Efficacy of acupuncture in the management of post-apoplectic aphasia: a systematic review and meta-analysis of randomized controlled trials

He-yong Tang, Wei Tang, Feng Yang, Wei-wei Wu and Guo-ming Shen

Abstract
Background: Aim of this study was to evaluate the effectiveness of scalp, tongue, and Jin’s 3-needle acupuncture for the improvement of postapoplectic aphasia.

Method: PubMed, Cochrane, Embase databases were searched using index words to identify qualifying randomized controlled trials (RCTs). Meta-analyses of odds ratios (OR) or standardized mean differences (SMD) were performed to evaluate the outcomes between investigational (scalp / tongue / Jin’s 3-needle acupuncture) and control (traditional acupuncture; TA and/or rehabilitation training; RT) groups.

Results: Thirty-two RCTs (1310 participants in investigational group and 1270 in control group) were included. Compared to TA, (OR 3.05 [95% CI: 1.77, 5.28]; p<0.00001), tongue acupuncture (OR 3.49 [1.99, 6.11]; p<0.00001), and Jin’s 3-needle therapy (OR 2.47 [1.10, 5.53]; p = 0.03) had significantly better total effective rate. Compared to RT, scalp acupuncture (OR 4.24 [95% CI: 1.68, 10.74]; p = 0.002) and scalp acupuncture with tongue acupuncture (OR 7.36 [3.33, 16.23]; p<0.00001) had significantly better total effective rate. In comparison with TA/RT, scalp acupuncture, tongue acupuncture, scalp acupuncture with tongue acupuncture, and Jin’s three-needling significantly improved ABC, oral expression, comprehension, writing and reading scores.

Conclusion: As treatments to postapoplectic aphasia, scalp / tongue acupuncture and Jin’s Three-needling are found better than TA and/or RT in yielding total effective rate and improving ABC, oral expression, comprehension, reading and writing scores.

Keywords: Acupuncture, Postapoplectic aphasia, Meta-analysis, Randomized controlled trial

Background
Stroke is a major cause of disability and mortality worldwide [1–3]. In 2013, stroke was the second most common cause of death (11.8% of all deaths) worldwide when the incidence of stroke was 3.28 million in women and 3.62 million in men [4]. Worldwide, stroke is the second leading cause of death and third biggest source of disability. Incidence of stroke has doubled in low-income countries in the last 4 decades whereas it has decreased by 42% in high-income countries. Stroke-related mortality is also much higher in low-income countries [5]. According to the American Heart Association, approximately 600,000 new individuals suffer stroke and 185,000 suffer recurrence of stroke each year. Healthcare and societal costs have increased from $53.6 billion to $68.9 billion from 2004 to 2009 (22% increase) [6].

Besides aging population, the contributory factors include work-related stress and dietary and lifestyle patterns due to which stroke morbidity is increasing even in younger individuals. Modifiable risk factors for the incidence of stroke include high blood pressure, diabetes, tobacco smoking, hyperlipidemia, obesity, unhealthy diet, and physical inactivity [7]. Besides significant mortality, post-
stroke disability causes a great burden to sufferers and caregivers.

Acupuncture is a prominent treatment in traditional Chinese medicine for over 3000 years for several diseases including post-stroke recovery. Among important methods, scalp acupuncture is carried out based on the functionality of brain areas to stimulate different scalp zones with needles which improves the reflexively of certain nerves [8]. Tongue acupuncture involves insertion of acupuncture needle into the acupoints on tongue followed by twisting it usually for multiple times [9]. In Jin’s 3 needles acupuncture, 3 needles are inserted on strategic acupoints depending on the target symptom such as slurred speech, difficulty swallowing, spastic paralysis, aphasia, etc. [10].

Previously, Zhang et al. [11] and Wu et al. [12] conducted meta-analyses of studies which performed acupuncture for poststroke rehabilitation and found that acupuncture provided significant benefits in stroke rehabilitation. The Ottawa Panel evidence-based clinical practice guidelines also identify acupuncture as one of the treatment options for post-stroke rehabilitation [13]. The purpose of this study was to comprehensively review the randomized controlled trials (RCTs) which evaluated the effectiveness of scalp acupuncture, tongue acupuncture, and Jin’s 3-needling by comparing with traditional acupuncture or rehabilitation training for participants with post-stroke aphasia and to perform meta-analyses of important endpoints in order to gain clinically meaningful synthesis of present day evidence.

Methods

Search strategy

Literature search was conducted in PubMed, Cochrane and Embase databases for RCTs which evaluated the effectiveness of acupuncture for the rehabilitation of post-apoplectic aphasia. Important keywords used for the search included acupuncture, scalp, tongue, intradermal, acupoint, apoplexy, stroke, cerebral apoplexy, cerebral infarction, cerebral hemorrhage, aphasia, cerebral infarction, randomized controlled trial, and RCT. Full literature search strategy is given as Additional file 1: Appendix S1 (online Supporting information file). Reference lists of important research and review articles were also screened. The search was completed by October 2018. Two reviewers conducted literature survey independently and then unified their outputs. For the screening and selection of studies, help of an additional researcher was sought who participated in eligibility criteria formulation earlier.

Inclusion and exclusion criteria

A study was included if (i) was RCT, (ii) subjects had post-apoplectic aphasia without further severe disease diagnosed with WHO criteria using appropriate radiological methods (iii) the interventions included scalp acupuncture, tongue acupuncture, or Jin’s 3-needling, (iv) had a control group with traditional acupuncture and/or rehabilitation training interventions, and (v) published in English or Chinese language.

A study was excluded if: (i) the outcomes were based on acupuncture or related therapies other than scalp acupuncture, tongue acupuncture, Jin’s 3-needling or traditional acupuncture, (ii) the interventions of either treatment or control group included medication, (iii) outcomes were reported in formats that could not be used in meta-analyses of odds ratio (OR) or standardized mean difference (SMD), or (iv) was systematic review, meta-analysis, theoretical research, expert commentary, conference report, economic analysis, or was a case report/study.

Data extraction, quality assessment and data analysis

Demographic data including the name of author (study identity), publication year, age, gender sample size, study quality (Jadad score), interventions of the treatment and control groups, and outcome data including total effective rate, the Aphasia Battery of Chinese (ABC) score, and the scores of comprehension, oral expression, repetition, denomination, reading and writing tests were extracted from the individual studies and organized in datasheets. Data were extracted by 2 reviewers independently which were then unified and validated with the involvement of a third reviewer.

The Jadad checklist was employed to assess the quality of the included studies. All RCTs were evaluated on the basis of five items: (i) statement of randomization, (ii) use of double-blind methods, (iii) appropriateness of generating randomized sequence, (iv) details of withdrawals and dropouts and (v) description of double blinding method. Studies with a score of < 3 signified low-quality and a high bias risk.

Meta-analyses of SMD were performed using STATA software (version 10.0; Stata Corporation, Texas, USA). Outcome variables reported as means and standard deviations (the ABC scores and subscale scores of comprehension / oral expression / repetition / denomination / reading / writing) were used to calculate SMD between treatment and control groups and then inverse variances weighted overall effect sizes were generated. Categorical outcomes (total effective rates) were used to perform meta-analyses of odds ratios (OR) using Review Manager (version 5.1.3, Cochrane) software.

The statistical heterogeneity of clinical trial data was assessed with Chi-squared and I² indices. A Chi-squared P values of ≤0.05 and I² > 50% was considered to show high heterogeneity of outcome data. In subgroup analysis, four subgroups were: (1) scalp acupuncture, (2) tongue acupuncture, (3) Jin’s 3-needling, and (4) scalp acupuncture with tongue acupuncture.
Results
Database search identified 1921 publications of which 1832 were excluded after title and/or abstract screening and 89 were subjected to eligibility criteria application. After review of the complete manuscripts, 57 articles were further excluded because 18 were reporting non-RCT studies as 15 involved theoretical or economic research, 13 had unqualified interventions, and 11 had no clinical outcomes. Finally, 32 RCTs were selected with 1310 participants in investigational group and 1270 in control group [14–45]. The screening and selection procedure is presented in Fig. 1. The study characteristics are presented in Additional file 1: Table S1.

The quality of the included studies was low, in general, as the overall Jadad score of included studies was 2.4. The funnel plot for total effective rate was symmetrical, depicting a lack of publication bias (Fig. 2). Begg’s and Mazumdar’s rank test (Z = 1.36, p = 0.173) and the Egger’s test (p = 0.362) also endorsed this finding.

Total effective rate
Total effective rate was reported by 29 studies with 1148 participants in investigational and 1108 in control groups. Compared to traditional acupuncture (TA), scalp acupuncture (OR 3.05 [95% CI: 1.77, 5.28]; p<0.00001), tongue acupuncture (OR 3.49 [1.99, 6.11]; p<0.00001), and Jin’s 3-needle therapy (OR 2.47 [1.10, 5.53]; p = 0.03 had significantly better total effective rate (Fig. 3a). Compared to rehabilitation training (RT), scalp acupuncture (OR 4.24 [95% CI: 1.68, 10.74]; p = 0.002) and scalp acupuncture with tongue acupuncture (OR 7.36 [3.33, 16.23]; p<0.00001) had significantly better total effective rate (Fig. 3b).

The ABC score
The ABC score was reported by 10 studies with 407 in investigational and 383 participants in control group. Investigational interventions improved the ABC score significantly more than control interventions (SMD: 2.05 [95% CI: 1.38, 2.73]; Fig. 4). In the subgroup analysis, we found that in comparison with control interventions, the ABC score improved significantly more with scalp acupuncture (SMD: 2.45 [95% CI: 0.84, 4.07]), tongue acupuncture (SMD 1.96 [95% CI: 0.41, 3.51]), Jin’s 3-needling (SMD: 0.84 [95% CI: 0.29, 1.38]), and scalp acupuncture with tongue acupuncture (SMD 2.53 [95% CI: 1.75, 3.32]) (Additional file 1: Figure S1).

The comprehension score
The comprehension score was reported by 7 studies with 283 participants in investigational and 278 in control group. Overall, the comprehension score was significantly higher in investigational group compared to the control group (SMD: 0.54 [95% CI, 0.21, 0.86]; Additional file 1: Figure S2). In subgroup analysis, comprehension score was significantly higher after tongue acupuncture (SMD 0.85 [95% CI: 0.32, 1.38]) and scalp acupuncture with tongue acupuncture (SMD 0.50 [95% CI: 0.04, 0.97]) in comparison with control group Additional file 1: Figure S2). However, in other subgroups, there was no significant difference between the groups.

The oral expression score
The oral expression score was reported by 6 studies with 254 participants in the investigational group and 250 in the control group. Compared to the control
group, investigational group had significantly higher oral expression score (SMD 0.80 [95% CI: 0.44, 1.16]; Additional file 1: Figure S3). In the subgroup analysis, compared to control interventions, tongue acupuncture (SMD 0.89 [95% CI: 0.45, 1.32]), scalp acupuncture (SMD 1.92 [95% CI: 1.31, 2.54]), Jin’s 3-needling (SMD 0.96 [95% CI: 0.48, 1.44]) and scalp acupuncture with tongue acupuncture (SMD 0.71 [95% CI: 0.38, 1.03] significantly improved the oral expression score (Additional file 1: Figure S3).

The repetition score
The repetition score was reported by 7 studies with 267 participants in investigational group and 263 in control group. Investigational group had significantly higher repetition score compared to the control group (SMD 1.05 [95% CI: 0.64, 1.47]; Additional file 1: Figure S4). In the subgroup analysis, in comparison with control interventions, the repetition score was significantly higher after tongue acupuncture (SMD 1.95 [95% CI: 1.51, 2.39]), scalp acupuncture with tongue acupuncture (SMD 1.03 [95% CI: 0.34, 1.72]), and Jin’s 3-needling (SMD 0.78 [95% CI: 0.19, 1.36]) (Additional file 1: Figure S4).

The denomination score
The denomination score was reported by 6 studies with 231 participants in the investigational group and 228 in the control group. Compared to the control group, investigational group had significantly higher denomination score (SMD 0.82 [95% CI: 0.20, 1.44]; Additional file 1: Figure S5). In the subgroup analysis, in comparison with control interventions, tongue acupuncture (SMD 1.38 [95% CI: 0.42, 2.34]), scalp acupuncture with tongue acupuncture (SMD 1.16 [95% CI: 0.60, 1.71]), and Jin’s 3-needling (SMD 1.25 [95% CI: 0.88, 1.62] were associated with significantly higher denomination score (Additional file 1: Figure S5).

The reading score
The reading score was reported by 4 studies with 147 participants in the investigational group and 147 in the control group. Investigational group had significantly higher reading score compared to the control group (SMD 0.90 [95% CI: 0.51, 1.30]; Additional file 1: Figure S6). In subgroup analysis, the reading score was significantly higher with scalp acupuncture (SMD 0.48 [95% CI 0.06, 0.90], tongue acupuncture (SMD 1.53 [95% CI: 0.95, 2.10], scalp acupuncture with tongue acupuncture (SMD 0.75 [95% CI: 0.28, 1.23]), and Jin’s 3-needling (SMD 1.08 [95% CI: 0.52, 1.64]) in comparison with control interventions (Additional file 1: Figure S6).

The writing score
The writing score was reported by 4 studies with 147 participants in the investigational group and 147 in the control group. Compared to the control group, investigational group had significantly higher writing score after therapy (SMD 1.56 [95% CI: 0.62, 2.49]; Additional file 1: Figure S7). In subgroup analysis, the writing score was significantly higher in scalp acupuncture (SMD 1.20 [95% CI: 0.76, 1.65]), tongue acupuncture (SMD 3.72 [95% CI: 2.88, 4.57]), and scalp acupuncture with tongue acupuncture (SMD 1.36 [95% CI: 0.82, 1.91] in comparison with control interventions (Additional file 1: Figure S8). However, there was no statistically significant difference in the writing score after Jin’s 3-needling (SMD 0.29 [95% CI: −0.23, 0.81] in comparison with control intervention (Additional file 1: Figure S7).
Fig. 3 Forest graph showing the meta-analysis of odds ratios of the total effective rate in the (a) investigational group compared to the (b) control group.

### (a) Scalp acupuncture
- **Cai 2014**: 42 events, 48 control events, total 50 events, weight 8.5%, odds ratio 5.00 [1.79, 14.01]
- **Jiang 2001**: 26 events, 30 control events, total 56 events, weight 5.8%, odds ratio 3.82 [1.03, 14.18]
- **Wang S 2009**: 45 events, 50 control events, total 95 events, weight 7.5%, odds ratio 3.86 [1.23, 12.12]
- **Xu 2005**: 30 events, 30 control events, total 60 events, weight 1.1%, odds ratio 7.76 [0.28, 157.71]
- **Zheng X 2010**: 33 events, 40 control events, total 73 events, weight 14.4%, odds ratio 0.63 [0.25, 2.74]
- **Subtotal (95% CI)**: 103 events, 188 control events, total 291 events, weight 37.3%, odds ratio 3.03 [1.17, 7.98]

**Total events**: 176, 134

*Heterogeneity: $\chi^2 = 6.10, df = 4 (P = 0.19); I^2 = 34%*

*Test for overall effect: Z = 4.01 (P < 0.0001)*

### (b) Tongue acupuncture
- **Liu X 2009**: 26 events, 30 control events, total 56 events, weight 7.1%, odds ratio 2.36 [0.63, 8.92]
- **Li Z 2005**: 43 events, 46 control events, total 89 events, weight 4.4%, odds ratio 6.31 [1.61, 24.78]
- **Luo 2007**: 49 events, 52 control events, total 101 events, weight 4.9%, odds ratio 6.17 [1.79, 25.21]
- **M 2004**: 37 events, 46 control events, total 83 events, weight 13.5%, odds ratio 1.90 [0.70, 5.15]
- **Zheng X 2005**: 30 events, 32 control events, total 62 events, weight 3.7%, odds ratio 3.76 [0.69, 20.20]
- **Subtotal (95% CI)**: 206 events, 382 control events, total 590 events, weight 33.7%, odds ratio 3.49 [1.49, 8.11]

**Total events**: 185, 131

*Heterogeneity: $\chi^2 = 3.43, df = 4 (P = 0.49); I^2 = 0%*

*Test for overall effect: Z = 4.37 (P < 0.0001)*

### (c) Jin's three-needle therapy
- **Cui 2013**: 26 events, 29 control events, total 55 events, weight 5.6%, odds ratio 2.86 [1.03, 7.69]
- **Han 2013**: 31 events, 30 control events, total 61 events, weight 8.5%, odds ratio 1.21 [0.29, 4.97]
- **Jiang 2008**: 37 events, 40 control events, total 77 events, weight 4.7%, odds ratio 4.74 [1.19, 18.94]
- **Subtotal (95% CI)**: 100 events, 97 control events, total 197 events, weight 19.1%, odds ratio 2.47 [1.16, 5.33]

**Total events**: 60, 76

*Heterogeneity: $\chi^2 = 1.84, df = 2 (P = 0.40); I^2 = 0%*

*Test for overall effect: Z = 2.20 (P = 0.03)*

*Total (95% CI)**: 581, 521

*Favours Control: 390, Favours Treatment: 191*

### Odds ratio

- **Total (95% CI)**: 3.00 [2.15, 4.20]
- **Total events**: 503, 388

*Heterogeneity: $\chi^2 = 12.19, df = 14 (P = 0.59); I^2 = 0%*

*Test for overall effect: Z = 6.42 (P < 0.0001)*

*Test for subgroup differences: $\chi^2 = 0.79, df = 3 (P = 0.65), I^2 = 0%*
Discussion

This meta-analysis found that compared to traditional acupuncture, scalp / tongue / Jin’s 3-needle acupuncture yield significantly better total effective rate, whereas compared to rehabilitation training, scalp acupuncture and scalp acupuncture with tongue acupuncture yield significantly better total effective rate. The ABC score also improved significantly more in scalp acupuncture, tongue acupuncture, Jin’s 3-needling, and scalp acupuncture with tongue acupuncture in comparison with traditional acupuncture or rehabilitation training. Moreover, comprehension, oral expression, repetition, denomination, reading, and writing also improved more with scalp, tongue and Jin’s 3-needle acupuncture, in general.

Previously, Zhang et al. [11] who conducted a meta-analysis of studies which used acupuncture for poststroke aphasia found that acupuncture provides benefits in swallowing (RR 1.61 [95% CI 0.73, 3.58]), neurological function (RR 6.55 [95% CI 1.89, 22.76]) and disability (RR 12.5 [95% CI 43.1,36.2]). Wu et al. [12] also conducted a meta-analysis of 38 trials and found an OR of 4.33 [95% CI 3.09, 6.08] in favor of acupuncture for providing improvement in poststroke rehabilitation but they concluded that quality of included studies was inadequate, and possibility of publication could not be ruled out.

In the present study, not only the total effective rate was better with scalp acupuncture, but it was also associated with significantly improved ABC score and its sub-scores. Scalp acupuncture stimulate different scalp zones to improve the reflexively of nervous tissue [8]. Litscher et al. [46] after monitoring cerebral oxygen saturation in 12 subjects during and after acupuncture have shown that cerebral oxygen saturation increased from 69.9% before to 70.3% during and 70.2% after acupuncture (p < 0.01). Studies in animals and humans have proven the effectiveness of scalp acupuncture in improving neurologic deficits by altering hormone levels in circulation and blood flow in the brain [47].

Tongue acupuncture was also associated with higher effectiveness rate and significantly improved ABC score and communication skills including comprehension, oral expression, repetition, denomination, reading and writing, in the present study. Tongue acupuncture targets acupoints on tongue for either single or repetitive stimulation [9]. Tongue acupuncture is found beneficial in improving symptoms in participants with chronic neurological conditions including autism, cerebral palsy, stroke, and drooling problems. Because tongue is an important part which is involved in many ingestion tasks and speech, it is rich in neural networks, vascular supply, and lymphatics. Thus, stimulation of key areas which link the vascularelymphatic neural networks by identifying acupoints to insert needles may trigger re-signaling of nerves and the potentiation of neurotransmission [48].
We have also found that Jin’s 3-needles acupuncture significantly improved ABC scores and communication skills. In the present study, total effective rate was statistically non-significantly higher, and the ABC score was significantly better in Jin’s 3-needles acupuncture group, besides oral expression, repetition, denomination, reading and writing improved significantly in Jin’s 3-needles acupuncture group. Jin’s 3 needles acupuncture is usually based on symptom such as slurred speech, difficulty swallowing, spastic paralysis, aphasia, etc. [10]. Several authors have reported that Jin’s 3-needle acupuncture was effective in improving cognitive and motor functions, hemiplegia and activities of daily living in stroke participants. Increase in cerebral blood flow coinciding with electrophysiological activities has been reported by many studies [49].

Although, the findings of the present study support the use of scalp, tongue, and Jin’s 3-needle acupuncture for post-stroke rehabilitation based on data obtained from 32 studies, many caveats are needed to be taken care of before clinical implications of this evidence could be reindeed. Quality of research is not adequate so far. There is less information available with regards to the follow-up of studies. There is also a need to use health-related quality of life tools to appraise participants for their outcomes with reasonable follow-up. If such a survey could be inculcated with a large sized randomized controlled trial, it could be the most useful.

In stroke rehabilitation there is essential and urgent need that patient should be involved in a total complex rehabilitation work, which should be built according to most recent situation and symptoms of individual patients. Moving exercises, professional logopedics intervention are much needed, but daily gymnastic (active/passive) is necessary. Acupuncture is a valuable and less expensive additional link to have better output from rehabilitation process. The number of acupuncture clients is increasing and therefore this modality can be benefitted with research and feedback.

Some limitations of the present study should be considered while interpreting the results. These include: (i) only randomized controlled trails were used, (ii) variables with regards to previous disease condition and treatments were unavailable for many studies, (iv) some studies of low quality and with a low Jadad score were included, and (v) treatment time and protocols may have varied across the included studies.

Conclusions
In this meta-analysis, we have found that, compared to traditional acupuncture and/or rehabilitation training, scalp acupuncture, tongue acupuncture, and Jin’ 3-needle acupuncture can better improve post-apoplectic aphasia as depicted by the total effective rate, the ABC score, and comprehension, oral expression, repetition, denomination, reading and writing scores. However, quality of the included studies was inadequate and therefore further high-quality studies with larger samples and longer follow-up times and with patient outcomes are necessary to verify the results presented herein. In future studies, researchers should also explore the efficacy and differences between scalp acupuncture, tongue acupuncture and Jin’s 3-needling in the treatment of postapoplectic aphasia.

Supplementary information
Supplementary information accompanies this paper at https://doi.org/10.1186/s12906-019-2687-1.

Additional file 1: Table 1. Characteristics of the included studies.

Figure S1. Forest graph showing the subgroup analyses of the SMDs in ABC scores between investigational and control groups. Figure S2. Forest plot of the score of comprehension in the investigational group compared to the control group in the subgroup analysis. Figure S3. Forest plot of the score of oral expression in the investigational group compared to the control group in the subgroup analysis. Figure S4. Forest plot of the score of repetition in the investigational group compared to the control group in the subgroup analysis. Figure S5. Forest plot of the score of denomination in the investigational group compared to the control group in the subgroup analysis. Figure S6. Forest plot of the score of reading in the investigational group compared to the control group in the subgroup analysis. Figure S7. Forest plot of the score of writing in the investigational group compared to the control group in the subgroup analysis. Appendix S1 (Literature search strategy).

Abbreviations
ABC: Aphasia Battery of Chinese; MD: Mean difference; OR: Odds ratio; RR: Relative risk.

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Not applicable.

Authors’ contributions
HYT and GMS designed experiments; WT carried out experiments; FY and WWW analyzed experimental results. HYT wrote the manuscript; GMS approved the manuscript. All authors read and approved the final manuscript.

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Availability of data and materials
The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request.

Ethics approval and consent to participate
Not applicable.

Consent for publication
Not applicable.

Competing interests
The authors declare that they have no competing interests.

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