of random numbers"
Allocation concealment (selection bias)	Low risk	"Randomisation was computerized"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias)	High risk	Unblinded outcome assessment

Organised inpatient (stroke unit) care for stroke: network meta-analysis (Review)

Orpington 1993 (GMW) *(Continued)*

All outcomes

Incomplete outcome data (attrition bias) All outcomes	Unclear risk	2 intervention and 5 control participants lost to follow-up
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Orpington 1993 (MRW)
Study characteristics

Methods	RCT Subgroup of Orpington 1993 (stroke unit vs mixed rehabilitation ward)
Participants	People who survived stroke for 2 weeks Suitable for transfer to rehabilitation ward
Interventions	Stroke rehabilitation ward (n = 71) vs conventional care in geriatric (mixed rehabilitation) ward (n = 73) Organised care provided for months if required
Outcomes	Death, Barthel Index, place of residence, length of initial hospital stay at end of follow-up
Notes	Stroke severity subgroup data inferred from distribution in whole group

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomised with the use of Geigy table of random numbers"
Allocation concealment (selection bias)	Low risk	"Randomisation was computerized"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	2 intervention and 5 control participants lost to follow-up
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Orpington 1995

Study characteristics

Methods	RCT
Participants	People who had a poor prognosis 2 weeks after stroke Suitable for transfer to rehabilitation ward
Interventions	Stroke rehabilitation ward in geriatric medicine department (n = 36) vs general medical ward (n = 37) Organised care provided for months if required
Outcomes	Death, Barthel Index, place of residence, length of hospital stay at end of follow-up
Notes	Variable duration of follow-up (hospital discharge) 2 control participants lost to follow-up; assumed to be alive and independent (ITT analysis)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	"Were randomized" "The process of randomization was not limited by bed availability"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment not described
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Unclear
Incomplete outcome data (attrition bias) All outcomes	Low risk	ITT analysis
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Orpington 2000

Study characteristics

Methods	RCT Blinded outcome assessment
Participants	People with acute stroke (meeting WHO definition of stroke) from a community stroke register Intermediate stroke severity
Interventions	3-arm comparison of: <ul style="list-style-type: none"> comprehensive stroke ward (co-ordinated multi-disciplinary team care) (n = 152);

Orpington 2000 (Continued)

- general ward with input from hospital mobile stroke team (comprising medical, physiotherapy, occupational therapy, speech therapy but not nursing or medical specialists) (n = 152); and
- domiciliary multi-disciplinary stroke team (not relevant to this review)

Outcomes	Death, dependency (Barthel Index), place of residence, length of stay, resource use up to 12 months 3 control participants lost to follow-up
Notes	-

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Unstratified ... using the block randomization technique computer generated random numbers"
Allocation concealment (selection bias)	Low risk	"Allocation schedule prepared using computer generated random numbers"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	3 control participants lost to follow-up at 12 months
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Perth 1997
Study characteristics

Methods	RCT
Participants	People with acute stroke within 7 days of stroke onset
Interventions	Comprehensive stroke ward (dedicated stroke unit) (n = 29) vs general medical ward (n = 30) Organised care provided for months if required
Outcomes	Death, Barthel Index, place of residence, length of hospital stay up to 6 months after stroke
Notes	Most people screened did not enter trial

Risk of bias

Bias	Authors' judgement	Support for judgement
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Perth 1997 (Continued)

Random sequence generation (selection bias)	Unclear risk	"Were randomized"
Allocation concealment (selection bias)	Unclear risk	Method of allocation concealment not reported
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Blinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Low risk	Outcomes not clearly pre-specified but all expected outcomes reported

Svendborg 1995
Study characteristics

Methods	RCT by means of sealed envelopes (stratified by age and side of lesion)
Participants	People with acute stroke patients (within 8 days of symptoms) meeting WHO diagnostic criteria
Interventions	Comprehensive stroke ward (n = 31) vs conventional care in general medical ward (n = 34)
Outcomes	Death, dependency (Rankin score), place of residence, length of hospital stay at 6 months after randomisation
Notes	Staffing levels were higher in the stroke unit group

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Translation - randomised by the envelope method (drawing lots), stratified by age and side of lesion
Allocation concealment (selection bias)	Low risk	Sealed envelopes
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Unclear

Svendborg 1995 (Continued)

Incomplete outcome data (attrition bias) All outcomes	Low risk	No obvious missing outcome data
Selective reporting (reporting bias)	Unclear risk	Unclear

Tampere 1993
Study characteristics

Methods	RCT
Participants	People with acute stroke within 7 days of stroke (usually earlier)
Interventions	Acute (semi-intensive) stroke ward in neurology department (n = 98) vs conventional care in neurology department (mixed rehabilitation unit) (n = 113) Organised care provided for approximately 1 week only
Outcomes	Death, Rankin score, place of residence, length of hospital stay up to 1 year after stroke 1 intervention and 1 control participant removed due to non-stroke diagnosis
Notes	Short duration (1 week) in stroke unit before transfer to conventional service

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomisation was performed with the aid of a table of random numbers" "Randomly assigned using serially numbered, sealed, envelopes"
Allocation concealment (selection bias)	Low risk	"Serially numbered, sealed, envelopes"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	High risk	Unblinded outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	1 participant in intervention group and 1 participant in control group removed due to incorrect diagnosis
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

Trondheim 1991
Study characteristics

Methods	RCT
Participants	People with stroke within 7 days (usually within 24 hours) of stroke onset Exclusion of deeply unconscious patients and those previously residing in a nursing home
Interventions	Comprehensive stroke ward (dedicated stroke unit) (n = 110) vs general medical ward (n = 110) Organised care provided for a maximum of 6 weeks
Outcomes	Death, Barthel Index, place of residence, length of stay in hospital or institution up to 1 year after stroke
Notes	-

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	"Randomly assigned ... using serially numbered sealed envelopes"
Allocation concealment (selection bias)	Low risk	"Serially numbered sealed envelopes"
Blinding of participants and personnel (performance bias) All outcomes	Unclear risk	Difficult to conceal
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Both blinded and open assessments available for 50% of participants at 52 weeks; open assessments available for only 50% Correlation between blinded and open was high, but risk of bias remains unclear
Incomplete outcome data (attrition bias) All outcomes	Low risk	No missing outcome data
Selective reporting (reporting bias)	Low risk	All pre-specified outcomes reported

ADL: activity of daily living.
 CT: computerised tomography.
 FIM: Functional Independence Measure.
 GMW: general medical ward.
 ITT: intention-to-treat.
 LOS: length of stay.
 mRS: modified Rankin Scale.
 MRW: mixed rehabilitation ward.
 NIHSS: National Institutes of Health Stroke Scale.
 OHS: Oxford Handicap Scale.
 RCT: randomised controlled trial.
 SD: standard deviation.
 SPSS: Statistical Package for the Social Sciences.
 TIA: transient ischaemic attack.
 WHO: World Health Organization.

Characteristics of excluded studies [ordered by study ID]

Study	Reason for exclusion
Abissi 1995	Trial tested a care plan protocol only No other aspect of organisation was under evaluation
Akhtar 2015	Not an RCT
Al-Qahtany 2014	Not an RCT
Asplund 2000	Trial of a geriatric assessment unit
Cavallini 2003	Quasi-randomised treatment allocation
Davis 2000	Intervention and control arms of trial were treated within the same stroke unit
Di Lauro 2003	Intervention and control arms of trial were treated within the same stroke unit
Diagana 2008	Quasi-randomised treatment allocation
Durastanti 2005	Quasi-randomised treatment allocation
Felix 2016	Trial of stroke education
Fu 2006	Not an RCT
HAMLET 2009	Does not report outcomes for different medical treatment arms
Hamrin 1982	Quasi-randomised treatment allocation
He 2014	Not a fully randomised cluster-RCT
Inoue 2013	Not an RCT
Janssen 2014	Trial of enriched environment
Koton 2005	Treatment allocated by selection criteria
Langhorne 2001	Study tested a care plan protocol only No other aspect of organisation was under evaluation
Middleton 2006	Care pathway study only
Middleton 2018	Care pathway study
Moloney 1999	Care pathway study only
Pappa 2009	Non-randomised
Patel 2000	Quasi-randomised treatment allocation
Pearson 1988	No available outcome data
Rai 2016	Not a full RCT
Raiborirug 2017	Not an RCT

Study	Reason for exclusion
Ricauda 2004	Trial comparing home care team vs general medical ward
Ronning 1998	Quasi-randomised treatment allocation
Ronning 1998a	A portion of the data was collected retrospectively All prospective data are included in the Akershus study (Ronning 1998)
Ronning 1998b	Comparison of stroke rehabilitation ward vs discharge to community-based stroke rehabilitation
Shiraishi 2004	Non-randomised treatment allocation
Silva 2004	Treatment allocated by the study neurologist
Stone 1998	No available outcome data
Strand 1985	Quasi-randomised treatment allocation
Von Arbin 1980	Quasi-randomised treatment allocation
Walter 2005	Non-randomised treatment allocation
Wang 2004	No available outcome data
Yagura 2005	Quasi-randomised treatment allocation

RCT: randomised controlled trial.

Characteristics of studies awaiting classification *[ordered by study ID]*

Anhui 2008

Methods	RCT
Participants	People with acute stroke
Interventions	"Standardised tertiary rehabilitation" (n = 51) vs usual inpatient care (n = 51)
Outcomes	Functional outcome (unknown scale) and quality of life (WHOQOL-BREF) at 1, 3, and 6 months Cost analysis
Notes	Currently no useable data

China (Hao) 2010

Methods	Possible RCT
Participants	People with pneumonia (n = 159) after acute stroke (within 2 weeks)
Interventions	Management in comprehensive stroke unit versus general ward Allocated 'treatment' group depended on which ward the person was in when pneumonia developed

China (Hao) 2010 *(Continued)*

Outcomes	Death, NIHSS, Barthel Index at 21 days Length of stay Cost analysis
Notes	Method of randomisation unclear

China (Pei) 2011

Methods	RCT
Participants	People with stroke (n = 236)
Interventions	Randomly assigned to organised stroke care model with integrated Chinese medicine (n = 121) vs traditional care model (n = 115)
Outcomes	Death, NIHSS, Barthel Index, OHS score at 21 days
Notes	Currently no useable data

China (Wang) 2008

Methods	RCT
Participants	People with 'acute cerebral infarction'
Interventions	Randomly assigned to stroke rehabilitation unit group (n = 77) vs ordinary care group (n = 73)
Outcomes	NIHSS Barthel Index (duration of follow-up unclear) Length of stay
Notes	-

China (Wu) 2007

Methods	RCT
Participants	2367 people with acute stroke
Interventions	Randomly assigned to organised stroke ward vs general ward
Outcomes	Death, 'non-recovery', and 'improvement' over 5 years
Notes	Currently no useable data

Haikou 2007

Methods	RCT
Participants	People with acute ischaemic stroke randomised within 1 week
Interventions	Randomised to extended stroke unit vs general medical ward for 3 weeks
Outcomes	Discharge Barthel Index and NIHSS
Notes	Currently no useable data

Shanghai 2006

Methods	RCT
Participants	Cerebral stroke from 22 hospitals
Interventions	"Standardised tertiary rehabilitation" vs routine care
Outcomes	Functional recovery (unknown scale) Cost-effectiveness analysis
Notes	Currently no useable data

NIHSS: National Institutes of Health Stroke Scale.

OHS: Oxford Handicap Scale.

RCT: randomised controlled trial.

WHOQOL-BREF: World Health Organization Quality of Life Project.

Characteristics of ongoing studies *[ordered by study ID]*
ChiCTR-OCH-09000335

Study name	A study of the stroke unit of traditional Chinese and Western medicine in the treatment of ischaemic stroke
Methods	-
Participants	-
Interventions	-
Outcomes	-
Starting date	-
Contact information	Qiujuan Zhang; zqiyyy@hotmail.com
Notes	Yueyang Hospital, Shanghai

China (Wang) 2015

Study name	Rationale and design of a cluster-randomised multifaceted intervention trial to improve stroke care quality in China: the GOLDEN BRIDGE-Acute Ischemic Stroke
Methods	Cluster-RCT
Participants	40 hospitals in China
Interventions	Multi-faceted quality improvement intervention (experimental group) or routine standard of care (control group)
Outcomes	Measure of adherence to evidence-based performance measures: in-hospital death; new vascular event; disability; all-cause death at 3, 6, and 12 months after initial symptom onset
Starting date	-
Contact information	Wang Yilong, Tiantan Clinical Trial and Research Center for Stroke, Department of Neurology, Beijing Tiantan Hospital, Capital Medical University, Beijing, China; China National Clinical Research Center for Neurological Diseases, Beijing, China
Notes	ClinicalTrials.gov/NCT02212912

NCT00544622

Study name	Structured stroke management improves outcomes at 6 months
Methods	-
Participants	-
Interventions	-
Outcomes	-
Starting date	-
Contact information	-
Notes	Kantonsspital Baden

NCT00843765

Study name	Efficiency study of traditional Chinese medicine (TCM) versus Western medicine (WM) on ischaemic stroke
Methods	-
Participants	-
Interventions	-
Outcomes	-

Organised inpatient (stroke unit) care for stroke: network meta-analysis (Review)

NCT00843765 (Continued)

Starting date	-
Contact information	-
Notes	Dongzhimen Hospital and Beijing Tiantan Hospital

Russia 2017

Study name	Development of medical rehabilitation in Russia (DOME): rehabilitation in stroke units and rehabilitation centres
Methods	Large clinical trial
Participants	Acute stroke
Interventions	
Outcomes	Recovery of functions, activity, and participation assessed with modified Rankin scale (mRS)
Starting date	
Contact information	G Ivanova, Pirogov Russian National Medical Research University, Department of Medical and Social Rehabilitation, Ministry of Health of Russia, Moscow, Russian Federation
Notes	ClinicalTrials.gov Identifier: NCT02793934

RCT: randomised controlled trial.

DATA AND ANALYSES
Comparison 1. Organised stroke care versus alternative service

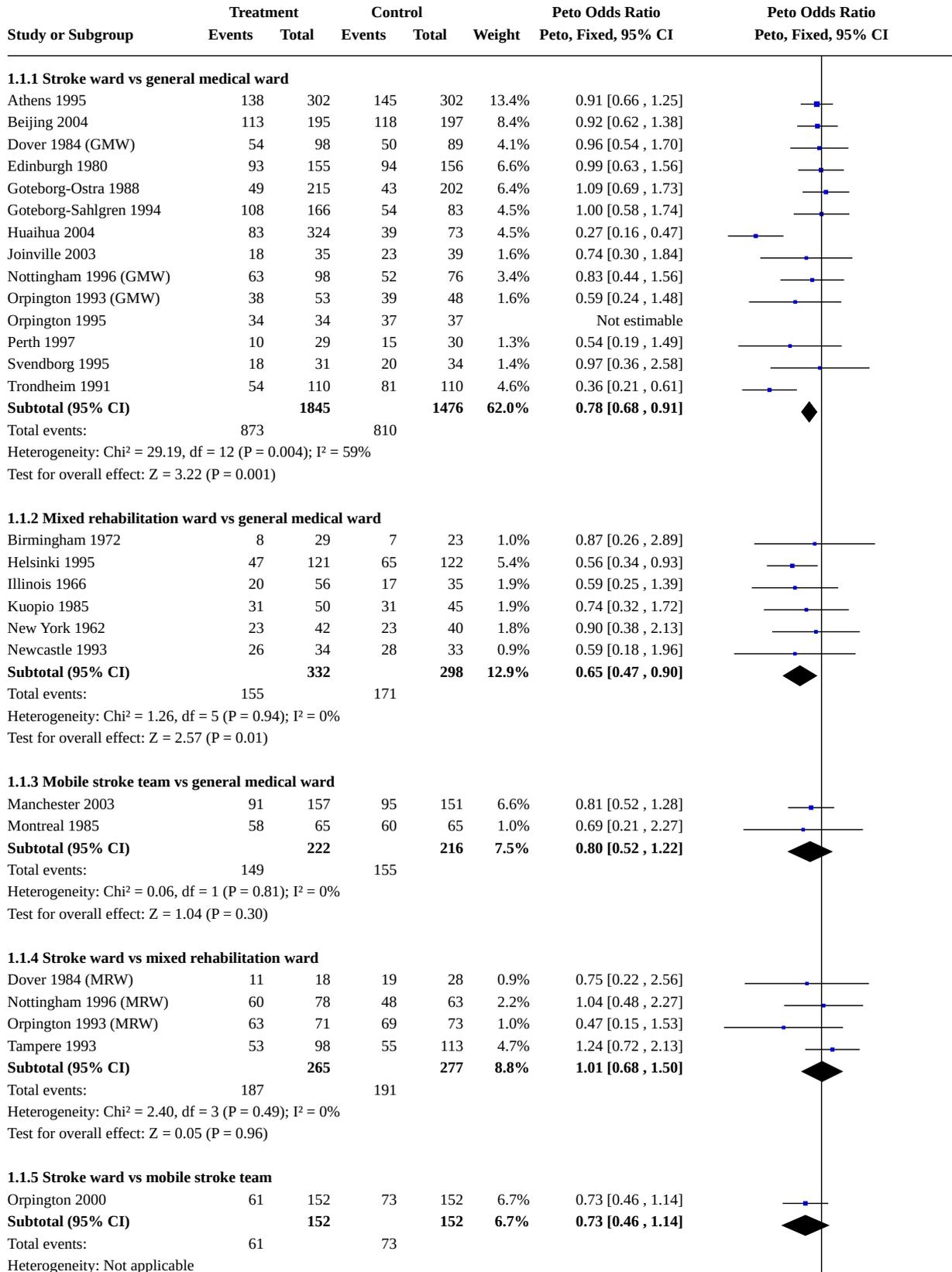
Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1.1 Poor outcome by the end of scheduled follow-up	29	5336	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.77 [0.69, 0.87]
1.1.1 Stroke ward vs general medical ward	14	3321	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.78 [0.68, 0.91]
1.1.2 Mixed rehabilitation ward vs general medical ward	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.65 [0.47, 0.90]
1.1.3 Mobile stroke team vs general medical ward	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.80 [0.52, 1.22]
1.1.4 Stroke ward vs mixed rehabilitation ward	4	542	Peto Odds Ratio (Peto, Fixed, 95% CI)	1.01 [0.68, 1.50]

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1.1.5 Stroke ward vs mobile stroke team	1	304	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.73 [0.46, 1.14]
1.1.6 Stroke ward vs stroke ward	2	101	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.47 [0.21, 1.04]
1.2 Death by the end of scheduled follow-up	32	5902	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.76 [0.66, 0.88]
1.2.1 Stroke ward vs general medical ward	15	3521	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.75 [0.63, 0.90]
1.2.2 Mixed rehabilitation ward vs general medical ward	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.91 [0.58, 1.42]
1.2.3 Mobile stroke team vs general medical ward	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	1.08 [0.71, 1.65]
1.2.4 Stroke ward vs mixed rehabilitation ward	4	542	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.82 [0.54, 1.24]
1.2.5 Stroke ward vs mobile stroke team	1	304	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.35 [0.19, 0.65]
1.2.6 Stroke ward vs stroke ward	4	467	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.36 [0.14, 0.94]
1.3 Death or institutional care by the end of scheduled follow-up	27	4887	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.76 [0.67, 0.85]
1.3.1 Stroke ward vs general medical ward	13	2924	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.74 [0.63, 0.87]
1.3.2 Mixed rehabilitation ward vs general medical ward	5	578	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.71 [0.51, 0.99]
1.3.3 Mobile stroke team vs general medical ward	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	1.27 [0.84, 1.93]
1.3.4 Stroke ward vs mixed rehabilitation ward	4	542	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.90 [0.64, 1.27]
1.3.5 Stroke ward vs mobile stroke team	1	304	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.40 [0.23, 0.68]
1.3.6 Stroke ward vs stroke ward	2	101	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.39 [0.16, 0.93]
1.4 Death or dependency by the end of scheduled follow-up	27	4854	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.75 [0.66, 0.85]
1.4.1 Stroke ward vs general medical ward	12	2839	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.75 [0.64, 0.88]

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1.4.2 Mixed rehabilitation ward vs general medical ward	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.65 [0.47, 0.90]
1.4.3 Mobile stroke team vs general medical ward	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.80 [0.52, 1.22]
1.4.4 Stroke ward vs mixed rehabilitation ward	4	542	Peto Odds Ratio (Peto, Fixed, 95% CI)	1.01 [0.68, 1.50]
1.4.5 Stroke ward vs mobile stroke team	1	304	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.73 [0.46, 1.14]
1.4.6 Stroke ward vs stroke ward	2	101	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.47 [0.21, 1.04]
1.5 Length of stay (days) in a hospital or institution or both	20	4162	Mean Difference (IV, Random, 95% CI)	-4.28 [-7.86, -0.71]
1.5.1 Stroke ward	17	3775	Mean Difference (IV, Random, 95% CI)	-4.76 [-8.46, -1.05]
1.5.2 Mixed rehabilitation ward	3	387	Mean Difference (IV, Random, 95% CI)	3.85 [-13.49, 21.18]
1.6 Length of stay (days) in a hospital or hospital plus institution	20		Mean Difference (IV, Random, 95% CI)	Subtotals only
1.6.1 Acute hospital stay only	7	1817	Mean Difference (IV, Random, 95% CI)	-2.78 [-6.13, 0.56]
1.6.2 Hospital and institution stay	13	2345	Mean Difference (IV, Random, 95% CI)	-4.84 [-14.52, 4.84]
1.7 Poor outcome at 5-year follow-up	2	535	Odds Ratio (M-H, Random, 95% CI)	0.54 [0.22, 1.34]
1.8 Death at 5-year follow-up	3	1139	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.74 [0.59, 0.94]
1.9 Death or institutional care at 5-year follow-up	2	535	Odds Ratio (M-H, Random, 95% CI)	0.59 [0.33, 1.05]
1.10 Death or dependency at 5-year follow-up	2	535	Odds Ratio (M-H, Random, 95% CI)	0.54 [0.22, 1.34]
1.11 Poor outcome at 10-year follow-up	2	535	Odds Ratio (M-H, Random, 95% CI)	0.70 [0.27, 1.80]
1.12 Death at 10-year follow-up	3	1139	Odds Ratio (M-H, Random, 95% CI)	0.66 [0.43, 1.03]
1.13 Death or institutional care at 10-year follow-up	2	535	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.57 [0.37, 0.88]

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1.14 Death or dependency at 10-year follow-up	2	535	Odds Ratio (M-H, Random, 95% CI)	0.70 [0.27, 1.80]

Analysis 1.1. Comparison 1: Organised stroke care versus alternative service, Outcome 1: Poor outcome by the end of scheduled follow-up



Analysis 1.1. (Continued)

Total events: 61 73
 Heterogeneity: Not applicable
 Test for overall effect: $Z = 1.38$ ($P = 0.17$)

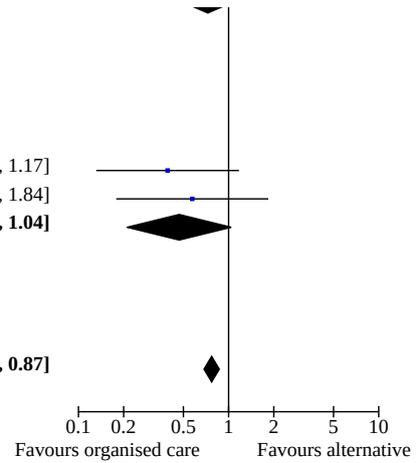
1.1.6 Stroke ward vs stroke ward

Groningen 2003	7	27	13	27	1.1%	0.39 [0.13, 1.17]
New South Wales 2014	8	25	10	22	1.0%	0.57 [0.18, 1.84]
Subtotal (95% CI)		52		49	2.1%	0.47 [0.21, 1.04]

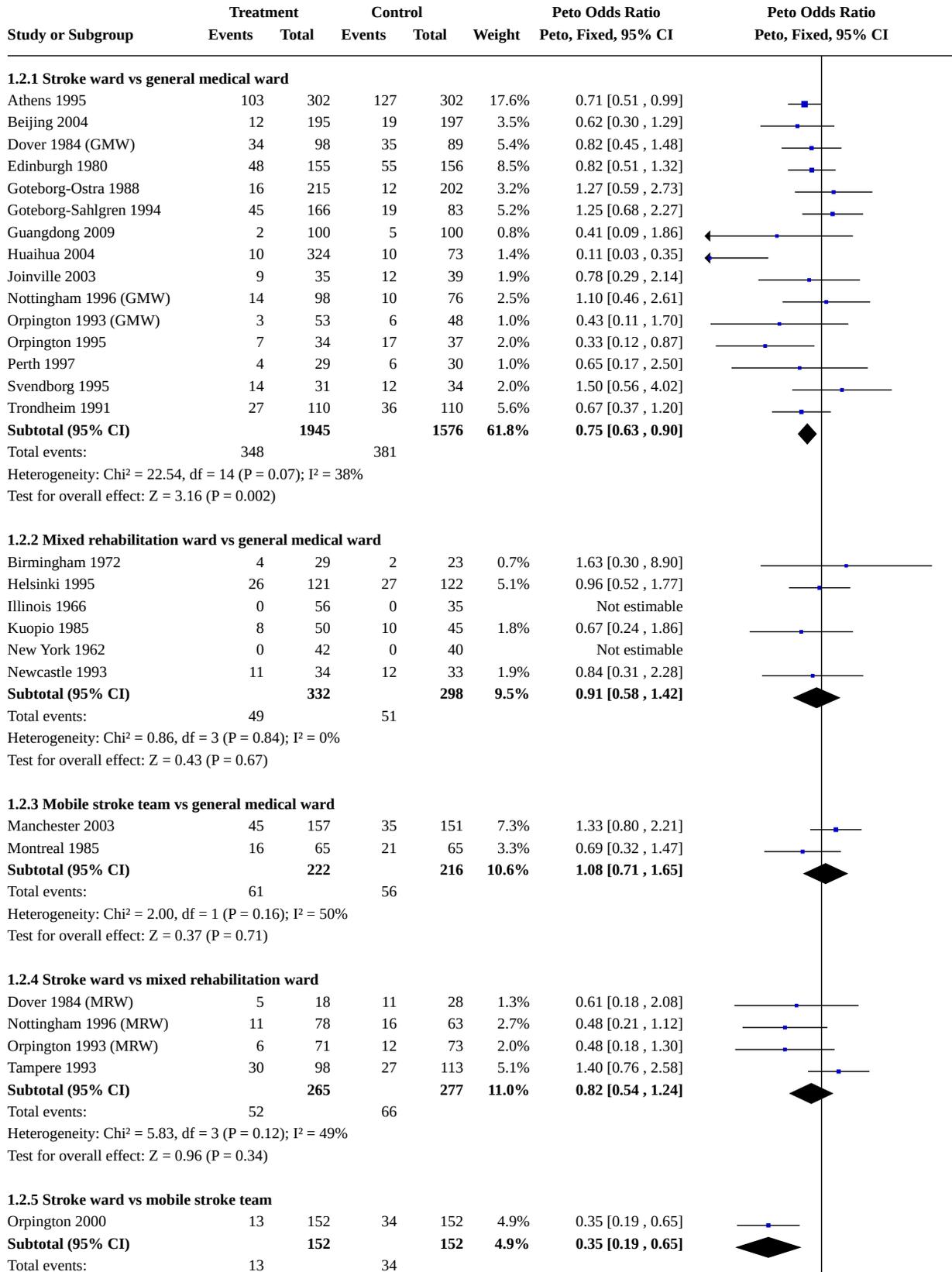
Total events: 15 23
 Heterogeneity: $\text{Chi}^2 = 0.21$, $df = 1$ ($P = 0.64$); $I^2 = 0\%$
 Test for overall effect: $Z = 1.86$ ($P = 0.06$)

Total (95% CI) 2868 2468 100.0% 0.77 [0.69, 0.87]

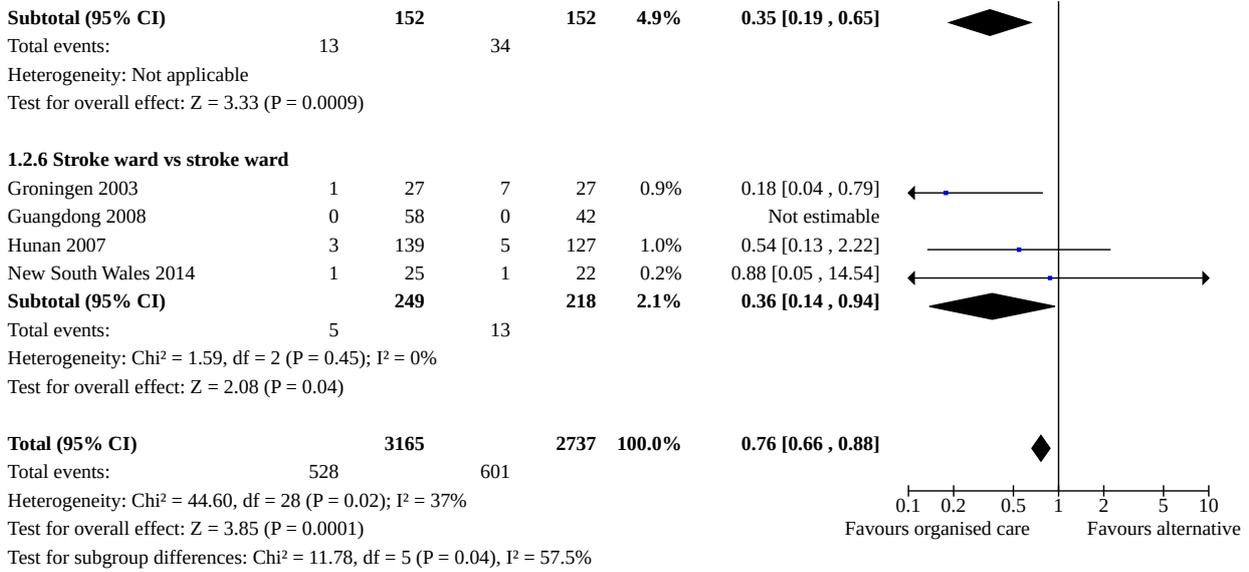
Total events: 1440 1423
 Heterogeneity: $\text{Chi}^2 = 37.58$, $df = 27$ ($P = 0.08$); $I^2 = 28\%$
 Test for overall effect: $Z = 4.36$ ($P < 0.0001$)
 Test for subgroup differences: $\text{Chi}^2 = 4.47$, $df = 5$ ($P = 0.48$), $I^2 = 0\%$



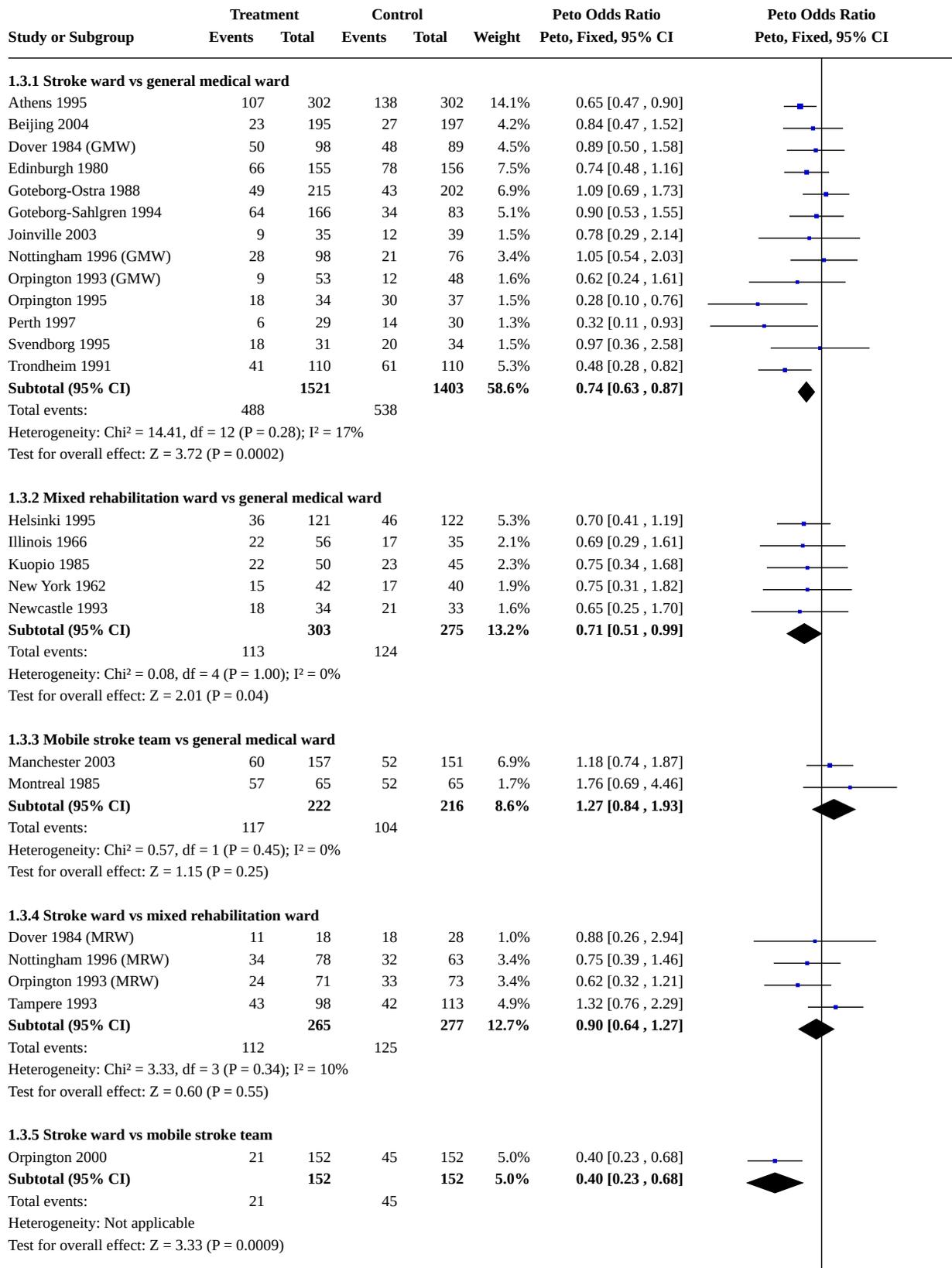
Analysis 1.2. Comparison 1: Organised stroke care versus alternative service, Outcome 2: Death by the end of scheduled follow-up



Analysis 1.2. (Continued)



Analysis 1.3. Comparison 1: Organised stroke care versus alternative service, Outcome 3: Death or institutional care by the end of scheduled follow-up



Analysis 1.3. (Continued)

Test for overall effect: $Z = 3.33$ ($P = 0.0009$)

1.3.6 Stroke ward vs stroke ward

Groningen 2003	13	27	18	27	1.3%	0.48 [0.16 , 1.38]
New South Wales 2014	2	25	6	22	0.7%	0.26 [0.06 , 1.19]
Subtotal (95% CI)		52		49	2.0%	0.39 [0.16 , 0.93]

Total events: 15 24

Heterogeneity: $\text{Chi}^2 = 0.39$, $\text{df} = 1$ ($P = 0.53$); $I^2 = 0\%$

Test for overall effect: $Z = 2.12$ ($P = 0.03$)

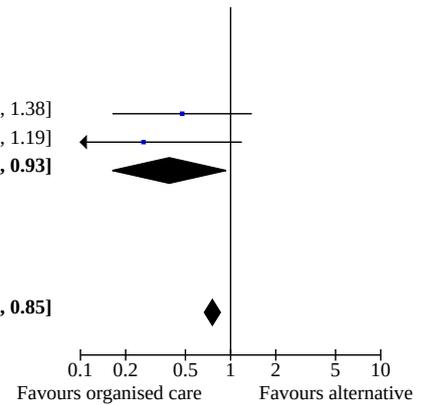
Total (95% CI) 2515 2372 **100.0%** **0.76 [0.67 , 0.85]**

Total events: 866 960

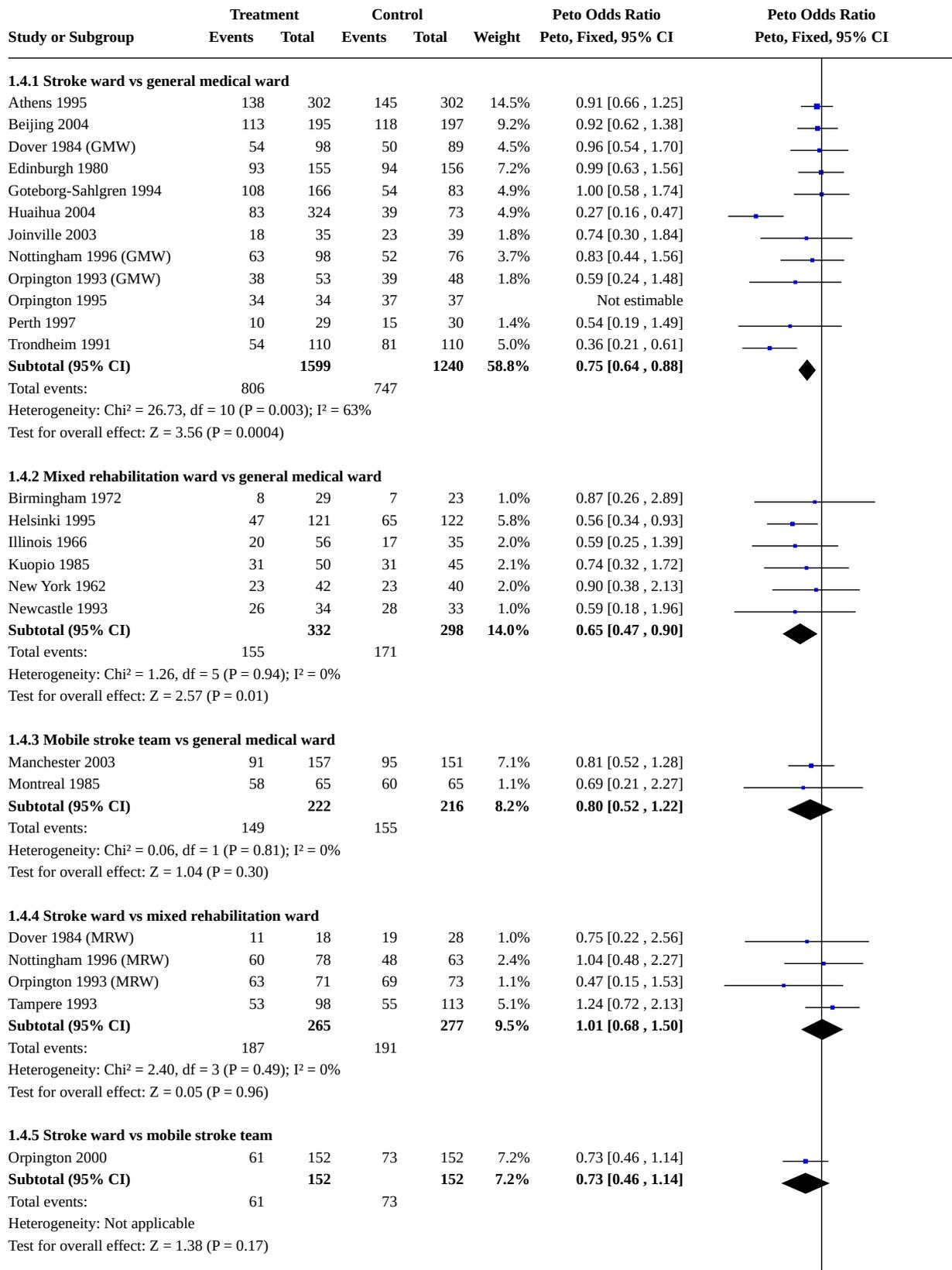
Heterogeneity: $\text{Chi}^2 = 33.70$, $\text{df} = 26$ ($P = 0.14$); $I^2 = 23\%$

Test for overall effect: $Z = 4.49$ ($P < 0.00001$)

Test for subgroup differences: $\text{Chi}^2 = 14.92$, $\text{df} = 5$ ($P = 0.01$), $I^2 = 66.5\%$



Analysis 1.4. Comparison 1: Organised stroke care versus alternative service, Outcome 4: Death or dependency by the end of scheduled follow-up



Analysis 1.4. (Continued)

Test for overall effect: $Z = 1.38$ ($P = 0.17$)

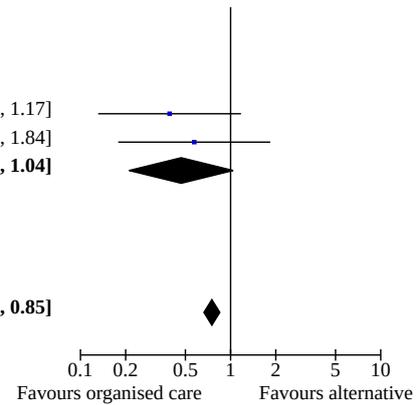
1.4.6 Stroke ward vs stroke ward

Groningen 2003	7	27	13	27	1.2%	0.39 [0.13 , 1.17]
New South Wales 2014	8	25	10	22	1.1%	0.57 [0.18 , 1.84]
Subtotal (95% CI)		52		49	2.3%	0.47 [0.21 , 1.04]

Total events: 15 23
Heterogeneity: $\text{Chi}^2 = 0.21$, $\text{df} = 1$ ($P = 0.64$); $I^2 = 0\%$
Test for overall effect: $Z = 1.86$ ($P = 0.06$)

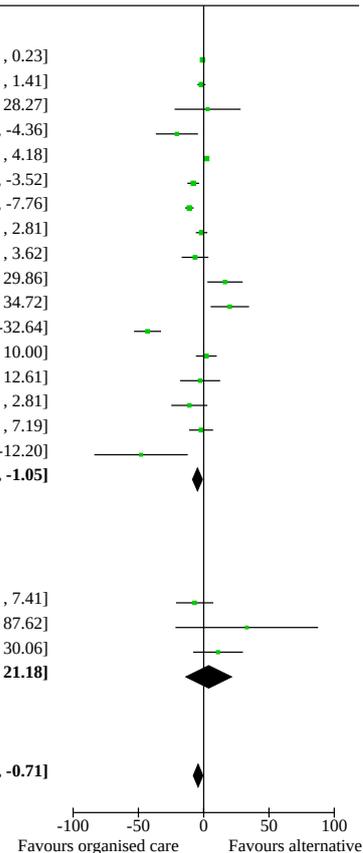
Total (95% CI)		2622		2232	100.0%	0.75 [0.66 , 0.85]
Total events:	1373		1360			

Heterogeneity: $\text{Chi}^2 = 35.01$, $\text{df} = 25$ ($P = 0.09$); $I^2 = 29\%$
Test for overall effect: $Z = 4.63$ ($P < 0.00001$)
Test for subgroup differences: $\text{Chi}^2 = 4.35$, $\text{df} = 5$ ($P = 0.50$), $I^2 = 0\%$

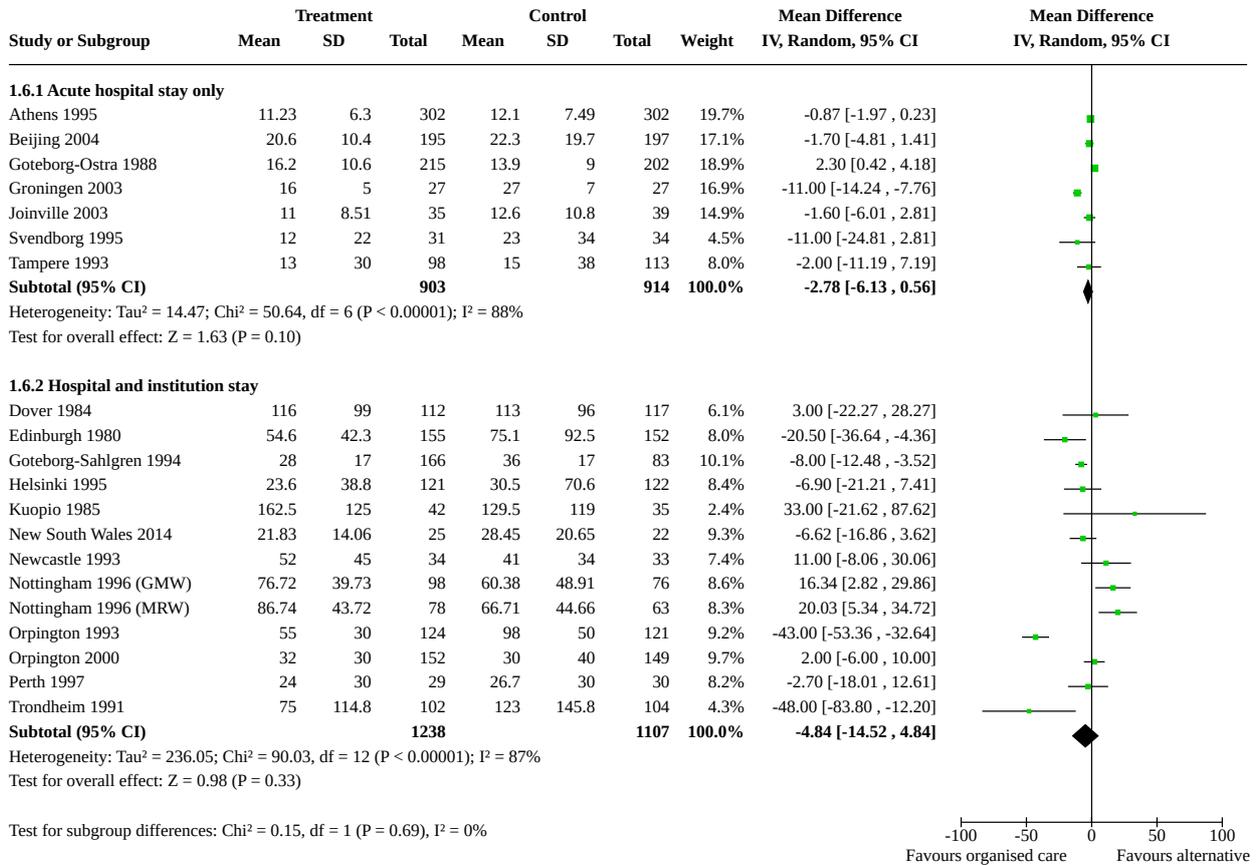


Analysis 1.5. Comparison 1: Organised stroke care versus alternative service, Outcome 5: Length of stay (days) in a hospital or institution or both

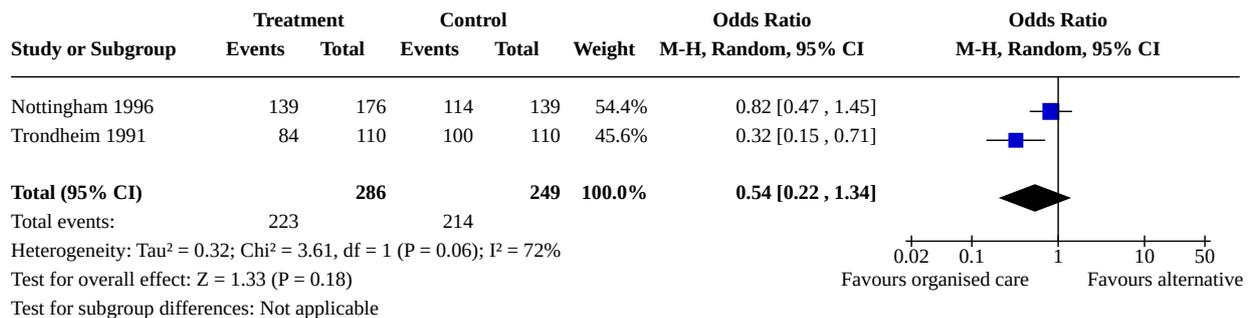
Study or Subgroup	Treatment			Control			Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
	Mean	SD	Total	Mean	SD	Total			
1.5.1 Stroke ward									
Athens 1995	11.23	6.3	302	12.1	7.49	302	9.0%	-0.87 [-1.97, 0.23]	
Beijing 2004	20.6	10.4	195	22.3	19.7	197	8.5%	-1.70 [-4.81, 1.41]	
Dover 1984	116	99	112	113	96	117	1.6%	3.00 [-22.27, 28.27]	
Edinburgh 1980	54.6	42.3	155	75.1	92.5	152	3.2%	-20.50 [-36.64, -4.36]	
Goteborg-Ostra 1988	16.2	10.6	215	13.9	9	202	8.8%	2.30 [0.42, 4.18]	
Goteborg-Sahlgren 1994	28	17	166	36	17	83	7.9%	-8.00 [-12.48, -3.52]	
Groningen 2003	16	5	27	27	7	27	8.4%	-11.00 [-14.24, -7.76]	
Joinville 2003	11	8.51	35	12.6	10.8	39	8.0%	-1.60 [-6.01, 2.81]	
New South Wales 2014	21.83	14.06	25	28.45	20.65	22	5.2%	-6.62 [-16.86, 3.62]	
Nottingham 1996 (GMW)	76.72	39.73	98	60.38	48.91	76	3.9%	16.34 [2.82, 29.86]	
Nottingham 1996 (MRW)	86.74	43.72	78	66.71	44.66	63	3.6%	20.03 [5.34, 34.72]	
Orpington 1993	55	30	124	98	50	121	5.1%	-43.00 [-53.36, -32.64]	
Orpington 2000	32	30	152	30	40	149	6.2%	2.00 [-6.00, 10.00]	
Perth 1997	24	30	29	26.7	30	30	3.4%	-2.70 [-18.01, 12.61]	
Svendborg 1995	12	22	31	23	34	34	3.9%	-11.00 [-24.81, 2.81]	
Tampere 1993	13	30	98	15	38	113	5.7%	-2.00 [-11.19, 7.19]	
Trondheim 1991	75	114.8	102	123	145.8	104	0.9%	-48.00 [-83.80, -12.20]	
Subtotal (95% CI)			1944			1831	93.4%	-4.76 [-8.46, -1.05]	
Heterogeneity: $\text{Tau}^2 = 36.88$; $\text{Chi}^2 = 150.03$, $\text{df} = 16$ ($P < 0.00001$); $I^2 = 89\%$ Test for overall effect: $Z = 2.52$ ($P = 0.01$)									
1.5.2 Mixed rehabilitation ward									
Helsinki 1995	23.6	38.8	121	30.5	70.6	122	3.7%	-6.90 [-21.21, 7.41]	
Kuopio 1985	162.5	125	42	129.5	119	35	0.4%	33.00 [-21.62, 87.62]	
Newcastle 1993	52	45	34	41	34	33	2.5%	11.00 [-8.06, 30.06]	
Subtotal (95% CI)			197			190	6.6%	3.85 [-13.49, 21.18]	
Heterogeneity: $\text{Tau}^2 = 100.29$; $\text{Chi}^2 = 3.55$, $\text{df} = 2$ ($P = 0.17$); $I^2 = 44\%$ Test for overall effect: $Z = 0.43$ ($P = 0.66$)									
Total (95% CI)			2141			2021	100.0%	-4.28 [-7.86, -0.71]	
Heterogeneity: $\text{Tau}^2 = 36.66$; $\text{Chi}^2 = 153.76$, $\text{df} = 19$ ($P < 0.00001$); $I^2 = 88\%$ Test for overall effect: $Z = 2.35$ ($P = 0.02$) Test for subgroup differences: $\text{Chi}^2 = 0.91$, $\text{df} = 1$ ($P = 0.34$), $I^2 = 0\%$									



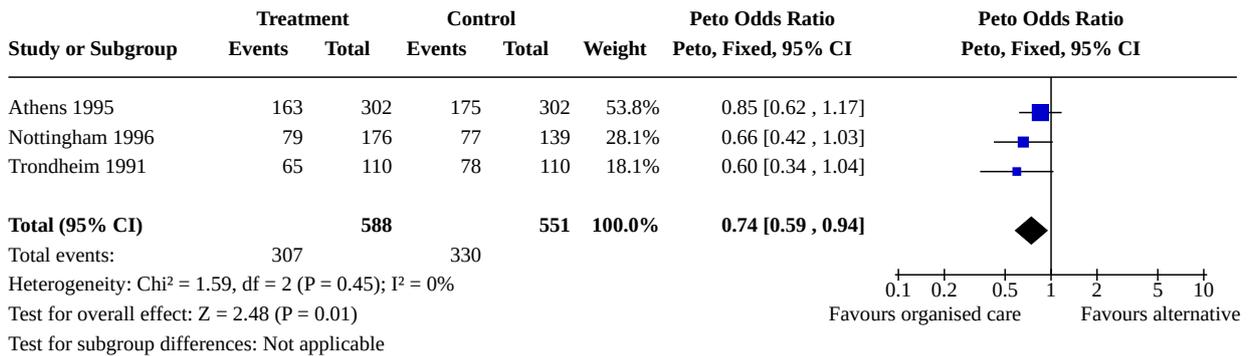
Analysis 1.6. Comparison 1: Organised stroke care versus alternative service, Outcome 6: Length of stay (days) in a hospital or hospital plus institution



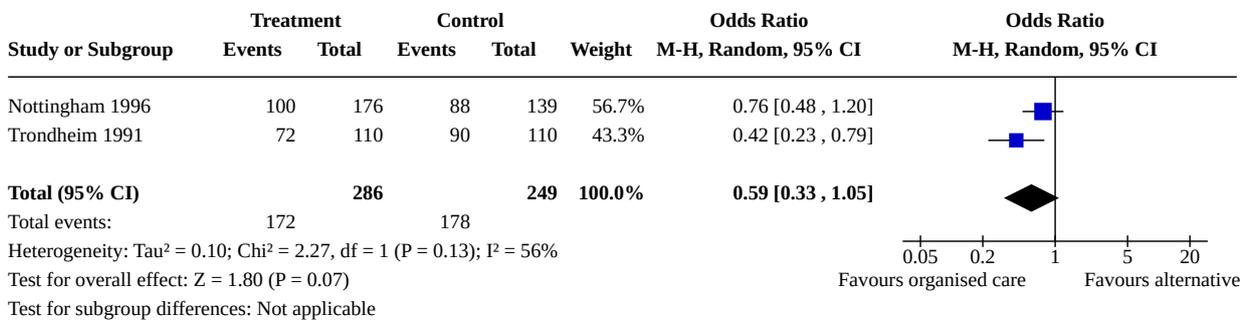
Analysis 1.7. Comparison 1: Organised stroke care versus alternative service, Outcome 7: Poor outcome at 5-year follow-up



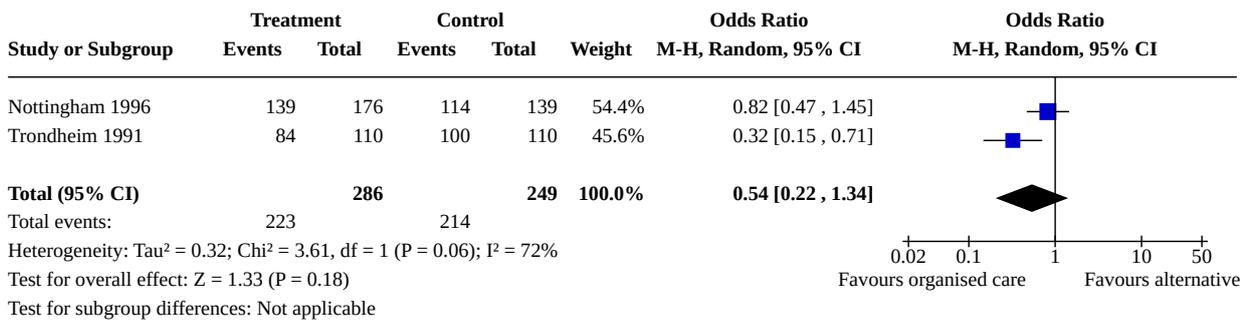
Analysis 1.8. Comparison 1: Organised stroke care versus alternative service, Outcome 8: Death at 5-year follow-up



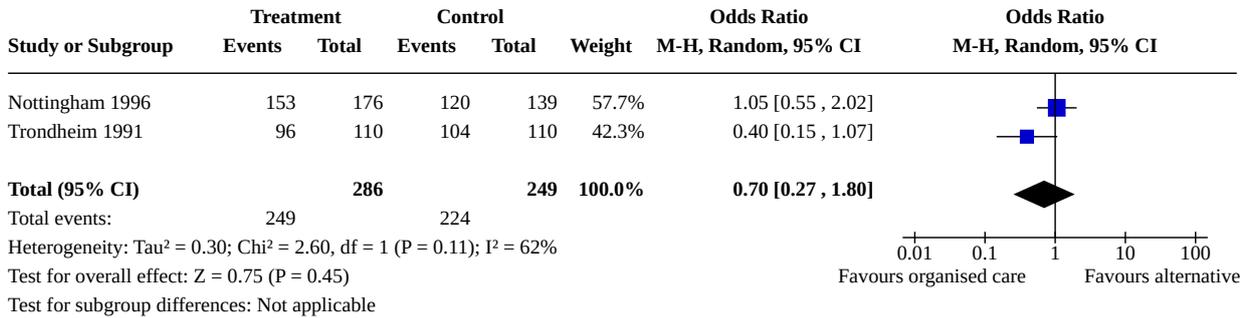
Analysis 1.9. Comparison 1: Organised stroke care versus alternative service, Outcome 9: Death or institutional care at 5-year follow-up



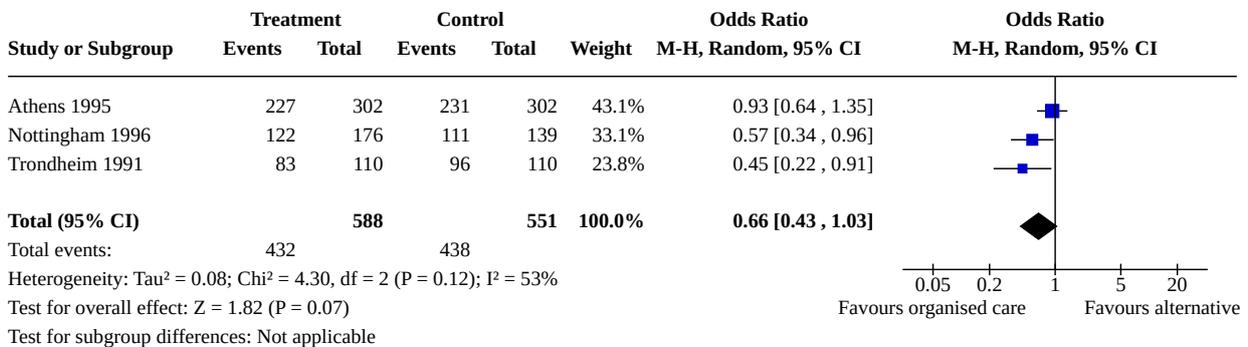
Analysis 1.10. Comparison 1: Organised stroke care versus alternative service, Outcome 10: Death or dependency at 5-year follow-up



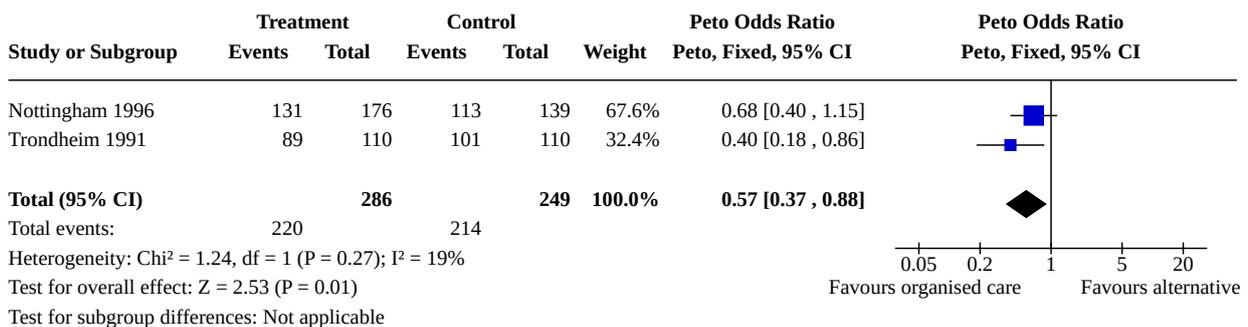
Analysis 1.11. Comparison 1: Organised stroke care versus alternative service, Outcome 11: Poor outcome at 10-year follow-up



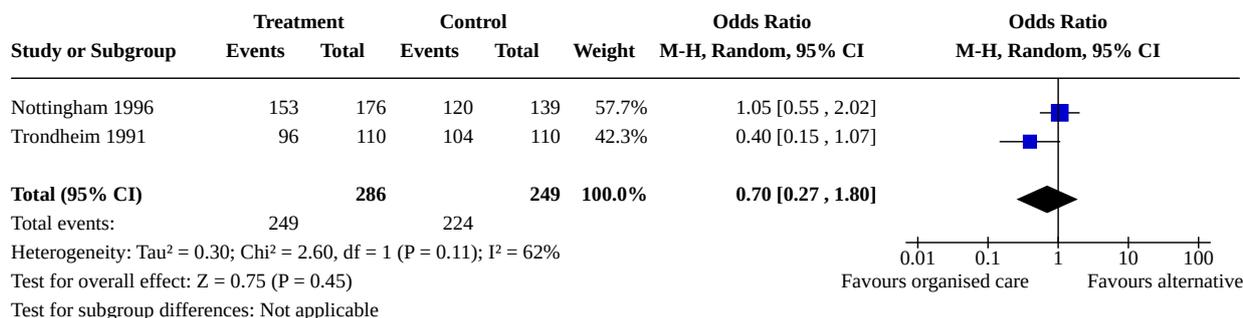
Analysis 1.12. Comparison 1: Organised stroke care versus alternative service, Outcome 12: Death at 10-year follow-up



Analysis 1.13. Comparison 1: Organised stroke care versus alternative service, Outcome 13: Death or institutional care at 10-year follow-up



Analysis 1.14. Comparison 1: Organised stroke care versus alternative service, Outcome 14: Death or dependency at 10-year follow-up

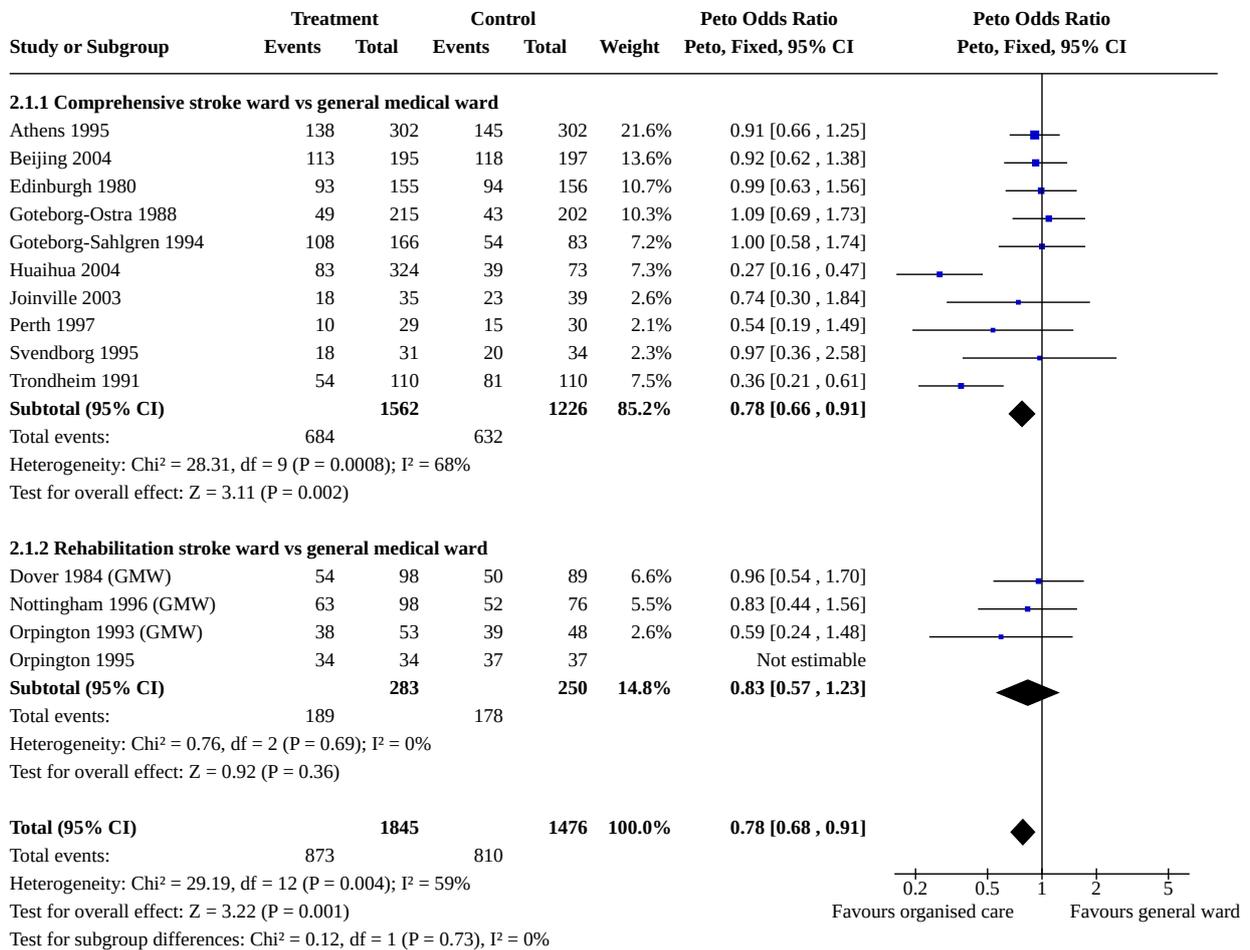


Comparison 2. Stroke ward versus general medical ward

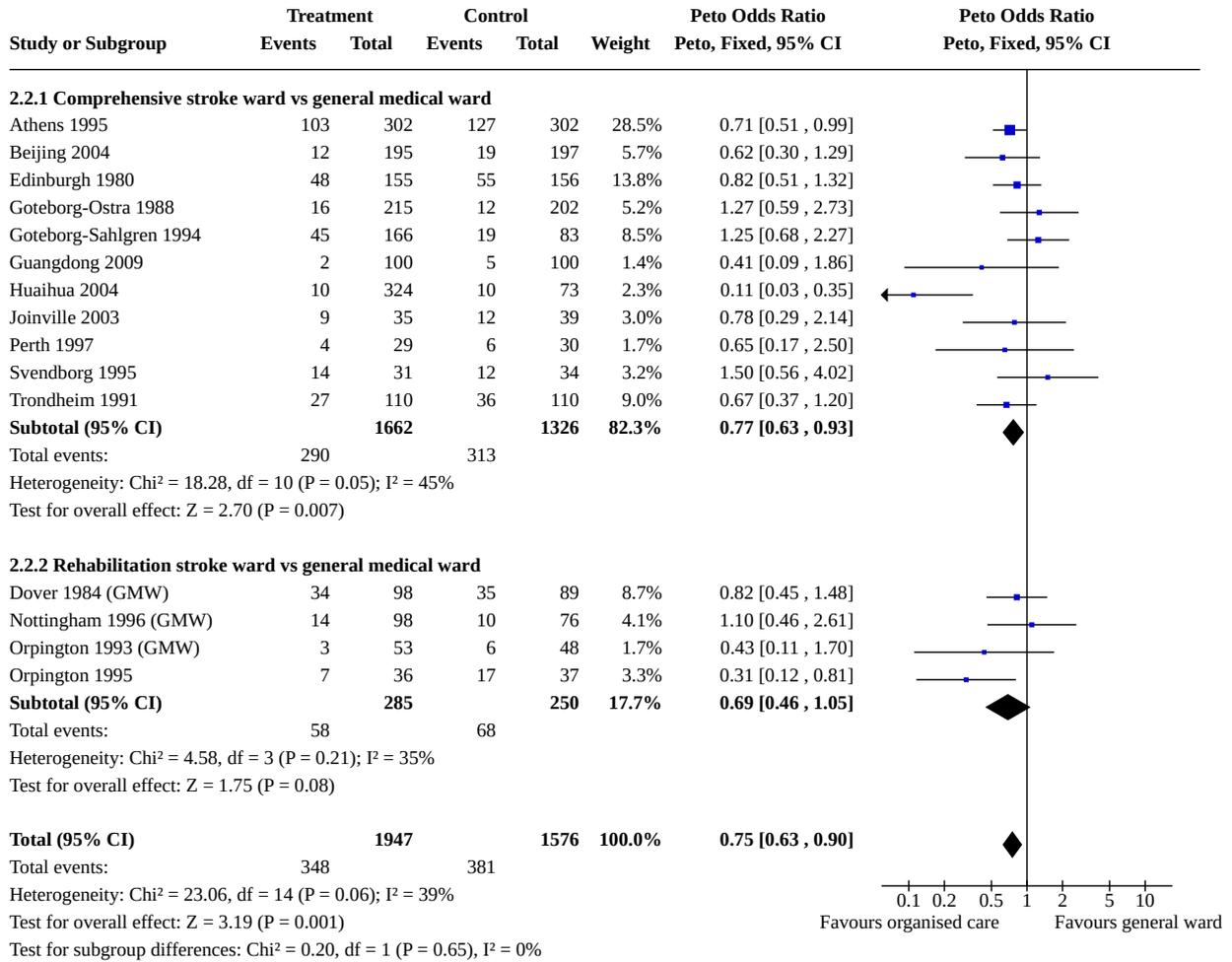
Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
2.1 Poor outcome by the end of scheduled follow-up	14	3321	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.78 [0.68, 0.91]
2.1.1 Comprehensive stroke ward vs general medical ward	10	2788	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.78 [0.66, 0.91]
2.1.2 Rehabilitation stroke ward vs general medical ward	4	533	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.83 [0.57, 1.23]
2.2 Death by the end of scheduled follow-up	15	3523	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.75 [0.63, 0.90]
2.2.1 Comprehensive stroke ward vs general medical ward	11	2988	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.77 [0.63, 0.93]
2.2.2 Rehabilitation stroke ward vs general medical ward	4	535	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.69 [0.46, 1.05]
2.3 Death or institutional care by the end of scheduled follow-up	13	2924	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.74 [0.63, 0.87]
2.3.1 Comprehensive stroke ward vs general medical ward	9	2391	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.74 [0.62, 0.88]
2.3.2 Rehabilitation stroke ward vs general medical ward	4	533	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.76 [0.52, 1.09]
2.4 Death or dependency by the end of scheduled follow-up	12	2839	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.75 [0.64, 0.88]
2.4.1 Comprehensive stroke ward vs general medical ward	8	2306	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.73 [0.62, 0.87]
2.4.2 Rehabilitation stroke ward vs general medical ward	4	533	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.83 [0.57, 1.23]

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
2.5 Length of stay (days) in a hospital or institution	10	2547	Mean Difference (IV, Random, 95% CI)	-2.19 [-5.19, 0.82]
2.5.1 Comprehensive stroke ward vs general medical ward	9	2373	Mean Difference (IV, Random, 95% CI)	-2.79 [-5.68, 0.10]
2.5.2 Rehabilitation stroke ward vs general medical ward	1	174	Mean Difference (IV, Random, 95% CI)	16.34 [2.82, 29.86]

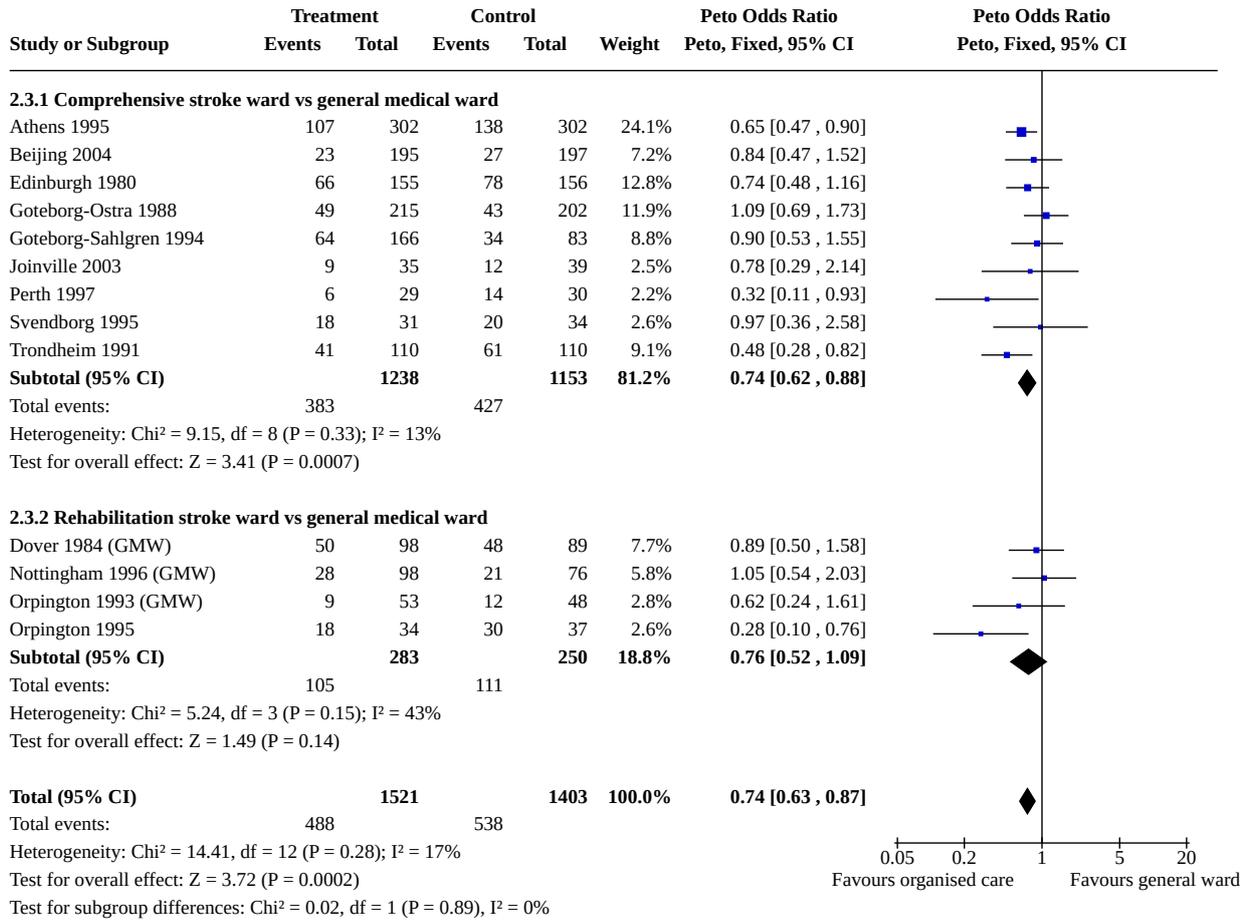
Analysis 2.1. Comparison 2: Stroke ward versus general medical ward, Outcome 1: Poor outcome by the end of scheduled follow-up



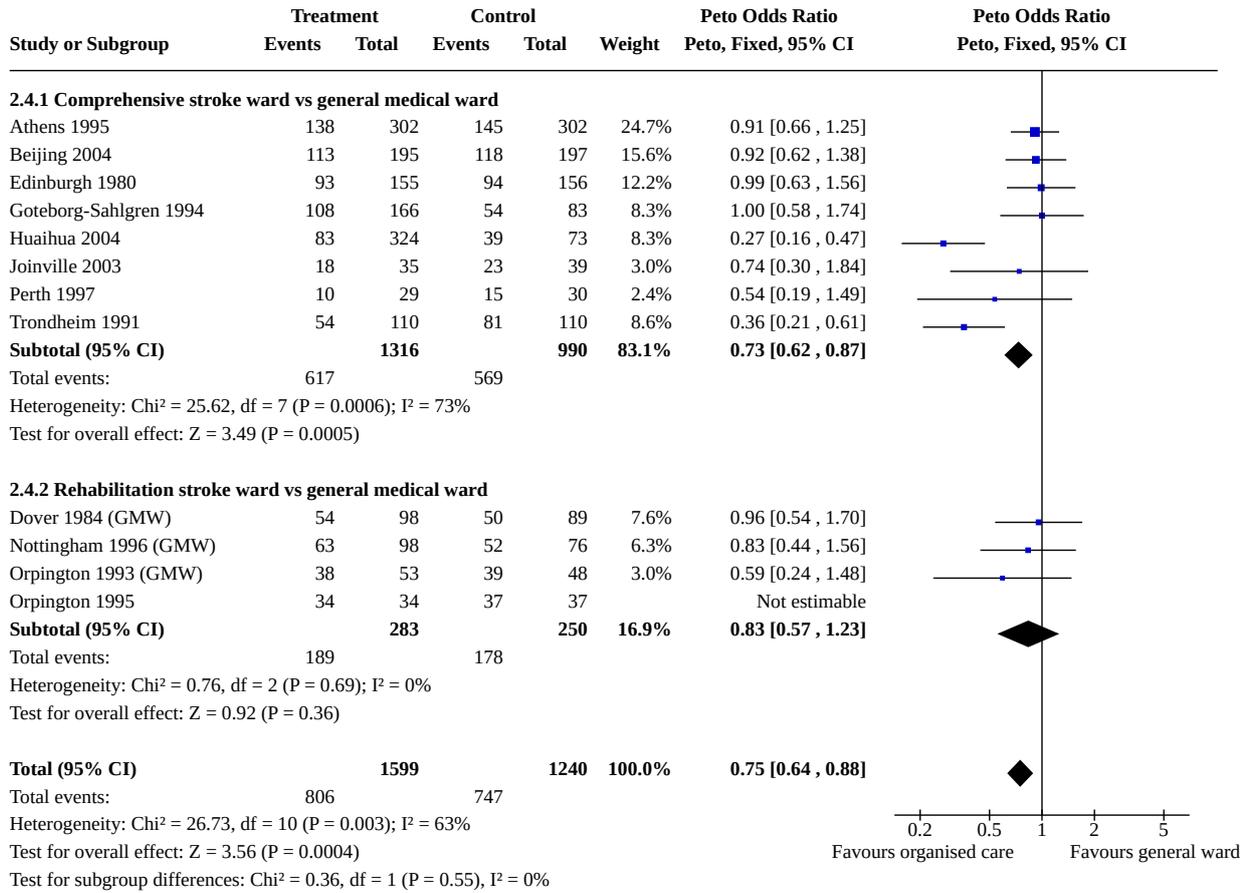
Analysis 2.2. Comparison 2: Stroke ward versus general medical ward, Outcome 2: Death by the end of scheduled follow-up



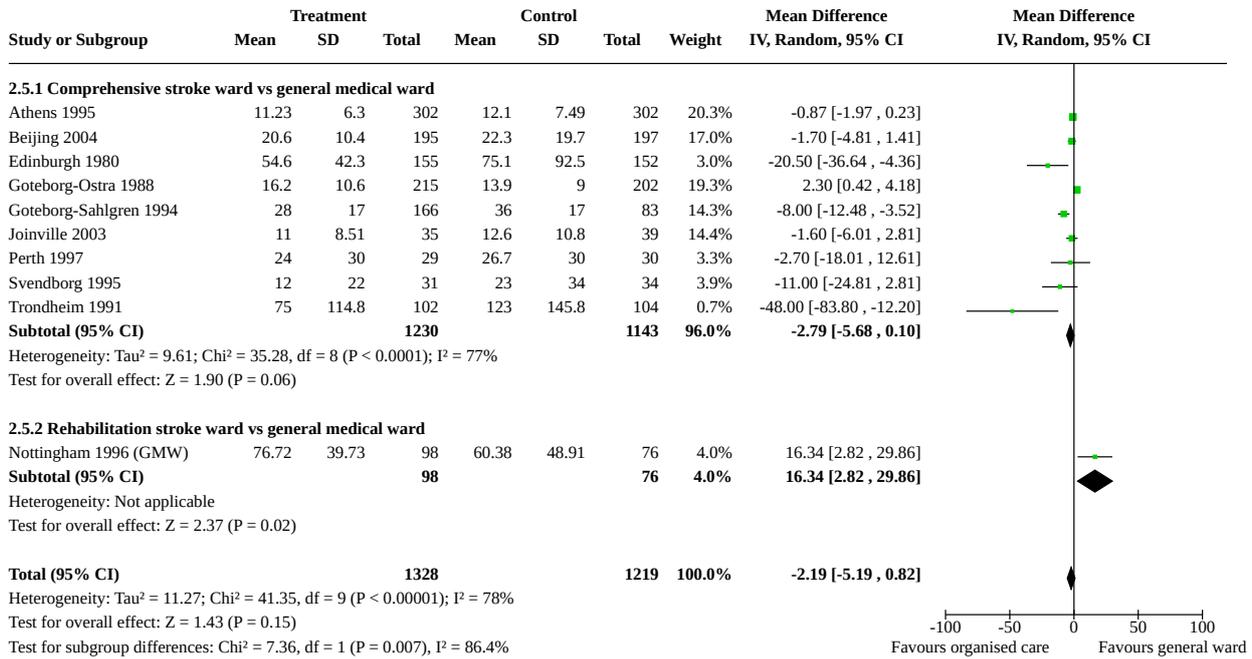
**Analysis 2.3. Comparison 2: Stroke ward versus general medical ward,
Outcome 3: Death or institutional care by the end of scheduled follow-up**



**Analysis 2.4. Comparison 2: Stroke ward versus general medical ward,
Outcome 4: Death or dependency by the end of scheduled follow-up**



Analysis 2.5. Comparison 2: Stroke ward versus general medical ward, Outcome 5: Length of stay (days) in a hospital or institution

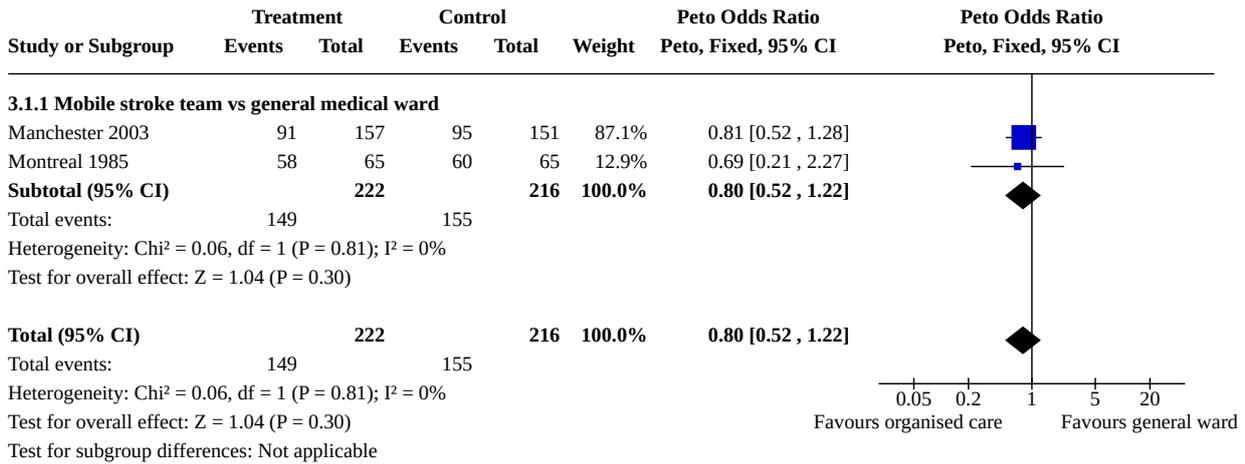


Comparison 3. Mobile stroke team versus general medical ward

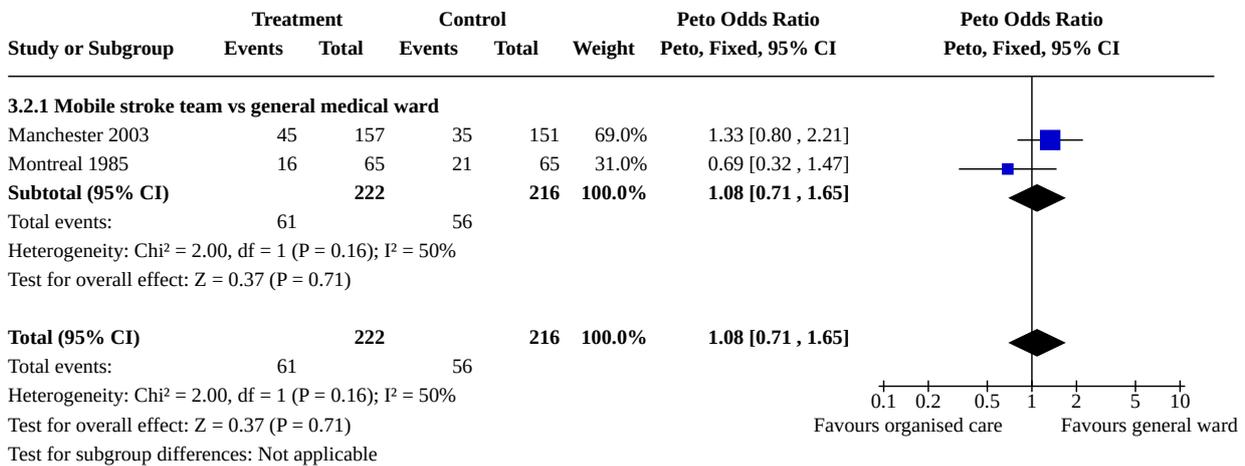
Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
3.1 Poor outcome by the end of scheduled follow-up	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.80 [0.52, 1.22]
3.1.1 Mobile stroke team vs general medical ward	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.80 [0.52, 1.22]
3.2 Death by the end of scheduled follow-up	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	1.08 [0.71, 1.65]
3.2.1 Mobile stroke team vs general medical ward	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	1.08 [0.71, 1.65]
3.3 Death or institutional care by the end of scheduled follow-up	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	1.27 [0.84, 1.93]
3.3.1 Mobile stroke team vs general medical ward	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	1.27 [0.84, 1.93]
3.4 Death or dependency by the end of scheduled follow-up	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.80 [0.52, 1.22]
3.4.1 Mobile stroke team vs general medical ward	2	438	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.80 [0.52, 1.22]
3.5 Length of stay (days) in a hospital or institution	0	0	Std. Mean Difference (IV, Random, 95% CI)	Not estimable

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
3.5.1 Mobile stroke team vs general medical ward	0	0	Std. Mean Difference (IV, Random, 95% CI)	Not estimable

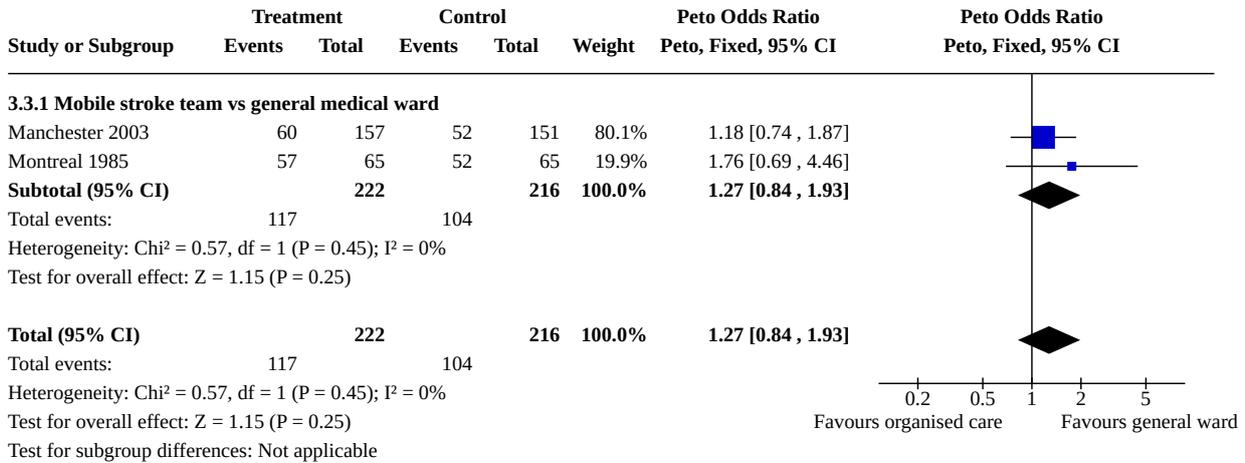
Analysis 3.1. Comparison 3: Mobile stroke team versus general medical ward, Outcome 1: Poor outcome by the end of scheduled follow-up



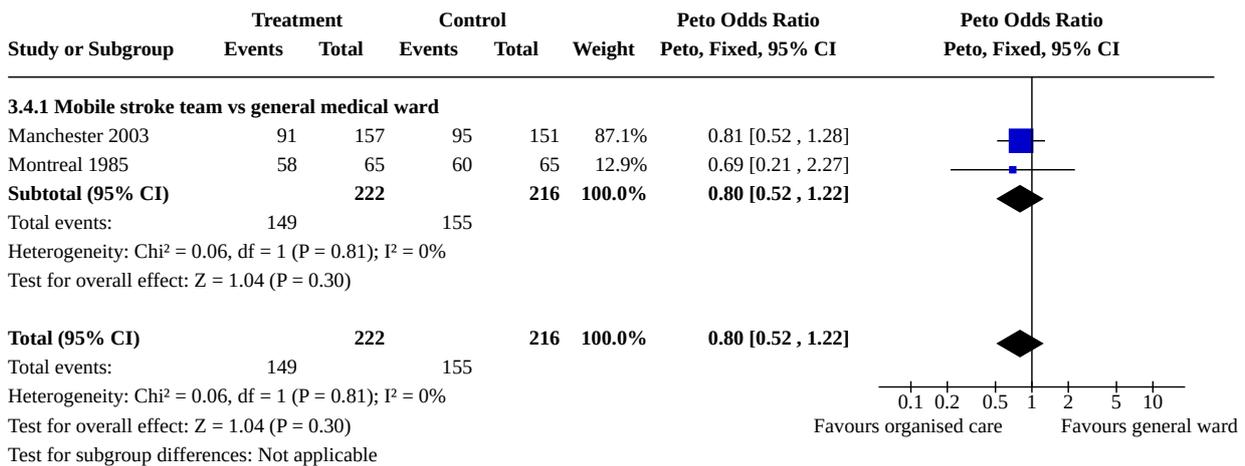
Analysis 3.2. Comparison 3: Mobile stroke team versus general medical ward, Outcome 2: Death by the end of scheduled follow-up



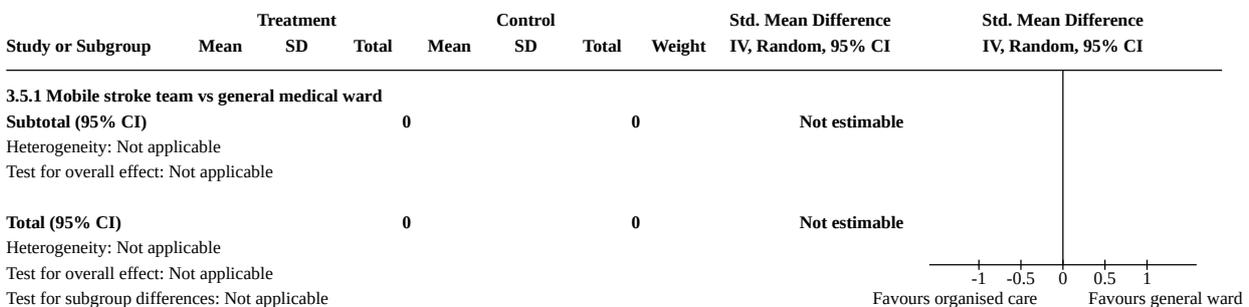
Analysis 3.3. Comparison 3: Mobile stroke team versus general medical ward, Outcome 3: Death or institutional care by the end of scheduled follow-up



Analysis 3.4. Comparison 3: Mobile stroke team versus general medical ward, Outcome 4: Death or dependency by the end of scheduled follow-up



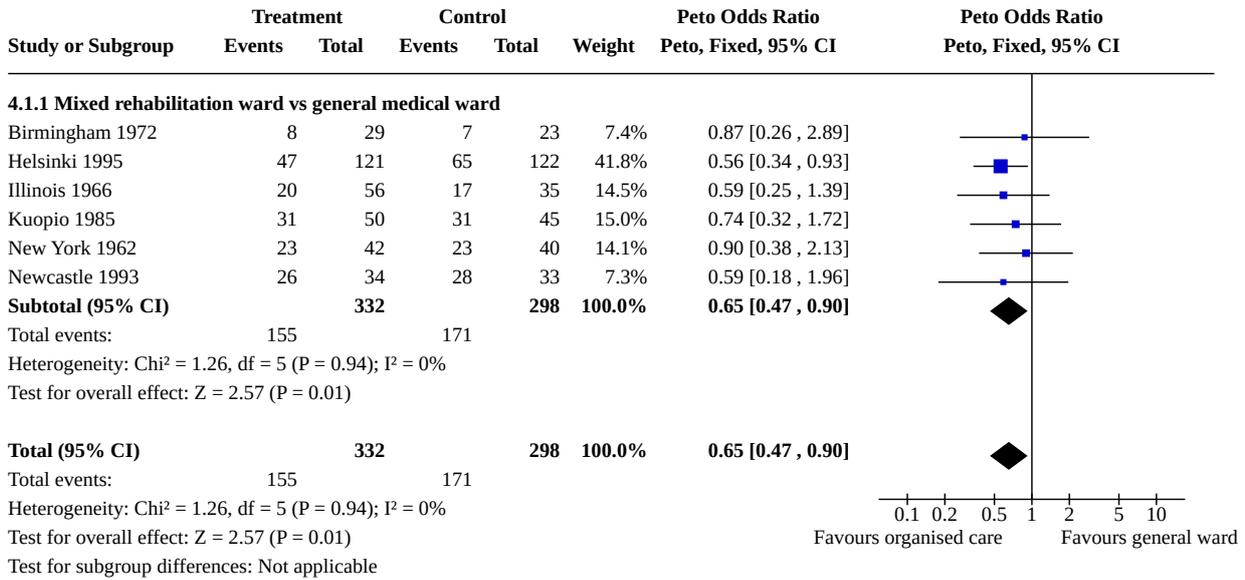
Analysis 3.5. Comparison 3: Mobile stroke team versus general medical ward, Outcome 5: Length of stay (days) in a hospital or institution



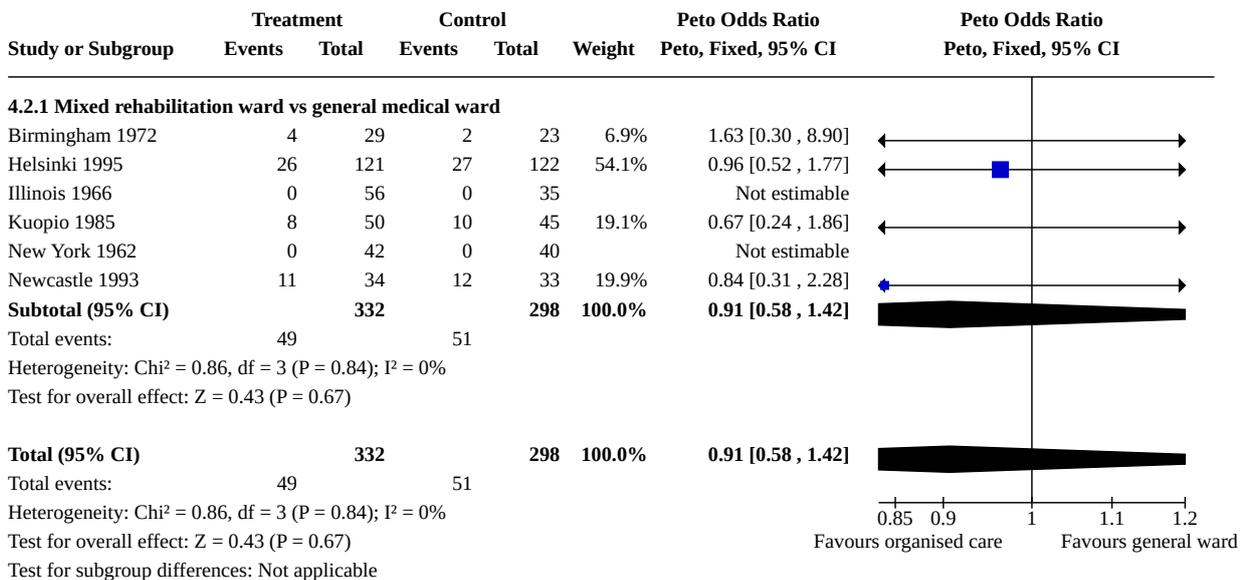
Comparison 4. Mixed rehabilitation ward versus general medical ward

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
4.1 Poor outcome by the end of scheduled follow-up	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.65 [0.47, 0.90]
4.1.1 Mixed rehabilitation ward vs general medical ward	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.65 [0.47, 0.90]
4.2 Death by the end of scheduled follow-up	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.91 [0.58, 1.42]
4.2.1 Mixed rehabilitation ward vs general medical ward	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.91 [0.58, 1.42]
4.3 Death or institutional care by the end of scheduled follow-up	5	578	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.71 [0.51, 0.99]
4.3.1 Mixed rehabilitation ward vs general medical ward	5	578	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.71 [0.51, 0.99]
4.4 Death or dependency by the end of scheduled follow-up	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.65 [0.47, 0.90]
4.4.1 Mixed rehabilitation ward vs general medical ward	6	630	Peto Odds Ratio (Peto, Fixed, 95% CI)	0.65 [0.47, 0.90]
4.5 Length of stay (days) in a hospital or institution	3	387	Mean Difference (IV, Random, 95% CI)	3.85 [-13.49, 21.18]
4.5.1 Mixed rehabilitation ward vs general ward	3	387	Mean Difference (IV, Random, 95% CI)	3.85 [-13.49, 21.18]

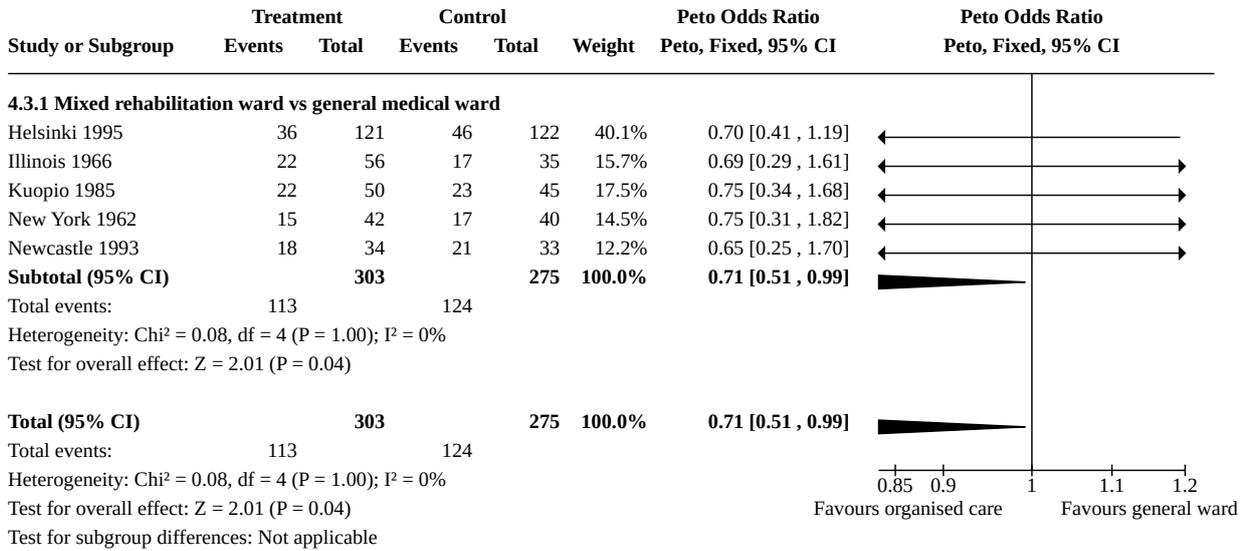
Analysis 4.1. Comparison 4: Mixed rehabilitation ward versus general medical ward, Outcome 1: Poor outcome by the end of scheduled follow-up



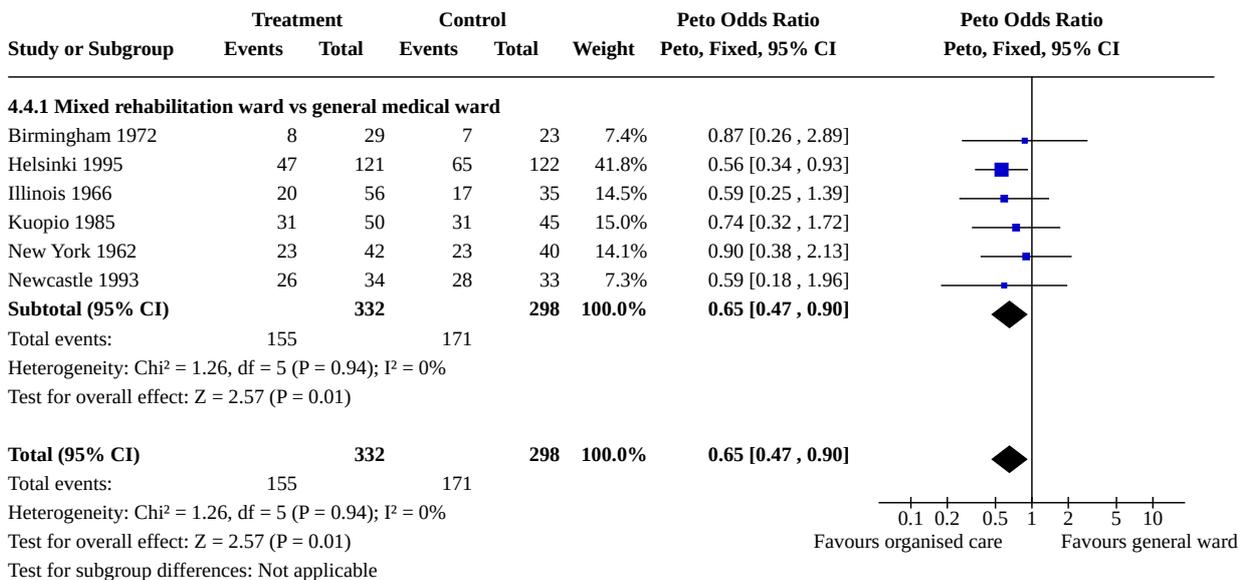
Analysis 4.2. Comparison 4: Mixed rehabilitation ward versus general medical ward, Outcome 2: Death by the end of scheduled follow-up



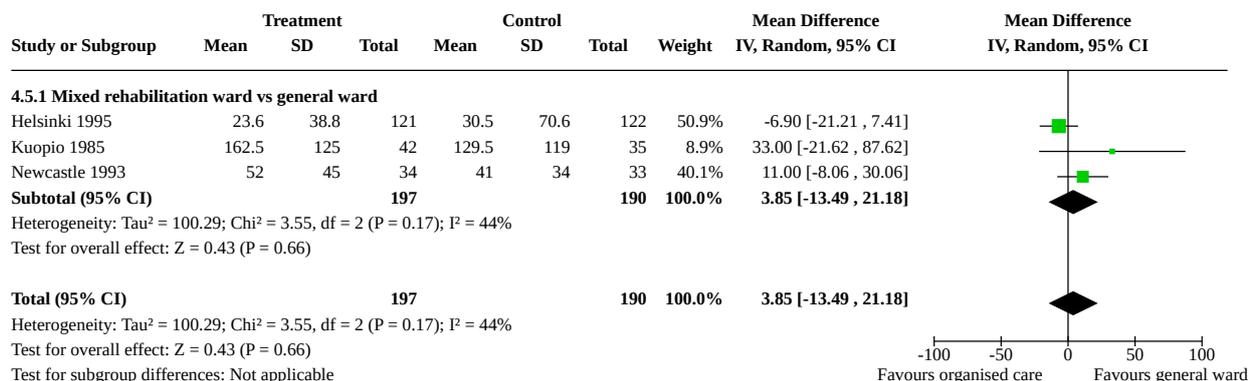
Analysis 4.3. Comparison 4: Mixed rehabilitation ward versus general medical ward, Outcome 3: Death or institutional care by the end of scheduled follow-up



Analysis 4.4. Comparison 4: Mixed rehabilitation ward versus general medical ward, Outcome 4: Death or dependency by the end of scheduled follow-up



Analysis 4.5. Comparison 4: Mixed rehabilitation ward versus general medical ward, Outcome 5: Length of stay (days) in a hospital or institution

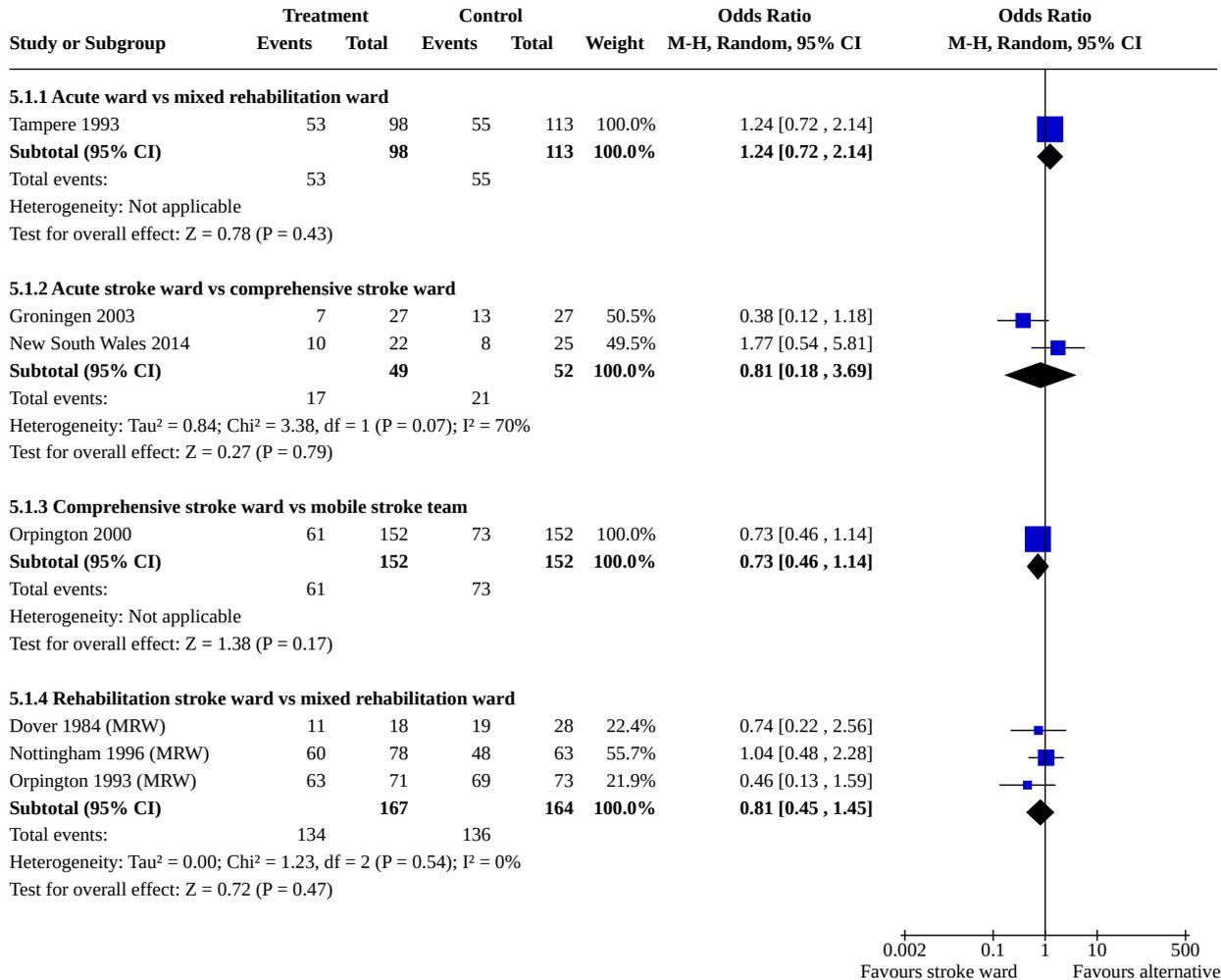


Comparison 5. Different systems of organised care: stroke ward versus alternative organised care

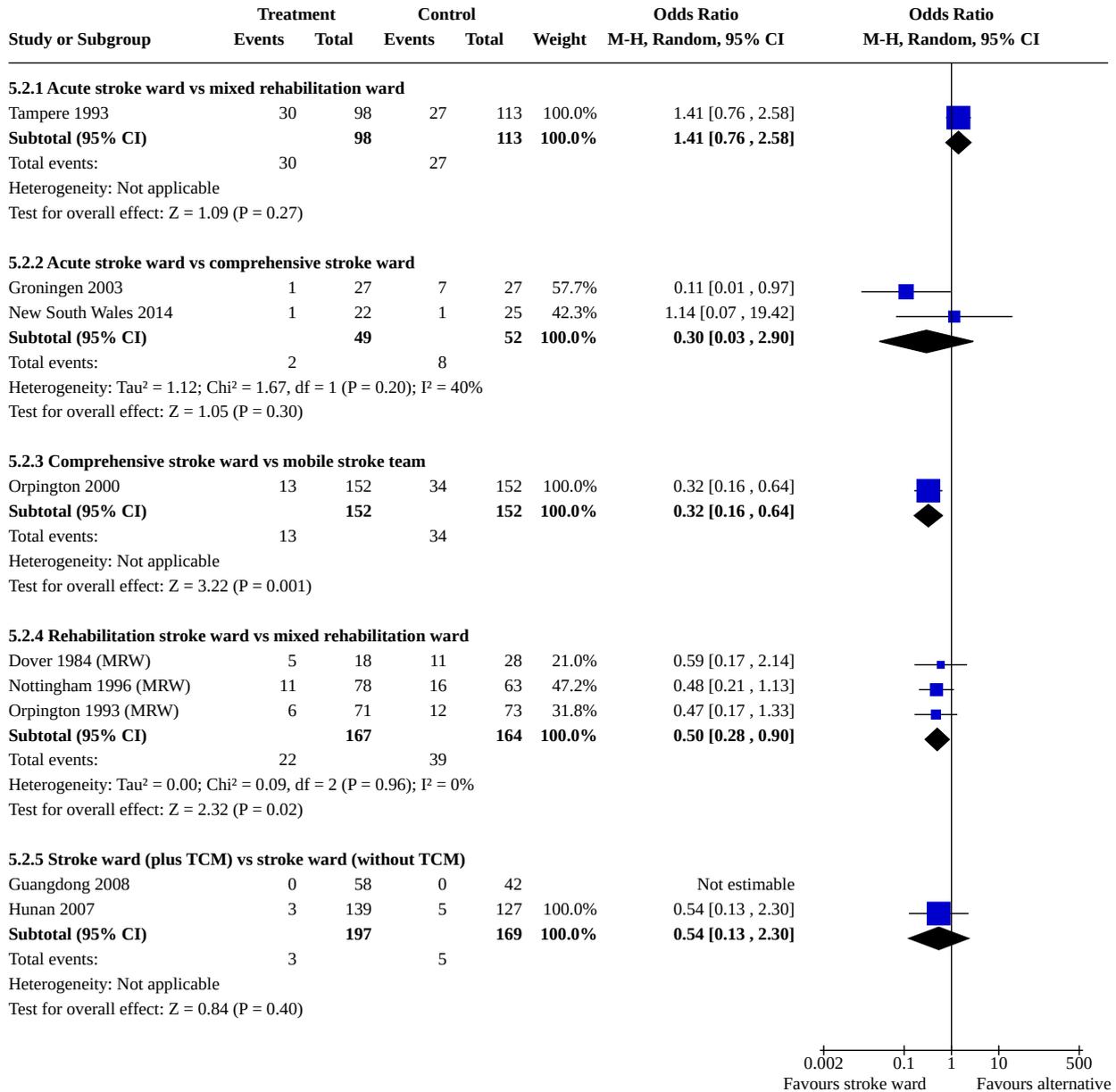
Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
5.1 Poor outcome by the end of scheduled follow-up	7		Odds Ratio (M-H, Random, 95% CI)	Subtotals only
5.1.1 Acute ward vs mixed rehabilitation ward	1	211	Odds Ratio (M-H, Random, 95% CI)	1.24 [0.72, 2.14]
5.1.2 Acute stroke ward vs comprehensive stroke ward	2	101	Odds Ratio (M-H, Random, 95% CI)	0.81 [0.18, 3.69]
5.1.3 Comprehensive stroke ward vs mobile stroke team	1	304	Odds Ratio (M-H, Random, 95% CI)	0.73 [0.46, 1.14]
5.1.4 Rehabilitation stroke ward vs mixed rehabilitation ward	3	331	Odds Ratio (M-H, Random, 95% CI)	0.81 [0.45, 1.45]
5.2 Death by the end of scheduled follow-up	9		Odds Ratio (M-H, Random, 95% CI)	Subtotals only
5.2.1 Acute stroke ward vs mixed rehabilitation ward	1	211	Odds Ratio (M-H, Random, 95% CI)	1.41 [0.76, 2.58]
5.2.2 Acute stroke ward vs comprehensive stroke ward	2	101	Odds Ratio (M-H, Random, 95% CI)	0.30 [0.03, 2.90]
5.2.3 Comprehensive stroke ward vs mobile stroke team	1	304	Odds Ratio (M-H, Random, 95% CI)	0.32 [0.16, 0.64]
5.2.4 Rehabilitation stroke ward vs mixed rehabilitation ward	3	331	Odds Ratio (M-H, Random, 95% CI)	0.50 [0.28, 0.90]
5.2.5 Stroke ward (plus TCM) vs stroke ward (without TCM)	2	366	Odds Ratio (M-H, Random, 95% CI)	0.54 [0.13, 2.30]
5.3 Death or institutional care by the end of scheduled follow-up	7		Odds Ratio (M-H, Random, 95% CI)	Subtotals only

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
5.3.1 Acute ward vs mixed rehabilitation ward	1	211	Odds Ratio (M-H, Random, 95% CI)	1.32 [0.76, 2.30]
5.3.2 Acute stroke ward vs comprehensive stroke ward	2	101	Odds Ratio (M-H, Random, 95% CI)	1.28 [0.14, 11.31]
5.3.3 Comprehensive stroke ward vs mobile stroke team	1	304	Odds Ratio (M-H, Random, 95% CI)	0.38 [0.21, 0.68]
5.3.4 Rehabilitation stroke ward vs mixed rehabilitation ward	3	331	Odds Ratio (M-H, Random, 95% CI)	0.70 [0.45, 1.09]
5.4 Death or dependency by the end of scheduled follow-up	7		Odds Ratio (M-H, Random, 95% CI)	Subtotals only
5.4.1 Acute ward vs mixed rehabilitation ward	1	211	Odds Ratio (M-H, Random, 95% CI)	1.24 [0.72, 2.14]
5.4.2 Acute stroke ward vs comprehensive stroke ward	2	101	Odds Ratio (M-H, Random, 95% CI)	0.81 [0.18, 3.69]
5.4.3 Comprehensive stroke ward vs mobile stroke team	1	304	Odds Ratio (M-H, Random, 95% CI)	0.73 [0.46, 1.14]
5.4.4 Rehabilitation stroke ward vs mixed rehabilitation ward	3	331	Odds Ratio (M-H, Random, 95% CI)	0.81 [0.45, 1.45]
5.5 Length of stay (days) in a hospital or institution	7		Mean Difference (IV, Random, 95% CI)	Subtotals only
5.5.1 Acute stroke ward vs mixed rehabilitation ward	1	211	Mean Difference (IV, Random, 95% CI)	-2.00 [-11.19, 7.19]
5.5.2 Acute (±rehabilitation) stroke ward vs comprehensive stroke ward	2	101	Mean Difference (IV, Random, 95% CI)	-2.89 [-20.10, 14.33]
5.5.3 Comprehensive stroke ward vs mobile stroke team	1	304	Mean Difference (IV, Random, 95% CI)	2.50 [-5.42, 10.42]
5.5.4 Rehabilitation stroke ward vs mixed rehabilitation ward	3	331	Mean Difference (IV, Random, 95% CI)	15.80 [-45.71, 77.31]

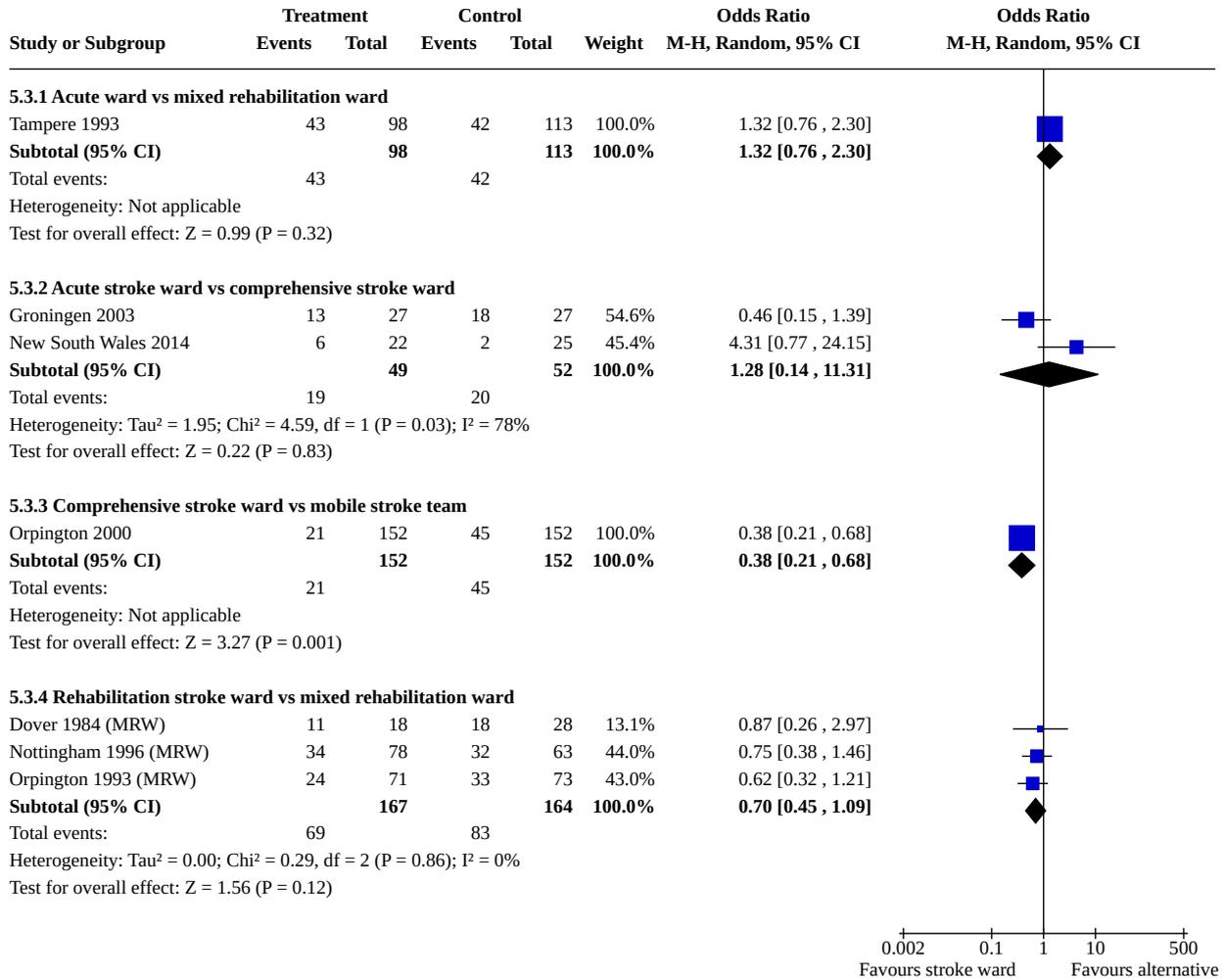
Analysis 5.1. Comparison 5: Different systems of organised care: stroke ward versus alternative organised care, Outcome 1: Poor outcome by the end of scheduled follow-up



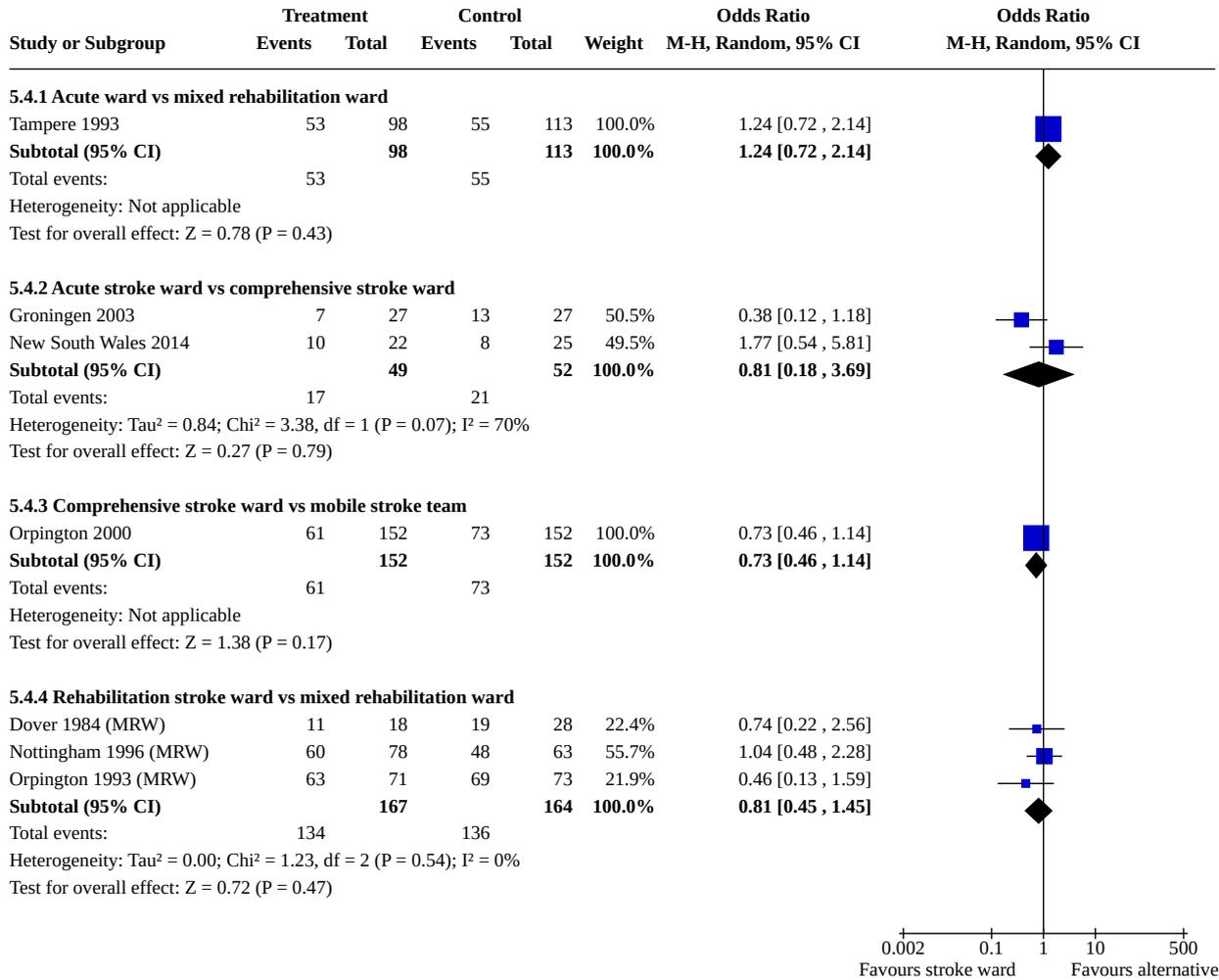
Analysis 5.2. Comparison 5: Different systems of organised care: stroke ward versus alternative organised care, Outcome 2: Death by the end of scheduled follow-up



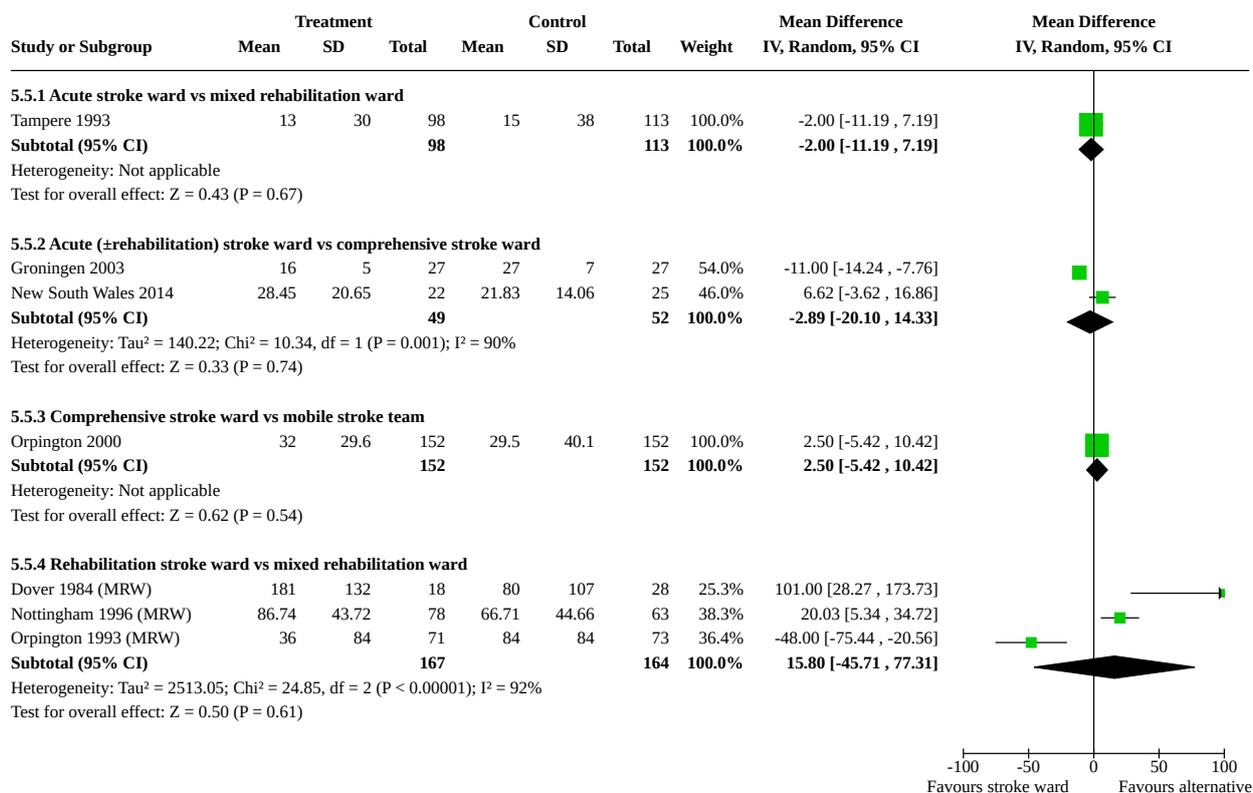
Analysis 5.3. Comparison 5: Different systems of organised care: stroke ward versus alternative organised care, Outcome 3: Death or institutional care by the end of scheduled follow-up



Analysis 5.4. Comparison 5: Different systems of organised care: stroke ward versus alternative organised care, Outcome 4: Death or dependency by the end of scheduled follow-up



Analysis 5.5. Comparison 5: Different systems of organised care: stroke ward versus alternative organised care, Outcome 5: Length of stay (days) in a hospital or institution



ADDITIONAL TABLES

Table 1. Typical characteristics of different models of organised stroke care

Name	Structure	Patients	Service type	Admission	Discharge	Features
Stroke ward	Ward	Stroke	Various (see below)	Various (see below)	Various (see below)	Various (see below)
Acute stroke ward	Ward	Stroke	Acute	Acute (hours)	Days	Close physiological monitoring, often followed by care in separate rehabilitation ward if required
Comprehensive stroke ward ^a	Ward	Stroke	Acute, rehabilitation	Acute (hours)	Days to weeks	Acute care and rehabilitation; conventional staffing levels
Rehabilitation stroke ward	Ward	Stroke	Rehabilitation	Delayed (days)	Weeks	Focus on rehabilitation
Mobile stroke team	Mobile team	Stroke	Mobile stroke team	Variable	Days to weeks	Peripatetic care to patients in general wards; medical and rehabilitation advice
Mixed rehabilitation ward	Ward	Mixed	Rehabilitation	Variable	Weeks	Mixed patient group; focus on rehabilitation

Table 1. Typical characteristics of different models of organised stroke care (Continued)

Intensive care	Ward	Mixed	Acute, intensive	Acute (hours)	Days	High nurse staffing; life support facilities
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^aTwo trials tested a comprehensive stroke ward incorporating traditional Chinese medicine (TCM) such as acupuncture.

Table 2. Service comparisons in standard analyses

Trials	Participants	Index (stroke unit) care	Conventional care	Reference
15	3521	Stroke ward	General medical ward	Athens 1995, Beijing 2004, Dover 1984 (GMW), Edinburgh 1980, Goteborg-Ostra 1988, Goteborg-Sahlgren 1994, Guangdong 2009, Huaihua 2004, Joinville 2003, Nottingham 1996 (GMW), Orpington 1993 (GMW), Orpington 1995, Perth 1997, Svendborg 1995, Trondheim 1991
6	630	Mixed rehabilitation ward	General medical ward	Birmingham 1972, Helsinki 1995, Illinois 1966, Kuopio 1985, New York 1962; Newcastle 1993
2	438	Mobile stroke team (peripatetic care)	General medical ward	Manchester 2003, Montreal 1985
4	542	Stroke ward	Mixed rehabilitation ward	Dover 1984 (MRW), Nottingham 1996 (MRW), Orpington 1993 (MRW), Tampere 1993
1	304	Stroke ward (comprehensive)	Mobile stroke team	Orpington 2000
1	54	Stroke ward (acute)	Stroke ward (comprehensive unit)	Groningen 2003
1	47	Stroke ward (comprehensive)	Stroke wards (acute+rehabilitation)	New South Wales 2014
2	366	Stroke ward (plus TCM)	Stroke ward (without TCM)	Guangdong 2008, Hunan 2007

TCM: traditional Chinese medicine.

Table 3. Service comparisons in network meta-analyses (NMAs)^a

Stroke ward		General medical ward	
15 trials (3521 participants)			
4 trials (542 participants)		6 trials (630 participants)	Mixed rehabilitation ward

Table 3. Service comparisons in network meta-analyses (NMAs)^a (Continued)

1 trial (304 participants)	2 trials (438 participants)	--	Mobile stroke team
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^aThe table shows the numbers of trials and participants for each service comparison (e.g. two trials (438 participants) featured a comparison of a mobile stroke team with a general medical ward).

Table 4. Inconsistency table for the network meta-analysis: poor outcome at the end of scheduled follow-up^a

Comparison	Number of studies	Log direct comparison OR	Log indirect comparison OR	Log difference (95% CI) between direct and indirect comparisons	P value of difference between direct and indirect comparisons
Mixed rehabilitation ward vs GMW	6	-0.413	-0.266	-0.147 (-0.776 to 0.483)	0.65
Mobile stroke team vs GMW	2	-0.238	0.032	-0.270 (-1.111 to 0.571)	0.53
Stroke ward vs GMW	13	-0.273	-0.478	0.205 (-0.317 to 0.728)	0.44
Mixed rehabilitation ward vs mobile stroke team	0	N/A	-0.237	N/A	N/A
Mixed rehabilitation ward vs stroke ward	4	0.022	-0.124	0.146 (-0.483 to 0.776)	0.65
Mobile stroke team vs stroke ward	1	0.321	0.051	0.270 (-0.571 to 1.111)	0.53

^aThe table shows the following for each service comparison: number of trials, log direct and indirect comparisons, difference between the two estimates, and P value of that difference. There was no significant inconsistency between direct and indirect estimates.

CI: confidence interval.

GMW: general medical ward.

N/A: not applicable.

OR: odds ratio.

Table 5. Inconsistency table for the network meta-analysis: death at the end of scheduled follow-up^a

Comparison	Number of studies	Log Direct comparison OR	Log Indirect comparison OR	Log difference (95% CI) between direct and indirect comparisons	P value of difference between direct and indirect comparisons
Mixed rehabilitation ward vs GMW	6	-0.101	0.617	-0.719	0.17

Table 5. Inconsistency table for the network meta-analysis: death at the end of scheduled follow-up^a (Continued)
 (-1.741 to 0.303)

Mobile stroke team vs GMW	2	0.003	0.678	-0.675 (-1.997 to 0.646)	0.32
Stroke ward vs GMW	15	-0.381	-1.132	0.750 (-0.084 to 1.586)	0.08
Mixed rehabilitation ward vs mobile stroke team	0	N/A	-0.023	N/A	N/A
Mixed rehabilitation ward vs stroke ward	4	1.037	0.318	0.719 (-0.303 to 1.741)	0.17
Mobile stroke team vs stroke ward	1	1.125	0.449	0.675 (-0.646 to 1.997)	0.32

^aThe table shows the following for each service comparison: number of trials, log direct and indirect comparisons, difference between the two estimates, and P value of that difference. There was no significant inconsistency between direct and indirect estimates.

CI: confidence interval.

GMW: general medical ward.

N/A: not applicable.

OR: odds ratio.

Table 6. Inconsistency table for the network meta-analysis: death or institutional care at the end of scheduled follow-up^a

Comparison	Number of studies	Log Direct comparison OR	Log Indirect comparison OR	Log difference (95% CI) between direct and indirect comparisons	P value of difference between direct and indirect comparisons
Mixed rehabilitation ward vs GMW	5	-0.347	-0.215	-0.132 (-0.640 to 0.375)	0.61
Mobile stroke team vs GMW	2	0.242	0.649	-0.406 (-1.136 to 0.322)	0.27
Stroke ward vs GMW	13	-0.299	-0.537	0.237 (-0.193 to 0.668)	0.28
Mixed rehabilitation ward vs mobile stroke team	0	NA	-0.665	N/A	N/A
Mixed rehabilitation ward vs stroke ward	4	0.105	-0.027	0.132 (-0.375 to 0.640)	0.61
Mobile stroke team vs stroke ward	1	0.964	0.557	0.406	0.27

Table 6. Inconsistency table for the network meta-analysis: death or institutional care at the end of scheduled follow-up^a (Continued)

(-0.322 to 1.136)

^aThe table shows the following for each service comparison: number of trials, log direct and indirect comparisons, difference between the two estimates, and P value of that difference. There was no significant inconsistency between direct and indirect estimates.

CI: confidence interval.

GMW: general medical ward.

N/A: not applicable.

OR: odds ratio.

Table 7. Inconsistency table for the network meta-analysis: death or dependency at the end of scheduled follow-up^a

Comparison	Number of studies	Log Direct comparison OR	Log Indirect comparison OR	Log difference (95% CI) between direct and indirect comparisons	P value of difference between direct and indirect comparisons
Mixed rehabilitation ward vs GMW	6	-0.413	-0.317	0.096 (-0.731 to 0.540)	0.77
Mobile stroke team vs GMW	2	-0.238	-0.016	-0.220 (-1.070 to 0.620)	0.61
Stroke ward vs GMW	12	-0.325	-0.473	0.150 (-0.380 to 0.680)	0.57
Mixed rehabilitation ward vs mobile stroke team	0	NA	-0.234	N/A	N/A
Mixed rehabilitation ward vs stroke ward	4	0.022	-0.073	0.100 (-0.540 to 0.735)	0.77
Mobile stroke team vs stroke ward	1	0.321	0.099	0.220 (-0.620 to 1.070)	0.61

^aThe table shows the following for each service comparison: number of trials, log direct and indirect comparisons, difference between the two estimates, and P value of that difference. There was no significant inconsistency between direct and indirect estimates.

CI: confidence interval.

GMW: general medical ward.

N/A: not applicable.

OR: odds ratio.

Table 8. Summary of findings table for the network meta-analysis of different types of organised inpatient (stroke unit) care

Intervention or-organised inpatient (stroke unit) care	Comparison GMW	Number of studies (participants)	Direct comparison evidence OR (95% CI)	Quality of the evidence (GRADE)	Direct plus indirect evidence in NMA OR (95% CI)	Quality of the evidence
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Table 8. Summary of findings table for the network meta-analysis of different types of organised inpatient (stroke unit) care (Continued)

			with direct compar- ison evi- dence		for direct compar- isons		(GRADE) for NMA
Poor outcome at end of scheduled follow-up							
Stroke ward	GMW	14 (3321)	0.78 (0.68 to 0.91)		Moderate ^a	0.74 (0.62 to 0.89)	Moderate ^a
Mobile stroke team	GMW	2 (438)	0.80 (0.52 to 1.22)		Low ^{a,b}	0.88 (0.58 to 1.34)	Low ^{a,b}
Mixed rehabilitation ward	GMW	6 (630)	0.65 (0.47 to 0.90)		Moderate ^a	0.70 (0.52 to 0.95)	Low ^{a,b}
Death at end of scheduled follow-up							
Stroke ward	GMW	15 (3523)	0.75 (0.63 to 0.90)		Moderate ^a	0.62 (0.47 to 0.82)	Moderate ^a
Mobile stroke team	GMW	2 (438)	1.08 (0.71 to 1.65)		Low ^{a,b}	1.23 (0.67 to 2.27)	Low ^{a,b}
Mixed rehabilitation ward	GMW	6 (630)	0.91 (0.58 to 1.42)		Low ^{a,b}	1.20 (0.73 to 1.99)	Low ^{a,b}
Death or institutional care at end of scheduled follow-up							
Stroke ward	GMW	13 (2924)	0.74 (0.63 to 0.87)		Moderate ^a	0.72 (0.62 to 0.83)	Moderate ^a
Mobile stroke team	GMW	2 (438)	1.27 (0.64 to 1.27)		Low ^{a,b}	1.46 (1.03 to 2.05)	Low ^{a,b}
Mixed rehabilitation ward	GMW	5 (578)	0.71 (0.51 to 0.99)		Low ^{a,b}	0.75 (0.58 to 0.96)	Low ^{a,b}
Death or dependency at end of scheduled follow-up							
Stroke ward	GMW	12 (2839)	0.75 (0.64 to 0.88)		Moderate ^a	0.71 (0.58 to 0.86)	Moderate ^a
Mobile stroke team	GMW	2 (438)	0.80 (0.52 to 1.22)		Low ^{a,b}	0.87 (0.57 to 1.32)	Low ^{a,b}
Mixed rehabilitation ward	GMW	6 (630)	0.65 (0.47 to 0.90)		Moderate ^a	0.69 (0.51 to 0.93)	Low ^{a,b}

The characteristics of different models of care are outlined in Additional Table 1.

^aDowngraded for risk of performance bias.

^bDowngraded for imprecision.

CI: confidence interval.

GMW: general medical ward.

NMA: network meta-analysis.

OR: odds ratio.

APPENDICES

Appendix 1. CENTRAL search strategy

CENTRAL search strategy (April 2019)

- #1 MeSH descriptor: [Cerebrovascular Disorders] this term only
- #2 MeSH descriptor: [Basal Ganglia Cerebrovascular Disease] explode all trees
- #3 MeSH descriptor: [Brain Ischemia] explode all trees
- #4 MeSH descriptor: [Carotid Artery Diseases] explode all trees
- #5 MeSH descriptor: [Intracranial Arterial Diseases] explode all trees
- #6 MeSH descriptor: [Intracranial Embolism and Thrombosis] explode all trees
- #7 MeSH descriptor: [Intracranial Hemorrhages] explode all trees
- #8 MeSH descriptor: [Stroke] this term only
- #9 MeSH descriptor: [Brain Infarction] explode all trees
- #10 MeSH descriptor: [Vertebral Artery Dissection] explode all trees
- #11 ((stroke or cerebrovasc* or brain vasc* or cerebral vasc* or cva* or apoplex*)):ti,ab,kw
- #12 (((brain* or cerebr* or cerebell* or vertebrobasilar or hemispher* or intracran* or intracerebral or infratentorial or supratentorial or MCA or anterior circulation or posterior circulation or basal ganglia) NEAR/5 (isch?emi* or infarct* or thrombo* or emboli*)):ti,ab,kw
- #13 (((brain* or cerebr* or cerebell* or intracerebral or intracran* or parenchymal or intraventricular or infratentorial or supratentorial or basal gangli*) NEAR/5 (haemorrhage* or hemorrhage* or haematoma* or hematoma* or bleed*)):ti,ab,kw
- #14 {or #1-#13}
- #15 MeSH descriptor: [Hospital Units] this term only
- #16 MeSH descriptor: [Patient Care Team] this term only
- #17 ((stroke NEAR/3 (unit or units or ward or wards or hospital or hospitals or centre* or team or teams)):ti,ab,kw
- #18 (((organi?ed or structured) NEAR/3 care)):ti,ab,kw
- #19 ((rehabilitation NEAR/3 (unit or units or ward or wards or hospital or hospitals or centre* or team or teams)):ti,ab,kw
- #20 ((multidisciplinary NEAR/3 (team or teams or staff* or care or rehabilitation or unit or units or ward or wards)):ti,ab,kw
- #21 (((dedicated or discrete or comprehensive) NEAR/5 (ward or wards or unit or units or stroke care)):ti,ab,kw
- #22 (((specialist or specialized or specialised) NEAR/5 (nurs* or staff* or care or unit or units or ward or wards)):ti,ab,kw
- #23 ((organi?ed NEAR/3 (unit or units or ward or wards)):ti,ab,kw
- #24 (focus* care):ti,ab,kw
- #25 ((package* NEAR/3 care)):ti,ab,kw
- #26 ((intensive NEAR/3 stroke NEAR/3 care)):ti,ab,kw
- #27 MeSH descriptor: [Intensive Care Units] this term only
- #28 MeSH descriptor: [Critical Care] this term only
- #29 {or #15-#28}
- #30 #14 and #29

Appendix 2. MEDLINE search strategy

MEDLINE (Ovid) search strategy

1. cerebrovascular disorders/ or exp basal ganglia cerebrovascular disease/ or exp brain ischemia/ or exp carotid artery diseases/ or exp intracranial arterial diseases/ or exp "intracranial embolism and thrombosis"/ or exp intracranial hemorrhages/ or stroke/ or exp brain infarction/ or exp vertebral artery dissection/
2. (stroke or cerebrovasc\$ or brain vasc\$ or cerebral vasc\$ or cva\$ or apoplex\$).tw.
3. ((brain\$ or cerebr\$ or cerebell\$ or vertebrobasilar or hemispher\$ or intracran\$ or intracerebral or infratentorial or supratentorial or MCA or anterior circulation or posterior circulation or basal ganglia) adj5 (isch?emi\$ or infarct\$ or thrombo\$ or emboli\$)).tw.
4. ((brain\$ or cerebr\$ or cerebell\$ or intracerebral or intracran\$ or parenchymal or intraventricular or infratentorial or supratentorial or basal gangli\$) adj5 (haemorrhage\$ or hemorrhage\$ or haematoma\$ or hematoma\$ or bleed\$)).tw.
5. 1 or 2 or 3 or 4
6. hospital units/ or patient care team/
7. (stroke adj3 (unit or units or ward or wards or hospital or hospitals or centre\$ or team or teams)).tw.
8. ((organi?ed or structured) adj3 care).tw.
9. (rehabilitation adj3 (unit or units or ward or wards or hospital or hospitals or centre\$ or team or teams)).tw.
10. (multidisciplinary adj3 (team or teams or staff\$ or care or rehabilitation or unit or units or ward or wards)).tw.
11. ((dedicated or discrete or comprehensive) adj5 (ward or wards or unit or units or stroke care)).tw.
12. ((specialist or specialized or specialised) adj5 (nurs\$ or staff\$ or care or unit or units or ward or wards)).tw.
13. (organi?ed adj3 (unit or units or ward or wards)).tw.
14. focus\$ care.tw.
15. (package\$ adj care).tw.
16. (intensive adj3 stroke adj3 care).tw.
17. Intensive Care Units/ or critical care/ or intensive care/
18. or/6-17
19. 5 and 18
20. Randomized Controlled Trials as Topic/
21. random allocation/
22. Controlled Clinical Trials as Topic/

23. control groups/
24. clinical trials as topic/
25. double-blind method/
26. single-blind method/
27. Research Design/
28. Program Evaluation/
29. randomised controlled trial.pt.
30. controlled clinical trial.pt.
31. clinical trial.pt.
32. random\$.tw.
33. (controlled adj5 (trial\$ or stud\$)).tw.
34. (clinical\$ adj5 trial\$).tw.
35. ((control or treatment or experiment\$ or intervention) adj5 (group\$ or subject\$ or patient\$)).tw.
36. (quasi-random\$ or quasi random\$ or pseudo-random\$ or pseudo random\$).tw.
37. ((control or experiment\$ or conservative) adj5 (treatment or therapy or procedure or manage\$)).tw.
38. ((singl\$ or doubl\$ or tripl\$ or trebl\$) adj5 (blind\$ or mask\$)).tw.
39. (assign\$ or allocat\$).tw.
40. controls.tw.
41. trial.ti.
42. or/20-41
43. 19 and 42
44. exp animals/ not humans.sh.
45. 43 not 44

Appendix 3. Embase search strategy

Embase (Ovid) search strategy

1. cerebrovascular disease/ or basal ganglion hemorrhage/ or exp brain hematoma/ or exp brain hemorrhage/ or exp brain infarction/ or exp brain ischemia/ or exp carotid artery disease/ or cerebral artery disease/ or cerebrovascular accident/ or exp intracranial aneurysm/ or exp occlusive cerebrovascular disease/ or stroke/
2. stroke patient/
3. (stroke or cerebrovasc\$ or brain vasc\$ or cerebral vasc\$ or cva\$ or apoplex\$).tw.
4. ((brain\$ or cerebr\$ or cerebell\$ or vertebrobasilar or hemispher\$ or intracran\$ or intracerebral or infratentorial or supratentorial or MCA or anterior circulation or posterior circulation or basal ganglia) adj5 (isch?emi\$ or infarct\$ or thrombo\$ or emboli\$)).tw.
5. ((brain\$ or cerebr\$ or cerebell\$ or intracerebral or intracran\$ or parenchymal or intraventricular or infratentorial or supratentorial or basal gangli\$) adj5 (haemorrhage\$ or hemorrhage\$ or haematoma\$ or hematoma\$ or bleed\$)).tw.
6. 1 or 2 or 3 or 4 or 5
7. "hospital subdivisions and components"/
8. ward/ or emergency ward/ or nursing unit/
9. intensive care unit/
10. exp intensive care/
11. (stroke adj3 (unit or units or ward or wards or hospital or hospitals or centre\$ or team or teams)).tw.
12. ((organi?ed or structured) adj3 care).tw.
13. (rehabilitation adj3 (unit or units or ward or wards or hospital or hospitals or centre\$ or team or teams)).tw.
14. (multidisciplinary adj3 (team or teams or staff\$ or care or rehabilitation or unit or units or ward or wards)).tw.
15. ((dedicated or discrete or comprehensive) adj5 (ward or wards or unit or units or stroke care)).tw.
16. ((specialist or specialized or specialised) adj5 (nurs\$ or staff\$ or care or unit or units or ward or wards)).tw.
17. (organi?ed adj3 (unit or units or ward or wards)).tw.
18. focus\$ care.tw.
19. (package\$ adj care).tw.
20. (intensive adj3 stroke adj3 care).tw.
21. or/7-20
22. 6 and 21
23. stroke unit/
24. 22 or 23
25. Randomized Controlled Trial/
26. Randomization/
27. Controlled Study/
28. control group/
29. clinical trial/
30. Double Blind Procedure/
31. Single Blind Procedure/ or triple blind procedure/

32. Parallel Design/
33. random\$.tw.
34. (controlled adj5 (trial\$ or stud\$)).tw.
35. (clinical\$ adj5 trial\$).tw.
36. ((control or treatment or experiment\$ or intervention) adj5 (group\$ or subject\$ or patient\$)).tw.
37. (quasi-random\$ or quasi random\$ or pseudo-random\$ or pseudo random\$).tw.
38. ((control or experiment\$ or conservative) adj5 (treatment or therapy or procedure or manage\$)).tw.
39. ((singl\$ or doubl\$ or tripl\$ or trebl\$) adj5 (blind\$ or mask\$)).tw.
40. (assign\$ or alternate or allocat\$ or counterbalance\$ or multiple baseline).tw.
41. controls.tw.
42. trial.ti.
43. or/25-42
44. 24 and 43
45. heart stroke volume/ or heat stroke/ or stroke volume.tw. or heat stroke.tw.
46. 44 not 45

Appendix 4. CINAHL search strategy

- S44 .S28 and S43
 S43 .S29 or S30 or S31 or S32 or S33 or S34 or S35 or S36 or S38 or S39 or S40 or S41 or S42
 S42 .TI trial
 S41 .TI controls OR AB controls
 S40 .TI (assign* or allocat*) OR AB (assign* or allocat*)
 S39 .TI ((singl* or doubl* or tripl* or trebl*) N5 (blind* or mask*)) OR AB ((singl* or doubl* or tripl* or trebl*) N5 (blind* or mask*))
 S38 .TI ((control or experiment* or conservative) N5 (treatment or therapy or procedure or manage*)) OR AB ((control or experiment* or conservative) N5 (treatment or therapy or procedure or manage*))
 S37 .TI (quasi-random* or quasi random* or pseudo-random* or pseudo random*) OR AB (quasi-random* or quasi random* or pseudo-random* or pseudo random*)
 S36 .TI ((control or treatment or experiment* or intervention) N5 (group* or subject* or patient*)) OR AB ((control or treatment or experiment* or intervention) N5 (group* or subject* or patient*))
 S35 .TI clinical* N5 trial* OR AB clinical* N5 trial*
 S34 .TI (controlled N5 (trial* or stud*)) OR AB (controlled N5 (trial* or stud*))
 S33 .TI random* OR AB random*
 S32 .(MH "Program Evaluation")
 S31 .(MH "Random Assignment")
 S30 .(ZT "clinical trial") or (ZT "randomised controlled trial")
 S29 .(MH "Clinical Trials") OR (MH "Double-Blind Studies") OR (MH "Intervention Trials") OR (MH "Randomized Controlled Trials") OR (MH "Single-Blind Studies") OR (MH "Therapeutic Trials") OR (MH "Triple-Blind Studies")
 S28 .S1 or S27
 S27 .S11 and S26
 S26 .S12 or S13 or S14 or S15 or S16 or S17 or S18 or S19 or S20 or S21 or S22 or S23 or S24 or S25
 S25 .TI intensive N3 stroke N3 care OR AB intensive N3 stroke N3 care
 S24 .TI package* N3 care OR AB package* N3 care
 S23 .TI focus* care OR AB focus* care
 S22 .TI (organi?ed N3 (unit or units or ward or wards)) OR AB (organi?ed N3 (unit or units or ward or wards))
 S21 .TI ((specialist or specialized or specialised) N5 (nurs* or staff* or care or unit or units or ward or wards)) OR AB ((specialist or specialized or specialised) N5 (nurs* or staff* or care or unit or units or ward or wards))
 S20 .TI ((dedicated or discrete or comprehensive) N5 (ward or wards or unit or units or stroke care)) OR AB ((dedicated or discrete or comprehensive) N5 (ward or wards or unit or units or stroke care))
 S19 .TI (multidisciplinary N3 (team or teams or staff* or care or rehabilitation or unit or units or ward or wards)) OR AB (multidisciplinary N3 (team or teams or staff* or care or rehabilitation or unit or units or ward or wards))
 S18 .TI (rehabilitation N3 (unit or units or ward or wards or hospital or hospitals or centre* or team or teams)) OR AB (rehabilitation N3 (unit or units or ward or wards or hospital or hospitals or centre* or team or teams))
 S17 .TI ((organi?ed or structured) N3 care) OR AB ((organi?ed or structured) N3 care)
 S16 .TI (stroke N3 (unit or units or ward or wards or hospital or hospitals or centre* or team or teams)) OR AB (stroke N3 (unit or units or ward or wards or hospital or hospitals or centre* or team or teams))
 S15 .(MH "Critical Care Nursing")
 S14 .(MH "Critical Care")
 S13 .(MH "Multidisciplinary Care Team")
 S12 .(MH "Hospital Units") OR (MH "Intensive Care Units")
 S11 .S2 or S3 or S4 or S7 or S10
 S10 .S8 and S9

S9.TI (haemorrhage* or hemorrhage* or haematoma* or hematoma* or bleed*) or AB (haemorrhage* or hemorrhage* or haematoma* or hematoma* or bleed*)

S8.TI (brain brain* or cerebr* or cerebell* or intracerebral or intracran* or parenchymal or intraventricular or infratentorial or supratentorial or basal gangli*) or AB (brain* or cerebr* or cerebell* or intracerebral or intracran* or parenchymal or intraventricular or infratentorial or supratentorial or basal gangli*)

S7 .S5 and S6

S6.TI (ischemi* or ischaemi* or infarct* or thrombo* or emboli*) or AB (ischemi* or ischaemi* or infarct* or thrombo* or emboli*)

S5.TI (brain* or cerebr* or cerebell* or vertebrobasilar or hemispher* or intracran* or intracerebral or infratentorial or supratentorial or MCA or anterior circulation or posterior circulation or basal ganglia) or AB (brain* or cerebr* or cerebell* or vertebrobasilar or hemispher* or intracran* or intracerebral or infratentorial or supratentorial or MCA or anterior circulation or posterior circulation or basal ganglia)

S4.TI (stroke or cerebrovasc* or brain vasc* or cerebral vasc* or cva* or apoplex*) or AB (stroke or cerebrovasc* or brain vasc* or cerebral vasc* or cva* or apoplex*)

S3.(MH "Stroke Patients")

S2.(MH "Cerebrovascular Disorders") OR (MH "Basal Ganglia Cerebrovascular Disease+") OR (MH "Carotid Artery Diseases+") OR (MH "Cerebral Ischemia+") OR (MH "Cerebral Vasospasm") OR (MH "Intracranial Arterial Diseases+") OR (MH "Intracranial Embolism and Thrombosis") OR (MH "Intracranial Hemorrhage+") OR (MH "Stroke") OR (MH "Vertebral Artery Dissections")

S1.(MH "Stroke Units")

Appendix 5. ClinicalTrials.gov search strategy

US National Institutes of Health Ongoing Trials Register ClinicalTrials.gov (www.clinicaltrials.gov)
 (Hospital Unit OR Intensive Care OR Critical Care Or Stroke Unit) AND INFLECT EXACT "Interventional" [STUDY-TYPES] AND (Brain Infarction OR Intracranial Hemorrhages OR Carotid Artery Diseases OR Brain Ischemia OR Cerebral Hemorrhage OR Cerebrovascular Disorders OR Stroke) [DISEASE] AND INFLECT ("09/01/2012": "08/13/2018") [STUDY-FIRST-POSTED]

Appendix 6. World Health Organization International Clinical Trials Registry search strategy

World Health Organization International Clinical Trials Registry Platform (apps.who.int/trialsearch)

Basic search/ Phases are: ALL:

stroke AND inpatient OR stroke AND hospital unit OR stroke AND patient care or stroke AND "stroke unit".

FEEDBACK

Patient subgroups,

Summary

The 95% CI includes 1.0 for patients with mild stroke. I would conclude that for this subgroup, there is no significant benefit insofar as preventing death or institutional care. I certify that I have no affiliations with or involvement in any organisation or entity with a direct financial interest in the subject matter of my criticisms.

Don Hess 2000-09-12 16:05

Criticism editor summary

Regarding the outcome 'death or institutional care' for patients with mild stroke, the 95% CIs around the odds ratio suggest that stroke unit care is not beneficial in this subgroup of patients. This is not made clear in the review's abstract, results, and discussion.

Reply

Thank you for your comment. The proper test in a subgroup analysis is not whether a subgroup result is statistically different from zero, but whether there is statistically significant heterogeneity between the estimates of effect in each of the relevant subgroups. In our subgroup analysis, the mild stroke patient group does indeed have CIs that include no effect (odds ratio = 1.0). However, we do not believe we can at present conclude that this subgroup of patients have a different result from the totality of patients. First, the statistical power of this analysis is limited because the mild stroke subgroup had relatively few outcome events (death or institutional care). Second, the mild stroke subgroup result is not significantly different from that of the moderate and severe subgroups. These analyses are explored in greater detail in the Stroke Unit Trialists' Collaboration. How do stroke units improve patient outcomes? A collaborative systematic review of the randomised trials. Stroke 1997;28:2139-44.

Contributors

Peter Langhorne 07/03/2001.

Numerical error, June 2014

Summary

Possible typo? The abstract states "... odds of death recorded at final (median one year) follow-up (odds ratio (OR) 0.87..." but text on page 15 (Comparisons 2.1, 2.2, 2.3 ...) and forest plot for 2.1 report OR = 0.81.

Reply

Dr Zekowski is correct. There appears to have been an error when updating the review. In the abstract, the correct OR for death should be (OR 0.81, 95% CI 0.69 to 0.94; P = 0.005). This has now been corrected in the text.

Contributors

Feedback: Steven Zekowski, MD.

Response: Peter Langhorne.

Long-term health outcome, December 2016

Summary

To my knowledge, there are no published randomised controlled trials directly comparing long-term health outcomes in patients managed at defined stroke units with those of conventional care. I would much appreciate if you can comment on this and also refer me to scientific articles estimating the 'number needed to care for' (analogous to number needed to treat in RCTs) regarding various health outcomes as well as health care utilization effects in stroke patients.

Reply

Thank you for your recent feedback submission on the stroke unit review. I will address your queries in turn.

1. Long-term health outcomes: to my knowledge, three randomised trials ([Athens 1995](#); [Nottingham 1996](#); [Trondheim 1991](#)) carried out 5- and 10-year follow-up of all participants. However, this is limited to a few fundamental outcomes (death, place of residence, disability). As you can imagine, almost all participants were dead or disabled at 10-year follow-up. There is very limited information on other outcomes.
2. Estimation of 'number needed to care for': I cannot reference a recent scientific article addressing your specific question. However, it is possible to calculate this information relatively easily from the current Cochrane Review. For instance, in Table 2.1 (Organised stroke unit care versus general medical ward), the absolute risk difference in deaths is -3 per 100 cared for (95% confidence interval -6 to -1). This translates into a number needed to care for of 33. The equivalent numbers needed to treat to avoid death or institutional care and death or disability are 20 and 17, respectively.
3. Other health outcomes (such as ADL score or quality of life) often are not reported in the trials. When they are, they tend to favour stroke unit care.
4. Information on health utilisation: once again, healthcare utilisation has been measured different ways in different trials. Overall, stroke unit patients tended to have a shorter length of stay in hospital, and length of stay is the main driver of costs in hospital. Several independent analyses have modelled potential healthcare utilisation effects of stroke unit care. Most have concluded that stroke unit costs are equivalent to or slightly lower than general medical costs.

I hope these comments are useful.

Contributors

Comment: Gunnar Akner, MD, PhD

Response: Peter Langhorne, Professor of Stroke Care

WHAT'S NEW

Date	Event	Description
2 April 2019	New search has been performed	This update incorporated (1) a revised literature search (updated to 2 April 2019), (2) 1 new included trial, (3) a single primary outcome (poor outcome: death or dependency or requiring institutional care) in addition to the previous outcomes, (4) a revision of the data presentation, (5) a new network meta-analysis, and (6) new 'Summary of findings' tables with GRADE quality of evidence classifications. The review now includes 29 studies involving 5902 participants
2 April 2019	New citation required but conclusions have not changed	The conclusions have not changed and remain similar to the previous version

HISTORY

Protocol first published: Issue 1, 1995

Review first published: Issue 1, 1995

Date	Event	Description
6 January 2017	Feedback has been incorporated	Feedback has been incorporated
14 July 2014	Feedback has been incorporated	Feedback has been incorporated and numerical error in the Abstract has been corrected with no change to the conclusions
29 January 2013	New search has been performed	This updated review identified 4 new trials (763 participants). We have excluded 7 previously included quasi-randomised prospective controlled clinical trials. This review now incorporates an individual patient data meta-analysis of 28 randomised controlled trials (5855 participants). More recent stroke unit trials have addressed different ways of providing organised care. This update contains data from trials comparing stroke unit care with care given in general medical wards and comparing 2 different forms of organised (stroke unit) care
29 January 2013	New citation required but conclusions have not changed	The conclusions of the review have not changed
9 September 2008	Amended	The review has been converted to new review format
28 November 2006	New search has been performed	New data on 2027 participants from 8 new trials (Athens, Beijing, Cape Town, Groningen, Joinville, Manchester, Osaka, and Pavia) have become available. More recent stroke unit trials have addressed different ways of providing organised care. This update contains new information and data from trials comparing stroke unit care with care provided on general medical wards and comparing 2 different forms of organised (stroke unit) care

CONTRIBUTIONS OF AUTHORS

Peter Langhorne initiated and co-ordinated the original review project, was principal grant holder, and revised the updated report.

For this version of the review, Peter Langhorne and Samantha Ramachandra selected trials and extracted data. Peter Langhorne performed updated literature searches, re-analysed the data, and re-drafted the manuscript.

For the previous version of the review, Patricia Fearon performed the updated literature searches, selected trials and extracted data, assisted with data analysis, and re-drafted the manuscript.

The following collaborators provided original data, advice, and comment, and assisted with re-drafting of the report: C Blomstrand (Goteborg, Sweden); NL Cabral (Joinville, Brazil); A Cavallini (Pavia, Italy); P Dey (Manchester, England); E Hamrin (Uppsala, Sweden); Graeme J Hankey (Perth, Australia); B Indredavik (Trondheim, Norway); L Kalra (Orpington, England); M Kaste (Helsinki, Finland); SO Laursen (Svendborg, Denmark); RH Ma (Beijing, China); N Patel (Cape Town, South Africa); H Rodgers (Newcastle, England); MO Ronning (Akershus, Norway); J Sivenius (Kuopio, Finland); G Sulter (Groningen, Netherlands); A Svensson (Goteborg, Sweden); K Vemmos (Athens, Greece); S Wood-Dauphinee (Montreal, Canada); and H Yagura (Osaka, Japan).

Previous versions of the review also received data, advice, and comment from K Asplund (Umea, Sweden); P Berman (Nottingham, England); M Britton (Stockholm, Sweden); J Douglas (Administrator); T Eriila (Tampere, Finland); M Garraway (Edinburgh, Scotland); M Ilmavirta (Tampere, Finland); R Stevens (Dover, England); SP Stone (London, England); and B Williams (Glasgow, Scotland).

Important contributions were also made by the following individuals, who supplied useful information and comment: D Deleo (Perth, Australia); A Drummond (Nottingham, England); R Fogelholm (Jyvaskyla, Finland); N Lincoln (Nottingham, England); H Palomaki (Helsinki, Finland); J Slattery (London, England); T Strand (Umea, Sweden); CP Warlow (Edinburgh, Scotland); and L Wilhelmsen (Goteborg, Sweden).

DECLARATIONS OF INTEREST

Most of the Stroke Unit Trialists Collaboration members carried out trials that are included in the review.

Peter Langhorne: none known.

Samantha Ramachandra: none known.

SOURCES OF SUPPORT

Internal sources

- University of Glasgow, UK
- University of Edinburgh, UK

External sources

- Chest, Heart and Stroke Scotland, UK

DIFFERENCES BETWEEN PROTOCOL AND REVIEW

This review update incorporated four new features. First, a minor revision of the data presentation such that comparisons are made at three levels: (1) organised inpatient (stroke unit) care versus any conventional service, (2) organised inpatient (stroke unit) care in a stroke ward versus care in a general medical ward, and then (3) organised inpatient (stroke unit) care in a stroke ward versus an alternative form of organised care (e.g. mobile stroke team).

Second, changes in reporting expectations required us to define a single primary outcome. We have therefore used 'poor outcome' - death or dependency or requiring institutional care (if dependency data were not available). This allowed us to keep the primary focus of the review while optimising the quantity of data available.

Third, we added an exploratory network meta-analysis (NMA) at the first level of comparison (see Comparison 1 above) to complement the previous analysis, which used a series of individual service comparisons. Both approaches are discussed here.

Finally, we have included a 'Summary of findings' table (plus GRADE) for both conventional and network meta-analyses.

INDEX TERMS

Medical Subject Headings (MeSH)

*Hospital Units; *Hospitalization; Length of Stay; *Network Meta-Analysis; Outcome Assessment, Health Care; *Patient Care Team; Prognosis; Randomized Controlled Trials as Topic; Stroke [mortality] [*therapy]; Stroke Rehabilitation; Treatment Outcome

MeSH check words

Humans