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Action observation for upper limb rehabilitation after stroke (Review)

Borges LRDM, Fernandes ABGS, Oliveira dos Passos J, Rego IAO, Campos TF

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[Intervention Review]

Action observation for upper limb rehabilitation after stroke

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ABSTRACT

Background

Action observation (AO) is a physical rehabilitation approach that facilitates the occurrence of neural plasticity through the activation of the mirror-neural system, promoting motor recovery in people with stroke.

Objectives

To assess whether AO enhances upper limb motor function in people with stroke.

Search methods

We searched the Cochrane Stroke Group Trials Register (last searched 18 May 2021), the Cochrane Central Register of Controlled Trials (18 May 2021), MEDLINE (1946 to 18 May 2021), Embase (1974 to 18 May 2021), and five additional databases. We also searched trial registries and reference lists.

Selection criteria

Randomized controlled trials (RCTs) of AO alone or associated with physical practice in adults after stroke. The primary outcome was upper limb (arm and hand) motor function. Secondary outcomes included dependence on activities of daily living (ADL), motor performance, cortical activation, quality of life, and adverse effects.

Data collection and analysis

Two review authors independently selected trials according to the predefined inclusion criteria, extracted data, assessed risk of bias using RoB 1, and applied the GRADE approach to assess the certainty of the evidence. The reviews authors contacted trial authors for clarification and missing information.

Main results

We included 16 trials involving 574 individuals. Most trials provided AO followed by the practice of motor actions. Training varied between 1 day and 8 weeks of therapy, 10 to 90 minutes per session. The time of AO ranged from 1 minute to 10 minutes for each motor action, task or movement observed. The total number of motor actions ranged from 1 to 3. Control comparisons included sham observation, physical therapy, and functional activity practice. Primary outcomes: AO improved arm function (standardized mean difference (SMD) 0.39, 95% confidence interval (Cl) 0.17 to 0.61; 11 trials, 373 participants; low-certainty evidence); and improved hand function (mean difference (MD) 2.76, 95% Cl 1.04 to 4.49; 5 trials, 178 participants; low-certainty evidence). Secondary outcomes: AO did not improve ADL performance (SMD 0.37, 95% Cl -0.34 to 1.08; 7 trials, 302 participants; very low-certainty evidence), or quality of life (MD 5.52, 95% Cl -30.74 to 41.78; 2 trials, 30 participants; very low-certainty evidence). We were unable to pool the other secondary outcomes (motor performance and cortical activation). Only two trials reported adverse events without significant adverse effects.



Authors' conclusions

The effects of AO are small for arm function compared to any control group; for hand function the effects are large, but not clinically significant. For both, the certainty of evidence is low. There is no evidence of benefit or detriment from AO on ADL and quality of life of people with stroke; however, the certainty of evidence is very low. As such, our confidence in the effect estimate is limited because it will likely change with future research.

PLAIN LANGUAGE SUMMARY

Action observation for arm rehabilitation after stroke

Review question

We sought to compare the effects of action observation on arm and hand function after stroke with an alternative intervention or no intervention. In addition, we observed the effects of this therapy on upper extremity performance, everyday activities, quality of life, and activation of brain areas.

Background

Individuals who survive a stroke often have difficulty moving their arms, which can lead to problems with everyday activities and reduced participation in daily situations. Action observation is a physical rehabilitation approach proposed for arm rehabilitation, in which the person with stroke observes a healthy individual performing a task, either on video or in person, followed or not by execution of the same task. This safe technique can be performed without expensive and complicated equipment and requires minimal therapist supervision. Trials show that action observation activates brain areas similar to those activated when performing the same action, and may favor movement recovery after stroke.

Trial characteristics

We identified 16 trials involving 574 individuals after stroke. Most used video sequences and action observation followed by some form of motor practice, using a range of activities, with task complexity increased over the course of training or when it was easy for the participant to carry out. The evidence is current to May 2021.

Key results

Trials tested whether the use of action observation compared with an alternative intervention or no intervention resulted in participants' improved ability to use their arms and hands, and found that action observation might have a small effect on arm function (11 trials) and a large effect on hand function (five trials). There is no evidence of benefit or detriment from this therapy on everyday activities and quality of life of stroke patients. It was not possible to evaluate the results of upper extremity performance and activation of brain areas.

Certainty of the evidence

The certainty of the evidence was low for arm function and hand function, and very low for everyday activities and quality of life. Participants could engage in this therapy safely, since adverse events were not significant in scale or magnitude. The certainty of the evidence for each outcome was limited due to the small number of study participants, low study quality, and poor reporting of study details.

Action observation for upper limb rehabilitation after stroke (Review) Copyright © 2022 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd. SUMMARY OF FINDINGS

Summary of findings 1. Action observation versus control: effect on upper limb rehabilitation after stroke

Action observation versus control: effect on upper limb rehabilitation after stroke

Patient or population: upper limb rehabilitation after stroke **Setting:** hospital, clinic, research laboratory or home Intervention: action observation

Comparison: control

Outcomes	Anticipated absolute effects [*] (95% CI)		(95% CI) pants	№ of partici- pants (trials)	nts the evidence	Comments
	Risk with con- trol	Risk with action observation		(,	()	
Arm function assessed by: Action Research Arm Test, Fugl-Meyer Assessment, Wolf Motor Function Test Follow-up: range 16 to 24 weeks	The mean arm function was 0 SD	SMD 0.39 higher (0.17 higher to 0.61 higher)	-	373 (11 RCTs)	⊕⊕⊝⊝ Lowa,b	Action observation may result in a slight increase in arm function.
Hand function assessed by: Box and Block Test Follow-up: range 12 to 20 weeks	The mean hand function was 0	MD 2.76 higher (1.04 higher to 4.49 higher)	-	178 (5 RCTs)	⊕⊕⊝⊝ Low a,b	Action observation may result in a slight increase in hand function.
Dependence on ADL assessed by: Barthel Index, Func- tional Independence Measure Follow-up: range 12 to 20 weeks	The mean de- pendence on activities of dai- ly living was 0	SMD 0.37 higher (0.34 lower to 1.08 higher)	-	302 (7 RCTs)	⊕⊝⊝⊝ Very low ^{a,b,c}	Action observation may in- crease/have little to no effect on de- pendence on activities of daily living but the evidence is very uncertain.
Quality of life assessed by: Stroke Impact Scale Follow-up: range 8 to 12 weeks	The mean qual- ity of life was 0	SMD 0.13 higher (1.76 lower to 2.02 higher)	-	30 (2 RCTs)	⊕⊙⊙⊙ Very low ^{a,b,c}	The evidence is very uncertain about the effect of action observation on quality of life.
Cortical activation assessed by: fMRI, MEP, EEG	See comment	See comment	-	102 (4 RCTs)	-	Trials could not be pooled due to the variety of measures and the absence of data in some groups.

·IIII. Cochrane Library

Action obconvotion fo	Motor performance assessed by: Accelerometer, 3-di- mensional motion analysis sys- tem, number of acting Follow-up: mean 1 week	See comment	See comment	- 91 - (4 RCTs)	Trials could not be pooled due to the variety of measures. Experimen- tal groups showed improvements in movement time, peak acceleration and number of repetitions of a task in 1 minute.
in the second second	Adverse effects	See comment	See comment	- 16 - (2 RCTs)	Trials could not be pooled due to subjectivity of the outcome assess- ment

*The risk in the intervention group (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).

ADL: activities of daily living; CI: confidence interval; EEG: electroencephalography; fMRI: functional magnetic resonance imaging; MD: mean difference; MEP: motor evoked potential; RCTs: randomized controlled trials; SD: standard deviation; SMD: standardized mean difference.

GRADE Working Group grades of evidence

High certainty: we are very confident that the true effect lies close to that of the estimate of the effect.

Moderate certainty: we are moderately confident in the effect estimate; the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low certainty: our confidence in the effect estimate is limited; the true effect may be substantially different from the estimate of the effect.

Very low certainty: we have very little confidence in the effect estimate; the true effect is likely to be substantially different from the estimate of effect.

^aDowngraded one level due to several ratings with 'unclear' or even 'high' risk of bias in allocation concealment, incomplete outcome data, blinding of outcome assessment or selective reporting.

^bDowngraded one level due to small total population size (< 400).

^cDowngraded one level due to moderate or high heterogeneity (> 50%).

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BACKGROUND

Description of the condition

Stroke is considered the second most common cause of death and the leading cause of adult neurological disability, according to the Global Burden of Diseases, Injuries and Risk factors study (GBD 2017; Katan 2018). It is predicted that in 2030 there will be 70 million stroke survivors worldwide (Feigin 2014).

People who survive the initial episode exhibit long-term motor impairment, limited functional activities, and reduced participation in daily situations (Langhorne 2011). Motor damage to the upper limb has a significant impact on quality of life (Lieshout 2020). Even 12 months after the stroke, daily use of the affected upper limb is three times less than the unaffected one (Rand 2015). In addition, sensorimotor function and paretic upper limb capacity can predict increased use of the arm and hand in daily life (Lundquist 2021). Thus, upper limb recovery after stroke is crucial for executing activities of daily living (ADL).

Description of the intervention

Motor recovery after stroke occurs due to neural plasticity mechanisms, allowing for lost functions to be regained. Some mechanisms enable the person with stroke to relearn motor skills, leading to a permanent change in motor behavior. This rehabilitation process exemplifies motor learning (Gregor 2021). A range of neurorehabilitation techniques aim to facilitate neural plasticity to compensate for functional impairments in affected people. Recovery of upper limb function after stroke has been the subject of numerous studies using different treatment techniques and modalities (Pollock 2014).

In this sense, besides the physical practice of action, alternative methods can be used without movement execution, such as through action observation (AO). The visual perception of movement performed by others can stimulate the observer's motor areas. This cortical motor facilitation maintains some specific properties of the observed action, such as the temporal structure and muscular organization (Bassolino 2015).

Considered a multisensory approach encompassing motor somatosensory and cognitive rehabilitation (Johansson 2011), this approach has demonstrated an important role in stroke motor recovery by activating the mirror-neural system of the brain (Buccino 2014). AO forms the basis of learning through imitation and consists of one person observing the performance of a motor task, either on a video or a real demonstration (Kim 2018). For example, the stroke patient is instructed to watch a video showing an adult stretching out their hand to pick up a cup, bringing the cup to their mouth, and then returning the cup to its initial position – the act of drinking. After observing the video sequence for a time, the individuals may or may not be asked to perform the same action.

AO can be performed from different perspectives. When it is done in a first-person perspective it involves visual information that is seen by the person performing the movement and is related to the centrality of the subjective multidimensional and multimodal experience space in one's own body. In contrast, from a thirdperson perspective, action is observed as if another person is performing it or observing themselves moving in a mirror. In both perspectives, the same brain areas are activated, but the firstperson perspective causes more extensive and stronger activation than the other (Ge 2018).

AO has been applied alone or in association with other practices, such as imitation and engagement in motor practice and training of functional activities aimed at stimulating motor relearning (Bazzini 2022; Small 2012).

How the intervention might work

There is growing evidence that motor areas (including the primary motor cortex) can be recruited not only when actions are executed, but also when they are exercised mentally or simply observed (Zhu 2019). The neurophysiological basis for this finding is the mirror-neuron system, formed by the rostral portion of the inferior parietal lobe, pars opercularis of the inferior frontal gyrus, and ventral portion of the premotor cortex (Garrison 2010; Kemmerer 2021).

For a long time, it was believed that the motor area engagement that occurs in the real execution of the action is the same that occurs during the observation of this action (Garrison 2013). However, a meta-analysis found consistent activations in a bilateral premotor, parietal, and sensorimotor network across AO and movement execution (Hardwick 2018). AO recruited a mainly bilateral network of premotor, parietal, and occipital regions, while movement execution spanned a mainly bilateral cortical sensorimotor network.

Despite these differences in activated areas in movement execution and AO, some areas are activated in common. Added to this, mirror-neuron system activation, given its distribution in the brain, provides multiple access to different brain areas; thus, AO might facilitate movement execution and motor learning by facilitating the excitability of the motor system (Huntley 2018; Small 2012) even in post-stroke patients (Tani 2018). For this reason, it is suggested that this mirror-neuron system activation may serve as an alternative means to rearrange damaged, but not completely lost, circuits, thereby rebuilding voluntary motor function. In addition, AO would therefore induce neural plasticity in people with stroke by promoting activation of the damaged motor circuits (Garrison 2013).

Why it is important to do this review

Stroke may lead to several neurological disorders, requiring longterm restorative and rehabilitative treatment. It is important to seek cheap and easy-to-apply therapies that are accessible to this population, in addition to promoting better and shorter recovery. This review is important because it involves current therapy with easy access, whose neurophysiological basis is neural plasticity, to provide evidence of its effectiveness. Given the evidence of plasticity through mirror-neural system activation during AO, it is necessary to determine the effect of this process on the acquisition of new motor skills or relearning of lost motor skills, resulting in improved upper limb performance in people with stroke.

Motor learning is a change in an individual's ability to perform a skill. The change emerges because of motor practice and is characterized by a relatively permanent improvement in performance (Krakauer 2019; Magill 1989). Motor skill acquisition occurs in three stages (cognitive, associative, and autonomous), during which cognitive demand decreases gradually, while physical demand is constant throughout the process (Fitts 1967; Krakauer 2019). Since learning a motor skill is conceptually dependent on

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motor practice, the question arises as to how AO (and its different applications) influences the learning process.

Thus, this review may clarify relevant aspects on how AO should be applied to promote significant improvements in upper limb motor function, whether it should be applied separately or in association with motor practice, in which stage of stroke, and the optimum dosage to be prescribed, among other questions. To that end, experimental trials that use this approach to promote motor learning in people with stroke must be pooled and systematically reviewed. Given that randomized clinical trials provide better evidence, the results of a number of investigations on the therapy in question must be carefully assessed, to assess the effectiveness and effects of this intervention. This would assist the rehabilitation therapist's decision making when treating upper limb motor function.

OBJECTIVES

To assess whether AO enhances upper limb motor function in people with stroke.

METHODS

Criteria for considering studies for this review

Types of studies

We planned to review published and unpublished randomized controlled trials (RCTs), including those published only as abstracts. We also examined cluster-RCTs and cross-over trials with random allocation, analyzing data from the first period. We did not include quasi-randomized or non-randomized trials, but rather those where random allocation was accomplished by using a randomnumber generator, referring to a random-number table, and using sequentially-numbered opaque sealed envelopes. We accepted trials in any language and from any year.

Types of participants

We included trials whose participants were clinically diagnosed with stroke and upper limb motor deficit. Participants were 18 years or older, any gender, with any degree of stroke impairment severity, and at any stage of the condition.

Types of interventions

We selected trials that included the following interventions.

- 1. AO alone or associated with motor practice, imitation or training of functional tasks versus other therapies.
- 2. AO alone or associated with motor practice, imitation or training of functional tasks versus conventional physiotherapy.
- 3. AO alone or associated with motor practice, imitation or training of functional tasks versus placebo.
- 4. AO alone or associated with motor practice, imitation or training of functional tasks versus no therapy.

We considered AO as the observation of an individual performing a motor action through videos or in real time. We did not include trials using mirror therapy in this review.

Types of outcome measures

We extracted the outcomes of interest from the baseline and the evaluation at the end of the intervention period (short term) and follow-up (long term). Our list of outcome measures was not exhaustive; when we found trials with other relevant outcome measures, we included them and documented these findings. Moreover, when we identified a study in which more than one measurement instrument found the same outcome, we used the one included on our list.

Primary outcomes

- 1. Upper limb motor function, measured by the following.
 - a. Arm function
 - i. Motor Assessment Scale (upper limb component)
 - ii. Frenchay Arm Test
 - iii. Motor Activity Log
 - iv. Wolf Motor Function Test
 - v. Action Research Arm Test
 - vi. Fugl-Meyer Assessment
 - b. Hand function
 - i. Motor Assessment scale (hand component)
 - ii. Box and Block Test
 - iii. Jebsen Test of Hand Function and Peg Test

Secondary outcomes

- 1. Dependence on activities of daily living (ADL), measured by the following.
 - a. Functional Independence Measure
 - b. Barthel Index
 - c. Rankin Scale
- 2. Motor performance, measured by kinematic analysis, e.g.
 - a. velocity
 - b. angular analysis of movement
- 3. Cortical activation, measured by the following.
 - a. Functional Magnetic Resonance
 - b. Transcranial Magnetic Stimulation
 - c. Electroencephalography
 - d. Positron Emission Tomography
- 4. Quality of life, restricted participation, or both, measured by the following.
 - a. London Handicap Scale
 - b. 36-Item Short Form Health Survey (SF-36)
 - c. EuroQoL 5 dimensions (EQ-5D)
 - d. Stroke Impact Scale
- 5. Adverse effects
 - a. pain
 - b. muscle weakness
 - c. fatigue
 - d. death

Search methods for identification of studies

See the Cochrane Stroke Group's search methods. We searched for trials in all languages and arranged for the translation of relevant articles where necessary.



Electronic searches

We searched the Cochrane Stroke Group Trials Register and the following electronic bibliographic databases and trial registers.

- Cochrane Central Register of Controlled Trials (CENTRAL; 2021, Issue 5) in the Cochrane Library (searched May 2021; Appendix 1).
- 2. MEDLINE (Ovid) (from 1946 to May 2021; Appendix 2).
- 3. Embase (Ovid) (from 1980 to May 2021; Appendix 3).
- 4. CINAHL EBSCO (Cumulative Index to Nursing and Allied Health Literature (from 1982 to May 2021; Appendix 4).
- 5. LILACS (Latin American and Caribbean Health Sciences Literature to May 2021; Appendix 5).
- 6. Allied and Complementary Medicine Database (AMED) (Ovid) (from 1985 to 18 May 2021; Appendix 6).
- Physiotherapy Evidence Database (PEDro: www.pedro.org.au; searched May 2021; Appendix 7)

We developed the MEDLINE search strategy with the help of the Cochrane Stroke Group Information Specialist (Appendix 2), and adapted it for the other databases (Appendix 1; Appendix 3; Appendix 4; Appendix 5; Appendix 6; Appendix 7).

We searched the following trial registries for ongoing trials.

- 1. US National Institutes of Health Ongoing Trials Register ClinicalTrials.gov (www.clinicaltrials.gov; searched May 2021; Appendix 8).
- ISRCTN Registry (www.isrctn.com; searched May 2021 included on WHO International Clinical Trials Registry; Appendix 9).
- 3. Australian New Zealand Clinical Trials Registry (www.anzctr.org.au; searched May 2021 - included on WHO International Clinical Trials Registry; Appendix 10).
- World Health Organization (WHO) International Clinical Trials Registry Platform (trialsearch.who.int/Default.aspx; searched May 2021; Appendix 11).

Searching other resources

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

- 1. screened reference lists of all relevant articles;
- 2. used Science Citation Index Cited Reference search for forward tracking of important articles;
- 3. contacted trialists, experts and researchers in our field of study;
- handsearched journals (International Journal of Neurorehabilitation, Neurorehabilitation and Neural Repair, and Stroke);
- 5. searched for PhD and MSc theses (using Latin American and Caribbean Health Sciences Literature: LILACS).

Data collection and analysis

Selection of studies

Two review authors (JP and IR) independently screened the titles and abstracts of records obtained from the electronic searches and excluded those that were obviously irrelevant. We obtained the full text of the remaining records and the same two review authors selected trials for inclusion according to the predefined inclusion criteria. If any methodological question raised doubts

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about whether the study met the inclusion criteria, we contacted the study authors for clarification. If there was disagreement regarding the selection of trials, we attempted to reach a consensus through discussion. If this was impossible, we asked another review author (TC) to decide if the study should be included. We recorded reasons for exclusion and completed a PRISMA flowchart (Page 2021).

Data extraction and management

Two review authors (AF and LB) were responsible for data extraction. To record these data, we used a form we created, based on the Cochrane Consumers and Communication Group data extraction template for included studies (Ryan 2016), which we tested (piloted) in two trials. Where there was incomplete or unclear data, we contacted the study authors for clarification. The same two review authors discussed any disagreements encountered in order to reach a consensus; and if we could not reach consensus, another review author (TC) ruled on the issue. One review author (AF) entered all extracted data into Review Manager 5 (RevMan 5) (Review Manager 2020), and another (LB) working independently checked the data to confirm accuracy. These data refer to the following.

- 1. General information: title of the review, name of the review author who completed the form, and study ID.
- Methods used: objectives, study design, instruments used, study duration, type of randomisation, allocation concealment, blind assessors, inclusion and exclusion criteria, institutions or study centers involved, study site, removal and abandonment of participants, and year of the study.
- 3. Participants: population description, sample size, age, sex, initial upper limb impairment, severity level of stroke and type of stroke, diagnostic criteria, phase (acute, subacute, and chronic), presence of communication or cognitive impairments.
- 4. Intervention: therapies used in association and in comparison, type of task, number and duration of the sessions and time of the sessions, methods used in the control group, and the profession of the person that applied the therapy.
- 5. Results: primary and secondary outcomes for each assessment and reassessment, methods and instrumentation for assessment, timing of outcome assessment, and adverse events.
- Notes: contact with authors (information obtained or not), article in a language other than English, study financing and noteworthy conflicts of interest of study authors.

Assessment of risk of bias in included studies

We assessed risk of bias using the RoB 1 tool described in the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2017). Two review authors (AF and LB) independently assessed the risk of bias and discussed disagreements and, if necessary, we asked another review author (TC) to come to a conclusion. We used a form containing the following risk of bias criteria.

- 1. Random sequence generation
- 2. Allocation concealment
- 3. Blinding of participants and personnel
- 4. Blinding of outcome assessment
- 5. Incomplete outcome data
- 6. Selective outcome reporting



7. Other bias

We classified each criterion, characterizing it as high, low, or uncertain risk of bias. We inserted this information into the risk of bias table produced for each study, along with the reason for each decision. We used Table 8.5.d from the *Cochrane Handbook for Systematic Reviews of Interventions*, which provides criteria for making judgements regarding risk of bias in each of the seven domains of the tool (Higgins 2017). We contacted trial authors for clarification and to request missing information. We considered the risk of bias of the trials and their contribution to the treatment effect.

Measures of treatment effect

We measured treatment effect using mean difference (MD) and standardized mean difference (SMD) for the continuous outcomes, with 95% confidence intervals (CIs). According to Higgins 2022, if trials are clinically diverse then a meta-analysis may be meaningless, and genuine differences in effects may be obscured. Often it is nonsensical to combine all included trials in a single meta-analysis: if there is a mix of comparisons of different treatments with different comparators, each combination may need to be considered separately (Higgins 2022). We performed a meta-analysis using RevMan 5 only if there was clinical and methodological similarity between trials, so that they could be pooled for analysis (Review Manager 2020). One review author (LB) conducted this judgment. In case of doubt, a third review author (TC) made the final decision. We based clinical similarity on population characteristics, such as age range, type of stroke, stroke severity, and stage of stroke (acute, subacute and chronic). We considered similar methodology when the type of intervention and outcomes (motor function, dependence of daily living, and others), even if measured by different instruments, were repeated between trials. We used the random-effects model in our analysis.

Unit of analysis issues

Cross-over trials

We included one cross-over study, using only the first period for analysis.

Cluster-randomized trials

There were no cluster-randomized trials.

Dealing with missing data

We contacted study authors when possible to verify key study characteristics and to obtain missing numerical outcome data. When this was not possible, and the missing data were thought to introduce serious bias, we performed a sensitivity analysis to explore the impact of including such trials in the overall assessment of results.

Assessment of heterogeneity

We assessed heterogeneity visually by observing the nonoverlapping of confidence intervals (CIs) in the forest plots. Once identified, we quantified heterogeneity by the I^2 statistic. When heterogeneity was caused by one or two trials with peripheral results that conflicted with the rest of the trials, we carried out analyses with and without these trials as part of the sensitivity analysis. We considered values of < 30% as low heterogeneity, from 30% to 60% moderate heterogeneity, 61% to 75%, representing substantial heterogeneity, and > 75% suggestive of considerable heterogeneity (Higgins 2022).

Assessment of reporting biases

The search for clinical trials registers helped in reducing publication bias. We compared the trial registry with the included trials methods. In the absence of registration, it was observed if the outcomes evaluated were surrogate or clinically relevant. We also investigated selective outcome reporting by the comparison of the methods section of papers with the results reported. We inspected the funnel plot for arm function analysis, the only one with more than 10 trials.

Data synthesis

We conducted a meta-analysis when the trials could be combined. The review authors used the random-effects model. We planned to do an analysis of AO (alone or associated with motor practice) versus any other control (active or inactive control). When it was not possible to perform a meta-analysis, we demonstrated the results with tables and a narrative synthesis, where we presented outcomes and results or objectives, according to each treatment category.

Subgroup analysis and investigation of heterogeneity

We planned to carry out the following subgroup analyses.

- 1. Age
- 2. Type of stroke
- Time poststroke: acute (< 1 month poststroke), subacute (1 month and ≤ 6 months poststroke) and chronic (> 6 months poststroke)
- 4. Length of treatment period or dose of treatment
- 5. Type of treatment: AO alone and associated with motor practice (activity, imitation or training of functional tasks)

Sensitivity analysis

As previously explained, we performed sensitivity analyses when we suspected that missing data could introduce important bias, and also to assess heterogeneity caused by trials with peripheral results. Furthermore we planned to carry out the following sensitivity analyses, excluding trials with a high risk of bias. We considered a study as having a high risk of bias if the following criteria were not met.

- 1. Allocation concealment
- 2. Blinding of outcome assessment

Summary of findings and assessment of the certainty of the evidence

We created a summary of findings table including the following outcomes: upper limb motor function (arm function and hand function), dependence on ADL, quality of life, cortical activation, motor performance, and adverse effects. We used the five GRADE considerations (study limitations, consistency of effect, imprecision, indirectness and publication bias) to assess the certainty of a body of evidence as it relates to the trials contributing data to the review for the outcomes. In particular, we downgraded the certainty of evidence by one level when trials exhibited high risk of bias, the total sample size was small (n < 400), or when heterogeneity was inexplicably moderate or high (> 50%). Two



review authors (LB and TC) independently assessed the certainty of the evidence.

We used the GRADEpro Guideline Development Tool to prepare the summary of findings table (GRADEpro GDT).

RESULTS

Description of studies

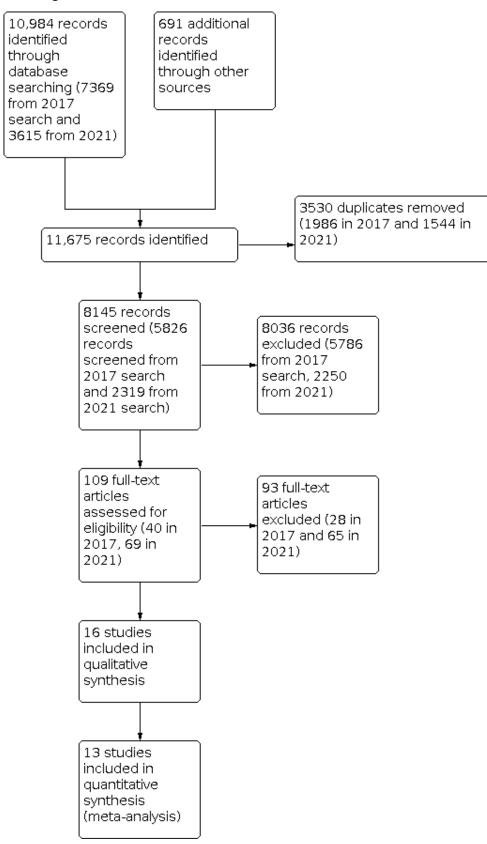
See Characteristics of included studies; Characteristics of excluded studies

Results of the search

The searches of electronic databases and trial registers produced 8145 unique references for screening. After excluding non-relevant citations, we obtained the full texts of 109 papers; of these, we included 16 trials in the qualitative analysis and 13 trials in the quantitative analysis of the review; Figure 1 shows the study flow diagram of the selected trials.



Figure 1. Study flow diagram





Included studies

We identified 16 randomized controlled trials (RCTs) with a total of 574 participants that met the inclusion criteria: 12 (478 participants) in the previous version (Celnik 2008; Cowles 2012; Dettmers 2014; Ertelt 2007; Franceschini 2012; Fu 2017; Harmsen 2015; Kim 2015; Kim 2016a; Kuk 2016; Lee 2013; Zhu 2015), and four (96 participants) in the updated review (Hsieh 2020; Jin-Woo 2011; Mancuso 2021; Younghwa 2013). The trials were all randomized; one was a cross-over trial (Celnik 2008). We contacted eight authors for clarification concerning methodology (how allocation concealment, random sequence generation, and blinding were performed) and results data; four authors replied.

We found two publications that we judged to be two reports of the same study (see study references for Franceschini 2012); while there were some differences in reported outcome measures and numbers of participants (Franceschini 2012 article with 102 participants; Sale 2012 article with 67 participants), we considered these were the same population of participants and therefore did not enter them as separate trials, since the use of duplicate data may lead to overestimating the intervention effects. We opted to use the study report with the largest sample size, considering this to provide the most comprehensive results.

Sample characteristics

We included 16 trials involving a total of 574 participants in the review. Samples ranged between nine (Celnik 2008), and 102 participants (Franceschini 2012), averaging approximately 35 participants per study. Only Kim 2015 did not characterize the sample. The mean age of participants ranged from $51.16 (\pm 12.77)$ to 77.2 (±10.4) years. The sample consisted of 273 men and 208 women (Kim 2015 data not included). Seven trials included participants in the acute and/or subacute phase of stroke (Cowles 2012; Franceschini 2012; Fu 2017; Hsieh 2020; Kim 2016a; Mancuso 2021; Zhu 2015), and six in the chronic phase (Celnik 2008; Ertelt 2007; Harmsen 2015; Jin-Woo 2011; Kuk 2016; Lee 2013). Dettmers 2014 and Younghwa 2013 included participants in the acute, subacute, and chronic phases. Initial upper limb impairment was severe (Cowles 2012; Franceschini 2012; Zhu 2015), and moderate (Ertelt 2007; Harmsen 2015; Hsieh 2020; Kim 2016a), but two trials included participants with mild, moderate, and severe paresis (Dettmers 2014; Mancuso 2021); the remaining trials did not specify initial upper limb impairment. Severity classification of upper limb impairment was performed using different instruments and methodologies, considering the Motricity Index, Fugl-Meyer Assessment, Wolf Motor Function Test, and ability to move the upper limb.

Nine trials specified stroke etiology: two recruited only participants with ischemic stroke (Ertelt 2007; Fu 2017), and seven recruited participants with ischemic and hemorrhagic stroke (Franceschini 2012; Younghwa 2013; Harmsen 2015; Hsieh 2020; Jin-Woo 2011; Kim 2016a; Kuk 2016). Five were carried out in a hospital setting or in rehabilitation centers (Cowles 2012; Ertelt 2007; Fu 2017; Kim 2016a; Zhu 2015); the others did not record treatment location.

Interventions

Intervention approaches

The following comparisons were used for the trials (Table 1).

Participants in seven trials did not undergo physiotherapy or occupational therapy in addition to the procedures proposed for control and experimental groups (Celnik 2008; Ertelt 2007; Harmsen 2015; Jin-Woo 2011; Kuk 2016; Lee 2013; Younghwa 2013). In all trials, AO was followed by some type of motor activity, largely functional tasks. Most of the investigations used variable practice, increasing the complexity of the tasks throughout training or when the participant demonstrated ease in carrying out the action. Constant practice of a motor activity was observed in two trials, practicing by repeating the same variant of the skill, one for upper limb reach and the other for water-drinking action (Harmsen 2015; Lee 2013). Hsieh 2020 added to functional activities, observation and execution of upper limb range of motion exercises. In another study, after the observation, the execution of motor tasks was carrying wooden blocks from one box to another, similar to the Box and Block Test (Kuk 2016).

In one study, the individuals did not undergo functional task training but did perform thumb movements. Furthermore, the movements were carried out simultaneously to observation and considered congruent when performed in the same direction as the observed movement, and incongruent when the movement was in the opposite direction to that observed (Celnik 2008). Kim 2015 did not clearly explain whether the individuals performed the observed task after AO. However this likely occurred, since the study authors reported the experimental group as having undergone a training program ("purposeful action observation training program"), which can be understood as being some type of motor practice after the observed activity.

Comparison interventions

In some trials, the control group underwent rehabilitation treatment through conventional physical therapy and/or occupational therapy (Cowles 2013; Franceschini 2012; Fu 2017; Hsieh 2020; Kim 2015; Kim 2016a; Mancuso 2021; Zhu 2015), with or without other therapies. In all of these trials, physical therapy and occupational therapy was provided to the control and experimental group.

In Cowles 2013, the control group performed only conventional physical therapy. Some trials used activities other than conventional physical therapy (Hsieh 2020; Kim 2015; Kim 2016a; Mancuso 2021), such as the practice of functional tasks (Hsieh 2020; Kim 2015; Kim 2016a), in addition to active range of motion exercises, reaching movement or object manipulation (Hsieh 2020). Mancuso 2021 added Task-oriented training to conventional physical. This intervention uses motor learning principles such as specificity of training, constrained use of impaired limbs, mass practice (repetition), shaping of skill, the saliency of a task, and knowledge of performance and result (Harvey 2009).

Some trials explored the content of conventional physical therapy applied. The control group in Cowles 2013 underwent soft tissue mobilization, joint mobilization, facilitating muscle activity, positioning, specific sensory input, splinting, exercise to increase strength, balance, mobility, upper limb functional tasks, and education. Franceschini 2012 used dexterity and gait training. Fu 2017 used Bobath/Brunnstrom, proprioceptive neuromuscular facilitation, and daily activity ability training. Mancuso 2021 submitted the control group to training for transfers, mobility, walking up and down steps, balance tasks, functional tasks for the



upper limbs, joint and soft tissue mobilization, and specific sensory stimulation exercises to increase strength.

In two trials, patients received medication in addition to rehabilitation treatment (Fu 2017; Zhu 2015). In addition to rehabilitation, some control groups performed placebo observation plus some upper limb action or movement practice (Franceschini 2012; Fu 2017).

Placebo observation was the choice of several trials (Dettmers 2014; Ertelt 2007; Franceschini 2012; Fu 2017; Harmsen 2015; Kuk 2016). The observed content was text (Dettmers 2014), geometric symbols, letters or symbols (Ertelt 2007; Fu 2017), static images of objects (Franceschini 2012), and photographs and landscapes (Harmsen 2015; Kuk 2016). After this observation, the participants underwent some physical or motor training, such as functional activities training (Dettmers 2014; Ertelt 2007; Franceschini 2012), reach training (Harmsen 2015), upper limb movements (Fu 2017), and action training similar to Box and Block Test (Kuk 2016).

In a few trials, the control group practiced functional tasks (Jin-Woo 2011; Lee 2013; Younghwa 2013), and specific movements, such as thumb movements (Celnik 2008). In only two trials (Dettmers 2014; Hsieh 2020), the exercises were individualized for the participants or the task was adjusted according to the participant's motor skills. In Franceschini 2012, the researcher provided physical help to support or mobilize the upper limb.

Two trials presented two control groups (Dettmers 2014; Lee 2013), one of which did not undergo specific training. To conduct the meta-analysis, we did not use this group in order to reduce heterogeneity between trials.

Resources

The resources used to provide observation of action for the experimental group were television (TV) (Ertelt 2007; Fu 2017; Zhu 2015) and computer monitor (Harmsen 2015; Hsieh 2020; Jin-Woo 2011; Kim 2016a; Kuk 2016; Younghwa 2013). The other authors did not specify the resource used for exposure (Celnik 2008; Cowles 2013; Dettmers 2014; Franceschini 2012; Kim 2015; Lee 2013; Mancuso 2021). Other resources used were objects, such as dish, towel, book, cup, coins, cubes, wooden blocks, clothes and others, materials used to carry out the training of tasks or functional activities, as detailed in seven trials (Ertelt 2007; Hsieh 2020; Jin-Woo 2011; Kim 2016a; Kuk 2016; Lee 2013; Zhu 2015).

For the materials used for control groups, two trials referred to the use and manipulation of objects, such as towels, magazines, tables, drawers, bottles, and caps (Hsieh 2020; Kim 2016a). Others used more specific objects such as blocks and boxes (Kuk 2016), a cup (Lee 2013), clothes, and a wheelchair (Younghwa 2013). Another resource used was the TV or monitor (Kuk 2016) for the video presentation. The other trials did not specify the material used but reported that individuals watched videotapes and slide shows, possibly using TV or computer (Dettmers 2014; Ertelt 2007; Franceschini 2012; Fu 2017; Harmsen 2015).

Delivery

All the trials used video sequences to apply AO except Cowles 2012, which used real demonstrations of the task prescribed by the therapist, and the Kim 2015 trial, which did not report how therapy was applied. Six trials used first-person perspective in

applying AO (Celnik 2008; Cowles 2012; Franceschini 2012; Harmsen 2015; Hsieh 2020; Mancuso 2021), and three used third-person perspective (Dettmers 2014; Fu 2017; Lee 2013). Two trials did not specify the perspective used (Kim 2015; Younghwa 2013). Other trials reported the position for the demonstration (in this case, third-person perspective), which could be front, side, top, and back views of motor action. Kuk 2016 used two different views. Ertelt 2007, Jin-Woo 2011, Kim 2016a; Zhu 2015 used three views.

In four trials, some type of verbal correction or feedback was provided after training in the experimental group (Cowles 2013; Franceschini 2012; Kim 2016a; Mancuso 2021). Other trials mentioned verbal instruction, explaining the motor task or movement (Franceschini 2012; Jin-Woo 2011; Lee 2013; Mancuso 2021; Zhu 2015). In one study, the choice of activities was performed according to the physical capacity of the participants (Cowles 2013). In the Franceschini 2012 trial, the occupational therapist could offer physical help to complete the task or movement.

Two trials reported providing control therapy using video (Dettmers 2014; Harmsen 2015), and one using a slide show (Kuk 2016). Regarding the type of instruction, verbal instruction on the task or movement was used in five trials by the researchers or clinicians who applied the control intervention (Ertelt 2007; Franceschini 2012; Jin-Woo 2011; Lee 2013; Younghwa 2013). The Younghwa 2013 trial included outcome feedback. Dettmers 2014 used textual instruction.

Setting

The clinical setting seems to have been the location of most trials, although only five reported this information. One trial carried out its research in an inpatient stroke unit (Cowles 2013), while four trials were carried out in rehabilitation centers (Ertelt 2007; Fu 2017; Kim 2016a; Zhu 2015). The Dettmers 2014 trial was carried out at the participants' home.

Amount of therapy provided

Intervention time in the experimental and control groups was 3 weeks in five trials (Cowles 2013; Ertelt 2007; Hsieh 2020; Jin-Woo 2011; Lee 2013); 4 weeks, 5 times/week (Franceschini 2012; Kim 2016a; Mancuso 2021); 6 weeks, raging from four to six times/week (Dettmers 2014; Kim 2015; Younghwa 2013); and 8 weeks, six times/ week (Fu 2017; Zhu 2015). Other trials performed the intervention in one day (Celnik 2008; Harmsen 2015; Kuk 2016).

The time of AO in the experimental group was variable between trials, ranging from a minimum of 1 minute (Cowles 2013; Kuk 2016), to a maximum of 10 minutes (Celnik 2008; Fu 2017), for each motor action, task or movement observed. The number of motor actions offered in each demonstration also varied between one (Celnik 2008; Lee 2013), two (Jin-Woo 2011; Kim 2015; Younghwa 2013), and three (Ertelt 2007; Franceschini 2012; Mancuso 2021). Furthermore, in some trials, the observation of action was provided more than once: twice in the Ertelt 2007 trial, five times in Kuk 2016, and three times in the Hsieh 2020 and Celnik 2008 trials.

The time of one session in the experimental group lasted 10 minutes (Lee 2013), 20 minutes (Fu 2017), 30 minutes (Celnik 2008; Franceschini 2012; Jin-Woo 2011; Kim 2015; Kim 2016a; Mancuso 2021; Younghwa 2013; Zhu 2015), 40 minutes (Ertelt 2007), and 60 minutes (Cowles 2013; Dettmers 2014; Hsieh 2020; Kuk 2016). One



study did not specify the duration of the intervention (Harmsen 2015).

Training varied between 1 day and 8 weeks of therapy, with 10 to 90 minutes per session. Total treatment duration was 20 minutes in Kuk 2016, 30 minutes in Celnik 2008 and Harmsen 2015, 150 minutes in Lee 2013, 270 minutes in Jin-Woo 2011, 600 minutes in Franceschini 2012 and Mancuso 2021, 720 minutes in Younghwa 2013, 800 minutes in Kim 2016a, 900 minutes in Cowles 2013, Kim 2015 and Hsieh 2020, 960 minutes in Fu 2017, 1440 minutes in Zhu 2015, 1620 minutes in Ertelt 2007, and 2520 minutes in Dettmers 2014. The protocol used in most of the trials divided the functional tasks into smaller parts and provided between 1 and 6 minutes of observation for each motor sequence, followed by two to six minutes of practice for the action observed. Other details regarding the intervention are presented in Table 2.

Session duration in the control group was 30 minutes in some trials (Celnik 2008; Franceschini 2012; Jin-Woo 2011; Kim 2015; Kim 2016a; Mancuso 2021; Younghwa 2013). Three trials did not specify the exact duration of the session (Cowles 2013; Harmsen 2015; Zhu 2015). The others lasted: 10 minutes (Lee 2013); 20 minutes (Fu 2017; Kuk 2016); 60 minutes (Dettmers 2014); and 90 minutes (Ertelt 2007). Details on the timing of each component of control therapy are available in Table 3.

Providers

Some authors mentioned that therapists provided the therapy without specifying the qualification or specialty of the professionals. Therefore, we used the same term as that provided by the study authors.

The professionals in charge of the control group therapy were therapists in seven trials (Cowles 2013; Ertelt 2007; Jin-Woo 2011; Kim 2016a; Lee 2013; Mancuso 2021; Younghwa 2013), and occupational therapists in three trials (Dettmers 2014; Franceschini 2012; Hsieh 2020). Therapists applied the intervention in the experimental group in five trials (Cowles 2013; Jin-Woo 2011; Kim 2016a; Mancuso 2021; Younghwa 2013). In three, the intervention was provided by occupational therapists (Dettmers 2014; Franceschini 2012; Hsieh 2020). In Ertelt 2007, individuals in this group were assisted by a psychologist. The remaining trials did not report who provided the intervention. One author recorded the time of professional experience of the therapist, from 5 to 8 years (Kim 2016a).

Outcomes

In relation to the outcomes used, 11 trials included upper limb motor function (Cowles 2012; Dettmers 2014; Ertelt 2007; Franceschini 2012; Fu 2017; Hsieh 2020; Jin-Woo 2011; Kim 2015; Kim 2016a; Mancuso 2021; Zhu 2015); and only seven exhibited functional dependence for ADL as an outcome (Franceschini 2012; Fu 2017; Hsieh 2020; Kim 2016a; Mancuso 2021; Zhu 2015; Younghwa 2013). Motor performance was assessed in three trials involving several variables (Celnik 2008; Harmsen 2015; Lee 2013). Cortical activation was observed in four trials, using transcranial magnetic stimulation (TMS) (Celnik 2008; Fu 2017), functional magnetic resonance imaging (fMRI) (Ertelt 2007), and electroencephalography (EEG) (Kuk 2016).

Three trials included quality of life as an outcome (Dettmers 2014; Ertelt 2007; Hsieh 2020); one had adverse events, attention level,

and fatigue as outcomes (Celnik 2008); while another monitored pain in the experimental group, but without quantifying it (Cowles 2013).

Some outcomes not foreseen in the review protocol were used. Mancuso 2021 evaluated spasticity. Jin-Woo 2011 measured handgrip and specific pinch strength.

A number of instruments were used to quantify primary and secondary outcomes. With respect to the primary outcome (upper limb motor function), subdivided into the arm and hand, the following were used for the arm: Action Research Arm Test (Cowles 2012); Wolf Motor Function Test (Dettmers 2014; Ertelt 2007; Fu 2017; Jin-Woo 2011; Kim 2015); Motor Activity Log (Dettmers 2014); Frenchay Arm Test (Ertelt 2007; Franceschini 2012); and Fugl-Meyer Assessment (Franceschini 2012; Fu 2017; Kim 2016a; Hsieh 2020; Mancuso 2021; Zhu 2015). The motor function outcome of the hand was assessed by the Nine-Hole Peg Test (Dettmers 2014), and Box and Block Test (Franceschini 2012; Hsieh 2020; Kuk 2016; Mancuso 2021).

All secondary outcomes specified in this review were present in the included trials. The following instruments were used to measure dependence in ADL: Functional Independence Measure - motor items (Franceschini 2012; Hsieh 2020; Mancuso 2021), and Modified Barthel Index/Barthel Index (Fu 2017; Kim 2016a; Younghwa 2013; Zhu 2015). Motor performance, measured by kinematic analysis and an accelerometer was determined in the Celnik 2008 study, which considered angular difference, angular dispersion, and peak acceleration as measures. Harmsen 2015 also used an accelerometer to measure motor performance considering reaching time, whereas Kim 2015 considered average velocity, trajectory ratio, and motion angle, measured by a 3dimensional motion analysis system. In Lee 2013, the number of times the drinking task was performed in 1 minute was measured. Heterogeneity of the variables used to characterize motor performance precluded combining them in analysis and comparing them.

Cortical activation was determined by motor-evoked potential (MEP) amplitudes, latency and center-motion conduction time provoked by TMS (Celnik 2008; Fu 2017). Ertelt 2007 used fMRI to analyze the effects of AO. Kuk 2016 used EEG for investigating the mirror neuron system activation during AO. The Stroke Impact Scale was applied to measure quality of life (Dettmers 2014; Ertelt 2007; Hsieh 2020).

Spasticity was evaluated by the Modifed Ashworth Scale (Mancuso 2021), and handgrip strength was evaluated by manual dynamometer (Jin-Woo 2011).

Adverse effects were quantified in one of the trials using the visual analogue scale, which focused on the level of attention and fatigue during the interventions (Celnik 2008). Other adverse effects, such as upper limb pain (overuse syndrome), were monitored for verbal or behavioral manifestations (e.g. grimacing, postural guarding), and for a decrease of at least two measurement levels in the Motricity Index, the scale that quantifies muscle weakness (Cowles 2012).

Three trials assessed the short-term effects of AO, with assessment and reassessment on the same day (Celnik 2008; Harmsen 2015; Kuk 2016). Five trials assessed the effects after conclusion of



therapy via follow-up at 1 week (Lee 2013), 2 months (Ertelt 2007), 3 months (Hsieh 2020), 4 to 5 months (Franceschini 2012), and 6 months (Dettmers 2014). The Dettmers 2014 and Ertelt 2007 trials conducted follow-up only for the experimental group.

Excluded studies

We excluded 25 trials for various reasons (Characteristics of excluded studies). Five trials are ongoing.

Risk of bias in included studies

Two review authors independently assessed the methodological quality of the included trials using RoB 1: see Characteristics of included studies, risk of bias summary (Figure 2), and risk of bias graph (Figure 3). Not all the trials followed CONSORT recommendations, a guideline for reporting randomized trials (Schulz 2010). For this reason, we emailed study authors for clarification in the event of questions concerning trial methodology.





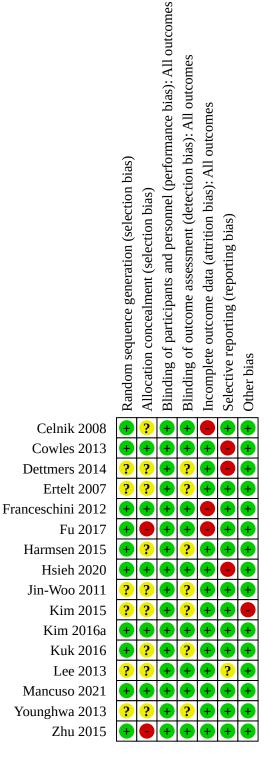
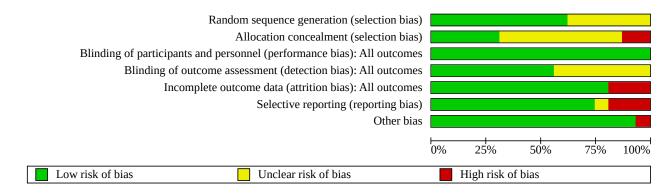




Figure 3. Risk of bias graph: review authors' judgments about each risk of bias item presented as percentages across all included trials.



Allocation

We judged random sequence generation as adequate in 10 trials (Celnik 2008; Cowles 2013; Franceschini 2012; Fu 2017; Harmsen 2015; Hsieh 2020; Kim 2016a; Kuk 2016; Mancuso 2021; Zhu 2015), while the other six exhibited unclear risk of bias for this criterion (Dettmers 2014; Ertelt 2007; Jin-Woo 2011; Kim 2015; Lee 2013; Younghwa 2013).

Five of the 16 trials appropriately described the allocation concealment of participants to groups and we deemed these to be at low risk (Cowles 2013; Franceschini 2012; Hsieh 2020; Kim 2016a; Mancuso 2021); we considered only two trials at high risk (Fu 2017; Zhu 2015).

Blinding

No trials were able to blind participants or personnel. However, this type of blinding is impossible to apply with this therapy, and since therapist and participant were aware of the treatment they were performing, we considered there to be low potential to negatively influence the effect of the therapy and, therefore, judged them to be at low risk of bias.

Nine trials reported blinding the outcome assessor (Celnik 2008; Cowles 2013; Franceschini 2012; Fu 2017; Hsieh 2020; Kim 2016a; Lee 2013; Mancuso 2021; Zhu 2015); we considered the other trials to be at unclear risk of bias.

Incomplete outcome data

We deemed 13 trials to have low risk of bias in relation to this criterion (Cowles 2013; Dettmers 2014; Ertelt 2007; Harmsen 2015; Hsieh 2020; Jin-Woo 2011; Kim 2015; Kim 2016a; Kuk 2016; Lee

2013; Mancuso 2021; Younghwa 2013; Zhu 2015), and considered three at high risk due to an imbalance in the number of participants in each group considered for analysis as well as the amount of losses in the sample (Celnik 2008; Franceschini 2012; Fu 2017).

Selective reporting

Three trials exhibited a high risk of selective reporting: Cowles 2013 and Hsieh 2020 did not include all outcomes and questionnaires provided in the trial registry, and Dettmers 2014 did not include data from a relevant questionnaire at follow-up. We considered one study to be unclear (Lee 2013). The remaining 12 were at low risk of bias: five were recorded on trial registry sites and followed their registered protocol; the others had no trial record, but the outcomes were clinically relevant for the population.

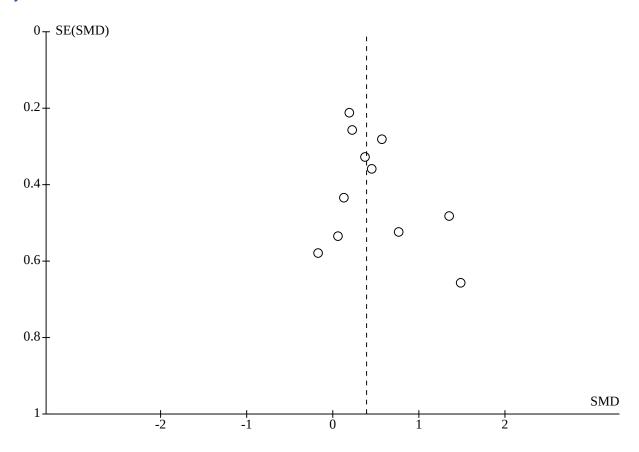
Other potential sources of bias

Fifteen trials were free of other potential sources of bias and we considered these to be at low risk. We considered one study to be at high risk of bias (Kim 2015), related to the lack of clarity about outcome assessment time.

We evaluated funnel plots for Analysis 1.1, for arm function outcome. The graph appears slightly asymmetrical, with a greater concentration of trials in the center and top of the graph. Publication bias can lead to asymmetry. Some articles may not have been found in the search strategy used. In addition, as only seven trials had a study record, it may be that non-significant results of some outcomes were not published by included trials that did not have a study registry. Furthermore, poor methodological quality can lead to spuriously inflated effects in smaller trials (Higgins 2022; Figure 4).



Figure 4. Funnel plot of comparison: 1 Action observation therapy versus control: effect on arm function, outcome: Analysis 1.1. Arm function.



Effects of interventions

See: Summary of findings 1 Action observation versus control: effect on upper limb rehabilitation after stroke

See Summary of findings 1.

We could use data from 13 trials in meta-analysis (Cowles 2013; Dettmers 2014; Ertelt 2007; Franceschini 2012; Fu 2017; Hsieh 2020; Jin-Woo 2011; Kim 2015; Kim 2016a; Kuk 2016; Mancuso 2021; Younghwa 2013; Zhu 2015). The remaining trials evaluated parameters from the same outcome that were not comparable to each other in meta-analysis (Celnik 2008; Harmsen 2015; Lee 2013).

Action observation versus control: effect on arm function

1.1 Arm function

Eleven trials (373 participants) provided postintervention assessment data for arm motor function (Cowles 2013; Dettmers 2014; Ertelt 2007; Fu 2017; Hsieh 2020; Jin-Woo 2011; Kim 2015; Kim 2016a; Franceschini 2012; Mancuso 2021; Zhu 2015). Dettmers 2014 used two scales to assess upper limb motor function: Motor Activity Log and Wolf Motor Function Test. We could only use the Wolf Motor Function Test data (the only data provided by the study authors) for meta-analysis. Ertelt 2007 used the measure of time for the Wolf Motor Function Test (lower values indicate better motor function), explaining the negative value in the analysis. We opted to use the Wolf Motor Function Test data instead of the Frenchay Arm Test because the first assesses upper limb motor function using 15 functional tasks (measuring the time to perform activities) and the score ranges from 0 to 30. The Frenchay Arm Test contains fewer tasks (5) and displays a lower range of scores (total score varies from 0 to 5).

Data used for meta-analysis of this outcome refer to Action Research Arm Test (Cowles 2013), Wolf Motor Function Test (Jin-Woo 2011; Kim 2015; Mancuso 2021), and Fugl-Meyer Assessment (Franceschini 2012; Fu 2017; Hsieh 2020; Kim 2016a; Zhu 2015).

In the meta-analysis, the impact of AO on arm function showed a small effect (standardised mean difference (SMD) 0.39, 95% confidence interval (CI) 0.17 to 0.61; P < 0.001, $I^2 = 6\%$; 11 trials, 373 participants; low-certainty evidence; Analysis 1.1).

1.2 Subgroup analysis: age

We analyzed the subgroups considering the mean age of the study participants. We compared trials in which the mean age was \geq 60 years (5 trials, 210 participants) with those in which participants were < 60 years (6 trials, 163 participants). There was no difference between the groups (P = 0.95, I² = 0%; Analysis 1.2).

1.3 Subgroup analysis: type of stroke

We compared trials in which the sample consisted only of participants diagnosed with ischemic stroke (3 trials, 101 participants) to others with samples of participants with either ischemic or hemorrhagic stroke (4 trials, 139 participants), and with a group containing trials in which the study authors did not

specify the type of stroke (4 trials, 133 participants). There was no difference between the groups (P = 0.32, $I^2 = 12.7\%$; Analysis 1.3).

1.4 Subgroup analysis: time poststroke

We conducted subgroup analyses between trials that recruited participants in the acute or subacute (6 trials, 280 participants), chronic phases (3 trials, 43 participants), and those with any phase of the condition (1 trial, 38 participants). There was no intergroup difference (P = 0.74, $I^2 = 0\%$; Analysis 1.4).

1.5 Subgroup analysis: treatment dose

We compared trials that provided \ge 1000 minutes (5 trials, 161 participants) and < 1000 minutes of therapy (6 trials, 103 participants), and observed no intergroup difference (P = 0.55, I² = 0%; Analysis 1.5).

1.6 Subgroup analysis: type of control group

Concerning the groups used to compare AO, six trials (231 participants) compared this therapy with placebo — that is, the participants watched images of geometric figures and texts, among others (Dettmers 2014; Ertelt 2007; Franceschini 2012; Fu 2017; Kim 2015; Kim 2016a). Two trials (83 participants) used conventional physiotherapy for comparison purposes (Cowles 2012; Zhu 2015), and one compared AO only with movements like those performed in the experimental group (Celnik 2008). Three trials (59 participants) used observed functional activity practice (Hsieh 2020; Jin-Woo 2011; Mancuso 2021). Thus, we decided to conduct subgroup analysis, comparing the types of control groups used. There was no intergroup difference (P = 0.57, $I^2 = 0\%$; Analysis 1.6).

Regarding the definition of the categories of control groups, we used the same terminology reported by study authors. Some study authors did not describe the conventional physical therapy protocol used. Furthermore, conventional physical therapy is a more global approach. The practice of the functional activity previously observed is more specific and, when used in the control group, makes this group more similar compared to the experimental group. Therefore, we chose to use different terms for conventional physical therapy and the observed functional activity practice.

1.7 Subgroup analysis: duration of observation

We compared trials that provided > 3 minutes of observation for each motor action (5 trials, 142 participants) with those that provided \leq 3 minutes (4 trials, 158 participants). We observed a difference between the groups (P=0.04, I²=75.3%). Trials providing > 3 minutes of intervention had a greater effect on upper limb function (Analysis 1.7).

1.8. Subgroup analysis: upper limb impairment

We performed a subgroup analysis considering upper limb impairment. We compared three groups: trials that included individuals with severe upper limb impairment (3 trials, 173 participants), trials composed of individuals with moderate upper limb impairment (3 trials, 52 participants), and those that included individuals with mild, moderate, and severe upper limb impairment (2 trials, 70 participants). The test for subgroup differences did not indicate significance (P = 0.35, $I^2 = 5.5\%$).

Arm function – sensitivity analyses: without high risk of bias for allocation concealment

Two trials showed a high risk of bias for arm function, considering the allocation concealment criterion (Fu 2017; Zhu 2015). Thus we performed sensitivity analysis, considering this bias, as proposed in the protocol. The effect increased (SMD 0.46, 95% CI 0.15 to 0.76; P < 0.001, $I^2 = 27\%$; 9 trials, 268 participants; Table 4).

Arm function – sensitivity analysis: without high risk of bias for incomplete outcome data

Additionally, two trials exhibited high risk of bias for incomplete outcome data (Franceschini 2012; Fu 2017), three for selective reporting (Cowles 2013; Dettmers 2014; Hsieh 2020), and one for other bias (Kim 2015). Thus we carried out sensitivity analysis to determine whether the effect of AO was influenced by these biases (including allocation concealment), and the effect increased (SMD 0.88, 95% Cl 0.39 to 1.36; P < 0.001, I² = 8%; 5 trials, 142 participants; Table 4).

Arm function – sensitivity analysis: home-based AO training (video therapy)

We conducted another sensitivity analysis excluding only Dettmers 2014, since it used a different methodology. In this study, treatment occurred at the participant's home (home-based AO training with video therapy), without researcher supervision. Effect size increased slightly (SMD 0.41, 95% CI 0.16 to 0.66; P = 0.001, I² = 16%; 10 trials, 335 participants; Table 4).

Arm function - sensitivity analysis: real demonstration

Cowles 2013 was the only study in which AO was demonstrated live and not via video presentation. We conducted sensitivity analysis excluding this study to observe whether there was a change in effect size, noting a slight increase from 0.39 to 0.42 (SMD 0.42, 95% CI 0.18 to 0.66, P < 0.001, $I^2 = 13\%$; 10 trials, 351 participants; Table 4).

1.9 Arm function follow-up

Four trials assessed the lasting effects of therapy on upper limb function at follow-up (Dettmers 2014; Ertelt 2007; Franceschini 2012; Hsieh 2020). However, two trials were not included in metaanalysis as they assessed only the experimental group (Dettmers 2014; Ertelt 2007).

In the Dettmers 2014 study, follow-up was carried out 6 months after the end of treatment using questionnaires (Motor Activity Log and Wolf Motor Function Test) only for the group submitted to AO (video and placebo groups). Participants from the conventional care group could not be included in follow-up analysis. Fourteen of 18 participants from the video group (AO) and 11 of 18 from the text group maintained their gains in quality and extent of movement, as measured by Motor Activity Log. Study authors did not record Wolf Motor Function Test data in follow-up.

Ertelt 2007 applied two scales in the follow-up, 8 weeks after treatment: Wolf Motor Function Test and Frenchay Arm Test. A comparison between the results of these clinical scales at the end of treatment and at follow-up in a subgroup of seven participants (experimental) showed no decline in clinical status (P = 0.63 for Wolf Motor Function Test; P = 1 for Frenchay Arm Test).

Franceschini 2012 conducted follow-up 4 to 5 months after treatment, and Hsieh 2020, 3 months. Both used Fugl-Meyer

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Assessment and were included in a meta-analysis with 93 participants. There was no effect of AO in follow-up (MD 1.06, 95% CI -8.38 to 10.50, P = 0.83, $I^2 = 17\%$; 2 trials, 93 participants; Analysis 1.9).

Action observation versus control: effect on hand function

2.1 Hand function

Six trials assessed motor hand function after intervention (Dettmers 2014; Franceschini 2012; Hsieh 2020; Kim 2016a; Kuk 2016; Mancuso 2021), but Dettmers 2014 presented the data in graphs, making it difficult to establish the average and standard deviation (SD) and, even after contacting the author, we could not obtain the data related to this outcome.

We pooled the data presented by five study authors (Franceschini 2012; Hsieh 2020; Kim 2016a; Kuk 2016; Mancuso 2021; 178 participants). All used the Box and Block Test to assess this outcome. The impact of AO on hand function showed a large effect (mean difference (MD 2.76, 95% CI 1.04 to 4.49; P < 0.001, $I^2 = 0\%$; 5 trials, 178 participants; low-certainty evidence; Analysis 2.1).

2.2 Subgroup analysis: age

In subgroup analysis considering age, there was no intergroup difference between patients with a mean age \ge 60 years (2 trials, 122 participants) with those in which participants were < 60 years (3 trials, 56 participants) (P = 0.94, l² = 0%; Analysis 2.2).

2.3 Subgroup analysis: time poststroke

In subgroup analysis of the effect of therapy on hand function, there was no intergroup difference in stroke phases: acute/subacute (3 trials, 144 participants) and chronic phase (2 trials, 34 participants) (P = 0.77, $I^2 = 0\%$; Analysis 2.3).

2.4 Subgroup analysis: duration of observation

Regarding the duration of observation, there was no difference between subgroups that provided > 3 minutes of observation for each motor action (1 trial, 22 participants) with those that provided \leq 3 minutes (4 trials, 156 participants) (P = 0.71, I² = 0%; Analysis 2.4).

Hand function – sensitivity analysis: without high risk of bias for incomplete outcome data

We conducted sensitivity analysis for hand function, disregarding Franceschini 2012, which showed high risk of bias for incomplete outcome data. The effect of therapy decreased (MD 2.59, 95% CI 0.81 to 4.37; P < 0.001, $I^2 = 0\%$; 4 trials, 88 participants; Table 5).

Hand function – sensitivity analysis: without high risk of bias for selective reporting

We performed another sensitivity analysis for hand function excluding Hsieh 2020, which showed high risk of bias for selective reporting and Franceschini 2012, which exhibited high risk of bias for incomplete outcome data. The effect of therapy decreased (MD 2.61, 95% CI 0.81, 4.41; P < 0.001, $I^2 = 0\%$; 3 trials, 76 participants; Table 5).

2.5 Hand function – follow-up

Two trials with 93 participants examined the effects of AO on hand function at the end of the follow-up, 4 months to 5 months

(Franceschini 2012), and 3 months after the intervention (Hsieh 2020). This analysis reported no evidence of effect of AO on hand function in follow-up (MD 5.19, 95% CI -1.82 to 12.21; P = 0.15, $I^2 = 0\%$).

Action observation versus control: effect on ADL

3.1 Dependence on ADL

Seven trials (302 participants) presented dependence in ADL as an outcome using different scales: Functional Independence Measure (Franceschini 2012; Hsieh 2020; Mancuso 2021); Modified Barthel Index (Fu 2017; Kim 2016a; Younghwa 2013); and Barthel Index (Zhu 2015). There was no effect: SMD 0.37, 95% CI -0.34 to 1.08; P = 0.31, I² = 87%; very low-certainty evidence; Analysis 3.1).

3.2 Subgroup analysis: age

There was no difference in subgroup analysis, considering the mean age of ≥ 60 years (4 trials, 205 participants) or < 60 years (3 trials, 97 participants) (P = 0.50, I² = 0%; Analysis 3.2).

3.3 Subgroup analysis: type of stroke

Subgroup analysis did not find any difference between subgroups of patients with ischemic stroke (2 trials, 85 participants) and subgroups with ischemic and hemorrhagic stroke (4 trials, 156 participants) (P = 0.21; $I^2 = 35.9\%$; Analysis 3.3).

3.4 Subgroup analysis: time poststroke

There was a difference in subgroup analysis regarding post-stroke time, considering the acute/subacute (5 trials, 258 participants) and chronic phase subgroups (2 trials, 44 participants). Patients in the acute/subacute phase had a greater impact of AO on the dependence on ADL; however, with considerable heterogeneity (SMD, 0.85, 95% CI 0.23 to 1.46, P < 0.001, I² = 91.2%; Analysis 3.4).

3.5 Subgroup analysis: treatment dose

The comparison between patients who underwent \ge 1000 minutes (3 trials, 107 participants) and < 1000 minutes of therapy (4 trials, 195 participants) was not significant for this outcome (P = 0.85, I² = 0%; Analysis 3.5).

3.6 Subgroup analysis: type of control group

Three subgroups were considered to analyze the types of control groups: placebo (3 trials, 165 participants); conventional physical therapy (1 trial, 61 participants); and observed functional activity practice, including task-oriented training, range of motion exercises, reaching, and object manipulation (3 trials, 76 participants). In Mancuso 2021, the control and experimental groups underwent conventional physical therapy; thus, this study was classified in the functional activities subgroup. There was no difference between the three subgroups (P = 0.29, $I^2 = 20\%$).

3.7 Subgroup analysis: duration of observation

We compared trials that provided > 3 minutes of observation for each motor action (3 trials, 105 participants) with those that provided \leq 3 minutes (3 trials, 136 participants). There was no difference (P = 0.70, I² = 0%; Analysis 3.7).

3.8 Subgroup analysis: upper limb impairment

Subgroup analysis found no difference between groups of patients with 1) severe impairment (2 trials, 151 participants), 2) moderate



impairment (2 trials, 36 participants), and 3) mild, moderate, and severe upper limb impairment (1 trial, 32 participants) (P = 0.22; $I^2 = 34.6\%$; Analysis 3.6).

Sensitivity analysis: without high risk of bias for allocation concealment

We performed sensitivity analysis for the dependence on ADL outcome, excluding trials with a high risk for allocation concealment (Fu 2017; Zhu 2015). The effect of the therapy increased (SMD 0.24, 95% CI -0.85 to 1.32; P = 0.67, $I^2 = 90\%$; 5 trials, 188 participants; Table 6).

Sensitivity analysis: without high risk of bias for incomplete outcome data

Two trials that assessed this outcome exhibited high risk of bias for incomplete outcome data (Franceschini 2012; Fu 2017) and one for selective reporting (Hsieh 2020). We carried out sensitivity analysis and observed that the effect of therapy increased (SMD 0.52, 95% CI -0.82 to 1.86; P = 0.452, l^2 = 92%; 4 trials, 145 participants; Table 6).

Sensitivity analysis: removing peripheral trials

In meta-analysis related to this outcome, CIs did not overlap on the forest plots, which were quantified, showing considerable heterogeneity (I² = 87%). After removing the peripheral trials, heterogeneity decreased, becoming low (I² = 26%); however, no effect of therapy was observed (SMD 0.13, 95% CI -0.25 to 0.51; P = 0.51, I² = 26%; 3 trials, 165 participant; Table 6).

3.9 Dependence on ADL - follow-up

Two trials conducted a follow-up for this outcome with 93 participants (Franceschini 2012; Hsieh 2020). There was no evidence of effect from the intervention (MD -0.77, 95% CI -5.43 to 3.88; P = 0.74; $I^2 = 0\%$; Analysis 3.9).

Action observation versus control: effect on motor performance

Three trials assessed motor performance (83 participants), but the fact that different kinematic variables were used precluded comparing them in analysis. One study assessed motor performance using the variable 'movement time' and was the only one that used the patient to make the video. Reaching movements from the unaffected arm were videotaped and mirrored. Thus, the filming and video demonstration methodology resembled the mirror therapy protocols. The authors found that the decrease in movement time was significantly more for the AO group compared with the control group (P = 0.026) (Harmsen 2015).

In Kim 2015, average velocity, trajectory ratio, and motion angle were the variables selected to measure the kinematic patterns of the upper limb before and after therapy. There was an improvement between pre and postintervention average velocity and trajectory ratio values in the experimental group. However, there was no intergroup difference.

One study assessed the number of times the drinking task was performed in 1 minute (Lee 2013). The experimental group watched a video of the task, the motor practice group performed the action, the combined group watched a video of the task and performed the action, and the control group performed neither AO nor physical execution. After the intervention, the number of drinking motions increased the most in the combination group.

Action observation versus control: effect on cortical activation

Four trials evaluated cortical activation after AO. Ertelt 2007 applied fMRI to analyze the effects of AO on motor system reorganization, using an independent sensorimotor task consisting of handling an object. The control group (placebo observation followed by hand and arm movements) showed practically no change in brain activity between pretest and post-test treatment. By contrast, the experimental group exhibited numerous differences between pretreatment and post-treatment brain activations on sensorimotor network (supplementary motor area, bilateral ventral premotor cortex, bilateral superior and inferior parietal areas, and bilateral cerebellum).

Celnik 2008 measured motor cortical excitability, recording the MEP amplitudes of muscles mediating movements in the trained (MEP agonist) and baseline (MEP antagonist) directions. The authors used two approaches to observe the action to be performed later: congruent (same direction) and incongruent (opposite direction). At postintervention, the MEP agonist showed a slight increase while the MEP antagonist decreased for the congruent group. This difference in excitability is reflected by a change in the MEP post/ preintervention ratio. Thus the study authors found that observing another participant performing training motions in the same direction and in phase with physically trained individuals enhanced motor memory formation compared with physical training alone.

One study used the evaluation of MEP, which was determined using a transcranial magnetic stimulator (Fu 2017). The study author calculated latency, amplitude, and central motor conduction time (CMCT - difference between cortex and spinal latency), finding an increase in amplitude and decrease in latency and CMCT. These changes were higher in the experimental than in the control group.

The MEP results reported by Celnik 2008 and Fu 2017 could not be pooled in meta-analysis, because data from the first period of the cross-over were not available for this outcome in the Celnik 2008 study.

Kuk 2016 analyzed cortical activation using EEG data from poststroke participants, only in the experimental group, in each session of AO, while they were watching the video clip. Sequential executions of the observed action after AO were not EEGmonitored. The results revealed selective activation of the mirrorneuron system, with the middle frontal gyrus less active.

Action observation versus control: effect on quality of life

4.1 Quality of life

Quality of life was assessed in three trials (30 participants) using the Stroke Impact Scale. We were unable to obtain data (mean and SD) from the Dettmers 2014 study to carry out analysis. However, the video group (video AO) improved on Stroke Impact Scale score (P < 0.001) from pre to post-treatment. We conducted a metaanalysis with the remaining two trials (Ertelt 2007; Hsieh 2020), with no evidence of benefit or detriment from the intervention (MD 5.52, 95% Cl -30.74 to 41.78; P = 0.77, I² = 88%; very low-certainty evidence; Analysis 4.1).

4.2 Quality of life follow-up

In the Dettmers 2014 study, 14 of 18 participants from the video group and 11 of 18 from the text group completed the questionnaires after 6 months. Quality of life subjective self-



assessment improved further in the video group, but not in the text group (P value not reported).

Ertelt 2007 assessed the long-term effects of AO only in the experimental group (seven participants). Follow-up was conducted 8 weeks after the intervention and comparisons between the results at the end of treatment and at follow-up in the subgroup of seven participants showed no decline in clinical status (P = 0.73).

In the Hsieh 2020 study, at 3-month follow-up, Stroke Impact Scale scores had been maintained in both groups, indicating retention effects.

Action observation versus control: adverse effects

This outcome was investigated in two trials (16 participants). The visual analogue scale was used in the Celnik 2008 trial to measure attention and fatigue levels, with scores varying from 1 to 7, where 1 equaled the worst response and 7 equaled the best response (least fatigue or best attention). The author reported that attention and fatigue level did not influence the findings, since there was no difference between attention (P = 0.17) or fatigue scores (P = 0.40) during the different intervention sessions. One participant did not conclude the experimental protocol due to a headache caused by TMS, not by the therapy.

In the Cowles 2012 study, participants were monitored during the experiment in order to detect the overuse syndrome. To that end participant accounts of upper limb pain — either verbal or behavioral, (e.g. grimacing, postural guarding) — and decreases of at least 2 measurement levels in the Motricity Index were recorded. No adverse events occurred in either group. Both groups (AO followed by motor practice and conventional physiotherapy) improved their Motricity Index score between baseline and outcome, but there was no intergroup difference.

DISCUSSION

Summary of main results

The aim of this review was to evaluate the effect of AO on motor function and upper limb motor performance in poststroke patients. We included 16 trials with 574 participants in this review. Overall, the certainty of the evidence for outcomes was low to very low. The main results are presented in Summary of findings 1.

Overall, AO increased the arm function of individuals after stroke compared with other physiotherapy interventions or placebo. However, the effect size is small, and, considering the certainty of the GRADE evidence, AO may improve the arm motor function of patients with stroke (low certainty). The gain in arm motor function at the end of AO did not persist at follow-up. We performed arm function analysis using multiple instruments. Accordingly, we were unable to consider the clinically-important difference for this outcome.

Furthermore, the effect of therapy was not influenced by issues such as age, type of stroke, poststroke time, treatment dose, and type of control group. The evidence suggests that AO results in little to no difference in arm motor function in subgroups considering upper limb impairment. The use of > 3 minutes of observation, the supervision of a therapist/researcher, and the observation of a video presentation (not a real demonstration) increased the effect of the intervention. The effect of AO on hand function was large. However, it does not appear to be clinically relevant, given that the minimum detectable difference is six blocks/minute for the Box and Block Test (Chen 2009). Furthermore, the evidence was considered low according to GRADE, that is, our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect. The effect does not appear to be influenced by issues related to the stroke or the intervention.

There was no effect of AO on the performance of ADL when compared to control groups. According to GRADE, we considered the certainty of the evidence to be very low that AO may not improve this outcome in stroke patients. However, this therapy has a higher impact on patients in the acute/subacute phases in performing ADL, although with considerable heterogeneity. Younghwa 2013 results may have caused considerable heterogeneity because the control group protocol included aspects not addressed in the experimental group. These aspects are associated with larger effect sizes, such as distributed practice, feedback, random practice, and the use of clear functional goals (Timmermans 2010). The motor practice of the experimental group could address these aspects. Then, we can say that there was a discrepancy between the protocols used in the control and experimental groups. We observed no difference in the effect of AO on the performance of ADL considering upper limb impairment. That is, the fact that it is an applicable therapy for patients with a severe upper limb impairment will not necessarily promote the motor and functional improvement of this individual.

Only two trials were included in the quality of life meta-analysis. There was no effect but considerable heterogeneity. According to GRADE, there is no evidence of benefit or detriment from AO in the quality of life of stroke patients.

We were unable to pool the results of motor performance, cortical activation, and adverse effects.

Despite being the outcome of a number of trials, the variables selected to measure motor performance were different between trials precluding intergroup comparison performing the metaanalysis. A decline in movement time and rise in the number of drinking tasks completed in 1 minute were the motor performance gains obtained by the AO compared to control therapies. The following variables did not improve in the experimental intervention: average velocity, trajectory ratio, motion angle, angular difference, and angular dispersion. Despite the positive findings for some of the variables, there is not enough evidence to confirm that AO has an effect on motor performance.

Considering cortical activation and changes in the nervous system, trials demonstrated that AO: 1) improves motor performance by reactivation of motor areas containing the AO/action execution matching system, 2) could improve motor nerve excitability, 3) enhanced motor memory formation, and 4) might alter cortical activation patterns.

Only two trials considered adverse effects, which could not be pooled in the meta-analysis. Nevertheless, these trials showed no adverse effects.

One of the causes of heterogeneity in this review refers to the different approaches to therapy application. The content offered in the AO was heterogeneous among the trials. In most trials, the

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content observed was functional tasks, composed of one to three motor actions, with or without increasing complexity of the tasks throughout training. Range of motion exercises, and performing active movements such as thumb movements were also offered for observation.

The time of AO was quite variable between trials, ranging from 1 minute to 10 minutes for each motor action, task, or movement observed. Adding a total observation time of 4 to 36 minutes per session.

The motor action performed after observing the action was the practice of the observed movement or task and ranged from 2 to 30 minutes of training for each motor action, totaling a minimum of 5 minutes and a maximum of 36 minutes of exercise per session.

Other causes of heterogeneity can be cited, such as duration of the treatment period, treatment frequency, and time points measured.

Overall completeness and applicability of evidence

We found only 16 RCTs that tested the effectiveness of AO; furthermore, the trials had a relatively small sample size. Since none of the trials included only participants with hemorrhagic stroke, we were unable to form a subgroup with only this population of participants. The mean age of the participants varied widely.

The most common outcome was arm motor function, and few trials assessed other important outcomes for people with stroke, such as hand motor function, dependence on ADL, quality of life, and adverse effects. Although cortical activation is not an outcome directly expressed by patients, it is particularly important to researchers and clinicians since it provides evidence that motor gains result from better brain activation. There is also a need to better monitor adverse effects such as pain, fatigue, and attention deficit, in order to determine the effectiveness of AO — that is, whether the benefits outweigh any adverse effect.

Nearly all the trials used motor practice after the observation of a task. This restricts the applicability of therapy for people with some degree of motricity in the affected upper limb; and although some trials included participants in the acute phase, it was not possible to observe the effect of AO on individuals who were unable to perform the movements. Therefore, it would be advisable to test whether applying AO alone, with no motor practice, would show any benefit for totally paralyzed patients. Moreover, the outcome of AO alone versus followed by motor practice should be compared, in order to determine whether the gains derive only from motor practice that occurs after observation.

Certainty of the evidence

The number of trials (only 13 trials in meta-analyses) and sample sizes included in this review were small. Some trials showed a high risk of bias for allocation concealment, incomplete outcome data, selective reporting, and other types of bias related to lack of information. We observed unclear risk of bias in many trials with important criteria such as randomisation, allocation concealment, and blinded researchers, resulting from poor reporting and lack of clarification from the authors. The effect estimate was also inaccurate. Nevertheless, most of the results were consistent (low heterogeneity) and exhibited good external validity. According to the GRADE system, the certainty of evidence remains 'low' or 'very low' for most of the outcomes (Summary of findings 1).

Potential biases in the review process

Two review authors independently reviewed the trials, obtaining and extracting data, with a third review author available to resolve disagreements as needed, thereby minimizing bias. We are confident that we have identified all relevant trials; however, there is a small possibility that we failed to identify additional (published or unpublished) papers. Moreover, some analyses could not determine the effect of AO on adverse effects, since not all study authors could provide the required data. Another limitation of this review is that some of the trials had methodological shortcomings or even did not report important procedures in the manuscripts, such as allocation concealment, incomplete outcome data, and selective reporting. These biases can lead to underestimation or overestimation of the true intervention effect, according to the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2022).

Agreements and disagreements with other studies or reviews

We found four systematic reviews on AO for stroke patients. Sarasso 2015 analyzed trials investigating the effects of this therapy on different motor abilities in diseases such as stroke (upper and lower limb), Parkinson's disease, cerebral palsy, and postsurgical orthopedic conditions, in 663 people (20 RCTs). This search included articles up to July 2015. Six trials involving stroke patients with upper limb impairment were also included in our review (Cowles 2012; Ertelt 2007; Franceschini 2012; Harmsen 2015; Kim 2015; Lee 2013). The conclusion of the Sarasso 2015 review suggested the efficacy of AO in improving motor functions both in neurological and orthopedic diseases. The study did not cover all the literature found in our review (only 6 of the 16 trials included) and did not pool the data in meta-analyses. Despite this, Sarasso 2015 obtained similar conclusions regarding the overall effect of the therapy, but without estimating the effect size.

Peng 2019 summarized trials (17 RCTs, 600 participants) that investigated the effectiveness of AO in the arm and hand motor function, walking ability, gait performance, and ADL in stroke patients. The search included articles up to January 2019. Seven trials were also included in our review (Cowles 2012; Ertelt 2007; Franceschini 2012; Fu 2017; Kim 2015; Kim 2016a; Zhu 2015). Compared to control treatments, AO had a moderate effect size on arm and hand motor function and a moderate to large effect size on ADL performance. Peng 2019 concluded that AO is an effective approach for stroke patients to improve arm and hand motor function and daily activity performance. Effect sizes were different from our study, as was the outcome for ADL. This increase in effect size may have been due to: 1) only seven of the 16 articles included in our review were included, and these trials had more favorable results for the studied therapy; 2) the inclusion of Sale 2012 as a different study than Franceschini 2012 generated duplication of results (Sale's sample is a part of Franceschini's sample).

Zhang 2019 analyzed trials (7 RCTs, 276 participants) investigating the effects of AO on improving upper limb motor functions in people with stroke. The search included manuscripts up to March 2019. Seven trials were also included in our review (Cowles 2012; Ertelt 2007; Franceschini 2012; Fu 2017; Kim 2015; Kim 2016a; Zhu

2015). The results revealed an effect favoring the therapy over the control. Zhang 2019 concluded that AO is an effective method for improving upper limb motor function after stroke, corroborating our findings, despite having included few trials.

Ryan 2021 summarized trials (36 RCTs, 1405 participants) that investigated the effects of AO in amputees, orthopedic disorders, dementia, cerebral palsy, multiple sclerosis, Parkinson's disease, and stroke (upper and lower limb). The search included manuscripts up to July 2020. Seven trials involving AO therapy for upper limb impairment of stroke patients were also included in our review (Cowles 2012; Franceschini 2012; Fu 2017; Hsieh 2020; Kim 2015; Kim 2016a; Zhu 2015). In meta-analyses, they observed an effect for hand function; and no effect for arm function. However, results related to Fugl-Meyer Assessment and Modified Barthel Index are not comparable, and the authors included trials that performed lower limb training. The conclusion of Ryan 2021 suggested that the benefit of incorporating AO into rehabilitation programs is strongly supported in stroke patients. Our findings do not support such a strong recommendation.

We believe that due to our more recent and comprehensive search strategy, we identified more trials evaluating the effects of AO on upper limb function in poststroke patients than other systematic reviews already published.

AUTHORS' CONCLUSIONS

Implications for practice

The results of this review show low-certainty evidence that AO may improve arm motor function of poststroke patients compared with any control group. This effect seems to be increased by adopting observations > 3 minutes, professional supervision, and video presentations. AO improves hand motor function, but the evidence showed that the improvements were not clinically relevant.

There is no convincing evidence that the intervention affects dependence on ADL, motor performance, cortical activation, or quality of life after stroke. There is little evidence in the literature of any adverse effects.

The certainty of GRADE-based evidence in this review varied from very low to low, due to the high risk of bias and small sample sizes. This means that our confidence in the estimated effect is limited and that the true effect may be substantially different. Thus, AO may improve arm and hand motor function. However, confidence in this effect is limited, mainly for arm function, but may be confirmed in future clinical trials.

Implications for research

To improve the certainty of the evidence (GRADE) regarding the effect of AO on upper limb rehabilitation in people with stroke, new RCTs should be conducted, with greater methodological rigor in order to reduce the risk of bias, using a larger sample and standardized outcome assessment instruments to increase consistency. A guideline specifying the main evaluation tools for each outcome would be useful in standardizing the findings, given the variety of existing instruments. The Standardizing Measurement in Arm Rehabilitation Trials (SMART) Toolbox provides a selection of measures of importance to stroke survivors, carers, clinicians, and researchers in future RCTs, whilst considering psychometric properties, the ICF classification, and outcomes important to stroke survivors, carers, and clinicians (Millar 2021).

Furthermore, trials have exhibited methodological problems, such as lack of clarity, which hinders assessment of bias and quality. In order to minimize this shortcoming, research should follow the CONSORT recommendations (Schulz 2010), which contain guidelines for writing RCTs in the area of health. Data presentation should also be clear to allow easy interpretation and pooling of quantitative data.

It is important that the authors of future trials characterize their samples in more detail in order to attest the applicability of AO in specific situations, according to the etiology and phase of stroke, upper limb impairment, and brain hemisphere affected. Other information regarding the form of therapy application is also necessary for assessing its effects, such as whether therapy was administered through videos or live demonstrations, which perspective was used, treatment setting, and dose. Trials should include follow-up assessment to determine if the effects are longlasting.

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CHARACTERISTICS OF STUDIES

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* Indicates the major publication for the study

Celnik 2008	
Study characteristic	S
Methods	Randomized cross-over trial
Participants	Methods of recruitment were not reported by authors
	9 participants in a cross-over design with 3 arms:



Bias	Authors' judgement	Support for judgement	
Risk of bias			
Notes	The authors declared r	no potential conflicts of interest.	
	1. Corticomotor excita	bility: MEP of agonist and antagonist muscles	
Outcomes	Outcomes recorded before and after intervention		
	Sessions were 30 minu	tes, 3 blocks of 10 minutes each separated by 2-minute rest	
		incongruent AO group : observed a video displaying thumb movements in a di It physically practiced simultaneously	
		congruent AO group : observed a video displaying thumb movements in the physically practiced simultaneously	
Interventions	Physical training group: performed voluntary thumb movements		
	Stroke phase: chronic		
	Stroke details: not rep	ported by authors	
	Mean (SD) age: not rep	orted by authors	
	Exclusion criteria: the	re were no exclusion criteria described in the study	
	Inclusion criteria: stro	ke patients with single unilateral cortical or subcortical lesions	
		g and congruent AO group; g and incongruent AO group.	
	1. 9 in physical training		

Random sequence genera- tion (selection bias)	Low risk	Quote: "The allocation was generated randomly using Excel creating a random session order for each of the subjects and conditions."
Allocation concealment (selection bias)	Unclear risk	The author did not make it clear whether there was allocation concealment.
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Quote: "The outcome measures were analyzed by a blinded individual who did not participate in the data acquisition."
Incomplete outcome data (attrition bias) All outcomes	High risk	One participant could not complete the protocol due to TMS-related headache. The reasons were provided, but the exclusion of 1/9 participants in a cross-over design, with 3 arms, may have affected the reliability of study re- sults.
Selective reporting (re- porting bias)	Low risk	There is no trial registration, but the outcomes are significant, and there is no selective reporting within the study.
Other bias	Low risk	No other potential risk of bias was found.



Cowles 2013

Study characteristics					
Methods	Randomized, controlled, observer-blind efficacy trial				
Participants	All participants were recruited from an inpatient stroke unit				
	29 participants:				
	 15 in experimental § 14 in control group. 				
	an intact premotor are in a substantially paret able to produce a grip	Its who had suffered a stroke between 3 and 31 days before recruitment but with a (ability to encode observed human actions); able to produce some movement ic upper limb as assessed by a score of at least 18 on the Motricity Index but un- force of more than 65% of that of the ipsilesional side; free from observable con- movement deficits attributable to pathology other than stroke			
	Exclusion criteria : patients who had no ability to imitate action with their ipsilesional limb. This ability was assessed by the research therapist who performed an upper limb activity 3 times, and potential participants were asked to observe and then perform the activity. This assessment was digitally filmed and saved onto a computer. An independent assessor viewed the recordings and judged the accuracy of imitation.				
	Mean (SD) age: 77.2 (SD 10.4) years				
	Stroke details: not reported by authors				
	Stroke phase: acute				
Interventions	Experimental group : additionally watched functional tasks with intent to imitate for 1 to 2 minutes; and then for 4 to 6 minutes performed the activity simultaneously with the therapist				
	Control group: received conventional physical therapy				
	Sessions were 2 × 30-minute sessions, with 10-minute rest, for 15 working days				
Outcomes	Outcomes recorded at	baseline and within 3 working days of the end of intervention			
	 Ability to voluntarily contract paretic muscle: Motricity Index Upper limb motor function: Action Research Arm Test 				
Notes	The authors declared r	no potential conflicts of interest.			
Risk of bias					
Bias	Authors' judgement	Support for judgement			
Random sequence genera- tion (selection bias)	Low risk	Group allocation was computer generated			
Allocation concealment (selection bias)	Low risk	Details of group allocation were placed into sequentially-numbered, opaque, sealed envelopes			
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding			

Cowles 2013 (Continued)		
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	All measures were made by an assessor blinded to treatment allocation
Incomplete outcome data (attrition bias) All outcomes	Low risk	29 participants were randomized. Outcome measures were completed on 13 participants in the control group and 9 in the experimental group (22 participants). Reasons for attrition were provided: moved out of area (one in control group and 4 in experimental group); withdrew from trial (1 in experimental group); medically unwell (1 in experimental group); 1 of the participants in the control group failed to fully comprehend the instructions for the Motricity Index at baseline so this score was missing. Reasons for missing outcome data unlikely to be related to true outcome
		ed in the analysis, and imputation was not used
Selective reporting (re- porting bias)	High risk	The study protocol is available but not all of the study's prespecified pri- mary outcomes have been reported. Registered on the Current Controlled Trials Database — ISCRTN 51553998: www.controlled-trials.com/ISR CTN51553998/51553998
Other bias	Low risk	No other potential risk of bias was found.

Dettmers 2014

Study characteristics	
Methods	Pilot randomized controlled trial
Participants	Participants were recruited from 2 hospitals
	56 participants:
	 19 in video group; 19 in text group; 18 in usual care group.
	Inclusion criteria : minimal function of the hand (ability to grip a small object and to release it (extension against gravity at wrist = 20° and at metacarpophalangeal and interphalangeal joints of each of the fingers = 10°), sufficient language abilities to complete the study's questionnaires, and sufficient cognitive ability to understand and follow instructions for training, distance to the participant's home restricted to 300 km to allow for a follow-up examination
	Exclusion criteria : prominent cognitive deficits (Mini Mental State Exam < 24), major depression (Beck Depression Index > 5), major aphasia (Token Test), apraxia (Florida Apraxia Screening Test), neglect (Albert's Neglect Test), hemianopia, prior infarct, leucoencephalopathy and age below 18 or over 75 years, severe psychiatric disease, severe pulmonary or cardiovascular disease, epilepsy, severe joint deformity (neurological examination) and severe pain
	Mean (SD) age: video group: 62.79 (no SD reported), text group: 53.89 (14.74), usual care group: 58.83 (11.25).
	Stroke details:
	 video group: 6 embolic, 4 lacunar, 2 hemodynamic, and 7 unknown etiology; text group: 9 embolic, 4 lacunar, 1 hemodynamic and 5 unknown etiology; usual care group: 7 embolic, 5 lacunar, 2 hemodynamic and 4 unknown etiology.

Dettmers 2014 (Continued)	Stroke phase: acute, subacute, and chronic			
Interventions	Video group : home-based training. Participants received a DVD displaying 10 object-related motor tasks of varying difficulty, each lasting 5 minutes and were requested to imitate the motor tasks, time not reported			
	Text group : home-based training. Participants performed the same tasks with written instructions without observation/imitation			
	Usual care group: discharged without specific homework			
	Participants were instructed to train 1 hour per day for 6 weeks (video and text groups)			
Outcomes	Outcomes recorded at baseline, postintervention, and at 6 months from treatment conclusion			
	1. Upper limb motor function: Wolf Motor Function Test – time and quality of movements			
	2. Hand function: Nine Hole Peg Test			
	3. ADL: Motor Activity Log – quality of movement and amount of use			
	4 Quality of life: Strake Impact Scale			

4. Quality of life: Stroke Impact Scale

Notes

The authors did not provide any declaration of interests.

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Randomization was done, but there was no information of how it was done
Allocation concealment (selection bias)	Unclear risk	There was no information about allocation concealment
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome was not likely to be influenced by lack of blind- ing
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Wolf Motor Function Test assessment was performed by a blind physiothera- pist (low risk). There was no information about blinding other assessment in- struments
Incomplete outcome data (attrition bias) All outcomes	Low risk	1 participant was excluded during the intervention because of a recurrent stroke (video group). 1 participant from the text group was unwilling to come to the re-evaluation. Another 2 participants refused to come to re-evaluation (usual care group). Nevertheless, the reasons were provided and they had no direct relationship with the outcomes studied, and there was no substantial imbalance in final number of participants in each group considered for analy- sis.
Selective reporting (re- porting bias)	High risk	Considering the description of the methodology, the follow-up investigation was scheduled 12 months after the training, using Motor Activiy Log and Wolf Motor Function Test. However, the results concerning the follow-up showed that the data were assessed 6 months after the end of the treatment and used as questionnaires the Stroke Impact Scale and Motor Activity Log, when the data of Wolf Motor Function Test were not available.
Other bias	Low risk	No other potential risk of bias was found.

Action observation for upper limb rehabilitation after stroke (Review)

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Ertelt 2007

Study characteristics		
Methods	Randomized clinical trial	
Participants	Participants were recruited from the local Rehabilitation Center	
	16 participants:	
	 8 in experimental; 8 in control. 	
	Inclusion criteria : confirmed diagnosis of a first-ever ischemic stroke in the territory of the medial cerebral artery, sustained more than 6 months prior to entering the study	
	Exclusion criteria : patients older than 76 years, with hemorrhagic stroke or ischemic lesions in the territory of posterior or anterior cerebral artery, impaired level of consciousness (confusion, stupor, coma), severe to moderate aphasia, anosognosia or neglect, amnesia or dementia, depression	
	Mean (SD) age: experimental group: 57.16 (8.73) years and control group: 55.40 (10.77) years	
	Stroke details: first ischemic stroke in the territory of the medial cerebral artery	
	Stroke phase: chronic	
Interventions	Experimental group: ("action observation therapy"): carefully watched video sequences containing daily life hand and arm actions for 6 minutes that were followed by repetitive practice of the observed actions for another 6 minutes using the same objects as shown in the video film	
	Control group : matched the experimental treatment with the exception that the participants watched sequences of geometric symbols and letters instead of action sequences	
	Sessions were 90 minutes, 18 rehabilitation sessions on consecutive working days	
Outcomes	Outcomes recorded at baseline, postintervention, and 8 weeks after the end of intervention	
	1. Upper limb function: Frenchay Arm Test, Wolf Motor Function Test	
	 Quality of life: Stroke Impact Scale Functional imaging: fMRI measurements 	
	5. Functional imaging.	
Notes	The authors did not provide any declaration of interests.	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Randomization was done, but no information of how it was done
Allocation concealment (selection bias)	Unclear risk	Authors did not report if there was allocation concealment
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding

Ertelt 2007 (Continued)

Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Authors did not report if there was blinding of outcome assessment
Incomplete outcome data (attrition bias) All outcomes	Low risk	There was no exclusion of participants after randomisation. fMRI measure- ments were analyzed in 7 participants in the experimental group and 6 partici- pants in the control group.
Selective reporting (re- porting bias)	Low risk	There is no trial registration, but the outcomes were significant, and there was no selective reporting within the study.
Other bias	Low risk	No other potential risk of bias was found.

Franceschini 2012

Study characteristic	s
Methods	Randomized controlled observer-blind trial
Participants	Eligible hemiparetic stroke survivors from 13 rehabilitation centers were recruited
	Franceschini 2012 paper:
	102 participants:
	 53 in experimental; 49 in control.
	Inclusion criteria : participants with first-ever stroke, enrolled 30 days (±7) after the event onset with is- chemia or primary hemorrhage, right-handed prior to stroke
	Exclusion criteria : posterior circulation infarction, subarachnoid hemorrhage, severe forms of ne- glect and anosognosia (number of errors in Bell Barrage test ≥ 15), impaired comprehension (Token test score ≤ 17), history of endogenous depression or serious psychiatric disorders, and severe visual deficits (restricting the access to visual stimuli)
	Mean (SD) age: experimental group: 67.0 (12.4) years; control group: 65.7 (11.9) years
	Stroke details : etiology – control group: 9 hemorrhagic, 40 ischemic; experimental group: 16 hemor- rhagic, 37 ischemic. Lesion side: control group: 18 right, 24 left; experimental group: 22 right, 26 left
	Stroke phase: acute
	Sale 2014 paper:
	67 participants:
	 33 in experimental; 34 in control.
	Inclusion criteria : moderate-to-severe upper limb paresis, first-ever ischemic stroke, 30 days (±7) after the event, right handed prior to stroke, unilateral brain lesions
	Exclusion criteria : posterior circulation infarction, subarachnoid hemorrhage, severe forms of neglect and anosognosia, impaired comprehension or dementia, history of endogenous depression or serious psychiatric disorders, severe visual deficits, bilateral motor impairment, severe sensory deficits in the paretic upper limb, refusal or inability to provide informed consent, other concomitant severe medical problems

Franceschini 2012 (Continued)	Mean (SD) age: 66.5 (12.7) years			
	Stroke details: ischemic			
	Stroke phase: acute			
Interventions	Experimental group : observation of 1 daily routine task (actions) carried out with the upper limb, each action consisted of 3 different meaningful motor sequences (3 minutes each). After observing the video, the participant should perform, with their paretic upper limb, the same movement (2 minutes each), with help when needed			
	Control intervention: observation of 5 static images displaying objects, without any animal or human being (different 3-minute sequences). Then, the participants had to perform limb movements for 2 minutes according to a standard sequence, simulating those performed by the other group			
	Sessions were 2 × 15 minutes, daily (with 60-minute interval), 5 days/week, for 4 consecutive weeks (in addition to 3 hours of standard rehabilitation)			
Outcomes	Franceschini 2012 paper:			
	Outcomes recorded at baseline, postintervention and at 4 to 5 months from treatment conclusion			
	1. Upper limb motor function: Fugl-Meyer Assessment - upper limb items, Frenchay Arm Test, and Box and Blocks Test			
	2. Muscle tone: Modified Ashworth Scale			
	3. Functional Independence: Functional Independence Measure - motor items			
	Sale 2014 paper:			
	Outcomes recorded at baseline, postintervention and at 4 to 5 months from treatment conclusion			
	1. Upper limb motor function: Fugl-Meyer Assessment - upper limb items and Box and Block Test			
Notes	There are 2 publications for this study. The authors declared no potential conflicts of interest.			

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Central randomisation
Allocation concealment (selection bias)	Low risk	Participants and investigators enrolling participants could not foresee assign- ment because central allocation was used to conceal allocation
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	All assessments were performed by trained professional not involved in the re- search treatment and blind to group allocation
Incomplete outcome data (attrition bias) All outcomes	High risk	12 participants were excluded postrandomization, 5 from experimental group and 7 from control group. Reasons were not reported. There was an imbalance of 6 participants in final number of participants in each group considered for analysis

Franceschini 2012 (Continued)

Selective reporting (re- porting bias)	Low risk	There is no trial registration, but the outcomes are significant, and there is no selective reporting within the study
Other bias	Low risk	No other potential risk of bias was found.

Fu 2017

Study characteristics	
Methods	Randomized clinical trial
Participants	Stroke patients who were hospitalized in Zhejiang Jiaxing Second Hospital Rehabilitation Center be- tween June 2014 and September 2016 were recruited in this study
	70 participants:
	 35 in experimental; 35 in control.
	Inclusion criteria: unilateral hemiplegia, first-episode of cerebral infarction determined by CT and MRI, stable vital signs, disease course of 2 to 6 months, age of 40 to 75 years; Mini Mental State Exam score ≥ 27 and treatment instructions can be performed, Fugl-Meyer Assessment score ≥ 20 for upper extremity motor function, binocular vision or corrected visual acuity ≥ 1.0, everyday treatment can be tolerated, and providing informed consent and willingness to participate in the study
	Exclusion criteria : cerebral hemorrhage, subarachnoid hemorrhage, venous sinus thrombosis, tran- sient ischemic attack, and progressive or reversible ischemic cerebral apoplexy; lesions located in bilat- eral cerebral hemisphere, cerebellum, or brain stem; joint and other diseases affecting patients sitting or active; heart, lung, liver, kidney, and other serious diseases; metal implantation in the body; a histo- ry of epilepsy; and skull defect
	Mean (SD) age: experimental group: 62.04 (9.93) years, control group: 59.76 (10.57) years
	Stroke details : etiology - ischemic; hemiplegia: control group: 13 right, 12 left; experimental group: 12 right, 16 left
	Stroke phase: subacute
Interventions	Experimental group : traditional rehabilitation treatment such as Bobath, Brunnstrom, proprioceptive neuromuscular facilitation, and daily activity ability training. In addition participants were requested to watch upper limbs motion video and required to imitate the action in the video. There were 30 actions in the video, including shoulder joint, elbow joint, wrist joints, forearm, and hand movements in all directions
	Control group : traditional rehabilitation treatment such as Bobath, Brunnstrom, proprioceptive neu- romuscular facilitation, and daily activity ability training. In addition they watched different geometric patterns and digit symbol, and did the action picked from the 30 actions in the video.
	In both groups sessions were 20 minutes, 6 days/week, for 8 weeks
Outcomes	Outcomes recorded at baseline and postintervention
	1. Motor function: Fugl-Meyer Assessment
	2. Upper limb motor function: Wolf Motor Function Test
	3. Ability of daily living activities: Modified Barthel Index
	4. Movement transmission mechanism: MEP – latency and amplitude, center-motion conduction time
Notes	The authors declared no potential conflicts of interest.

Action observation for upper limb rehabilitation after stroke (Review)



Fu 2017 (Continued)

Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Random number table
Allocation concealment (selection bias)	High risk	Authors did not report if there was allocation concealment, but randomisation type classifies the allocation concealment as high risk
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	There was no blinding of participants and personnel, but this item is not es- sential for this review
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Assessments were made by 2 trained professional therapists and the whole process was recorded. Subsequently, the average score from another 2 trained professional therapists was used for the analyses
Incomplete outcome data (attrition bias) All outcomes	High risk	There were 7 exclusions from the experimental group and 10 from the control group after randomisation, which corresponded to more than 10% and may compromise the results. Reasons were not reported
Selective reporting (re- porting bias)	Low risk	There is no trial registration, but the outcomes are significant, and there is no selective reporting within the study.
Other bias	Low risk	No other potential risk of bias was found.

Harmsen 2015

Study characteristics	5
Methods	Randomized clinical trial
Participants	Patients were recruited from the outpatient clinic of Rijndam Rehabilitation Centre in Rotterdam, the Netherlands
	37 participants:
	1. 18 in experimental group;
	2. 19 in control group.
	Inclusion criteria : Brunnstrom score for upper-extremity function between III and VI, a home-dwelling status, and at least 6 months poststroke
	Exclusion criteria : presence of neglect, comorbidities that influence voluntary upper-extremity func- tion, or multiple strokes
	Mean (SD) age: experimental group: 57 (10.4) years, control group: 60 (8.8) years
	Stroke details : experimental group: 13 ischemic, 5 hemorrhagic; control group: 12 ischemic, 7 hemor- rhagic
	Stroke phase: chronic
Interventions	Experimental group (mirror therapy–based AO): sets of affected-arm reaching movements, alternat- ed with periods of observation. Participants observed mirrored video tapes (the video screen was posi- tioned in the midsagittal plane) of reaching movements performed by their unaffected arm.

Action observation for upper limb rehabilitation after stroke (Review)



Harmsen 2015 (Continued)	Control group : sets of affected-arm reaching movements, alternated with periods of observation. Par- ticipants observed static photographs of landscapes (the video screen was positioned in the midsagit- tal plane).		
		r 1 day: 3 minutes of AO, 30 repetitions of reaching, 1 minute of AO, 20 repetitions of AO, 20 repetitions of reaching, and 1 last minute of AO	
Outcomes	Outcome recorded at baseline and postintervention		
	1. Movement time of reaching: accelerometer		
Notes	The authors declared no potential conflicts of interest.		
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Low risk	Computer-generated random code using blocks of 20 participants	
Allocation concealment (selection bias)	Unclear risk	Use of sealed envelopes. Authors did not report if envelopes were opaque or sequentially numbered.	
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding	
Blinding of outcome as-	Unclear risk	Authors did not report if there was blinding of outcome assessment.	

Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Authors did not report if there was blinding of outcome assessment.
Incomplete outcome data (attrition bias) All outcomes	Low risk	No withdraw was reported. There was no loss of data after intervention
Selective reporting (re- porting bias)	Low risk	The study protocol is available and the outcomes were reported in accordance with what has been proposed DOI 10.1186/ISRCTN40128145

No other potential risk of bias was found.

Hsieh 2020

Other bias

Study characteristics		
Methods	Pilot randomized clinical trial	
Participants	Subacute stroke patients (1 to 6 months from onset)	
	21 participants:	
	1. 7 in AO group;	
	2. 7 in control group;	
	3. 7 in mirror therapy group.	

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Low risk

Isieh 2020 (Continued)	
	Inclusion criteria : diagnosis of cerebral ischemic or hemorrhagic stroke, 1 to 6 months since unilater- al stroke onset, age between 20 and 80 years, baseline score of the Fugl-Meyer Assessment between 20 and 60, ability to follow the study instructions (assessed by the Taiwan version of the Montreal Cogni- tive Assessment), and ability to participate in study therapy and assessment sessions
	Exclusion criteria : global or receptive aphasia, severe neglect, or major medical problems or comor- bidities that influenced the usage of the upper limbs or caused severe pain
	Mean (SD) age: experimental group: 52.77 (11.25) years, control group: 54.30 (13.61) years
	Stroke details
	 Etiology: experimental group: 3 ischemic, 4 hemorrhagic; control group: 5 ischemic, 2 hemorrhagic Hemiparesis: experimental group: 5 right, 2 left; control group: 4 right, 3 left
	Stroke phase: subacute
Interventions	AO group:
	Phase 1 (10–15 minutes) - patients watched upper limb active range of motion exercises and executed observed exercises with both arms and hands simultaneously.
	Phase 2 (15–20 minutes) - patients watched one reaching movement or one object manipulation task for 2 minutes, and executed the movements that they had observed for 3 minutes; this sequence was repeated 3 times.
	Phase 3 (30 minutes) - patients watched one functional task in each session (folding a towel, wiping a table, drinking water, opening a small drawer, and using a mobile phone) for 2 minutes and executed the action they had observed for 3 minutes; this sequence was repeated 3 times
	Mirror therapy group : patients were seated in front of a mirror box placed at their midsagittal plane. The affected arm of the participants was placed inside the mirror box, and the unaffected arm was in front of the mirror. The patient was instructed to watch the mirror reflection of the movement per- formed by his/her unaffected hand carefully and to imagine that the movement was performed by the affected hand. The participant was also encouraged to move the affected arm and hand as much as they could. In the mirror therapy group, treatment activities also contained active range of motion ex- ercises (10–15 minutes), reaching movement or object manipulation (15–20 minutes), and functional task practice (30 minutes)
	Control group : dose-matched bilateral arm training, but no video input or mirror box was provided for this group. The same 3 categories of movements and tasks as provided in the AO group were used. Treatment programs also included active range of motion exercises (10–15 minutes), reaching movement or object manipulation (15–20 minutes), and functional task practice (30 minutes). During training, patients were required to move both of their arms and hands simultaneously as possible
	In all groups sessions were 5 days/week, for 3 weeks
Outcomes	Outcomes recorded at baseline, postintervention, and 3 months follow-up
	 Upper limb motor function: Fugl-Meyer Assessment - upper limb items (total, proximal and distal) and Box and Blocks Test
	 Functional Independence: Functional Independence Measure (total and motor) Quality of life: Stroke Impact Scale (total, physical function and recovery)
Notes	The authors declared no potential conflicts of interest.
Risk of bias	
Bias	Authors' judgement Support for judgement



Hsieh 2020 (Continued)

Other bias	Low risk	No other potential risk of bias was found.
		Authors justified a lack of resources. However, some of the outcomes were questionnaires, which did not need resources.
Selective reporting (re- porting bias)	High risk	This trial is registered with NCT02871700. Registry contains some outcomes not reported in manuscript: Modified Rankin scale, Wolf Motor function Test, Medical Research Council scale, Motor Activity Log, Chedoke Arm and Hand Activity Inventory, Revised Nottingham Sensory Assessment, ABILHAND ques- tionnaire, Actigraphy, Magnetoencephalography, Visual Analogue Scale for pain and fatigue.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Dropouts were recorded at follow-up (3 months after treatment), but none af- ter therapy. This study used an intention-to-treat analysis. For missing data, the last observation carried forward method was used.
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	All outcome measures were administered to the patients by the same rater, who was blinded to the group allocation.
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding
Allocation concealment (selection bias)	Low risk	For concealment of allocation, randomisation procedure and assignment were managed by an independent research assistant who was not involved in par- ticipants' screening or evaluation.
Random sequence genera- tion (selection bias)	Low risk	Randomization was carried out using an online web-based randomisation tool

Jin-Woo 2011

Study characteristics	
Methods	Randomized clinical trial
Participants	Chronical stroke patients
	13 participants:
	1. 7 in experimental group;
	2. 6 in control group.
	Inclusion criteria : stroke diagnosis by a medical rehabilitation specialist; more than 6 months from the event; more than 24 points in Mini Mental State Examination; more than 34 points on the Motor-Free Visual Perception Test; ability to pick up a 2.5 cm square with the affected hand, to communicate and follow instructions; not diagnosed with specific psychiatric findings of mental illness by a psychiatrist
	Exclusion criteria: not reported
	Mean (SD) age: experimental group: 63.2 (11.7) years, control group: 57.2 (10.2) years
	Stroke details
	 Etiology: experimental group: 2 ischemic, 5 hemorrhagic; control group: 2 ischemic, 4 hemorrhagic Hemiparesis: experimental group: 5 right, 2 left; control group: 4 right, 2 left

Jin-Woo 2011 (Continued)				
	All right-handed			
	Stroke phase: chronic			
Interventions	Experimental group : participants sat in a quiet room, looking at a computer screen (13 inches) 30 cm ahead. The therapist provided verbal commands about the characteristics and movements needed to carry out the tasks. After observation (5 minutes), the subjects repeatedly imitated the movement observed with the object shown in the video for about 10 minutes. 2 tasks per session were included			
	The tasks were upper limb movements related to a functional activity			
	Control group : participants sat on chairs in front of a table. The therapist presented a functional activ- ity task and verbally explained the characteristics and movements required for the task. Each task was repeated intensively for 15 minutes. 2 tasks were performed per session. The tasks were the same se- lected for the AO training			
	In both groups sessions were 30 minutes/day, 3 times/week, for 3 weeks			
Outcomes	Outcomes recorded at baseline and postintervention			
	 Upper limb motor function: Wolf Motor Function Test Handgrip, tip-to-tip pinch grip, lateral pinch grip, and three-point pinch grip strength – manual dy- namometer 			

Notes

The authors did not provide any declaration of interests.

Risk of bias

Bias Authors' judgement Support for judgement Unclear risk Authors do not describe how randomisation was done. Random sequence generation (selection bias) Unclear risk Authors did not report if there was allocation concealment. Allocation concealment (selection bias) No blinding, but the outcome is not likely to be influenced by lack of blinding **Blinding of participants** Low risk and personnel (performance bias) All outcomes Unclear risk Authors did not report if there was blinding of outcome assessment. Blinding of outcome assessment (detection bias) All outcomes Incomplete outcome data Low risk No withdraw was reported. There was no loss of data after intervention. (attrition bias) All outcomes Selective reporting (re-Low risk There is no trial registration. Outcomes are significant. There is no selective reporting bias) porting within the study. Other bias Low risk No other potential risk of bias was found.

Kim 2015

Study characteristics

Methods	Non-equivalent pretest	t/post-test control group design	
Participants	Methods of recruitment were not reported by authors		
	12 participants:		
	 6 in experimental gr 6 in control group. 	roup;	
	Inclusion criteria: part	ticipants diagnosed as having hemiplegia due to stroke	
	Exclusion criteria: the	re were no exclusion criteria described in the study	
	Mean (SD) age: not rep	orted by authors	
	Stroke details: not rep	ported by authors	
	Stroke phase: not repo	orted by authors	
Interventions		pants received traditional occupational treatment and topological treatment in purposeful AO program assignments without actually observing the purposeful	
	Experimental group: purposeful AO training	participants received traditional occupational treatment and also performed a program	
	AO training was 30 minutes/session, 5 sessions/week for 6 weeks, a total of 30 sessions		
Outcomes	Outcome recorded after and before intervention		
	 Upper limb function: Wolf Motor Function Test Kinematic variables: average velocity, trajectory ratio and motion angle by a 3D motion analysis system 		
Notes	The authors did not provide any declaration of interests.		
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Unclear risk	Randomization was done, but there was no information of how it was done.	
Allocation concealment (selection bias)	Unclear risk	Authors did not report if there was allocation concealment.	
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding	
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Authors did not report if there was blinding of outcome assessment.	
Incomplete outcome data (attrition bias)	Low risk	There was no loss of outcome data or dropouts.	

Action observation for upper limb rehabilitation after stroke (Review)

Kim 2015 (Continued) All outcomes

Selective reporting (re- porting bias)	Low risk	There is no trial registration, but the outcomes are significant, and there is no selective reporting within the study.
Other bias	High risk	The article does not say how long after the evaluation the treatment began nor how long after the intervention the revaluation was performed.

Kim 2016a

Study characteristics			
Methods	Randomized clinical trial		
Participants	Methods of recruitment were not reported by authors		
	22 participants:		
	 11 in AO training group; 11 in task-oriented training group. 		
	Inclusion criteria : aged between 18 and 70 years with a first-time ischemic or hemorrhagic stroke; stroke experienced more than 1 month and less than 6 months before the study; sufficient cognition to participate in the study: a Mini Mental State Exam scores of 24 or higher; no excessive spasticity, de- fined as a grade of 3 or higher on the Modified Ashworth Scale		
	Exclusion criteria : patients with any comorbidity or disability other than stroke that precluded up- per-extremity training; any uncontrolled health condition for which exercise was contraindicated		
	Mean (SD) age: AO training group: 60.77 (7.03); task-oriented training group: 59.11 (7.05)		
	Stroke details : AO training group: 9 ischemic, 2 hemorrhagic; task-oriented training group: 7 ischemic, 4 hemorrhagic		
	Stroke phase: subacute		
Interventions	AO training group: participants watched video tasks for 9 minutes, had a 1-minute break to organize their thoughts and practiced the task-oriented training for 30 minutes		
	Task-oriented training group : participants performed tasks based on ADL for 30 minutes, without watching the video		
	Sessions were 5 times/week for 4 weeks		
Outcomes	Outcomes recorded after and before intervention		
	 Upper limb function: upper extremity part of the Fugl-Meyer Assessment and Box and Block Test ADL: Modified Barthel Index Spasticity: Modified Ashworth Scale 		
Notes	The authors did not provide any declaration of interests.		
Risk of bias			
Bias	Authors' judgement Support for judgement		

Kim 2016a (Continued)

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Random sequence genera- tion (selection bias)	Low risk	A computer-generated random number was used to allocate participants.
Allocation concealment (selection bias)	Low risk	Authors did not report if there was allocation concealment, but randomisation type classifies the allocation concealment as low risk.
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	The evaluator was not aware of the treatment allocation.
Incomplete outcome data (attrition bias) All outcomes	Low risk	There was no exclusion of participants after randomisation.
Selective reporting (re- porting bias)	Low risk	There is no trial registration, but the outcomes are significant, and there is no selective reporting within the study.
Other bias	Low risk	No other potential risk of bias was found.

Kuk 2016

Study characteristics				
Methods	Randomized clinical trial			
Participants	Methods of recruitment were not reported by authors			
	20 participants:			
	 10 in experimental group; 10 in control group. 			
	Inclusion criteria : > 6 months since stroke onset, no other neurological and orthopedic impairments, no cognitive impairment (> 24 points in Mini Mental State Exam – Korean version) and ability to grasp s small cube (2.5 cm × 2.5 cm × 2.5 cm).			
	Exclusion criteria: there were no exclusion criteria described in the study			
	Mean (SD) age: experimental group: 60.00 (9.36) years; control group: 59.70 (6.58) years			
	Stroke details : experimental group: 5 ischemic, 5 hemorrhagic; control group: 6 ischemic, 4 hemor- rhagic			
	Stroke phase: chronic			
nterventions	Experimental group : participants observed the action of placing wooden blocks from 1 side of the b to another for 1 minute followed by performing the same motor task for 3 minutes			
	Control group : participants observed landscapes for 1 minute followed by performing motor task of placing wooden blocks from 1 box to another for 3 minutes			
	5 sessions with 10-minute rest intervals between the sessions			



Kuk 2016 (Continued)

Outcomes	Outcomes recorded at baseline and postintervention			
	1. Hand dexterity: Box and Block Test			
Notes	The authors declared no potential conflicts of interest.			
Risk of bias				
Bias	Authors' judgement	Support for judgement		
Random sequence genera- tion (selection bias)	Low risk	Random allocation was generated by flipping a coin		
Allocation concealment (selection bias)	Unclear risk	Authors did not report if there was allocation concealment.		
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding		
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Authors did not report if there was blinding of outcome assessment.		
Incomplete outcome data (attrition bias) All outcomes	Low risk	There was one exclusion for each group after randomisation; however, it was less than 10% of the sample, which did not compromise the results.		
Selective reporting (re- porting bias)	Low risk	The study protocol is not available but it is clear that the published reports in- clude all expected outcomes, including those that were prespecified.		
Other bias	Low risk	No other potential risk of bias was found.		

Lee 2013

Study characteristic	S
Methods	Randomized clinical trial
Participants	Methods of recruitment were not reported by authors
	33 participants:
	1. 8 in AO group;
	2. 9 in action practice group;
	3. 9 in combined AO and action practice group;
	4. 7 in control group.
	Inclusion criteria : patients with stroke at least 6 months earlier who were assessed as having recovered to Brunnstrom stage 5, scored at least 20 points in the Korean Mini Mental State Exam, were able to understand and perform instructions, had visual acuity that was sufficient for watching videos, and right hemiplegic patients
	Exclusion criteria: there were no exclusion criteria described in the study



ee 2013 (Continued)				
	Mean (SD) age: AO group: 63 (3.7) years, action practice group: 62 (1.5) years, combined Ao and action practice group: 61 (2.3) years, and control group: 60 (5.9) years Stroke details: not reported by authors			
	Stroke phase: chronic			
Interventions	AO group : watched a drinking task video 20 times in 10 minutes and imagined they were performing the same task at the same time as they watched the video			
	Action practice group	: repeatedly practiced a drinking task for 10 minutes		
	AO and action practic minutes	e group : watched the task video for 5 minutes and practiced the actions for 5		
	Control group: neither	r watched the video nor practiced the actions		
	Sessions were 10 minutes/day for 3 weeks, 15 days total			
Outcomes	Outcomes recorded at	baseline, postintervention and 1 week after the end of intervention		
	1. Number of drinking	motions: number of times the full drinking action was performed in 1 minute		
Notes	The authors did not provide any declaration of interests.			
Risk of bias				
Bias	Authors' judgement Support for judgement			
Random sequence genera- tion (selection bias)	Unclear risk	Randomization was cited in abstract, but there was no information of how it was done		
Allocation concealment (selection bias)	Unclear risk	Authors did not report if there was allocation concealment.		
Blinding of participants				
and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding		
and personnel (perfor- mance bias)	Low risk Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding Physical therapist who conducted the assessments did not have any informa- tion about the experimental group of participants		
and personnel (perfor- mance bias) All outcomes Blinding of outcome as- sessment (detection bias)		Physical therapist who conducted the assessments did not have any informa-		
and personnel (perfor- mance bias) All outcomes Blinding of outcome as- sessment (detection bias) All outcomes Incomplete outcome data (attrition bias)	Low risk	Physical therapist who conducted the assessments did not have any informa- tion about the experimental group of participants There was no exclusion of participants after randomisation and no loss of out-		

Mancuso 2021

Study characteristics



Mancuso 2021 (Continued)

Methods	Randomized clinical trial		
Participants	Acute stroke patients (at a maximum of 30 days from onset) clinically evaluated by a neurologist and that underwent a CT brain scan		
	32 participants:		
	 16 in experimental group; 16 in control group. 		
	unilateral cerebral lesio	n genders, any education level, aged from 18 to 90 years old, first-ever stroke, on, upper limb impairment, and at a maximum of 30 days from onset, and ability anguage (Token Test score higher than 8)	
ioural Inattention Test (a scor ken Test), ideomotor apraxia, ination), history of endogenou		arachnoid hemorrhage, severe neglect at the Star Cancellation from the Behav- (a score lower than 51), impaired comprehension (a score lower than 8 in the To- oraxia, cognitive decline (a score lower than 23.8 in the Mini Mental State Exam- ogenous depression or severe psychiatric disorders for which patients needed apy, or severe visual deficits (eye field examination of the National Institutes of	
	Median (IQR) age: expe	rimental group: 76.5 (13.7) years, control group: 64.5 (15.75)	
	Stroke details: Hemipa	aresis: experimental group: 7 right and 9 left; control group: 8 right and 8 left	
	Stroke phase: acute		
Interventions	 Experimental group: conventional therapy (60 minutes - training for transfers, mobility, walking up and down steps, balance tasks, tailored functional tasks for the upper limbs (unimanual and bimanual), joint and soft tissue mobilization, and specific sensory stimulation) + AO (30 minutes). Participants were asked to carefully observe the videos for 3 minutes in order to prepare themselves to imitate the presented action. At the end of each sequence, participants performed the same movement with the paretic upper limb over a time period of 2 minutes. 3 minutes of sequence observation and 2 minutes of action performance for 3 motor sequences, repeated twice Control group: conventional therapy (60 minutes) + task oriented therapy (30 minutes). Participants performed functional activities with the upper limbs, using the same objects as AO, in both unimanual and bimanual modalities, without watching the video beforehand 		
	In both groups, each session was 190 minutes (60 minutes of conventional treatment + 30 minutes o AO or task oriented training), 5 days per week, for 4 weeks		
Outcomes	Outcomes recorded at baseline and postintervention		
	 Upper limb motor function: Fugl-Meyer Assessment (upper extremity subscale), Box and Block Test Functional independence: Functional Independence Measure Spasticity: Modified Ashworth Scale 		
Notes	The authors declared no potential conflicts of interest.		
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Low risk	A computer-generated random number was used to allocate participants arbi- trarily to the AO training group or the control group.	
Allocation concealment (selection bias)	Low risk	The randomisation procedure and assignment were managed by an indepen- dent researcher who was not involved in the evaluation of the participants.	

Mancuso 2021 (Continued)

Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	All assessments were performed by trained researchers not involved in the treatment administration and blinded to the patient's allocation.
Incomplete outcome data (attrition bias) All outcomes	Low risk	No dropouts were recorded during the treatment, and all participants fulfilled the protocol.
Selective reporting (re- porting bias)	Low risk	The study was registered on ClinicalTrials.gov, with the number NCT04604171. Registry contains all reported outcomes
Other bias	Low risk	No other potential risk of bias was found.

Younghwa 2013

Study characteristics	5
Methods	Randomized clinical trial
Participants	Patients diagnosed with stroke by imaging (CT or MRI) who were admitted to Hospital D in Busan and received occupational therapy from January 2013 to April 2013
	30 participants:
	1. 15 in experimental;
	2. 15 in control.
	Inclusion criteria: patients who understood the purpose of this study and agreed to participate
	Exclusion criteria : no cognitive impairment with a score of 24 or greater on the Korean version of the Mini Mental State Examination, no other neurological disorders or orthopedic damage, and normal vision, hearing, and sensation
	Age. Experimental group: 30-39 years – 2 patients, 40-49 years – 1 patient, 50-59 years – 3 patients, over 60 years – 8 patients. Control group: 30-39 years – 2 patients, 40-49 years – 3 patients, 50-59 years – 3 patients, over 60 years – 7 patients
	Stroke details
	 Etiology: experimental group: 4 ischemic, 11 hemorrhagic; control group: 8 ischemic, 7 hemorrhagic Hemiparesis: experimental group: 4 right, 11 left; control group: 8 right, 7 left
	Stroke phase: acute, subacute and chronic
Interventions	Experimental group : participants watched a 3-4 minute video. After observation, the movement was imitated by the participant, including: turning on a faucet, opening a bottle cap, folding a towel, moving around, putting on and taking off clothes, washing hands and applying lotion. 10-15 minutes per movement (observing and imitating), 2 movements per day, totaling 20-30 minutes per day
	Control group : training consisted of four tasks: putting on and taking off clothes, eating, decorating and moving the wheelchair. During execution, verbal commands and guidance on the activities were given and, after the completion of a movement, feedback on the activity was given. One task was applied at a time and repeated for 30 minutes.



Younghwa 2013 (Continued)

	In both groups session	s were 4 days per week, for 6 weeks	
Outcomes	Outcomes recorded at baseline and postintervention		
	1. Ability of daily living activities: Korean Modified Barthel Index		
Notes	The authors did not pro	ovide any declaration of interests.	
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Unclear risk	Authors do not describe how randomisation was done.	
Allocation concealment (selection bias)	Unclear risk	Authors did not report if there was allocation concealment.	
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding	
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Authors did not report if there was blinding of outcome assessment.	
Incomplete outcome data (attrition bias) All outcomes	Low risk	No withdraw was reported. There was no loss of data after intervention.	
Selective reporting (re- porting bias)	Low risk	There is no trial registration. There is only one outcome, but significant, and there is no selective reporting within the study.	
Other bias	Low risk	No other potential risk of bias was found.	

Zhu 2015

Study characteristic	S
Methods	Randomized clinical trial
Participants	Stroke patients hospitalized in Jiaxing Rehabilitation Medical Center were recruited
	70 participants:
	1. 35 in experimental group;
	2. 35 in control group.
	Inclusion criteria : patients who met the diagnosis for stroke formulated in the 4th National Academic Conference on Cerebrovascular Disease in 1995, and were confirmed as having a stroke upon comput- ed tomography or magnetic resonance imaging; first-episode patients who began rehabilitation thera- py within 6 months, for whom the sitting balance was ≥ Level 1, and the Fugl-Meyer Assessment score was ≥ 15 for upper extremity motor function; patients with a stable condition; patients with a normal Kinesthetic and Visual Imagery Questionnaire score; patients who were 42 to 75 years of age; patients who provided informed consent and were willing to participate in the study



Zhu 2015 (Continued)	patients with a junior h	tients with cognitive impairment mini-mental state examination score of < 24 in high school education or < 17 if illiterate; patients with severe upper limb spas- vere bone joint malformation or myopathy; patients with severe diseases of the lney	
	Mean (SD) age: experin	nental group: 57.75 (15.57) years, control group: 56.89 (14.93) years	
	Stroke details: not rep	orted by authors	
	Stroke phase: acute		
Interventions	Experimental group: additionally watched a video showing a specific action of the upper limb and then performed the same exercise after watching 30 videos		
	Control group: received conventional drug treatment and traditional physical and occupational thera- py		
	Control group received conventional therapy 2 to 5 hours per day, 6 times/week for 8 weeks. Experi- mental group had additional 30 minutes/day, 6 times/week, for 8 weeks		
Outcomes	Outcomes recorded at	baseline and postintervention	
		w flexor and forearm pronator: Motor Activity Log n: Fugl-Meyer Assessment of the upper limb	
Notes	The authors declared r	no potential conflicts of interest.	
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Low risk	Random number table	
Allocation concealment (selection bias)	High risk	Authors did not report if there was allocation concealment, but randomisation type classifies the allocation concealment as high risk.	
Blinding of participants and personnel (perfor-	Low risk	No blinding, but the outcome is not likely to be influenced by lack of blinding	
mance bias) All outcomes			
mance bias)	Low risk	Evaluators were unaware of the participant grouping status and did not partic- ipate in the treatment.	
mance bias) All outcomes Blinding of outcome as- sessment (detection bias)	Low risk Low risk		
mance bias) All outcomes Blinding of outcome as- sessment (detection bias) All outcomes Incomplete outcome data (attrition bias)		 ipate in the treatment. 9 participants were excluded from analysis: 4 from experimental group and 5 from control group. Participants were excluded for absence on 3 consecutive days of intervention and aggravated condition of the participant. Nevertheless the reasons were reported and they have no direct relationship with the outcomes studied, and there is not a substantial imbalance in final number of par- 	

Action observation for upper limb rehabilitation after stroke (Review)

ADL: activities of daily living AO: action observation CT: computed tomography fMRI: functional magnetic resonance MEP: motor evoked potential MRI: magnetic resonance imaging SD: standard deviation TMS: transcranial magnetic stimulation IQR: interquartile range TMS: transcranial magnetic stimulation

Characteristics of excluded studies [ordered by study ID]

Study	Reason for exclusion
Brunner 2014	Study design: not a RCT
Cha 2015	Study design: not a RCT
Chang 2017	The intervention was mirror therapy
Emmerson 2017	The intervention was home exercise programs including techniques, such as constraint-induced movement therapy, repetitive task-specific training, mirror therapy, and bilateral training
Ertelt 2012	It is a protocol without results
Franceschini 2010a	Study design: not a RCT
Frenkel-Toledo 2014	Study design: not a RCT
Kim 2010	Study design: not a RCT
Kim 2013a	Did not assess upper limb, but trunk and lower limb function
Kim 2013b	Did not assess upper limb, only lower limb
Kim 2014	Study design: not a RCT
Kim 2016b	Used AO associated with interface-based electrical stimulation
Ko 2014	The control and experimental group performed AO
Lee 2016	Irrelevant intervention: experimental group performed remote movement observation training based on telerehabilitation
Liepert 2014	Study design: not a RCT
Lima 2020	Study design: not a RCT
Lubart 2017	The control and experimental group performed AO
Marangon 2014	Study design: not a RCT
Sale 2012	Study design: not a RCT
Shih 2017	It is a protocol without results



Study	Reason for exclusion
Sugg 2015	Study design: not a RCT
Sun 2016	Used AO associated with motor imagery
Szameitat 2012	Study design: not a RCT
Wright 2014	Used AO associated with motor imagery
Yun 2011	Study design: not a RCT

AO: action observation; RCT: randomized controlled trial

Characteristics of ongoing studies [ordered by study ID]

NCT02235350

Study name	Study on the effectiveness of action observation treatment as a rehabilitation tool in acute stroke patients and in chronic stroke patients with apraxia: a randomized controlled trial	
Methods	Randomized controlled trial	
Participants	First-ever ischemic or hemorrhagic stroke confirmed by CT/MRI, no later than 30 days from acute event, or chronic stroke patients (more than 3 months from the acute event) with apraxia. Aged 18 to 85 years. Upper limb plegia or paresis with muscle strength in grasping - MRC < 4. Ability to understand and carry out simple verbal instruction. Mini Mental State Examination = 20	
Interventions	Experimental group: AO	
	Control group : observation of videos with no motor content	
Outcomes		
	1. FMA of motor recovery after stroke	
	2. Functional Independence Measure	
	3. fMRI	
Starting date	June 2014	
Contact information	buccino@unicz.it	
Notes	Date accessed: October 2017	

The author did not provide any declaration of interests.

NCT04015271

Study name	Feasibility of action observation and repetitive task practice on upper extremity outcomes in chronic stroke survivors
Methods	Randomized controlled trial

NCT04015271 (Continued)	
Participants	Aged 18 to 85 years. Experienced a stroke resulting in a Upper Extremity Fugl Meyer score >17 < 49; experienced a poststroke > 6 months; > 24 on the Folstein Mini Mental Status Examination
	Experienced only one stroke; discharged from all forms of physical rehabilitation intervention
	Visual acuity of 20/50 or greater with or without corrective lenses; > 19 on the Hooper Visual Orga- nization Test
	Unilateral stroke only; cerebral stroke; age of onset of stroke greater than 18 years old
Interventions	Experimental group: AO + repetitive task practice
	Placebo comparator: placebo video + repetitive task practice
Outcomes	
Outcomes	 Change from baseline in: Upper extremity section of the Fugl-Meyer Scale; Arm Motor Ability Test; Motor Activity Log
Outcomes	
Outcomes	Motor Activity Log
Outcomes	Motor Activity Log2. Smoothness of movements measured by three-axis accelerometer3. Stroke Impact Scale 2.0. Survey to assess participants experience in the study; daily diary; number
Outcomes	 Motor Activity Log Smoothness of movements measured by three-axis accelerometer Stroke Impact Scale 2.0. Survey to assess participants experience in the study; daily diary; number of trials/repetitions per task
Outcomes	 Motor Activity Log Smoothness of movements measured by three-axis accelerometer Stroke Impact Scale 2.0. Survey to assess participants experience in the study; daily diary; number of trials/repetitions per task Change in movement time to complete task

Starting date	27 June 2019
Contact information	John Buford, PT, PhD, Ohio State University
Notes	Date accessed: May 2021
	The author did not provide any declaration of interests.

NCT04047134

Study name	Neuroimaging biomarkers toward a personalized upper limb action observation treatment in chronic stroke patients (BE-TOP)
Methods	Randomized controlled trial
Participants	Aged 18 to 80 years; chronic stroke (never experienced AO treatment); first-ever unilateral stroke due to ischemia provoking a clinically evident upper limb/hand deficit; diagnosis verified by brain imaging (MRI); cognitive function sufficient to understand the experimental instructions; Chedo- ke-McMaster stroke Assessment Scale score greater than 1; informed written consent to participate in the study
Interventions	Experimental group: AO treatment
	Control group : observation of videos with non-motor contents
Outcomes	
	1. Change in Frenchay Arm Test, Box and Block Test, Modified Ashworth Scale, Chedoke-McMaste

Change in Frenchay Arm Test, Box and Block Test, Modified Ashworth Scale, Chedoke-McMaster Stroke Assessment scale, Mini-Mental State Examination, electroencephalography EEG and sEMG



NCT04047134 (Continued)

Starting date	1 July 2019
Contact information	Francesco Infarinato, PhD
Notes	Date accessed: May 2021.
	The author did not provide any declaration of interests.

NCT04574687

Study name	Effects of action observation therapy on fine motor skills of upper limb functions in chronic stroke patients
Methods	Randomized controlled trial
Participants	Aged 30 to 70 years, first-ever stroke, enroled 6 months after the event onset with ischemia or pri- mary hemorrhage. All patients were right handed prior to stroke
Interventions	Experimental group: AO
	Control group: neurodevelopmental techniques
Outcomes	1. Fugl-Meyer Assessment (FMA)
	2. Box and Block Test (BBT)
	3. Frenchay Arm Test (FAT)
Starting date	15 September 2019
Contact information	Maryam Shabbir, PhD, Riphah International University
Notes	Date accessed: May 2021
	The author did not provide any declaration of interests.

NCT04943601

	Active comparator: conventional therapy group
Interventions	Experimental: AO training group
	Dominant hand
Participants	Male and female aged 40-75 years; acute phase of stroke (< 3 months); without cognitive impair- ments (Mini-Mental State Examination > 23); no visual or auditory abnormalities; preserved visual acuity; middle cerebral artery infarction; Fugl-Meyer assessment score ≥ 20 for upper extremity sta tus
Methods	Randomized controlled trial
Study name	Action observation therapy for stroke

Outcomes

NCT04943601 (Continued)

- 1. Fugel Meyer Assessment scale
- 2. Box and block test
- 3. Rating of Everyday Arm-use in the Community and Home (Reach) scale

Starting date	1 December 2020
Contact information	Imran Amjad, PhD imran.amjad@riphah.edu.pk
	Ayesha Afridi, PhD ayesha.afridi@riphah.edu.pk
Notes	Date accessed: November 2021.
	The author did not provide any declaration of interests.

AO: action observation CT: computed tomography EEG: electroencephalography fMRI: Functional Magnetic Resonance MEG: magnetoencephalography MRC: Medical Research Council MRI: magnetic resonance imaging sEMG: surface electromyography

DATA AND ANALYSES

Comparison 1. Action observation versus control: effect on arm function

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1.1 Arm function	11	373	Std. Mean Difference (IV, Random, 95% CI)	0.39 [0.17, 0.61]
1.2 Subgroup analysis: age	11	373	Std. Mean Difference (IV, Random, 95% CI)	0.39 [0.17, 0.61]
1.2.1 ≥ 60 years	5	210	Std. Mean Difference (IV, Random, 95% CI)	0.39 [0.09, 0.69]
1.2.2 < 60 years	6	163	Std. Mean Difference (IV, Random, 95% CI)	0.41 [0.04, 0.78]
1.3 Subgroup analysis: type of stroke	11	373	Std. Mean Difference (IV, Random, 95% CI)	0.39 [0.17, 0.61]
1.3.1 Ischemic stroke	3	101	Std. Mean Difference (IV, Random, 95% CI)	0.56 [0.16, 0.96]
1.3.2 Ischemic and hemor- rhagic stroke	4	139	Std. Mean Difference (IV, Random, 95% CI)	0.67 [-0.04, 1.38]
1.3.3 Type of stroke not stat- ed	4	133	Std. Mean Difference (IV, Random, 95% CI)	0.22 [-0.13, 0.56]

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Cochrane Database of Systematic Reviews

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1.4 Subgroup analysis: time poststroke	10	361	Std. Mean Difference (IV, Random, 95% CI)	0.41 [0.19, 0.64]
1.4.1 Acute or subacute phase	6	280	Std. Mean Difference (IV, Random, 95% CI)	0.39 [0.12, 0.66]
1.4.2 Chronic phase	3	43	Std. Mean Difference (IV, Random, 95% CI)	0.70 [-0.06, 1.47]
1.4.3 Acute, subacute and chronic phase	1	38	Std. Mean Difference (IV, Random, 95% CI)	0.37 [-0.27, 1.02]
1.5 Subgroup analysis: treat- ment dose	11	373	Std. Mean Difference (IV, Random, 95% CI)	0.39 [0.17, 0.61]
1.5.1 ≥ 1000 minutes of thera- py	5	161	Std. Mean Difference (IV, Random, 95% CI)	0.34 [0.03, 0.65]
1.5.2 < 1000 minutes of thera- 6 py		212	Std. Mean Difference (IV, Random, 95% CI)	0.50 [0.07, 0.93]
1.6 Subgroup analysis: type of control group	11	373	Std. Mean Difference (IV, Random, 95% CI)	0.39 [0.17, 0.61]
1.6.1 Placebo therapy	Placebo therapy 6		Std. Mean Difference (IV, Random, 95% CI)	0.45 [0.13, 0.78]
1.6.2 Conventional physical therapy	2	83	Std. Mean Difference (IV, Random, 95% CI)	0.20 [-0.23, 0.63]
1.6.4 Functional activities training	3	59	Std. Mean Difference (IV, Random, 95% CI)	0.56 [-0.12, 1.24]
1.7 Subgroup analysis: dura- tion of observation	9	300	Std. Mean Difference (IV, Random, 95% CI)	0.46 [0.21, 0.72]
1.7.1 > 3 minutes of observa- tion	5	142	Std. Mean Difference (IV, Random, 95% CI)	0.72 [0.35, 1.10]
1.7.2 ≤ 3 minutes of observa- tion	4	158	Std. Mean Difference (IV, Random, 95% CI)	0.22 [-0.09, 0.54]
1.8 Subgroup analysis: upper limb impairment	8	295	Std. Mean Difference (IV, Random, 95% CI)	0.34 [0.11, 0.57]
1.8.1 Severe impairment	3	173	Std. Mean Difference (IV, Random, 95% CI)	0.20 [-0.10, 0.50]
1.8.2 Moderate impairment	3	52	Std. Mean Difference (IV, Random, 95% Cl)	0.75 [0.02, 1.49]
1.8.3 Mild, moderate and se- vere impairments	2	70	Std. Mean Difference (IV, Random, 95% CI)	0.41 [-0.06, 0.88]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1.9 Arm function follow-up	2	93	Mean Difference (IV, Random, 95% CI)	1.06 [-8.38, 10.50]

Analysis 1.1. Comparison 1: Action observation versus control: effect on arm function, Outcome 1: Arm function

		AO			Control			Std. Mean Difference	Std. Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
Kim 2015	39.8	12.2	6	42	11.6	6	3.6%	-0.17 [-1.31 , 0.96]		
Hsieh 2020	47.43	13.38	7	46.71	8.44	7	4.2%	0.06 [-0.99 , 1.11]		
Cowles 2013	29.8	20	9	27.1	20.2	13	6.3%	0.13 [-0.72 , 0.98]		
Franceschini 2012	72.7	31.2	48	66.9	28.4	42	23.1%	0.19 [-0.22 , 0.61]	_ _	
Zhu 2015	35.5	12.4	31	32.8	11.2	30	16.6%	0.23 [-0.28 , 0.73]	_ _	
Dettmers 2014	47.4	11.8	19	43.5	8.3	19	10.7%	0.37 [-0.27 , 1.02]		
Mancuso 2021	47.37	14.88	16	39.87	17.34	16	9.0%	0.45 [-0.25 , 1.16]		
Fu 2017	42.32	12.56	28	35.08	12.44	25	14.1%	0.57 [0.02 , 1.12]		
Ertelt 2007	-7	6.8	8	-16.9	15.9	8	4.4%	0.77 [-0.26 , 1.79]		
Kim 2016a	52.7	3.3	11	48.4	2.8	11	5.1%	1.35 [0.41 , 2.30]	<u> </u>	
Jin-Woo 2011	56.9	12.4	7	38.8	9.9	6	2.8%	1.49 [0.20 , 2.77]		
Total (95% CI)			190			183	100.0%	0.39 [0.17 , 0.61]		
Heterogeneity: Tau ² = 0).01; Chi ² = 10).68, df =	10 (P = 0.3	8); I ² = 6%					•	
Test for overall effect: 2	Z = 3.52 (P =	0.0004)								
Test for subgroup differ	rences: Not ap	plicable							Favors control Favors AO	

Analysis 1.2. Comparison 1: Action observation versus control: effect on arm function, Outcome 2: Subgroup analysis: age

		AO			Control			Std. Mean Difference	Std. Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
1.2.1 ≥ 60 years										
Cowles 2013	29.8	20	9	27.1	20.2	13	6.3%	0.13 [-0.72 , 0.98]		
Franceschini 2012	72.7	31.2	48	66.9	28.4	42	23.1%	0.19 [-0.22 , 0.61]	_ _	
Fu 2017	42.32	12.56	28	35.08	12.44	25	14.1%	0.57 [0.02 , 1.12]		
Jin-Woo 2011	56.9	12.4	7	38.8	9.9	6	2.8%	1.49 [0.20 , 2.77]		
Mancuso 2021	47.37	14.88	16	39.87	17.34	16	9.0%	0.45 [-0.25 , 1.16]		
Subtotal (95% CI)			108			102	55.4%	0.39 [0.09 , 0.69]		
Heterogeneity: Tau ² = 0	0.01; Chi ² = 4.	46, df = 4	(P = 0.35)	; I ² = 10%					•	
Test for overall effect: 2	Z = 2.57 (P =	0.01)								
1.2.2 < 60 years										
Dettmers 2014	47.4	11.85	19	43.5	8.3	19	10.7%	0.37 [-0.27 , 1.02]		
Ertelt 2007	-7	6.8	8	-16.9	15.9	8	4.4%	0.77 [-0.26, 1.79]		
Hsieh 2020	47.43	13.38	7	46.71	8.44	7	4.2%	0.06 [-0.99 , 1.11]		
Kim 2015	39.8	12.2	6	42	11.6	6	3.6%	-0.17 [-1.31 , 0.96]		
Kim 2016a	52.7	3.3	11	48.4	2.8	11	5.1%	1.35 [0.41 , 2.30]		
Zhu 2015	35.5	12.4	31	32.8	11.2	30	16.6%	0.23 [-0.28 , 0.73]	_ _	
Subtotal (95% CI)			82			81	44.6%	0.41 [0.04 , 0.78]		
Heterogeneity: Tau ² = 0	0.04; Chi ² = 6.	22, df = 5	(P = 0.28)	; I ² = 20%						
Test for overall effect: 2	Z = 2.19 (P =	0.03)								
Total (95% CI)			190			183	100.0%	0.39 [0.17 , 0.61]		
Heterogeneity: $Tau^2 = 0$	0.01; Chi ² = 10).68, df =	10 (P = 0.3)	8); I ² = 6%						
Test for overall effect: 2	,	<i>,</i>							-2 -1 0 1 2	
Test for subgroup differ									Favors control Favors AO	

Action observation for upper limb rehabilitation after stroke (Review)

Analysis 1.3. Comparison 1: Action observation versus control: effect on arm function, Outcome 3: Subgroup analysis: type of stroke

		AO			Control			Std. Mean Difference	Std. Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
1.3.1 Ischemic stroke										
Ertelt 2007	-7	6.8	8	-16.9	15.9	8	4.4%	0.77 [-0.26 , 1.79]		
Fu 2017	42.32	12.56	28	35.08	12.44	25	14.1%	0.57 [0.02 , 1.12]	-	
Mancuso 2021	47.37	14.88	16	39.87	17.34	16	9.0%	0.45 [-0.25 , 1.16]		
Subtotal (95% CI)			52			49	27.5%	0.56 [0.16 , 0.96]	•	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 0.	.25, df = 2	(P = 0.88)	; I ² = 0%					•	
Test for overall effect: 2	Z = 2.76 (P =	0.006)								
1.3.2 Ischemic and her	norrhagic st	roke								
Franceschini 2012	72.7	31.2	48	66.9	28.4	42	23.1%	0.19 [-0.22 , 0.61]	_	
Hsieh 2020	47.43	13.38	7	46.71	8.44	7	4.2%	0.06 [-0.99 , 1.11]		
Jin-Woo 2011	56.9	12.4	7	38.8	9.9	6	2.8%	1.49 [0.20 , 2.77]		
Kim 2016a	52.7	3.3	11	48.4	2.8	11	5.1%	1.35 [0.41 , 2.30]		
Subtotal (95% CI)			73			66	35.3%	0.67 [-0.04 , 1.38]	•	
Heterogeneity: Tau ² = 0	.31; Chi ² = 7.	.98, df = 3	(P = 0.05)	; I ² = 62%					•	
Test for overall effect: 2	Z = 1.85 (P =	0.06)								
1.3.3 Type of stroke no	ot stated									
Cowles 2013	29.8	20	9	27.1	20.2	13	6.3%	0.13 [-0.72 , 0.98]	<u> </u>	
Dettmers 2014	47.4	11.8	19	43.5	8.3	19	10.7%	0.37 [-0.27 , 1.02]		
Kim 2015	39.8	12.2	6	42	11.6	6	3.6%	-0.17 [-1.31 , 0.96]		
Zhu 2015	35.5	12.4	31	32.8	11.2	30	16.6%	0.23 [-0.28 , 0.73]	-	
Subtotal (95% CI)			65			68	37.2%	0.22 [-0.13 , 0.56]	•	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 0.	.72, df = 3	(P = 0.87)	; I ² = 0%					ľ	
Test for overall effect: 2	Z = 1.24 (P =	0.22)								
Total (95% CI)			190			183	100.0%	0.39 [0.17 , 0.61]	•	
Heterogeneity: $Tau^2 = 0$	0.01; Chi ² = 1	0.68, df =	10 (P = 0.3	38); I ² = 6%					•	
Test for overall effect: 2	Z = 3.52 (P =	0.0004)							-4 -2 0 2 4	
Test for subgroup differ	ences: Chi ² =	2.29, df =	2 (P = 0.3	32), I ² = 12.	7%				Favors control Favors AC	

Analysis 1.4. Comparison 1: Action observation versus control: effect on arm function, Outcome 4: Subgroup analysis: time poststroke

		AO			Control			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
.4.1 Acute or subacut	e phase								
Cowles 2013	29.8	20	9	27.1	20.2	13	6.6%	0.13 [-0.72 , 0.98]	_
Franceschini 2012	72.7	31.2	48	66.9	28.4	42	23.8%	0.19 [-0.22 , 0.61]	
Fu 2017	42.32	12.56	28	35.08	12.44	25	14.6%	0.57 [0.02 , 1.12]	_ _
Kim 2016a	52.7	3.3	11	48.4	2.8	11	5.4%	1.35 [0.41 , 2.30]	
/Iancuso 2021	47.37	14.88	16	39.87	17.34	16	9.4%	0.45 [-0.25 , 1.16]	
Zhu 2015	35.5	12.4	31	32.8	11.2	30	17.2%	0.23 [-0.28 , 0.73]	_ _ _
ubtotal (95% CI)			143			137	76.9%	0.39 [0.12 , 0.66]	
Ieterogeneity: $Tau^2 = 0$.02; Chi ² = 6.	04, df = 5	(P = 0.30)	; I ² = 17%					•
est for overall effect: 2	Z = 2.81 (P =	0.005)							
.4.2 Chronic phase									
Ertelt 2007	-7	6.8	8	-16.9	15.9	8	4.6%	0.77 [-0.26 , 1.79]	<u> </u>
Isieh 2020	47.43	13.38	7	46.71	8.44	7	4.4%	0.06 [-0.99 , 1.11]	
in-Woo 2011	56.9	12.4	7	38.8	9.9	6	3.0%	1.49 [0.20 , 2.77]	
ubtotal (95% CI)			22			21	11.9%	0.70 [-0.06 , 1.47]	
Heterogeneity: $Tau^2 = 0$.14; Chi ² = 2.	88, df = 2	(P = 0.24)	; I ² = 30%					-
Test for overall effect: Z	Z = 1.80 (P =	0.07)							
.4.3 Acute, subacute a	and chronic j	ohase							
Dettmers 2014	47.4	11.85	19	43.5	8.3	19	11.1%	0.37 [-0.27 , 1.02]	 _
ubtotal (95% CI)			19			19	11.1%	0.37 [-0.27 , 1.02]	
leterogeneity: Not app	licable								
est for overall effect: 2	Z = 1.14 (P =	0.25)							
Fotal (95% CI)			184			177	100.0%	0.41 [0.19 , 0.64]	•
Heterogeneity: Tau ² = 0	.01; Chi ² = 9.	73, df = 9	(P = 0.37)	; I ² = 8%					•
est for overall effect: Z	Z = 3.63 (P =	0.0003)							-4 -2 0 2 4
est for subgroup differ	ences: Chi ² =	0.60, df =	2 (P = 0.7	74), $I^2 = 0\%$					Favors control Favors AO

Analysis 1.5. Comparison 1: Action observation versus control: effect on arm function, Outcome 5: Subgroup analysis: treatment dose

		AO			Control			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
1.5.1 ≥ 1000 minutes of	f therapy								
Dettmers 2014	47.4	11.85	19	43.5	8.3	19	10.7%	0.37 [-0.27 , 1.02]	
Ertelt 2007	-7	6.8	8	-16.9	15.9	8	4.4%	0.77 [-0.26 , 1.79]	
Hsieh 2020	47.43	13.38	7	46.71	8.44	7	4.2%	0.06 [-0.99 , 1.11]	
Mancuso 2021	47.37	14.88	16	39.87	17.34	16	9.0%	0.45 [-0.25 , 1.16]	
Zhu 2015	35.5	12.4	31	32.8	11.2	30	16.6%	0.23 [-0.28 , 0.73]	
Subtotal (95% CI)			81			80	44.9%	0.34 [0.03 , 0.65]	
Heterogeneity: Tau ² = 0	.00; Chi ² = 1.	24, df = 4	(P = 0.87)	; I ² = 0%					•
Test for overall effect: Z	Z = 2.14 (P =	0.03)							
1.5.2 < 1000 minutes o	f therapy								
Cowles 2013	29.8	20	9	27.1	20.2	13	6.3%	0.13 [-0.72, 0.98]	
Franceschini 2012	72.7	31.2	48	66.9	28.4	42	23.1%	0.19 [-0.22 , 0.61]	
Fu 2017	42.32	12.56	28	35.08	12.44	25	14.1%	0.57 [0.02 , 1.12]	
Jin-Woo 2011	56.9	12.4	7	38.8	9.9	6	2.8%	1.49 [0.20 , 2.77]	
Kim 2015	39.8	12.2	6	42	11.6	6	3.6%	-0.17 [-1.31, 0.96]	
Kim 2016a	52.7	3.3	11	48.4	2.8	11	5.1%	1.35 [0.41 , 2.30]	
Subtotal (95% CI)			109			103	55.1%	0.50 [0.07 , 0.93]	
Heterogeneity: Tau ² = 0	.12; Chi ² = 9.	30, df = 5	(P = 0.10)	; I ² = 46%					•
Test for overall effect: Z	z = 2.30 (P = 0.00)	0.02)							
Total (95% CI)			190			183	100.0%	0.39 [0.17 , 0.61]	
Heterogeneity: $Tau^2 = 0$.01; Chi ² = 10).68, df = 1	10 (P = 0.3	8); I ² = 6%					•
Test for overall effect: Z	z = 3.52 (P =	0.0004)							-2 -1 0 1 2
Test for subgroup differ	<u> </u>	0.25 46-	1 (D 0 5	E) 12 00/					Favors control Favors A

Analysis 1.6. Comparison 1: Action observation versus control: effect on arm function, Outcome 6: Subgroup analysis: type of control group

		AO			Control			Std. Mean Difference	Std. Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
.6.1 Placebo therapy										
Dettmers 2014	47.4	11.85	19	43.5	8.3	19	10.7%	0.37 [-0.27 , 1.02]	_ _	
Ertelt 2007	-7	6.8	8	-16.9	15.9	8	4.4%	0.77 [-0.26 , 1.79]		
Franceschini 2012	72.7	31.2	48	66.9	28.4	42	23.1%	0.19 [-0.22 , 0.61]	_ _ _	
Fu 2017	42.32	12.56	28	35.08	12.44	25	14.1%	0.57 [0.02 , 1.12]		
Kim 2015	39.8	12.2	6	42	11.6	6	3.6%	-0.17 [-1.31 , 0.96]		
Kim 2016a	52.7	3.3	11	48.4	2.8	11	5.1%	1.35 [0.41 , 2.30]		
Subtotal (95% CI)			120			111	61.1%	0.45 [0.13 , 0.78]		
Heterogeneity: Tau ² = 0	0.04; Chi ² = 6	.68, df = 5	(P = 0.25)	; I ² = 25%					•	
Test for overall effect:	Z = 2.73 (P =	0.006)								
.6.2 Conventional ph	ysical therap	v								
Cowles 2013	29.8	20	9	27.1	20.2	13	6.3%	0.13 [-0.72, 0.98]		
Zhu 2015	35.5	12.4	31	32.8	11.2	30	16.6%	0.23 [-0.28 , 0.73]	_ _	
Subtotal (95% CI)			40			43	22.9%	0.20 [-0.23 , 0.63]		
Heterogeneity: Tau ² = (0.00; Chi ² = 0.	.04, df = 1	(P = 0.85)	; I ² = 0%						
Test for overall effect:	Z = 0.91 (P =	0.36)								
1.6.4 Functional activi	ities training									
Hsieh 2020	47.43	13.38	7	46.71	8.44	7	4.2%	0.06 [-0.99 , 1.11]		
in-Woo 2011	56.9	12.4	7	38.8	9.9	6	2.8%	1.49 [0.20 , 2.77]		
Mancuso 2021	47.37	14.88	16	39.87	17.34	16	9.0%	0.45 [-0.25 , 1.16]		
Subtotal (95% CI)			30			29	16.1%	0.56 [-0.12 , 1.24]		
Heterogeneity: Tau ² = ().12; Chi ² = 2.	.94, df = 2	(P = 0.23)	; I ² = 32%						
Test for overall effect:	Z = 1.62 (P =	0.11)								
Total (95% CI)			190			183	100.0%	0.39 [0.17 , 0.61]		
Heterogeneity: Tau ² = (0.01; Chi ² = 1	0.68, df =	10 (P = 0.3)	38); I ² = 6%						
Test for overall effect:									-2 -1 0 1	
Test for subgroup diffe		,	-2(D - 0)	(7) $12 - 00/$					Favors control Favors A	

Analysis 1.7. Comparison 1: Action observation versus control: effect on arm function, Outcome 7: Subgroup analysis: duration of observation

Study or Subgroup	Mean	AO SD	Total	Mean	Control SD	Total	Weight	Std. Mean Difference IV, Random, 95% CI	Std. Mean Difference IV, Random, 95% CI
	Mican	50	Iotai	Wittan	30	Total	weight	1 v, Kandolii, 55 /0 C1	
1.7.1 > 3 minutes of ob	servation								
Dettmers 2014	47.4	11.8	19	43.5	8.3	19	13.6%	0.37 [-0.27 , 1.02]	_ _
Ertelt 2007	-7	6.8	8	-16.9	15.9	8	5.9%	0.77 [-0.26 , 1.79]	
Fu 2017	42.32	12.56	28	35.08	12.44	25	17.5%	0.57 [0.02 , 1.12]	
Jin-Woo 2011	56.9	12.4	7	38.8	9.9	6	3.8%	1.49 [0.20 , 2.77]	│ →
Kim 2016a	52.7	3.3	11	48.4	2.8	11	6.8%	1.35 [0.41 , 2.30]	
Subtotal (95% CI)			73			69	47.6%	0.72 [0.35 , 1.10]	
Heterogeneity: $Tau^2 = 0$	0.02; Chi ² = 4	47, df = 4	(P = 0.35)	; I ² = 11%					
Test for overall effect: 2	Z = 3.80 (P =	0.0001)							
1.7.2 ≤ 3 minutes of ot	servation								
Cowles 2013	29.8	20	9	27.1	20.2	13	8.3%	0.13 [-0.72 , 0.98]	_
Franceschini 2012	72.7	31.2	48	66.9	28.4	42	26.8%	0.19 [-0.22 , 0.61]	_ _
Hsieh 2020	47.43	13.38	7	46.71	8.44	7	5.6%	0.06 [-0.99 , 1.11]	
Mancuso 2021	47.37	14.88	16	39.87	17.34	16	11.6%	0.45 [-0.25 , 1.16]	
Subtotal (95% CI)			80			78	52.4%	0.22 [-0.09 , 0.54]	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 0.	57, df = 3	(P = 0.90)	; I ² = 0%					•
Test for overall effect: 2	Z = 1.39 (P =	0.16)							
Total (95% CI)			153			147	100.0%	0.46 [0.21 , 0.72]	
Heterogeneity: $Tau^2 = 0$	02: Chi ² = 9	14 df = 8		· I ² = 12%		14,	100.070		
Test for overall effect: 2	,	<i>,</i>	(1 0.00)	,1 12/0					
Test for subgroup differ			= 1 (P = 0.0)	(4) $I^2 = 75$	3%				-2 -1 0 1 2 Favors control Favors AO
rest for subgroup differ	ences. Chi	4.04, ui -	- 1 (1 - 0.0	+), 1 - 73.	.570				Favors control Favors AO

Analysis 1.8. Comparison 1: Action observation versus control: effect on arm function, Outcome 8: Subgroup analysis: upper limb impairment

		AO			Control			Std. Mean Difference	Std. Mean Difference		
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI		
1.8.1 Severe impairme	nt										
Cowles 2013	29.8	20	9	27.1	20.2	13	7.4%	0.13 [-0.72 , 0.98]			
Franceschini 2012	72.7	31.2	48	66.9	28.4	42	31.3%	0.19 [-0.22 , 0.61]			
Zhu 2015	35.5	12.4	31	32.8	11.2	30	21.2%	0.23 [-0.28 , 0.73]	_		
Subtotal (95% CI)			88			85	60.0%	0.20 [-0.10 , 0.50]	L		
Heterogeneity: Tau ² = 0	.00; Chi ² = 0.	04, df = 2	(P = 0.98)	; I ² = 0%					ľ		
Test for overall effect: Z	Z = 1.28 (P =	0.20)									
1.8.2 Moderate impair	ment										
Ertelt 2007	-7	6.8	8	-16.9	15.9	8	5.1%	0.77 [-0.26 , 1.79]			
Hsieh 2020	47.43	13.38	7	46.71	8.44	7	4.9%	0.06 [-0.99 , 1.11]			
Kim 2016a	52.7	3.3	11	48.4	2.8	11	6.0%	1.35 [0.41 , 2.30]	-		
Subtotal (95% CI)			26			26	16.1%	0.75 [0.02 , 1.49]			
Heterogeneity: Tau ² = 0	.16; Chi ² = 3.	22, df = 2	(P = 0.20)	; I ² = 38%					•		
Test for overall effect: Z	Z = 2.01 (P =	0.04)									
1.8.3 Mild, moderate a	nd severe in	pairment	s								
Dettmers 2014	47.4	11.8	19	43.5	8.3	19	13.1%	0.37 [-0.27 , 1.02]	-		
Mancuso 2021	47.37	14.88	16	39.87	17.34	16	10.9%	0.45 [-0.25 , 1.16]	-		
Subtotal (95% CI)			35			35	24.0%	0.41 [-0.06 , 0.88]	▲		
Heterogeneity: Tau ² = 0	.00; Chi ² = 0.	03, df = 1	(P = 0.87)	; I ² = 0%					·		
Test for overall effect: Z	Z = 1.69 (P =	0.09)									
Total (95% CI)			149			146	100.0%	0.34 [0.11 , 0.57]			
Heterogeneity: Tau ² = 0	.00; Chi ² = 6.	37, df = 7	(P = 0.50)	; I ² = 0%					ľ		
Test for overall effect: Z	Z = 2.87 (P =	0.004)							-10 -5 0 5 1		
Test for subgroup differ	oncos: Chi2 -	2 12 df =	2(P = 0.3)	(5) $I^2 = 5.59$	0/-				Favors control Favors AO		

Action observation for upper limb rehabilitation after stroke (Review)

Analysis 1.9. Comparison 1: Action observation versus control: effect on arm function, Outcome 9: Arm function follow-up

Study or Subgroup	Mean	AO SD	Total	Mean	Control SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
Franceschini 2012	79.6	33.4	40	72.4	31.1	39	38.1%	7.20 [-7.03 , 21.43]	
Hsieh 2020	46.71	11.91	7	49.43	7.93	7	61.9%	-2.72 [-13.32 , 7.88]	_
Total (95% CI)			47			46	100.0%	1.06 [-8.38 , 10.50]	
Heterogeneity: Tau ² = 8	.23; Chi ² = 1.	20, df = 1	(P = 0.27)	; I ² = 17%					Ť
Test for overall effect: Z	Z = 0.22 (P = 0)	0.83)							-100 -50 0 50 100
Test for subgroup differ	ences: Not ap	plicable							Favours control Favours AO

Comparison 2. Action observation versus control: effect on hand function

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
2.1 Hand function	5	178	Mean Difference (IV, Random, 95% CI)	2.76 [1.04, 4.49]
2.2 Subgroup analysis: age	5	178	Mean Difference (IV, Random, 95% CI)	2.76 [1.04, 4.49]
2.2.1 ≥ 60 years	2	122	Mean Difference (IV, Random, 95% CI)	3.00 [-4.02, 10.02]
2.2.2 < 60 years	3	56	Mean Difference (IV, Random, 95% CI)	2.71 [0.91, 4.51]
2.3 Subgroup analysis: time poststroke	5	178	Mean Difference (IV, Random, 95% CI)	2.76 [1.04, 4.49]
2.3.1 Acute or subacute phase	3	144	Mean Difference (IV, Random, 95% CI)	2.67 [0.83, 4.51]
2.3.2 Chronic phase	2	34	Mean Difference (IV, Random, 95% CI)	3.46 [-1.56, 8.47]
2.4 Subgroup analysis: dura- tion of observation	5	178	Mean Difference (IV, Random, 95% CI)	2.76 [1.04, 4.49]
2.4.1 > 3 minutes of observa- tion	1	22	Mean Difference (IV, Random, 95% CI)	2.60 [0.67, 4.53]
2.4.2 ≤ 3 minutes of observa- tion	4	156	Mean Difference (IV, Random, 95% CI)	3.40 [-0.45, 7.26]
2.5 Hand function follow-up	2	93	Mean Difference (IV, Random, 95% CI)	5.19 [-1.82, 12.21]

Analysis 2.1. Comparison 2: Action observation versus control: effect on hand function, Outcome 1: Hand function

	A				Control			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Mancuso 2021	12.18	11.9	16	14.31	19.77	16	2.3%	-2.13 [-13.44 , 9.18]	
Hsieh 2020	16	11.14	7	14.14	11.61	7	2.1%	1.86 [-10.06 , 13.78]	
Kim 2016a	24.6	2.5	11	22	2.1	11	80.0%	2.60 [0.67 , 4.53]	
Kuk 2016	24.4	5.4	10	20.6	7.1	10	9.7%	3.80 [-1.73 , 9.33]	
Franceschini 2012	20	19.2	48	14.5	15.3	42	5.8%	5.50 [-1.64 , 12.64]	
Total (95% CI)			92			86	100.0%	2.76 [1.04 , 4.49]	
Heterogeneity: Tau ² = 0	0.00; Chi ² = 1.	47, df = 4	(P = 0.83)	; I ² = 0%					•
Test for overall effect: 2	Z = 3.14 (P = 0	0.002)							-10 -5 0 5 10
Test for subgroup differ	rences: Not ap	plicable							Favors control Favors AO

Analysis 2.2. Comparison 2: Action observation versus control: effect on hand function, Outcome 2: Subgroup analysis: age

		AO			Control			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
2.2.1 ≥ 60 years									
Franceschini 2012	20	19.2	48	14.5	15.3	42	5.8%	5.50 [-1.64 , 12.64]	
Mancuso 2021	12.18	11.9	16	14.31	19.77	16	2.3%	-2.13 [-13.44 , 9.18]	
Subtotal (95% CI)			64			58	8.2%	3.00 [-4.02 , 10.02]	
Heterogeneity: Tau ² = 5	.84; Chi ² = 1.	25, df = 1	(P = 0.26)	; I ² = 20%					
Test for overall effect: Z	z = 0.84 (P = 0.000)	0.40)							
2.2.2 < 60 years									
Hsieh 2020	16	11.14	7	14.14	11.61	7	2.1%	1.86 [-10.06 , 13.78]	
Kim 2016a	24.6	2.5	11	22	2.1	11	80.0%	2.60 [0.67 , 4.53]	
Kuk 2016	24.4	5.4	10	20.6	7.1	10	9.7%	3.80 [-1.73 , 9.33]	
Subtotal (95% CI)			28			28	91.8%	2.71 [0.91 , 4.51]	
Heterogeneity: $Tau^2 = 0$.00; Chi ² = 0.	18, df = 2	(P = 0.91)	; I ² = 0%					•
Test for overall effect: Z	Z = 2.95 (P = 0	0.003)							
Total (95% CI)			92			86	100.0%	2.76 [1.04 , 4.49]	
Heterogeneity: $Tau^2 = 0$.00; Chi ² = 1.	47, df = 4	(P = 0.83)	; I ² = 0%					▼
Test for overall effect: Z	Z = 3.14 (P = 0	0.002)							-10 -5 0 5 10
Test for subgroup differ	ences: Chi ² =	0.01, df =	1 (P = 0.9	4), $I^2 = 0\%$					Favors control Favors AO

Analysis 2.3. Comparison 2: Action observation versus control: effect on hand function, Outcome 3: Subgroup analysis: time poststroke

		AO			Control			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
2.3.1 Acute or subacute	e phase								
Franceschini 2012	20	19.2	48	14.5	15.3	42	5.8%	5.50 [-1.64 , 12.64]	
Kim 2016a	24.6	2.5	11	22	2.1	11	80.0%	2.60 [0.67 , 4.53]	
Mancuso 2021	12.18	11.9	16	14.31	19.77	16	2.3%	-2.13 [-13.44 , 9.18]	
Subtotal (95% CI)			75			69	88.2%	2.67 [0.83 , 4.51]	
Heterogeneity: Tau ² = 0	.00; Chi ² = 1.	30, df = 2	(P = 0.52)	; I ² = 0%					-
Test for overall effect: Z	z = 2.84 (P =	0.004)							
2.3.2 Chronic phase									
Hsieh 2020	16	11.14	7	14.14	11.61	7	2.1%	1.86 [-10.06 , 13.78]	•
Kuk 2016	24.4	5.4	10	20.6	7.1	10	9.7%	3.80 [-1.73 , 9.33]	
Subtotal (95% CI)			17			17	11.8%	3.46 [-1.56 , 8.47]	
Heterogeneity: Tau ² = 0	.00; Chi ² = 0.	08, df = 1	(P = 0.77)	; I ² = 0%					
Test for overall effect: Z	z = 1.35 (P =	0.18)							
Total (95% CI)			92			86	100.0%	2.76 [1.04 , 4.49]	
Heterogeneity: $Tau^2 = 0$.00; Chi ² = 1.	47, df = 4	(P = 0.83)	; I ² = 0%					•
Test for overall effect: Z	= 3.14 (P =	0.002)							-10 -5 0 5 10
Test for subgroup differ	st for subgroup differences: $Chi^2 = 0.08$, $df = 1$ (P = 0.77), $I^2 = 0\%$								Favors control Favors AO

Analysis 2.4. Comparison 2: Action observation versus control: effect on hand function, Outcome 4: Subgroup analysis: duration of observation

		AO			Control			Mean Difference	Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
2.4.1 > 3 minutes of ob	servation								
Kim 2016a	24.6	2.5	11	22	2.1	11	80.0%	2.60 [0.67 , 4.53]	-
Subtotal (95% CI)			11			11	80.0%	2.60 [0.67 , 4.53]	
Heterogeneity: Not appl	licable								•
Test for overall effect: Z	Z = 2.64 (P = 1)	0.008)							
2.4.2 ≤ 3 minutes of ob	servation								
Franceschini 2012	20	19.2	48	14.5	15.3	42	5.8%	5.50 [-1.64 , 12.64]	
Hsieh 2020	16	11.14	7	14.14	11.61	7	2.1%	1.86 [-10.06 , 13.78]	•
Kuk 2016	24.4	5.4	10	20.6	7.1	10	9.7%	3.80 [-1.73 , 9.33]	
Mancuso 2021	12.18	11.9	16	14.31	19.77	16	2.3%	-2.13 [-13.44 , 9.18]	
Subtotal (95% CI)			81			75	20.0%	3.40 [-0.45 , 7.26]	
Heterogeneity: Tau ² = 0	.00; Chi ² = 1.	34, df = 3	(P = 0.72)	; I ² = 0%					-
Test for overall effect: Z	Z = 1.73 (P =	0.08)							
Fotal (95% CI)			92			86	100.0%	2.76 [1.04 , 4.49]	•
Heterogeneity: $Tau^2 = 0$.00; Chi ² = 1.	47, df = 4	(P = 0.83)	; I ² = 0%					•
Test for overall effect: Z	Z = 3.14 (P =	0.002)							-10 -5 0 5 10
Fest for subgroup differ	ences: Chi ² =	0.13, df =	1 (P = 0.7	(1), $I^2 = 0\%$					Favors control Favors AO

Analysis 2.5. Comparison 2: Action observation versus control: effect on hand function, Outcome 5: Hand function follow-up

Study or Subgroup	Mean	AO SD	Total	Mean	Control SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Diffe IV, Random,	
Franceschini 2012	25.6	20	40	18.7	17.7	39	71.1%	6.90 [-1.42 , 15.22]		ŀ
Hsieh 2020	18.71	10.08	7	17.71	14.43	7	28.9%	1.00 [-12.04 , 14.04]		
Total (95% CI)			47			46	100.0%	5.19 [-1.82 , 12.21]		
Heterogeneity: Tau ² = 0	.00; Chi ² = 0.	56, df = 1	(P = 0.45)	; I ² = 0%					•	
Test for overall effect: Z	Z = 1.45 (P = 0	0.15)							-100 -50 0	50 100
Test for subgroup differ	ences: Not ap	plicable							Favours control	Favours AO

Comparison 3. Action observation versus control: effect on ADL

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
3.1 Dependence on ADL	7	302	Std. Mean Difference (IV, Random, 95% CI)	0.37 [-0.34, 1.08]
3.2 Subgroup analysis: age	7	302	Std. Mean Difference (IV, Random, 95% CI)	0.37 [-0.34, 1.08]
3.2.1 ≥ 60 years	4	205	Std. Mean Difference (IV, Random, 95% CI)	0.14 [-0.83, 1.10]
3.2.2 < 60 years	3	97	Std. Mean Difference (IV, Random, 95% CI)	0.75 [-0.73, 2.23]
3.3 Subgroup analysis: type of stroke	6	241	Std. Mean Difference (IV, Random, 95% CI)	0.38 [-0.52, 1.27]
3.3.1 Ischemic stroke	2	85	Std. Mean Difference (IV, Random, 95% CI)	0.99 [0.54, 1.44]
3.3.2 Ischemic and hemor- rhagic stroke	4	156	Std. Mean Difference (IV, Random, 95% CI)	0.07 [-1.30, 1.44]
3.4 Subgroup analysis: time poststroke	7	302	Std. Mean Difference (IV, Random, 95% CI)	0.37 [-0.34, 1.08]
3.4.1 Acute or subacute phase	5	258	Std. Mean Difference (IV, Random, 95% CI)	0.85 [0.23, 1.46]
3.4.2 Chronic phase	2	44	Std. Mean Difference (IV, Random, 95% CI)	-1.16 [-2.16, -0.17]
3.5 Subgroup analysis: treatment dose	7	302	Std. Mean Difference (IV, Random, 95% CI)	0.37 [-0.34, 1.08]
3.5.1 ≥ 1000 minutes of ther- apy	3	107	Std. Mean Difference (IV, Random, 95% CI)	0.33 [-0.35, 1.01]
3.5.2 < 1000 minutes of ther- apy	4	195	Std. Mean Difference (IV, Random, 95% CI)	0.47 [-0.78, 1.73]

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Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
3.6 Subgroup analysis: type of control group	7	302	Std. Mean Difference (IV, Random, 95% CI)	0.37 [-0.34, 1.08]
3.6.1 Placebo therapy	3	165	Std. Mean Difference (IV, Random, 95% CI)	1.11 [-0.03, 2.26]
3.6.2 Conventional physical therapy	1	61	Std. Mean Difference (IV, Random, 95% CI)	0.38 [-0.12, 0.89]
3.6.3 Functional activities training	3	76	Std. Mean Difference (IV, Random, 95% Cl)	-0.42 [-2.02, 1.17]
3.7 Subgroup analysis: du- ration of observation	6	241	Std. Mean Difference (IV, Random, 95% Cl)	0.38 [-0.52, 1.27]
3.7.1 > 3 minutes of obser- vation	3	105	Std. Mean Difference (IV, Random, 95% CI)	0.64 [-1.51, 2.79]
3.7.2 ≤ 3 minutes of obser- vation	3	136	Std. Mean Difference (IV, Random, 95% CI)	0.20 [-0.50, 0.90]
3.8 Subgroup analysis: up- per limb impairment	5	219	Std. Mean Difference (IV, Random, 95% CI)	0.59 [-0.10, 1.28]
3.8.1 Severe	2	151	Std. Mean Difference (IV, Random, 95% Cl)	0.21 [-0.11, 0.53]
3.8.2 Moderate	2	36	Std. Mean Difference (IV, Random, 95% Cl)	0.98 [-2.12, 4.09]
3.8.3 Mild, moderate and se- vere	1	32	Std. Mean Difference (IV, Random, 95% Cl)	0.90 [0.17, 1.63]
3.9 Dependence on ADL fol- low-up	2	93	Mean Difference (IV, Random, 95% CI)	-0.77 [-5.43, 3.88]

Analysis 3.1. Comparison 3: Action observation versus control: effect on ADL, Outcome 1: Dependence on ADL

		AO		Control				Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Franceschini 2012	63.5	18.3	48	61.8	18.4	42	16.3%	0.09 [-0.32 , 0.51]	•
Fu 2017	73.5	10.1	28	62.2	11.2	25	15.5%	1.05 [0.47 , 1.62]	+
Hsieh 2020	79.71	8.73	7	84	4	7	12.3%	-0.59 [-1.67 , 0.49]	
Kim 2016a	78.5	3.4	11	70.5	2.5	11	11.6%	2.58 [1.39 , 3.76]	
Mancuso 2021	97.87	20.61	16	78.18	21.97	16	14.6%	0.90 [0.17 , 1.63]	-
Younghwa 2013	52.4	7.84	15	65.2	7.58	15	13.9%	-1.62 [-2.45 , -0.78]	-
Zhu 2015	75.3	15.5	31	69.5	14.3	30	15.8%	0.38 [-0.12 , 0.89]	-
Fotal (95% CI)			156			146	100.0%	0.37 [-0.34 , 1.08]	
Heterogeneity: Tau ² = 0	0.77; Chi ² = 46	5.82, df =	6 (P < 0.00	0001); I ² = 8	37%				•
Test for overall effect: 2	Z = 1.02 (P =	0.31)							-10 -5 0 5
Test for subgroup differences: Not applicable									Favors control Favors AO

Action observation for upper limb rehabilitation after stroke (Review)

	SD 18.3 10.1 20.61 7.84	Total 48 28 16 15 107	Mean 61.8 62.2 78.18 65.2	SD 18.4 11.2 21.97	Total 42 25	Weight 16.3% 15.5%	IV, Random, 95% CI 0.09 [-0.32 , 0.51] 1.05 [0.47 , 1.62]	IV, Random, 95% CI
73.5 97.87 52.4 ; Chi ² = 29	10.1 20.61 7.84	28 16 15	62.2 78.18	11.2 21.97	25			-
73.5 97.87 52.4 ; Chi ² = 29	10.1 20.61 7.84	28 16 15	62.2 78.18	11.2 21.97	25			+
97.87 52.4 ; Chi ² = 29	20.61 7.84	16 15	78.18	21.97		15.5%	1 05 [0 47 1 62]	
52.4 ; Chi² = 29	7.84	15			10		1.05 [0.47, 1.02]	-
; Chi² = 29			65.2		16	14.6%	0.90 [0.17 , 1.63]	
	99 df - 3	107		7.58	15	13.9%	-1.62 [-2.45 , -0.78]	-
	00 df - 3				98	60.2%	0.14 [-0.83 , 1.10]	•
	.00, ui – 3	B (P < 0.00	001); I ² = 9	0%				ľ
0.28 (P = 0)).78)							
79.71	8.73	7	84	4	7	12.3%	-0.59 [-1.67 , 0.49]	
78.5	3.4	11	70.5	2.5	11	11.6%	2.58 [1.39 , 3.76]	
75.3	15.5	31	69.5	14.3	30	15.8%	0.38 [-0.12 , 0.89]	-
		49			48	39.8%	0.75 [-0.73 , 2.23]	•
; Chi ² = 15	.94, df = 2	2 (P = 0.00)	03); I ² = 87	%				•
0.99 (P = 0).32)							
		156			146	100.0%	0.37 [-0.34 , 1.08]	
; Chi ² = 46	.82, df = 6	6 (P < 0.00	001); I ² = 8	7%				•
1.02 (P = 0)).31)							-10 -5 0 5 1
	· ·	1 (P = 0.5)	0), $I^2 = 0\%$					Favours control Favours AO
; (0.28 ($P = 0$ 79.71 78.5 75.3 Chi ² = 15 0.99 ($P = 0$ Chi ² = 46 1.02 ($P = 0$	0.28 (P = 0.78) 79.71 8.73 78.5 3.4 75.3 15.5 Chi ² = 15.94, df = 2 0.99 (P = 0.32) Chi ² = 46.82, df = 6 1.02 (P = 0.31)	$79.71 8.73 7$ $78.5 3.4 11$ $75.3 15.5 31$ 49 $Chi^2 = 15.94, df = 2 (P = 0.000)$ $0.99 (P = 0.32)$ 156 $Chi^2 = 46.82, df = 6 (P < 0.000)$ $1.02 (P = 0.31)$	$\begin{array}{cccccccccccccccccccccccccccccccccccc$	79.71 8.73 7 84 4 78.5 3.4 11 70.5 2.5 75.3 15.5 31 69.5 14.3 49 Chi ² = 15.94, df = 2 (P = 0.0003); I ² = 87% 0.99 (P = 0.32) 156 Chi ² = 46.82, df = 6 (P < 0.00001); I ² = 87% 1.02 (P = 0.31)	$79.71 8.73 7 84 4 7$ $78.5 3.4 11 70.5 2.5 11$ $75.3 15.5 31 69.5 14.3 30$ $49 48$ $(Chi^2 = 15.94, df = 2 (P = 0.0003); I^2 = 87\%$ $0.99 (P = 0.32)$ $156 146$ $(Chi^2 = 46.82, df = 6 (P < 0.00001); I^2 = 87\%$ $1.02 (P = 0.31)$	$\begin{array}{cccccccccccccccccccccccccccccccccccc$	$79.71 8.73 7 84 4 7 12.3\% -0.59 \ [-1.67 , 0.49]$ $78.5 3.4 11 70.5 2.5 11 11.6\% 2.58 \ [1.39 , 3.76]$ $75.3 15.5 31 69.5 14.3 30 15.8\% 0.38 \ [-0.12 , 0.89]$ $49 48 39.8\% 0.75 \ [-0.73 , 2.23]$ $Chi^2 = 15.94, df = 2 \ (P = 0.0003); \ I^2 = 87\%$ $0.99 \ (P = 0.32)$ $156 146 100.0\% 0.37 \ [-0.34 , 1.08]$ $Chi^2 = 46.82, df = 6 \ (P < 0.00001); \ I^2 = 87\%$ $1.02 \ (P = 0.31)$

Analysis 3.2. Comparison 3: Action observation versus control: effect on ADL, Outcome 2: Subgroup analysis: age

Analysis 3.3. Comparison 3: Action observation versus control: effect on ADL, Outcome 3: Subgroup analysis: type of stroke

		AO	m . 1		Control	m . 1		Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
3.3.1 Ischemic stroke									
Fu 2017	73.5	10.1	28	62.2	11.2	25	18.0%	1.05 [0.47 , 1.62]	-
Mancuso 2021	97.87	20.61	16	78.18	21.97	16	17.2%	0.90 [0.17 , 1.63]	-
Subtotal (95% CI)			44			41	35.1%	0.99 [0.54 , 1.44]	♦
Heterogeneity: Tau ² = 0.	00; Chi ² = 0.	09, df = 1	(P = 0.76)	; I ² = 0%					•
Test for overall effect: Z	= 4.28 (P <	0.0001)							
3.3.2 Ischemic and hem	orrhagic st	roke							
Franceschini 2012	63.5	18.3	48	61.8	18.4	42	18.6%	0.09 [-0.32 , 0.51]	•
Isieh 2020	79.71	8.73	7	84	4	7	15.2%	-0.59 [-1.67 , 0.49]	
Kim 2016a	78.5	3.4	11	70.5	2.5	11	14.5%	2.58 [1.39 , 3.76]	-
Younghwa 2013	52.4	7.84	15	65.2	7.58	15	16.6%	-1.62 [-2.45 , -0.78]	-
Subtotal (95% CI)			81			75	64.9%	0.07 [-1.30 , 1.44]	•
Heterogeneity: Tau ² = 1.	74; Chi ² = 33	8.66, df = 3	3 (P < 0.00	001); I ² = 9	1%				Ť
Test for overall effect: Z	= 0.10 (P =	0.92)							
Fotal (95% CI)			125			116	100.0%	0.38 [-0.52 , 1.27]	
Heterogeneity: Tau ² = 1.	08; Chi ² = 46	6.78, df = 5	5 (P < 0.00)	001); I ² = 8	9%				ľ
Test for overall effect: Z	= 0.82 (P =	0.41)							-10 -5 0 5 10
est for subgroup differe	ences: Chi ² =	1.56, df =	1 (P = 0.2	1), I ² = 35.9	9%				Favours control Favours AO

Analysis 3.4. Comparison 3: Action observation versus control: effect on ADL, Outcome 4: Subgroup analysis: time poststroke

		AO			Control			Std. Mean Difference	Std. Mean Difference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI	
3.4.1 Acute or subacut	e phase									
Franceschini 2012	63.5	18.3	48	61.8	18.4	42	16.3%	0.09 [-0.32 , 0.51]	+	
Fu 2017	73.5	10.1	28	62.2	11.2	25	15.5%	1.05 [0.47 , 1.62]	-	
Kim 2016a	78.5	3.4	11	70.5	2.5	11	11.6%	2.58 [1.39 , 3.76]		
Mancuso 2021	97.87	20.61	16	78.18	21.97	16	14.6%	0.90 [0.17 , 1.63]	-	
Zhu 2015	75.3	15.5	31	69.5	14.3	30	15.8%	0.38 [-0.12 , 0.89]	-	
Subtotal (95% CI)			134			124	73.8%	0.85 [0.23 , 1.46]		
Heterogeneity: Tau ² = 0	.37; Chi ² = 20	0.07, df = -	4 (P = 0.00	05); I ² = 80)%				•	
Test for overall effect: 2	Z = 2.71 (P =	0.007)								
3.4.2 Chronic phase										
Hsieh 2020	79.71	8.73	7	84	4	7	12.3%	-0.59 [-1.67 , 0.49]		
Younghwa 2013	52.4	7.84	15	65.2	7.58	15	13.9%	-1.62 [-2.45 , -0.78]	-	
Subtotal (95% CI)			22			22	26.2%	-1.16 [-2.16 , -0.17]		
Heterogeneity: Tau ² = 0	.28; Chi ² = 2.	15, df = 1	(P = 0.14)	; I ² = 54%					•	
Test for overall effect: 2	Z = 2.28 (P =	0.02)								
Total (95% CI)			156			146	100.0%	0.37 [-0.34 , 1.08]		
Heterogeneity: Tau ² = 0	0.77; Chi ² = 40	5.82, df =	6 (P < 0.00	001); I ² = 8	87%				T	
Test for overall effect: 2	Z = 1.02 (P =	0.31)							-10 -5 0 5	
Test for subgroup differ	ences: Chi ² =	11.33, df	= 1 (P = 0.	0008), I ² =	91.2%				Favours control Favours AO	

Analysis 3.5. Comparison 3: Action observation versus control: effect on ADL, Outcome 5: Subgroup analysis: treatment dose

Study or Subgroup	Mean	AO SD	Total	Mean	Control SD	Total	Weight	Std. Mean Difference IV, Random, 95% CI	Std. Mean Difference IV, Random, 95% CI
Study of Subgroup	wicali	30	IUtai	wicali	30	Iutai	weight	1 v , Kandoni, 5570 CI	Tv, Kalidolli, 95 % CI
3.5.1 ≥ 1000 minutes o	f therapy								
Hsieh 2020	79.71	8.73	7	84	4	7	12.3%	-0.59 [-1.67 , 0.49]	
Mancuso 2021	97.87	20.61	16	78.18	21.97	16	14.6%	0.90 [0.17 , 1.63]	-
Zhu 2015	75.3	15.5	31	69.5	14.3	30	15.8%	0.38 [-0.12 , 0.89]	-
Subtotal (95% CI)			54			53	42.7%	0.33 [-0.35 , 1.01]	
Heterogeneity: Tau ² = 0	.21; Chi ² = 5.	05, df = 2	(P = 0.08)	; I ² = 60%					
Test for overall effect: Z	Z = 0.96 (P =	0.34)							
3.5.2 < 1000 minutes o	f therapy								
Franceschini 2012	63.5	18.3	48	61.8	18.4	42	16.3%	0.09 [-0.32 , 0.51]	1
Fu 2017	73.5	10.1	28	62.2	11.2	25	15.5%	1.05 [0.47 , 1.62]	+
Kim 2016a	78.5	3.4	11	70.5	2.5	11	11.6%	2.58 [1.39 , 3.76]	
Younghwa 2013	52.4	7.84	15	65.2	7.58	15	13.9%	-1.62 [-2.45 , -0.78]	-
Subtotal (95% CI)			102			93	57.3%	0.47 [-0.78 , 1.73]	
Heterogeneity: Tau ² = 1	.48; Chi ² = 4	1.58, df = 1	3 (P < 0.00	0001); I ² = 9	3%				
Test for overall effect: 2	Z = 0.74 (P =	0.46)							
Total (95% CI)			156			146	100.0%	0.37 [-0.34 , 1.08]	
Heterogeneity: $Tau^2 = 0$.77; Chi ² = 46	5.82, df =	6 (P < 0.00	$(001); I^2 = 8$	7%			- / -	T
Test for overall effect: Z	Z = 1.02 (P =	0.31)							-10 -5 0 5
Test for subgroup differ		· ·	1 (P = 0.8)	(5), $I^2 = 0\%$					Favours control Favours AG

Analysis 3.6. Comparison 3: Action observation versus control: effect on ADL, Outcome 6: Subgroup analysis: type of control group

		AO			Control			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
3.6.1 Placebo therapy									
Franceschini 2012	63.5	18.3	48	61.8	18.4	42	16.3%	0.09 [-0.32 , 0.51]	+
Fu 2017	73.5	10.1	28	62.2	11.2	25	15.5%	1.05 [0.47 , 1.62]	+
Kim 2016a	78.5	3.4	11	70.5	2.5	11	11.6%	2.58 [1.39 , 3.76]	
Subtotal (95% CI)			87			78	43.4%	1.11 [-0.03 , 2.26]	
Heterogeneity: $Tau^2 = 0$.88; Chi ² = 18	3.76, df = 2	2 (P < 0.00	01); I ² = 89	%				•
Test for overall effect: Z	z = 1.91 (P =	0.06)							
3.6.2 Conventional phy	sical therap	y							
Zhu 2015	75.3	15.5	31	69.5	14.3	30	15.8%	0.38 [-0.12 , 0.89]	-
Subtotal (95% CI)			31			30	15.8%	0.38 [-0.12 , 0.89]	•
Heterogeneity: Not appl	icable								₹.
Test for overall effect: Z	z = 1.48 (P =	0.14)							
3.6.3 Functional activit	ties training								
Hsieh 2020	79.71	8.73	7	84	4	7	12.3%	-0.59 [-1.67 , 0.49]	
Mancuso 2021	97.87	20.61	16	78.18	21.97	16	14.6%	0.90 [0.17 , 1.63]	
Younghwa 2013	52.4	7.84	15	65.2	7.58	15	13.9%	-1.62 [-2.45 , -0.78]	-
Subtotal (95% CI)			38			38	40.8%	-0.42 [-2.02 , 1.17]	•
Heterogeneity: Tau ² = 1	.78; Chi ² = 20).04, df = 1	2 (P < 0.00	01); I ² = 90	1%				•
Test for overall effect: Z	a = 0.52 (P =	0.60)							
Total (95% CI)			156			146	100.0%	0.37 [-0.34 , 1.08]	
Heterogeneity: Tau ² = 0	.77; Chi ² = 46	5.82, df =	6 (P < 0.00	001); I ² = 8	7%				
Test for overall effect: Z	= 1.02 (P =	0.31)							-10 -5 0 5 1
Test for subgroup differ	ences: Chi ² =	2.50, df =	2 (P = 0.2	9), I ² = 20.0)%				Favours control Favours AO

Analysis 3.7. Comparison 3: Action observation versus control: effect on ADL, Outcome 7: Subgroup analysis: duration of observation

	AO			Control			Std. Mean Difference	Std. Mean Difference
Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
servation								
73.5	10.1	28	62.2	11.2	25	18.0%	1.05 [0.47 , 1.62]	+
78.5	3.4	11	70.5	2.5	11	14.5%	2.58 [1.39 , 3.76]	
52.4	7.84	15	65.2	7.58	15	16.6%	-1.62 [-2.45 , -0.78]	+
		54			51	49.0%	0.64 [-1.51 , 2.79]	-
.40; Chi ² = 39	9.64, df = 2	2 (P < 0.00	001); I ² = 9	95%				
= 0.59 (P =	0.56)							
servation								
63.5	18.3	48	61.8	18.4	42	18.6%	0.09 [-0.32 , 0.51]	+
79.71	8.73	7	84	4	7	15.2%	-0.59 [-1.67 , 0.49]	
97.87	20.61	16	78.18	21.97	16	17.2%	0.90 [0.17 , 1.63]	-
		71			65	51.0%	0.20 [-0.50 , 0.90]	•
.25; Chi ² = 5.	85, df = 2	(P = 0.05)	; I ² = 66%					ľ
= 0.56 (P =).57)							
		125			116	100.0%	0.38 [-0.52 , 1.27]	
.08; Chi ² = 46	6.78, df =	5 (P < 0.00	001); I ² = 8	39%				
								-10 -5 0 5 10
	· ·	1 (P = 0.7)	(0), $I^2 = 0\%$					Favours control Favours AO
	servation 73.5 78.5 52.4 40; Chi ² = 38 = 0.59 (P = 0 servation 63.5 79.71 97.87 25; Chi ² = 5. = 0.56 (P = 0 08; Chi ² = 46 = 0.82 (P = 0	Mean SD servation 73.5 10.1 78.5 3.4 52.4 7.84 40 ; Chi ² = 39.64, df = 2 $= 0.59$ (P = 0.56) $= 0.59$ (P = 0.56) servation 63.5 18.3 79.71 8.73 97.87 20.61 $= 0.56$ (P = 0.57) $= 0.56$ (P = 0.57) $= 0.82$ (P = 0.41)	Mean SD Total servation 73.5 10.1 28 78.5 3.4 11 52.4 7.84 15 54	MeanSDTotalMeanservation 73.5 10.1 28 62.2 78.5 3.4 11 70.5 52.4 7.84 15 65.2 5440; Chi ² = 39.64, df = 2 (P < 0.00001); I ² = 5e 0.59 (P = 0.56)servation63.5 18.3 48 61.8 79.71 8.73 7 84 97.87 20.61 16 78.18 7125; Chi ² = 5.85 , df = 2 (P = 0.05); I ² = 66% = 0.56 (P = 0.57) 125 08; Chi ² = 46.78 , df = 5 (P < 0.00001); I ² = 8 = 0.82 (P = 0.41)	Mean SD Total Mean SD servation 73.5 10.1 28 62.2 11.2 78.5 3.4 11 70.5 2.5 52.4 7.84 15 65.2 7.58 40; Chi ² = 39.64, df = 2 (P < 0.00001); I ² = 95% = 0.59 (P = 0.56) servation 63.5 18.3 48 61.8 18.4 79.71 8.73 7 84 4 97.87 20.61 16 78.18 21.97 71 25; Chi ² = 5.85, df = 2 (P = 0.05); I ² = 66% = 0.56 (P = 0.57) 125 08; Chi ² = 46.78, df = 5 (P < 0.00001); I ² = 89%	MeanSDTotalMeanSDTotalservation73.510.12862.211.22578.53.41170.52.51152.47.841565.27.5815545140; Chi ² = 39.64, df = 2 (P < 0.00001); I ² = 95%e 0.59 (P = 0.56)servation63.518.34861.818.44279.718.737844797.8720.611678.1821.9716716525; Chi ² = 5.85, df = 2 (P = 0.05); I ² = 66%E 0.56 (P = 0.57)1251160.8; Chi ² = 46.78, df = 5 (P < 0.00001); I ² = 89%= 0.82 (P = 0.41)	MeanSDTotalMeanSDTotalWeightservation73.510.12862.211.22518.0%78.53.41170.52.51114.5%52.47.841565.27.581516.6%545149.0%40; Chi ² = 39.64, df = 2 (P < 0.00001); I ² = 95%e 0.59 (P = 0.56)servation63.518.34861.818.44218.6%79.718.737844715.2%97.8720.611678.1821.971617.2%716551.0%25; Chi ² = 5.85, df = 2 (P = 0.05); I ² = 66%=0.56 (P = 0.57)116100.0%08; Chi ² = 46.78, df = 5 (P < 0.00001); I ² = 89%= 0.82 (P = 0.41)	Mean SD Total Mean SD Total Weight IV, Random, 95% CI servation 73.5 10.1 28 62.2 11.2 25 18.0% 1.05 [0.47, 1.62] 78.5 3.4 11 70.5 2.5 11 14.5% 2.58 [1.39, 3.76] 52.4 7.84 15 65.2 7.58 15 16.6% -1.62 [-2.45, -0.78] 54 51 49.0% 0.64 [-1.51, 2.79] 0.64 [-1.51, 2.79] 40; Chi ² = 39.64, df = 2 (P < 0.00001); I ² = 95% - - 0.59 (P = 0.56) servation 63.5 18.3 48 61.8 18.4 42 18.6% 0.09 [-0.32, 0.51] 79.71 8.73 7 84 4 7 15.2% -0.59 [-1.67, 0.49] 97.87 20.61 16 78.18 21.97 16 17.2% 0.90 [0.17, 1.63] 25; Chi ² = 5.85, df = 2 (P = 0.05); I ² = 66% - 51.0% 0.38 [-0.52, 1.27]

Analysis 3.8. Comparison 3: Action observation versus control: effect on ADL, Outcome 8: Subgroup analysis: upper limb impairment

Study or Subgroup	Mean	AO SD	Total	Mean	Control SD	Total	Weight	Std. Mean Difference IV, Random, 95% CI	Std. Mean Difference IV, Random, 95% CI
3.8.1 Severe									
Franceschini 2012	63.5	18.3	48	61.8	18.4	42	24.5%	0.09 [-0.32 , 0.51]	_
Zhu 2015	75.3	15.5	31	69.5	14.3	30	23.5%	0.38 [-0.12 , 0.89]	4
Subtotal (95% CI)			79			72	48.0%	0.21 [-0.11 , 0.53]	
Heterogeneity: Tau ² = 0.	.00; Chi ² = 0.	76, df = 1	(P = 0.38)	; I ² = 0%					
Test for overall effect: Z	= 1.28 (P =	0.20)							
3.8.2 Moderate									
Hsieh 2020	79.71	8.73	7	84	4	7	16.3%	-0.59 [-1.67 , 0.49]	4
Kim 2016a	78.5	3.4	11	70.5	2.5	11	15.0%	2.58 [1.39 , 3.76]	-
Subtotal (95% CI)			18			18	31.3%	0.98 [-2.12 , 4.09]	•
Heterogeneity: Tau ² = 4.	.69; Chi ² = 1	5.02, df =	1 (P = 0.00)	01); I ² = 93	8%				
Test for overall effect: Z	= 0.62 (P =	0.53)							
3.8.3 Mild, moderate a	nd severe								
Mancuso 2021	97.87	20.61	16	78.18	21.97	16	20.7%	0.90 [0.17 , 1.63]	L .
Subtotal (95% CI)			16			16	20.7%	0.90 [0.17 , 1.63]	
Heterogeneity: Not appl	icable								
Test for overall effect: Z	= 2.41 (P =	0.02)							
Total (95% CI)			113			106	100.0%	0.59 [-0.10 , 1.28]	
Heterogeneity: Tau ² = 0.	.46; Chi ² = 20).13, df =	4 (P = 0.00	05); I ² = 80)%				
Test for overall effect: Z	= 1.67 (P =	0.09)							-100 -50 0 50 10
Test for subgroup differe	ences: Chi ² =	3.06, df =	2 (P = 0.2	2), I ² = 34.6	5%				Favours control Favours AO

Analysis 3.9. Comparison 3: Action observation versus control: effect on ADL, Outcome 9: Dependence on ADL follow-up

Study or Subgroup	Mean	AO SD	Total	Mean	Control SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
Franceschini 2012	73.4	17.9	40	72.4	18	39	34.5%	1.00 [-6.92 , 8.92]	
Hsieh 2020	84.29	6.85	7	86	3.65	7	65.5%	-1.71 [-7.46 , 4.04]	
Total (95% CI)			47			46	100.0%	-0.77 [-5.43 , 3.88]	•
Heterogeneity: Tau ² = 0	.00; Chi ² = 0.	29, df = 1	(P = 0.59)	; I ² = 0%					
Test for overall effect: Z	Z = 0.33 (P = 0	0.74)							-10 -5 0 5 10
Test for subgroup differ	ences: Not ap	plicable							Favours control Favours AO

Comparison 4. Action observation versus control: effect on quality of life

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
4.1 Quality of life	2	30	Mean Difference (IV, Random, 95% CI)	5.52 [-30.74, 41.78]

Analysis 4.1. Comparison 4: Action observation versus control: effect on quality of life, Outcome 1: Quality of life

Study or Subgroup	Mean	AO SD	Total	Mean	Control SD	Total	Weight	Mean Difference IV, Random, 95% CI	Mean Difference IV, Random, 95% CI
Ertelt 2007	277.4	17	8	252.5	25.33	8	47.7%	24.90 [3.76 , 46.04]	-
Hsieh 2020	58.57	10.69	7	70.71	15.92	7	52.3%	-12.14 [-26.35 , 2.07]	-
Total (95% CI)			15			15	100.0%	5.52 [-30.74 , 41.78]	
Heterogeneity: Tau ² = 60	01.55; Chi ² =	8.12, df =	1 (P = 0.0	04); I ² = 88	3%				T
Test for overall effect: Z	= 0.30 (P = 0.00)).77)							-100 -50 0 50 100
Test for subgroup different	ences: Not ap	plicable							Favours control Favours AO

ADDITIONAL TABLES

Table 1. Table of comparisons

Included trials	AO group	Control group		
Celnik 2008	1. Congruent AO simultaneous to motor practice of thumb movements	Motor practice of thumb movements		
	2. Incongruent AO simultaneous to motor practice of thumb movements			
Cowles 2013	Conventional physical therapy + AO with observa- tion-to-imitate followed by motor practice	Conventional physical therapy		
Dettmers 2014	AO of typical ADL followed by motor practice (home- based intervention)	1. Placebo group: text observation followed by motor practice of typical ADL		
		2. Usual care: no specific training		
Ertelt 2007	AO of daily life hand and arm actions followed by motor practice	Placebo group: observation of geometric sym- bols and letters followed by motor practice of daily life hand and arm actions		
Franceschini 2012	Conventional physical therapy + AO of typical ADL fol- lowed by motor practice	Placebo group: conventional physical thera- py + observation of objects followed by limb movements (exact order as experimental group)		
Fu 2017	Traditional rehabilitation treatment + drug treatment + action observation with observation-to-imitate	Traditional rehabilitation treatment + drug treatment + observation of geometric patterns and digit symbol and performed one action		
Harmsen 2015	AO of mirrored arm-reaching activity from unaffect- ed arm, alternated with affected arm-reaching move- ments	Placebo group: observation of static pho- tographs of landscapes, alternated with affect- ed arm-reaching movements		
Hsieh 2020	1. AO of active range of motion exercises followed by motor practice	Motor practice of active range of motion exer- cises, reaching movement or object manipula-		
	2. AO of reaching movement or one object manipula- tion task followed by motor practice	tion, and functional task action		
	3. AO of one functional task (motor acts and complete action) followed by motor practice			

Table 1. Table of comparisons (Continued)

Jin-Woo 2011	AO of functional tasks with objects followed by motor practice of the same movements with the objects	Motor practice of the functional task (verbally explained)
Kim 2015	Occupational therapy + purposeful action observation program	Placebo group: occupational therapy + pur- poseful action observation program assign- ments without actually observing the purpose- ful actions
Kim 2016a	AO of functional tasks followed by motor practice of the same movements	Motor practice of functional tasks
Kuk 2016	AO of an action similar to Box and Block test followed by motor practice of the same movements	Placebo group: observation of landscapes fol- lowed by motor practice of an action similar to Box and Block test
Lee 2013	 AO: action observation of drinking behavior simultaneous to action imagination AO + action: action observation of drinking behavior followed by motor practice 	 Action: motor practice of drinking behavior Control: no specific training
Mancuso 2021	Physical therapy + AO of functional tasks followed by motor practice of the same movements	Task oriented training (verbal instructions and feedback)
Younghwa 2013	AO of daily life hand and arm actions followed by motor practice	Task oriented training (verbal instructions and feedback)
Zhu 2015	Physical therapy + occupational therapy + drug treat- ment + AO of upper limb movements followed by mo- tor practice	Physical therapy + occupational therapy + drug treatment

ADL: activities of daily living; AO: action observation

Table 2. Action observation application to experimental group

Included trials	Time of AO of each motor ac- tion (min)	Time of exercise or imitation (min) of each motor action	Total AO (min)	Total exer- cise (min)	Total session (min)
Celnik 2008	10	10	30	30	30
Cowles 2013	1 to 2	4 to 6 (2 to 4 rest)	4 to 5	16 to 18	2 × 30-min ses- sions (10 rest)
Dettmers 2014	5 + 4 after practice	Not reported	9	20	60
Ertelt 2007	6	6	36	36	90
Franceschini 2012	3	2	9	6	2 × 15-min ses- sions (60 rest)
Fu 2017	10	10	10	10	20
Harmsen 2015	3	30 repetitions of reaching*	6	60 repeti-	Not reported
	1	20 repetitions of reaching		tions* of	

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Table 2. Action observation application to experimental group (Continued)

	1	20 repetitions of reaching		reaching (time not informed)	
Hsieh 2020	Phase 1: 2 to 3	Phase 1: 2 to 4 minutes	25	35	60
	Phase 2: 2	Phase 2: 3 minutes			
	Phase 3: 2 + 2 (complete task)	Phase 3: 3 minutes			
Jin-Woo 2011	5	10	10	20	30
Kim 2015	Not reported	Not reported	Not reported	Not reported	30
Kim 2016a	9 (+1 to rest)	30	9	30	40
Kuk 2016	1	3 (10 rest)	5	15	60
Lee 2013	5	5	5	5	10
Mancuso 2021	3	2	18	12	90 (60 of physical therapy and 30 of AO therapy)
Younghwa 2013	3 to 4	7 to 11	6 to 8	14 to 22	20 to 30
Zhu 2015	Not reported	Not reported	Not reported	Not reported	30

Some of the trials did not provide all the values contained in the table, but based on those presented, it was possible to extrapolate the other values. AO: action observation

min: minutes

*Not minutes

Table 3. Interventions of control group

Included trials	Time of conventional physical therapy or occupational ther- apy/session	Time of con- trol interven- tion	Session/day	Time of placebo video/session	Time of motor activity*/ses- sion
Celnik 2008	Did not perform	30 min	1	Did not perform	30 min
Cowles 2013	Not reported	Not reported	Not reported	Did not perform	Not reported
Dettmers 2014	Did not perform	60 min	1	Not reported	Not reported
Ertelt 2007	Did not perform	90 min	1	36 min	36 min
Franceschini 2012	At least 3 hours/day of physio- therapy	30 min	2	9 min	6 min
Fu 2017	Not reported	20 min	1	Not reported	Not reported
Harmsen 2015	Did not perform	Not reported	1	6 min	70 repetitions o reaching

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Tuble 5. Interve	inclosed of control group (continue	u)			
Hsieh 2020	Not reported	60 min	1	Did not perform	60 min
Jin-Woo 2011	Did not perform	30 min	1	Did not perform	30 min
Kim 2015	Not reported	30 min	1	Did not perform	Not reported
Kim 2016a	Occupational therapy (1 hour/ day)	30 min	1	Did not perform	30 min
	Physical therapy (2 hours/day)				
Kuk 2016	Did not perform	20 min	1	5 min	15 min
Lee 2013	Did not perform	10 min	1	Did not perform	10 min
Mancuso 2021	1 hour	30 min	1	Did not perform	30 min
Younghwa 2013	Did not perform	30 min	1	Did not perform	30 min
Zhu 2015	2 to 5 hours	2 to 5 hours	1	Did not perform	2 to 5 hours

Table 3. Interventions of control group (Continued)

*Motor activities refers to imitation, task practice, movement or functional training. min: minutes

Table 4. Arm function - sensitivity analysis

Sensitivity analy- sis conducted	Trials included in analysis	Number of trials	Number of participants	Effect size	 2
Without high risk of bias for allocation concealment	Cowles 2013; Dettmers 2014; Ertelt 2007;	9	268	SMD 0.46, 95% CI 0.15 to 0.76,	27%
	Franceschini 2012; Hsieh 2020; Jin- Woo 2011; Kim 2015; Kim 2016a; Mancuso 2021			P < 0.001	
Without high risk of bias for incomplete outcome data	Ertelt 2007; Jin-Woo 2011; Kim 2016a;	4	83	SMD 0.88, 95% CI 0.39 to 1.36, P <	8%
	Mancuso 2021			0.001	
Home-based action observation train- ing (video-therapy)	Cowles 2013; Ertelt 2007; Frances- chini 2012; Fu 2017; Hsieh 2020; Jin- Woo 2011; Kim 2015; Kim 2016a; Mancuso 2021; Zhu 2015	10	335	SMD 0.41, 95% Cl 0.16 to 0.66, P = 0.001	16%
Real demonstration	Dettmers 2014; Ertelt 2007; Frances- chini 2012; Fu 2017; Hsieh 2020; Jin- Woo 2011; Kim 2015; Kim 2016a; Mancuso 2021; Zhu 2015	10	351	SMD 0.42, 95% CI 0.18 to 0.66, P < 0.001	13%

CI: confidence interval; SMD: standardized mean difference

Table 5. Hand function - sensitivity analysis

Sensitivity analysis con- ducted	Trials included in analysis	Number of trials	Number of participants	Effect size	²
Without high risk of bias for incomplete outcome data	Hsieh 2020; Kim 2016a;	4	,	MD 2.59, 95% CI	
	Kuk 2016; Mancuso 2021			0.81 to 4.37, P < 0.001	
Without high risk of bias for	Kim 2016a; Kuk 2016;	3	74	MD 2.61, 95% CI 0.81, 4.41, P < 0.001	0%
selective reporting	Mancuso 2021				

CI: confidence interval; MD: mean difference

Table 6. Dependence on ADL - sensitivity analysis

Sensitivity analysis conducted	Trials included in analysis	Number of trials	Number of participants	Effect size	 2
Without high risk of bias for allocation conceal- ment	Franceschini 2012; Hsieh 2020; Kim 2016a; Mancuso 2021; Younghwa 2013	5	188	SMD 0.24, 95% CI -0.85 to 1.32, P = 0.67	90%
Without high risk of bias for incomplete outcome data	Kim 2016a; Mancuso 2021; Younghwa 2013; Zhu 2015	4	145	SMD 0.52, 95% CI -0.82 to 1.86, P = 0.45	92%
Removing peripheral trials	Franceschini 2012; Hsieh 2020; Zhu 2015	3	165	SMD 0.13, 95% CI -0.25 to 0.51, P = 0.51	26%

CI: confidence interval; SMD: standardized mean difference

APPENDICES

Appendix 1. The Cochrane Central Register of Controlled Trials (CENTRAL) search strategy

#1MeSH descriptor: [Cerebrovascular Disorders] this term only

#2MeSH descriptor: [Basal Ganglia Cerebrovascular Disease] this term only

#3MeSH descriptor: [Brain Ischemia] explode all trees

#4MeSH descriptor: [Carotid Artery Diseases] explode all trees

#5MeSH descriptor: [Cerebral Small Vessel Diseases] explode all trees

#6MeSH descriptor: [Intracranial Arterial Diseases] explode all trees

#7MeSH descriptor: [Intracranial Embolism and Thrombosis] explode all trees

#8MeSH descriptor: [Intracranial Hemorrhages] explode all trees

#9MeSH descriptor: [Stroke] explode all trees

#10MeSH descriptor: [Vasospasm, Intracranial] this term only

#11MeSH descriptor: [Vertebral Artery Dissection] this term only

#12(stroke* or poststroke or apoplex* or cerebral vasc* or brain vasc* or cerebrovasc* or cva* or SAH):ti,ab,kw (Word variations have been searched)

#13((brain or cerebr* or cerebell* or vertebrobasil* or hemispher* or intracran* or intracerebral or infratentorial or supratentorial or middle cerebral artery or MCA* or anterior circulation or posterior circulation or basilar artery or vertebral artery or space-occupying) near/5 (isch? emi* or infarct* or thrombo* or emboli* or occlus* or hypoxi*)):ti,ab,kw (Word variations have been searched)

#14((brain* or cerebr* or cerebell* or intracerebral or intracran* or parenchymal or intraparenchymal or intraventricular or infratentorial or supratentorial or basal gangli* or putaminal or putamen or posterior fossa or hemispher* or subarachnoid) near/5 (h?emorrhag* or h? ematoma* or bleed*)):ti,ab,kw (Word variations have been searched)



#15MeSH descriptor: [Hemiplegia] this term only #16MeSH descriptor: [Paresis] explode all trees #17MeSH descriptor: [Gait Disorders, Neurologic] explode all trees #18(hemipleg* or hemipar* or paresis or paraparesis or paretic):ti,ab,kw (Word variations have been searched) #19{or #1-#18} 51283 #20MeSH descriptor: [Observation] this term only #21MeSH descriptor: [Psychomotor Performance] explode all trees #22MeSH descriptor: [Imitative Behavior] this term only #23MeSH descriptor: [Motion Perception] this term only #24MeSH descriptor: [Learning] this term only #25MeSH descriptor: [Anticipation, Psychological] this term only #26MeSH descriptor: [Photic Stimulation] this term only #27MeSH descriptor: [Mirror Neurons] this term only #28action observation or action-observation or AO or AOT:ti,ab,kw (Word variations have been searched) #29((observ* or watch8) near/10 (action* or movement* or reach* or activit* or task* or motion* or motor train* or perform* or gestur* or demonstrat* or video* or TV screen or television screen or computer screen)):ti,ab,kw (Word variations have been searched) #30((visual or action or motion) near/5 perception):ti,ab,kw (Word variations have been searched) #31(visual-motor matching):ti,ab,kw (Word variations have been searched) #32(mirror neur*):ti,ab,kw (Word variations have been searched) #33{or #20-#32} #34MeSH descriptor: [Upper Extremity] explode all trees #35(upper limb* or upper extremit* or arm or arms or shoulder or shoulders or hand or hands or axilla* or elbow* or forearm* or finger*

or wrist*):ti,ab,kw (Word variations have been searched) #36#34 or #35

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 cerebral small vessel diseases/ or exp intracranial arterial diseases/ or exp "intracranial embolism and thrombosis"/ or exp intracranial hemorrhages/ or stroke/ or exp brain infarction/ or stroke, lacunar/ or vasospasm, intracranial/ or vertebral artery dissection/ or brain injuries/ or brain injury, chronic/

2. (stroke\$ or poststroke or apoplex\$ or cerebral vasc\$ or brain vasc\$ or cerebrovasc\$ or cva\$ or SAH).tw.

3. ((brain or cerebr\$ or cerebell\$ or vertebrobasil\$ or hemispher\$ or intracran\$ or intracerebral or infratentorial or supratentorial or middle cerebral artery or MCA\$ or anterior circulation or posterior circulation or basilar artery or vertebral artery or space-occupying) adj5 (isch? emi\$ or infarct\$ or thrombo\$ or emboli\$ or occlus\$ or hypoxi\$)).tw.

4. ((brain\$ or cerebr\$ or cerebell\$ or intracerebral or intracran\$ or parenchymal or intraparenchymal or intraventricular or infratentorial or supratentorial or basal gangli\$ or putaminal or putamen or posterior fossa or hemispher\$ or subarachnoid) adj5 (h?emorrhag\$ or h? ematoma\$ or bleed\$)).tw.

5. hemiplegia/ or exp paresis/

6. (hemipleg\$ or hemipar\$ or paresis or paretic or brain injur\$).tw.

7. or/1-6

8. exp upper extremity/

9. (upper limb\$ or upper extremit\$ or arm or arms or shoulder or shoulders or hand or hands or axilla\$ or elbow\$ or forearm\$ or finger \$ or wrist\$).tw.

10. 8 or 9

11. observation/ or psychomotor performance/ or imitative behavior/ or imagination/

12. motion perception/ or visual perception/ or learning/

13. anticipation, psychological/ or photic stimulation/ or mirror neurons/

14. "Imagery (Psychotherapy)"/

15. (action observation or action-observation or AO or AOT).tw.

16. ((observ\$ or watch\$) adj10 (action\$ or movement\$ or reach\$ or activit\$ or task\$ or motion\$ or motor train\$ or perform\$ or gestur\$ or demonstrat\$ or video\$ or TV screen or television screen or computer screen)).tw.

17. ((visual or action or motion) adj5 perception).tw.

18. visual-motor matching.tw.

19. mirror neur\$.tw.

20. 11 or 12 or 13 or 14 or 15 or 16 or 17 or 18 or 19

21. Randomized Controlled Trials as Topic/

22. random allocation/

23. Controlled Clinical Trials as Topic/

24. control groups/

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- 25. clinical trials as topic/ or clinical trials, phase i as topic/ or clinical trials, phase ii as topic/ or clinical trials, phase iii as topic/ or clinical trials, phase iv as topic/
- 26. double-blind method/
- 27. single-blind method/
- 28. Placebos/
- 29. placebo effect/
- 30. cross-over studies/
- 31. randomized controlled trial.pt.
- 32. controlled clinical trial.pt.
- 33. (clinical trial or clinical trial phase i or clinical trial phase ii or clinical trial phase iii or clinical trial phase iv).pt.
- 34. (random\$ or RCT or RCTs).tw.
- 35. (controlled adj5 (trial\$ or stud\$)).tw.
- 36. (clinical\$ adj5 trial\$).tw.
- 37. ((control or treatment or experiment\$ or intervention) adj5 (group\$ or subject\$ or patient\$)).tw.
- 38. (quasi-random\$ or quasi random\$ or pseudo-random\$ or pseudo random\$).tw.
- 39. ((control or experiment\$ or conservative) adj5 (treatment or therapy or procedure or manage\$)).tw.
- 40. ((singl\$ or doubl\$ or tripl\$ or trebl\$) adj5 (blind\$ or mask\$)).tw.
- 41. (cross-over or cross over or crossover).tw.
- 42. (placebo\$ or sham).tw.
- 43. trial.ti.
- 44. (assign\$ or allocat\$).tw.
- 45. controls.tw.
- 46. or/21-45
- 47. 7 and 10 and 20 and 46
- 48. exp animals/ not humans/
- 49. 47 not 48

Appendix 3. Embase search strategy

1. cerebrovascular disease/ or brain disease/ or exp basal ganglion hemorrhage/ or exp brain hemangioma/ or exp brain hematoma/ or exp brain hemorrhage/ or exp brain infarction/ or exp brain ischemia/ or exp carotid artery disease/ or exp cerebral artery disease/ or exp cerebrovascular accident/ or exp cerebrovascular malformation/ or exp intracranial aneurysm/ or exp occlusive cerebrovascular disease/ or exp vertebrobasilar insufficiency/

2. (stroke\$ or poststroke or apoplex\$ or cerebral vasc\$ or brain vasc\$ or cerebrovasc\$ or cva\$ or SAH).tw.

3. ((brain or cerebr\$ or cerebell\$ or vertebrobasil\$ or hemispher\$ or intracran\$ or intracerebral or infratentorial or supratentorial or middle cerebral artery or MCA\$ or anterior circulation or posterior circulation or basilar artery or vertebral artery or space-occupying) adj5 (isch? emi\$ or infarct\$ or thrombo\$ or emboli\$ or occlus\$ or hypoxi\$)).tw.

4. ((brain\$ or cerebr\$ or cerebell\$ or intracerebral or intracran\$ or parenchymal or intraparenchymal or intraventricular or infratentorial or supratentorial or basal gangli\$ or putaminal or putamen or posterior fossa or hemispher\$ or subarachnoid) adj5 (h?emorrhag\$ or h? ematoma\$ or bleed\$)).tw.

- 5. exp hemiplegia/ or exp paresis/ or neurologic gait disorder/
- 6. (hemipleg\$ or hemipar\$ or paresis or paraparesis or paretic).tw.
- 7. or/1-6
- 8. exp upper limb/

9. (upper limb\$ or upper extremit\$ or arm or arms or shoulder or shoulders or hand or hands or axilla\$ or elbow\$ or forearm\$ or finger \$ or wrist\$).tw.

- 10. 8 or 9
- 11. observation/ or psychomotor performance/ or exp imitation/
- 12. movement perception/ or vision/ or learning/
- 13. anticipation/ or photostimulation/ or mirror neuron/
- 14. (action observation or action-observation or AO or AOT).tw.
- 15. ((observ\$ or watch\$) adj10 (action\$ or movement\$ or reach\$ or activit\$ or task\$ or motion\$ or motor train\$ or perform\$ or gestur\$ or demonstrat\$ or video\$ or TV screen or television screen or computer screen)).tw.
- 16. ((visual or action or motion) adj5 perception).tw.
- 17. visual-motor matching.tw.
- 18. mirror neur\$.tw.
- 19. or/11-18
- 20. Randomized Controlled Trial/ or "randomized controlled trial (topic)"/
- 21. Randomization/
- 22. Controlled clinical trial/ or "controlled clinical trial (topic)"/
- 23. control group/ or controlled study/

24. clinical trial/ or "clinical trial (topic)"/ or phase 1 clinical trial/ or phase 2 clinical trial/ or phase 3 clinical trial/ or phase 4 clinical trial/

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- 25. Crossover Procedure/
- 26. Double Blind Procedure/
- 27. Single Blind Procedure/ or triple blind procedure/
- 28. placebo/ or placebo effect/
- 29. (random\$ or RCT or RCTs).tw.
- 30. (controlled adj5 (trial\$ or stud\$)).tw.
- 31. (clinical\$ adj5 trial\$).tw.
- 32. ((control or treatment or experiment\$ or intervention) adj5 (group\$ or subject\$ or patient\$)).tw.
- 33. (quasi-random\$ or quasi random\$ or pseudo-random\$ or pseudo random\$).tw.
- 34. ((control or experiment\$ or conservative) adj5 (treatment or therapy or procedure or manage\$)).tw.
- 35. ((singl\$ or doubl\$ or tripl\$ or trebl\$) adj5 (blind\$ or mask\$)).tw.
- 36. (cross-over or cross over or crossover).tw.
- 37. (placebo\$ or sham).tw.
- 38. trial.ti.
- 39. (assign\$ or allocat\$).tw.
- 40. controls.tw.
- 41. (exp animal/ or animal.hw. or nonhuman/) not (exp human/ or human cell/ or (human or humans).ti.)
- 42. or/20-40
- 43. 7 and 10 and 19 and 42

Appendix 4. CINAHL search strategy

S1(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") OR (MH "Stroke Patients") OR (MH "Stroke Units")

S2TI (stroke or poststroke or post-stroke or cerebrovasc* or brain vasc* or cerebral vasc or cva or apoplex or SAH) or AB (stroke or poststroke or post-stroke or cerebrovasc* or brain vasc* or cva or apoplex or SAH)

S3TI (brain* or cerebr* or cerebell* or intracran* or intracerebral) or AB (brain* or cerebr* or cerebell* or intracran* or intracerebral)

S4TI (ischemi* or ischaemi* or infarct* or thrombo* or emboli* or occlus*) or AB (ischemi* or ischaemi* or infarct* or thrombo* or emboli* or occlus*)

S5S3 AND S4

S6TI (brain* or cerebr* or cerebell* or intracerebral or intracranial or subarachnoid) or AB (brain* or cerebr* or cerebell* or intracerebral or intracranial or subarachnoid)

S7TI (haemorrhage* or hemorrhage* or haematoma* or hematoma* or bleed*) or AB (haemorrhage* or hemorrhage* or haematoma* or hematoma* or bleed*)

S8S6 AND S7

S9TI transient ischaemic attack* or TI transient ischemic attack* or AB transient ischaemic attack* or AB transient ischemic attack* or TI TIA or TI TIA s or AB TIA or AB TIAs

S10S1 OR S2 OR S3 OR S6 OR S9 OR S10

S11(MH "Upper Extremity+")

S12TI (upper limb* or upper extremit* or arm or arms or shoulder or shoulders or hand or hands or axilla* or elbow* or forearm* or finger* or wrist*) OR AB (upper limb* or upper extremit* or arm or arms or shoulder or shoulders or hand or hands or axilla* or elbow* or forearm* or finger* or wrist*)

S13S11 OR S12

S14(MH "Psychomotor Performance+")

S15(MH "Imitative Behavior")

S16(MH "Visual Perception+")

S17(MH "Learning")

S18(MH "Neurons+")

S19TI ((action observation or action-observation or AO or AOT)) OR AB ((action observation or action-observation or AO or AOT))

S20TI (((observ* or watch*) N10 (action* or movement* or reach* or activit* or task* or motion* or motor train* or perform* or gestur* or demonstrat* or video* or TV screen or television screen or computer screen)) OR AB (((observ* or watch*) N10 (action* or movement* or reach* or activit* or task* or motion* or motor train* or perform* or gestur* or demonstrat* or video* or TV screen or television screen or computer screen)) OR AB ((observ* or watch*) N10 (action* or movement* or reach* or activit* or task* or motion* or motor train* or perform* or gestur* or demonstrat* or video* or TV screen or television screen or computer screen)) OR AB ((observ* or watch*) N10 (action* or motor train* or perform* or gestur* or demonstrat* or video* or TV screen or television screen or computer screen)) OF AB ((observ* or watch*) N10 (action* or motor train* or perform* or gestur* or demonstrat* or video* or TV screen or television screen or computer screen)) OF AB ((observ* or watch*) N10 (action* or motor train* or perform* or gestur* or demonstrat* or video* or TV screen or television screen or computer screen)) OF AB ((observ* or watch*) N10 (action* or motor train* or perform* or gestur* or demonstrat* or video* or TV screen or television screen or computer screen)) OF AB ((observ* or Video* or TV screen or television screen or computer screen)) OF AB ((observ* or Video* or TV screen or television screen or computer screen)) OF AB ((observ* or Video* or TV screen or television screen or computer screen)) OF AB ((observ* or Video* or TV screen or television screen or computer screen)) OF AB ((observ* or Video* or TV screen or television screen or computer screen)) OF AB ((observ* or Video* or TV screen or television screen or computer screen)) OF AB ((observ* or Video* or TV screen or television screen or computer screen)))

S21TI ((action observation or action-observation or AO or AOT)) OR AB ((action observation or action-observation or AO or AOT).)

S22TI (((visual or action or motion) N5 perception).) OR AB (((visual or action or motion) N5 perception).)

S23TI (visual-motor matching) OR AB (visual-motor matching)

S24TI mirror neur* OR AB mirror neur*

S25S14 OR S15 OR S16 OR S17 OR S18 OR S19 OR S20 OR S21 OR S22 OR S23 OR S24

S26(MH "Randomized Controlled Trials") or (MH "Random Assignment") or (MH "Random Sample+")

S27(MH "Clinical Trials") or (MH "Intervention Trials") or (MH "Therapeutic Trials")

S28(MH "Double-Blind Studies") or (MH "Single-Blind Studies") or (MH "Triple-Blind Studies")
S29(MH "Control (Research)") or (MH "Control Group") or (MH "Placebos") or (MH "Placebo Effect")
S30(MH "Crossover Design") OR (MH "Quasi-Experimental Studies")
S31PT (clinical trial or randomized controlled trial)
S32TI (random* or RCT or RCTs) or AB (random* or RCT or RCTs)
S33TI (controlled N5 (trial* or stud*)) or AB (controlled N5 (trial* or stud*))
S34TI (clinical* N5 trial*) or AB (clinical* N5 trial*)
S35TI ((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*)) or AB ((control or experiment* or conservative) N5 (treatment or therapy or procedure or manage*))
S37TI ((singl* or doubl* or tripl* or trebl*) N5 (blind* or mask*)) or AB ((singl* or doubl* or tripl* or trebl*) N5 (blind* or mask*))
S38TI (cross-over or cross over or crossover) or AB (cross-over or cross over or crossover)
S39TI (placebo* or sham) or AB (placebo* or sham)
S40TI trial

S41TI (assign* or allocat*) or AB (assign* or allocat*)

S42TI controls or AB controls

S43TI (quasi-random* or quasi random* or pseudo-random* or pseudo random*) or AB (quasi-random* or quasi random* or pseudo-random*)

S44S26 OR S27 OR S28 OR S29 OR S30 OR S31 OR S32 OR S33 OR S34 OR S35 OR S36 OR S37 OR S38 OR S39 OR S40 OR S41 OR S42 OR S43 S45S10 AND S13 AND S25 AND S44

Appendix 5. LILACS search strategy

(TW:"Brain Vascular Disorders" OR TW:"Intracranial Vascular Disorders" OR TW:"Transtornos Vasculares do Cérebro" OR TW:"Transtornos Vasculares Intracranianos" OR TW: "Doenças Vasculares Intracranianas" OR TW: "Trastornos Vasculares del Cerebro" OR TW: "Trastornos Vasculares Intracraneales" OR TW:"Enfermedades Intracraneales Vasculares" OR MH:C10.228.140.300 OR MH:C14.907.253 OR TW:"Lenticulostriate Vasculopathy" OR TW: "Doença Cerebrovascular dos Gânglios Basais" OR TW: "Doenças Vasculares Lenticulostratais" OR TW: "Vasculopatia Lenticulostriatal" OR TW: "Vasculopatia Lenticuloestriatal" OR TW: "Enfermedad de los Ganglios Basales Cerebrovascular" OR TW: "Enfermedades Vasculares Lenticulostriatales" OR MH:C10.228.140.079.127\$ OR MH:C10.228.140.300.100\$ OR MH:C14.907.253.061\$ OR TW:"Cerebral Ischemia" OR TW:"Ischemic Encephalopathy" OR TW:"Isquemia Cerebral" OR TW:"Encefalopatía Isquémica" OR MH:10.228.140.300.150\$ OR MH:C14.907.253.092\$ OR TW:"Carotid Atherosclerosis" OR TW:"Aterosclerosis de la Carótida " OR TW:"Aterosclerose Carotídea" OR TW:"Aterosclerose da Carótida" OR MH:C10.228.140.300.200\$ OR MH:C14.907.253.123\$ OR MH:C10.228.140.300.275\$ OR MH:C14.907.253.329\$ OR MH:C10.228.140.300.510\$ OR MH:C14.907.253.560\$ OR TW:"Cerebral Embolism and Thrombosis" OR TW:"Brain Embolism and Thrombosis" OR TW:"Embolia e Trombose Cerebral" OR TW:"Embolia e Trombose Encefálica" OR TW:"Embolia y Trombosis Cerebral" OR TW:"Embolia y Trombosis del Cerebro" OR MH:C10.228.140.300.525\$ OR MH:C14.907.253.566\$ OR MH:C14.907.355.590.213\$ OR TW:"Brain Hemorrhage" OR TW:"Hemorragia Encefálica" OR TW:"Hemorragia Intracraniana" OR TW: "Hemorragia del Cerebro" OR TW: "Hemorragia Intracraneal" OR MH:C10.228.140.300.535\$ OR MH:C14.907.253.573\$ OR MH:C23.550.414.913\$ OR TW:"Apoplexy" OR TW:"Cerebral Stroke" OR TW:"Cerebrovascular Accident" OR TW:"Cerebrovascular Apoplexy" OR TW:"Acidente Vascular Cerebral" OR TW:"Derrame Cerebral" OR TW:"Ictus Cerebral" OR TW:"AVC" OR TW:"Apoplexia" OR TW:"Acidente Cerebrovascular" OR TW:"Apoplexia Cerebral" OR TW:"Apoplexia Cerebrovascular" OR TW:"Icto Cerebral" OR TW:"Acidente Vascular Encefálico" OR TW:"AVE" OR TW:"Acidente Vascular do Cérebro" OR TW:"Acidente Cerebral Vascular" OR TW:"Acidentes Cerebrais Vasculares" OR TW:"Acidentes Cerebrovasculares" OR TW:"Acidentes Vasculares Cerebrais" OR TW:"Ataque" OR TW:"Ictus" OR TW:"Ictus Cerebral" OR TW:"Apoplejía" OR TW:"Ataque Cerebral" OR TW:"Apoplejía Cerebral" OR TW:"Apoplejía Cerebrovascular" OR TW:"Accidente Vascular Encefálico" OR TW:"Accidente Vascular del Cerebro" OR TW:"Accidente Vascular Cerebral" OR TW:"Accidente Cerebral Vascular" OR TW:"Accidentes Cerebrovasculares" OR MH:C10.228.140.300.775\$ OR MH:C14.907.253.855\$ OR TW:"Infarto Venoso Encefálico" OR TW:"Infarto Venoso Cerebral" OR TW:"Infarto Venoso do Encefalo" OR TW:"Infarto do Encéfalo" OR TW:"Infarto Venoso del Encéfalo" OR TW:"Infarto del Encéfalo" OR MH:C10.228.140.300.150.477\$ OR MH:C10.228.140.300.775.200\$ OR MH:C14.907.253.092.477\$ OR MH:C14.907.253.855.200\$ OR TW:"Infarto Lacunar" OR TW:"Acidente Vascular Encefálico Lacunar" OR MH:C10.228.140.300.275.800\$ OR MH:C10.228.140.300.775.600\$ OR MH:C14.907.253.329.800\$ OR MH:C14.907.253.855.600\$ OR MH:C10.228.140.300.90\$ OR MH:C14.907.253.951\$ OR TW:"Cerebral Vasospasm" OR TW:"Intracranial Vasospasm" OR TW:"Angioespasmo Intracraniano" OR TW:"Angiospasmo Intracraniano" OR TW:"Vasoespasmo Cerebral" OR TW:"Vasospasmo Cerebral" OR TW:"Vasospasmo Intracraniano" OR TW:"Angioespasmo Intracraneal" OR TW:"Vasoespasmo Cerebral" OR TW:"Dissecting Vertebral Artery Aneurysm" OR MH:C10.228.140.300.350.875\$ OR MH:C10.900.250.650\$ OR MH:C14.907.055.050.575\$ OR MH:C14.907.253.535.800\$ OR MH:C26.915.200.600\$ OR TW:"Aneurisma Disecante de la Arteria Vertebral" OR TW:"Dissecção da Artéria Vertebral" OR TW:"Aneurisma Dissecante da Artéria Vertebral" OR TW:"Brain Contusion" OR TW:"Brain Lacerations" OR TW:"Traumatic Brain Injury" OR MH:C10.228.140.199\$ OR MH:C10.900.300.087\$ OR MH:C26.915.300.200\$ OR TW:"Lesión Cerebral" OR TW:"Lesiones Encefálicas" OR TW:"Traumatismo Cerebral" OR TW:"Traumatismos Cerebrales" OR TW:"Contusión Encefálica" OR TW:"Lesiones Traumáticas del Encéfalo" OR TW:"Lesiones Encefálicas Traumáticas" OR TW:"Laceraciones del Encéfalo" OR TW:"Laceraciones Encefálicas" OR TW:"Traumatismo Encefálico" OR TW:"Lesiones del Encéfalo Traumáticas" OR TW:"Lesão Cerebral" OR TW:"Lesões Encefálicas" OR TW:"Traumatismo Cerebral" OR TW:"Traumatismos Cerebrais" OR TW:"Contusão Encefálica" OR TW:"Lesões Encefálicas Traumáticas" OR TW:"Lacerações Encefálicas" OR TW:"Traumatismo do Encéfalo" OR TW:"Lesão Encefálica



Traumática" OR MH:C10.228.140.140.127\$ OR MH:C10.228.140.199.500\$ OR MH:C10.900.300.087.250\$ OR MH:C26.915.300.200.200\$ OR tw:stroke* or tw:poststroke or tw:apoplex* or tw:cerebrovasc* or cva* or tw:SAH OR TW:Monoplegia OR MH:C10.597.622.295\$ OR MH:C23.888.592.636.312\$ OR TW:Hemiparesis OR TW:Muscle Paresis OR MH:C10.597.636\$ OR MH:C23.888.592.643\$ OR TW:"Hemiparesia" OR TW:"Paresia Muscular" OR TW:hemipleg* or TW:hemipar* or TW:paresis or TW:paretic) AND (MH:A01.378.800\$ OR TW:"Extremidade Superior" OR TW: "upper limb" or TW: "upper extremity" or TW:arm or TW:arms or TW:shoulder or TW:shoulders or TW:hand or TW:hands or TW:axilla* or TW:elbow* or TW:forearm* or TW:finger* or TW:wrist*) AND (TW:Observation OR TW:Observación OR TW:Observação OR MH:E05.581.249 OR TW: "Perceptual Motor Performance" OR TW: "Visual Motor Coordination" OR TW: "Sensory Motor Performance" OR MH:F02.808 OR MH:G11.427.700 OR MH:G11.561.623 OR TW:"Desempenho Perceptual-Motor" OR TW:"Coordenação Visiomotora" OR TW:"Desempenho Sensório-Motor" OR TW:"Desempeño Motor Perceptual" OR TW:"Coordinación Motora Visual" OR TW:"Desempeño Motor Sensorial" OR TW:"Imitative Behavior" OR TW:"Conducta Imitativa" OR TW:"Comportamento Imitativo" OR MH:F01.145.510\$ OR TW:"Motion Perception" OR TW:"Percepción de Movimiento" OR TW:"Percepção de Movimento" OR MH:F02.463.593.932.567 OR TW:"Percepción Visual" OR TW:"Percepção Visual" OR TW:"visual perception" OR MH:F02.463.593.932 OR TW:Learning OR TW:Aprendizaje OR TW:Aprendizagem OR TW:Phenomenography OR MH:F02.463.425 OR MH:F02.784.629.529 OR MH:SP4.006.047.453.604 OR TW:Aprendizado OR TW:Fenomenografia OR TW:"Psychological Anticipation" OR TW:"Anticipación Psicológica" OR TW:"Antecipação Psicológica" OR MH:F02.463.093 OR TW:"Photic Stimulation" OR TW:"Estimulación Luminosa" OR TW:"Estimulação Luminosa" OR TW:"Visual Stimulation" OR MH:E05.723.729 OR TW:"Estimulação Fótica" OR TW:"Estimulação Visual" OR TW:"Estimulación Fótica" OR TW:"Estimulación Visual" OR TW:"Mirror Neurons" OR TW:"Neurônios-Espelho" OR TW:"Neuronas Espejo" OR MH:A08.675.500 OR MH:A11.671.487 OR TW:"action observation" OR TW:"action-observation" OR TW:AO OR TW: AOT OR TW: "observación de la acción" OR TW:"observação da ação" OR TW:"visual-motor matching" OR TW:"correspondência visual-motora" OR TW:"coincidente visomotora") AND (((PT:"randomized controlled trial" OR PT:"controlled clinical trial" OR PT:"multicenter study" OR MH:"randomized controlled trials as topic" OR MH:"controlled clinical trials as topic" OR MH:"multicenter study as topic" OR MH:"random allocation" OR MH:"double-blind method" OR MH: "single-blind method") OR ((ensaio\$ OR ensayo\$ OR trial\$) AND (azar OR acaso OR placebo OR control\$ OR aleat\$ OR random\$ OR enmascarado\$ OR simpleciego OR ((simple\$ OR single OR duplo\$ OR doble\$ OR double\$) AND (cego OR ciego OR blind OR mask))) AND clinic\$)) AND NOT (MH:animals OR MH:rabbits OR MH:rats OR MH:primates OR MH:dogs OR MH:cats OR MH:swine OR PT:"in vitro"))

Update 2021:

(ti:(((stroke OR poststroke OR post-stroke OR cerebrovasc* OR brain vasc* OR cerebral vasc* OR cva* OR apoplex* OR sah) OR ((brain* OR cerebr* OR cerebell* OR intracerebral OR intracranial OR subarachnoid) near/5 (haemorrhage* OR hemorrhage* OR haematoma* OR hematoma* OR bleed*)) OR ((brain* OR cerebr* OR cerebell* OR intracrana* OR intracerebral) near/5 (isch*emi* OR infarct* OR thrombo* OR emboli* OR occlus*))))) AND ((action observation OR action-observation OR ao OR aot) OR (observ* OR watch*)) OR (action* OR learn* OR movement* OR reach* OR activit* OR task* OR motion* OR motor train* OR perform* OR gestur* OR demonstrat* OR video* OR tv screen OR television screen OR computer screen) AND (db:("LILACS") AND type_of_study:("clinical_trials")) AND (year_cluster:[2016 TO 2021])

Appendix 6. Allied and Complementary Medicine Database (AMED) search strategy

1. cerebrovascular disorders/ or cerebral hemorrhage/ or cerebral infarction/ or cerebral ischemia/ or cerebrovascular accident/ or stroke/

- 2. (stroke or poststroke or post-stroke or cerebrovasc\$ or brain vasc\$ or cerebral vasc\$ or cva\$ or apoplex\$ or SAH).tw.
- 3. ((brain\$ or cerebr\$ or cerebell\$ or intracran\$ or intracerebral) adj5 (isch?emi\$ or infarct\$ or thrombo\$ or emboli\$ or occlus\$)).tw.
- 4. ((brain\$ or cerebr\$ or cerebell\$ or intracerebral or intracranial or subarachnoid) adj5 (haemorrhage\$ or hemorrhage\$ or haematoma\$ or hematoma\$ or bleed\$)).tw.
- 5. hemiplegia/
- 6. (hemipleg\$ or hemipar\$ or paresis or paretic).tw.
- 7. or/1-6
- 8. exp arm/
- 9. (upper limb\$ or upper extremit\$ or arm or arms or shoulder or shoulders or hand or hands or axilla\$ or elbow\$ or forearm\$ or finger \$ or wrist\$).tw.
- 10. 8 or 9
- 11. psychomotor performance/ or Movement/ or Motion/
- 12. Visual perception/
- 13. Learning/
- 14. Photic stimulation/
- 15. (action observation or action-observation or AO or AOT).tw.
- 16. ((observ\$ or watch\$) adj10 (action\$ or learn\$ or movement\$ or reach\$ or activit\$ or task\$ or motion\$ or motor train\$ or perform\$ or gestur\$ or demonstrat\$ or video\$ or TV screen or television screen or computer screen)).tw.
- 17. ((visual or action or motion) adj5 perception).tw.
- 18. visual-motor matching.tw.
- 19. mirror neur\$.tw.
- 20. or/11-19
- 21. 7 and 10 and 20

Action observation for upper limb rehabilitation after stroke (Review)

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Appendix 7. PEDro search strategy

Abstract and Title: stroke*

Body part: upper arm, shoulder or shoulder girdle / forearm or elbow / hand or wrist.

Therapy: Neurodevelopmental therapy, neurofacilitation

Methods: Clinical trial

All search terms in the title or abstract were combined with body part descriptors using the AND operator.

Appendix 8. US National Institutes of Health Ongoing Trials Register (ClinicalTrials.gov)

action observation AND INFLECT EXACT "Interventional" [STUDY-TYPES] AND (Cerebrovascular Disorders OR Stroke OR Transient Ischaemic Attack OR Subarachnoid) [DISEASE]

Appendix 9. ISRCTN Registry

Text search: stroke

Interventions: action observation

Appendix 10. Australian New Zealand Clinical Trials Registry

Search terms: cerebrovascular or cerebral or stroke or "transient ischemic attack or TIA or SAH" Description of intervention(s) / exposure: "action observation"

Appendix 11. WHO International Clinical Trials Registry Platform search strategy

Condition: stroke

Intervention: action observation

WHAT'S NEW

Date	Event	Description
18 November 2021	New citation required but conclusions have not changed	The conclusions of the review have not changed.
18 November 2021	New search has been performed	We updated the searches to May 2021. We added four new trials, bringing the total number of included trials to 16, with a total of 574 particpiants.

HISTORY

Protocol first published: Issue 9, 2015 Review first published: Issue 10, 2018

CONTRIBUTIONS OF AUTHORS

Lorenna Borges (LB): conducted the review, assessed risk of bias and certainty of the evidence, performed statistical analyses, interpreted the results, and was responsible for writing the review.

Aline Fernandes (AF): data extraction and assessment of risk of bias. She also helped in methodological planning and contributed to writing the review.

Jacilda Passos (JP): study selection and data extraction. She also revised the manuscript.

Tania Campos (TC): was the reviewing judge, helped interpret the results. She also revised the manuscript.

Isabelle Rego (IR): study selection and data extraction. She also revised the manuscript.

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DECLARATIONS OF INTEREST

Lorenna RDM Borges: none known Aline Fernandes: none known Jacilda Passos: none known Isabelle Rego: none known Tania F Campos: none known

SOURCES OF SUPPORT

Internal sources

• Department of Physical Therapy, Federal University of Rio Grande do Norte, Brazil

External sources

• No sources of support provided

DIFFERENCES BETWEEN PROTOCOL AND REVIEW

There were a number of differences between what we planned in the protocol and the final review. Sensitivity analysis should be conducted after removing trials with high risk of allocation concealment and blinding of outcome assessment. We did this, disregarding only two trials that exhibited high risk of bias for allocation concealment. Furthermore we carried out two sensitivity analyses to determine whether the effect of AO changed with the mode of application: one removing trials in which therapy took place in the participants' home, and the other in which the action was demonstrated by the therapist and not through video.

We also intended to conduct subgroup analysis considering the type of treatment, as follows: one subgroup in which therapy was based on AO alone and another where observation was followed by physical exercise. However, there were no trials that applied AO alone to upper limb motor function; and therefore we decided to conduct subgroup analysis considering the types of control groups (placebo, conventional physiotherapy, and with no specific therapy). We performed another subgroup analysis considering the duration of observation.

Although not foreseen in the protocol, we also conducted a subgroup analysis based on upper limb motor impairment.

When updating this review, the search of other resources did not include conference proceedings, as the search in databases, newspapers, and citation indexes captured abstracts published in relevant events. We decided not to update Rehabdata due to the small number of trials found on this database in the first version of this review. It was necessary to use another search strategy for the Lilacs database (Latin American and Caribbean Health Sciences Literature; Appendix 5) to complete the search for theses, dissertations, and other records.

INDEX TERMS

Medical Subject Headings (MeSH)

Activities of Daily Living; Physical Therapy Modalities; Randomized Controlled Trials as Topic; *Stroke [complications]; *Stroke Rehabilitation; Upper Extremity

MeSH check words

Adult; Humans