

Research Article

Repetitive Transcranial Magnetic Stimulation in Aphasia and Communication Impairment in Post-Stroke: Systematic Review of Literature

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Keywords

- Aphasia
- Post-stroke
- Repetitive transcranial magnetic stimulation

Abstract

Background: Post-stroke aphasia is one of the major disabilities and a risk factor for other complications, is also associated with increased mortality, depression and impairments in communication. Repetitive Transcranial Magnetic Stimulation (rTMS) is an alternative. We propose the review of literature published until now of rTMS.

Methods: We included articles published on PubMed and EMBASE, we included studies if were: randomized controlled blinded clinical trials, meta-analyses or crossover designs of rTMS alone or with speech therapy or any other therapy tested with rTMS. One author included these if: report baseline assessment and one or more posterior assessments; overall results and subtest report; statistically significant results showed by p value. We also assessed the risk of bias of each article.

Main results: We included 15 articles, the average age was 62.42 ± 4.04 ; most used low frequency stimulation (1 Hz), most used a coil of 70mm, 77% applied 10 sessions, most did a Speech language therapy. From overall test result: 66% studies improved Aphasia; room subtest results: only 1 study improved listening, speaking and reading; 2 studies improved writing, 80% studies improved naming skills, Two studies improved writing and description, 3 studies improved compression and expression, 46% improved repetition. 47% of studies used Boston battery, 33% used Aachen Aphasia Test.

Conclusions: The articles included showed improvement of scale results for aphasia in post-stroke patients, mainly due to improvement in naming and repetition skills.

ABBREVIATIONS

rTMS: Repetitive Transcranial Magnetic Stimulation; SLT: Speech Language Therapy; BDAE: Boston Diagnostic Aphasia Examination; AAT: Aachen Aphasia Test; BNT: Boston Naming Test; CCAT: Criteria Cognitive Aptitude Test; KWAB: Western Aphasia Battery; CPNT: Computerized Picture Naming Test; ASRS: Aphasia Severity Rating Scale; ANELT: Amsterdam-Nijmegen Everyday Language Test; HSS: High Sensation Seeking Test; HRQL: Health-Related Quality of Life; SAQOL: Aphasia Quality of Life Scale-39; PRO: Patient Reported Outcomes

INTRODUCTION

Aphasia is one of the major disabilities and a risk factor for other complications in post-stroke patients. The worldwide incidence of stroke is about 217 per 100.000 person years and

its prevalence, 715 per 100.000. In high income countries, the aphasia incidence in first-ever ischemic stroke amounted to 43 per 100.000 inhabitants [1,2]. Unlike any other condition, disabilities due to stroke are multiple, affecting: walking, speech, balance; co-ordination, vision, spatial awareness, swallowing; bladder and bowel control [3]. Aphasia as sequel of stroke is associated with increased mortality (36% vs. 16%), depression (70% of aphasic at 3 months) and impairments in communication [4-7]. Social participation for people with aphasia is affected; they communicate with fewer friends, have poor social networks and negative impacts on-quality of life [8-10].

Aphasia is an acquired disorder that affects the ability to use and/or understand language; it involves various components: meaning (semantics) sounds (phonology) and structure (syntax/morphology) [11]. Aphasia has many classifications

and these can be used to guide treatment selection. The classic classifications are based on: fluency, language understanding and preserved repeated speech and syndromes (global, Broca, isolated, transcortical motor; Wernicke, transcortical sensory, conduction and anomic), but remain based on elementary clinical characteristics of dichotomies (motor-sensory, expressive-receptive, fluent or nonfluent) [12]. In recent years, researches showed more participation of sub cortical gray matter structures in brain that could explain language impairment in brain areas different of the lobes in post-stroke patients. Pedersen et al, found one year after stroke: global 7%, Broca's 13%, transcortical motor 1%, wernicke's 5%, transcortical, conduction 6% and anomic 29% [13]. Language deficits assessment is an important research area, clinical practice and rehabilitation [14]; standardised batteries for aphasia gathered information from different language subtest that assesses: spontaneous speech, comprehension, repetition and naming [14]. This is important to focus aphasia therapy in specific language impairment.

Post-stroke aphasia treatment is a challenge that includes: deal with own aphasia and its associated comorbidities as depression, deglutition dysfunction and cognitive deficit. Speech-language therapy is considered the mainstay treatment of aphasia. Medications (piracetam, bromocriptine, dexamfetamine, donezopil,) are also considered, but there is a lack of consensus about their effectiveness and the best time to start with speech language therapy (SLT). In this scenario, repetitive Transcranial Magnetic Stimulation (rTMS) is an alternative for aphasia treatment; however, it remains as an experimental alternative due to lack of epidemiological evidence about its general effectiveness or subtest efficacy, and how it is clinically significant for daily communication. We proposed to carry out a systematic review of literature of studies published for rTMS, with the objective of finding evidence of efficacy and sub-test results that could be useful for individualized patient treatment.

MATERIALS AND METHODS

We performed the search on May 2016, we included articles published until May 2016 on PubMed and EMBASE, using the following terms: (transcranial magnetic stimulation or rTMS) AND (aphasia) AND (Stroke OR post stroke) (clinical trial or meta-analysis). We did not restrict for age or other study characteristics. The studies were selected if they met the inclusion and exclusion criteria: clinical trials, meta-analyses or crossover designs of transcranial magnetic stimulation alone or with speech therapy or any other therapy tested with rTMS. Figure (1), shows the flow chart of the search and the included and excluded studies. The results were filtered to discard duplicate items.

One of the authors selected the articles by title; after that, we reviewed the articles pre-selected to include them in a first step; from this first step, we excluded 10 articles due to duplicate. Other author reviewed the rest of the studies included, and chose those articles considered relevant, and finally took off others that did not meet all of the inclusion criteria. Finally, the articles were put in a spreadsheet designed to describe, classify and check results from each study; all Results of each study were put there to describe general characteristics and the scales used. We included study data, if they were reported as: baseline assessment and one or two posterior assessments; overall results and subtest report;

statistically significant results showed by p value. We also did a risk of bias using the PEDro scale of clinical trials: the scale uses different criteria for overall evaluation. Results of risk bias were represented using REVMAN software of Cochrane systematic review Figures (2,3).

RESULTS

From the studies retrieved and included in the systematic review, (13 clinical trials and 2 meta-analyses) we described the general characteristics of each study. The number of subjects included ranged from 8 to 56. The types of aphasia included were fluent and no-fluent, the average age was 62.42 ± 4.04 , the mean time post-stroke included was 20.38 ± 20 months, and all studies stimulated contra lateral area. Most of studies did speech-language therapy additional to rTMS. All the studies included sham therapy as comparator and used an aphasia scale to measure outcomes.

We also described the therapeutic application of repetitive transcranial magnetic stimulation (Table 1,2), most of studies included, used low frequency stimulation (1 Hz) only Szaflarski study used 50 Hz every 200 milliseconds in two second trains, repeated every 10 seconds over 200 seconds for a total of 600 pulses. Related to the coil, most of studies used a 70mm coil; Khedr et al used a 90mm coil and one study did not report it. 11 of 13 (without meta-analyses) used the Magstim® stimulator; 2 studies reported use of magPro® stimulator. Related to the number of sessions, 77% (10/13) reported 10 stimulations every day (except weekends), two studies 15 days, and one study 20 sessions. Authors also reported length of each stimulation, 73% of studies stimulate 20 minutes in average; Hara et al reported 40 minutes of stimulation each session, Waldowski et al., stimulate 30 minutes, two studies did not report time of each stimulation session. Of stimulation sites, all of them stimulate the contra lateral area Broca's area. Sham strategy for stimulation were some: stimulation in vertex, same place but a sham coil; same localization but less than 5% of stimulation intensity and same site but without skull contact. All studies performed a SLT following the stimulation, the type, length and intensity depended of each investigator.

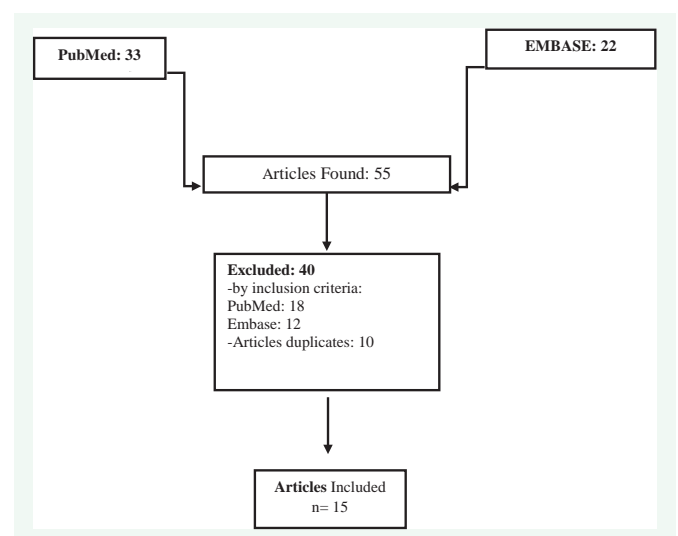


Figure 1 Studies included in the review.

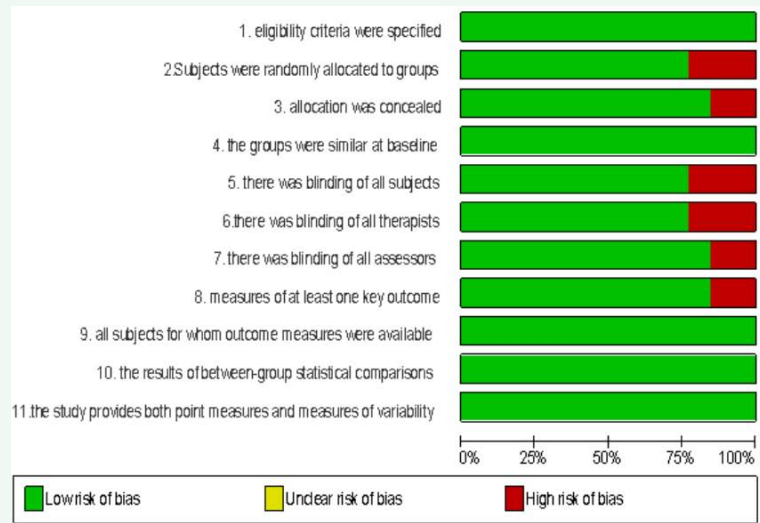


Figure 2 peDro scale risk of bias by criterion: these figures summarize Risks of bias, presented as percentage across all included studies.

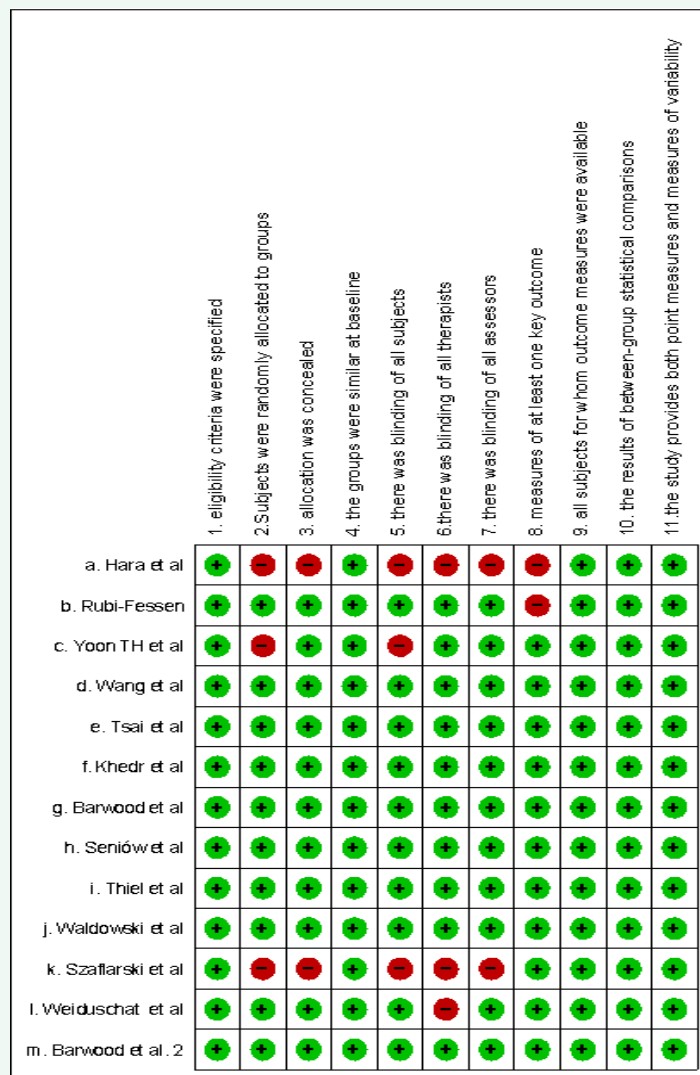


Figure 3 Risk of bias by study and criterion: Review authors' judgements about each risk of bias item for each.

Table 1: Characteristics of included studies.

Study	n	Aphasia Type	Mean age	Mean time post-stroke (months)	Study design	Site of stimulation	Experimental	Control	Outcome measurement
Hara et al [15].	50	FA / NFA	60.3	55.9±38.0	CT	CL-IL*	rTMS	—	SLTA
Rubi-fessen et al [16].	30	FA/ NFA	NI	NI	CT	CL	rTMS	Sham	AAT, FIM, Anelt A-scale
Li Y et al [19].	132	NFA/ ALL TYPE	NA	—	MA	CL	rTMS	Sham	BNT, BDAE, CCAT, AAT
Otal et al [42].	183	ALL TYPE	NI	NI	MA	CL	rTMS	Sham	AAT, BDAE, CPNT PNT
Yoon et al [21].	20	NFA	Active60.46 Sham 61.13	6,80 ± 2,39 5,20 ± 2,67	CT	CL	rTMS	Sham	K-WAB
Wang et al [20].	45	FA/NFA	TMSsyn 61.3 TMSsub 62.1 TMSsham 60.4	TMSsyn16.8±6.4 TMSsub15.7±8.5 TMSsham 16.1±7.3	CT	CL	Syn(rTMS + naming training) together/ Sub(rTMS + naming training)separate	Sham + sync	CCAT
Tsai et al [17].	56	NFA	Experimental 62.3 Sham 62.8	Experimental 17.8 ± 7.2 Sham 18.3 ± 8.2	CT	CL	rTMS	Sham	CCAT
Khedr et al [18].	30	NFA	57,3	Experimental: 2 Sham: 1	CT	CL	rTMS	Sham	HSSASRS
Barwood et al [22].	12	ALL TYPE	Active 60.8 Sham 67	Active 18.46 Sham 15.12	CT	CL	rTMS	Sham	BNT – BDAE
Seniow et al [23].	40	ALL TYPE	60.7	Active:1.10 Sham: 1.31	CT	CL	rTMS	Sham	BDAE
Thiel et al [44].	30	FA/ NFA	Active: 71.2 Sham: 69.8	Active: 1.23 Sham: 1.66	CT	CL	rTMS	Sham	AAT
Waldowski et al [45].	26	NFA	Active: 62.31 Sham: 60.15	NI	CT	CL	rTMS	Sham	CPNT ASRS BDAE
Szaflarski et al [46].	8	NFA	54.4	63.3	CT	IL	rTMS	Sham	SFT, COWAT BNT, BDAE
Weiduschat et al [47].	10	NFA	66.6	Active: 45.2 Sham:57.5	CT	CL	rTMS	Sham	AAT
Barwood et al [43].	12	NFA	Active 60.8 Sham 67	18.36	CT	CL	rTMS	Sham	BNT – BDAE

Abbreviations: FA: Fluent Aphasia; NFA: Nonfluent Aphasia; CT: Clinical Trial; CL: Contralesional, IL: Ipsilesional; SLTA: the Standard Language Test of Aphasia; AAT: Aachen Aphasia Test; FIM: The Functional Independence Measure; Anelt A-scale: Amsterdam-Nijmegen Everyday Language Test; BNT: Boston Naming Test; BDAE: Boston Diagnostic Aphasia Examination; CCAT: Criteria Cognitive Aptitude Test; AAT: Aachen Aphasia Test; PNT: Philadelphia Naming Test; K-WAB: Korean-version of the Western Aphasia Battery; ASRS: Aphasia Severity Rating Scale; HSS: High Sensation Seeking Test; CPNT: Computerized Picture Naming Test; SFT: Semantic Fluency Test; COWAT: Controlled Oral Word Association Test; Syn: Synchronous; Sub: Subsequent, rTMS: Repetitive Transcranial Magnetic Stimulation; NA: No Apply; NI: No information.

*Ipsilesional or contralesional depend on results by functional magnetic resonance imaging.

Table 2: Protocols for repetitive transcranial magnetic stimulation.

Study	1 Hz	Coil	Stimulator	Frequency	# Sessions	Length	Intensity*	Localization	sham localization	SLT After rTMS
Hara et al [15].	YES	70-mm	MagPro R30	Every day	10	40 min	90%	IFG-STG (left or right)	—	YES
Rubi-fessen et al [16].	YES	70-mm	Magstim Rapid	NI	15	20 min	90%	PT-IFG (right)	Vertex	YES
Yoon et al [21].	YES	NI	MagPro®	Every day	20	20 min	90%	IFG (right)	—	YES
Wang et al [20].	YES	70-mm	Magstim Rapid	Every day	10	20 min	90%	PT (right)	same-sham-coil	YES
Tsai et al [17].	YES	70-mm	Magstim Rapid	Every day	10	10min	90%	PT IFG(right)	Same less 5%	YES
Khedr et al [18].	YES	90-mm	Magstim Rapid	Every day	10	NI	80%	PT- Pop (right)	Same no-contact	YES
Barwood et al [22].	YES	70-mm	Magstim Rapid	Every day	10	20 min	90 %	PT- Pop (right)	same no contact	No
Seniow et al [23].	YES	70-mm	Magstim Rapid	Every day	10	20 min	90%	PT (right)	same-sham coil	YES

Thiel et al [44].	YES	70-mm	Magstim Rapid	Every day	10	20 min	90%	PT- PIG (right)	Vertex	YES
Waldowski et al [45].	YES	70-mm	Magstim Rapid	Everyday	15	30 min	90%	PT - POp (right)	same- Sham coil	YES
Szaflarski et al [46].	NO	70-mm	Magstim Rapid2	Every day	10	NI	80%	left Broca's	—	No
Weiduschat et al [47].	YES	70-mm	Magstim Rapid	Every day	10	20 min	90%	IFG (right)	Vertex	YES
Barwood et al [43].	YES	70-mm	Magstim Rapid	Every day	10	20 min	90%	PT(right)	Same-Sham coil	No

Abbreviations: % of each individual patient's motor threshold intensity; IFG: Inferior Frontal Gyrus; STG: Superior Temporal Gyrus; PT: Pars Triangularis; POp: Pars Operculus; PIG: Posterior Inferior Gyrus; NI: No Information.

Results of all studies included are shown in Table (3); by test results, 60% (9/15) studies showed overall improvement of aphasia and 6 did not. Results were also described for sub-scale tests in aphasia if they were described and had statistical significance. We included the following: listening, speaking, reading, writing, naming, comprehension, expression; repetition and description assessments. For listening, speaking and reading; only 1 study showed improvement [15] and 2 studies reported improvement in writing skills [15,16]. 80% (12/15) studies showed improvement in naming skills. Two studies reported improvement for writing and description [15,16] and [17,18], respectively, 3 studies showed improvement for comprehension and expression [16,18-19] and [16-17,20] respectively), 47% (7/15) reported improvement for repetition [16-19, 21-23].

Related to scales used on this studies, 47% (7/15) of studies used or reported results with Boston Diagnostic Aphasia Examination (BDAE), 33% of studies used Aachen Aphasia Test (AAT) or Boston Naming Test (BNT), 20% (3/15) used Criteria Cognitive Aptitude Test (CCAT) and 13% (2/15) used Korean-version of the Western Aphasia Battery (KWAB), Computerized Picture Naming Test (CPNT) and Aphasia Severity Rating Scale (ASRS). From overall test results the scales used were BDAE 30% (3/10), CCAT and AAT 20% (2/10), and 10% (1/10) with Amsterdam-Nijmegen Everyday Language Test (ANELT), SLTA and High Sensation Seeking Test (HSS). Listening, speaking and reading skills only improve in Hara's et al., study; using SLAT scale, writing results improved in Hara et al and Rubi-fessen et al., using SLTA and Anelt A scale.

DISCUSSION

Most of the studies reported improvement of aphasia post-stroke with repetitive transcranial magnetic stimulation. There were improvements in results of scales from baseline score to post-stimulation assessment score, which could be due to rTMS alone or synchronic naming training and complementary to speech therapy in aphasia post-stroke. Aphasia is one of the most challenging consequences of stroke. In this scenario, repetitive transcranial magnetic stimulation emerged as a promising alternative to improve communication skills in patients that until now only speech therapy has showed to be effective to maximize the natural recovery after stroke [24].

We identified a lack of standardization of the rTMS's protocols, which can lead to heterogeneity in the effect of the therapy or the dilution of it; in addition, the limited size of sample of studies contributes to the heterogeneity in results. We

also identified lack of homogenous uses of scales in the studies included, however; it is because the origin of each one, and the use of specific designed scale by language. Boston battery is a generic scale for aphasia, but CCAT and Anelt A scale were designed for specific languages (Chinese and Netherlands). Boston scale was used more frequently than others in this study; some authors have reported its limitations (poor psychometric properties, lack of standardization and inadequate norms), but it is still the most used test by neuropsychologists [25,26]. From subtest results the body of evidence shows improvement on naming results (12 studies), following by repetition (7 studies) and less by comprehension, expression and description. There is a lack of evidence by listening, speaking and reading (1 study); the split out of studies from sub-test results is useful to describe strength and weakness of rTMS, and possible language skills to improve with SLT. The overall test results are in the same line of other authors; Wong et al, in their systematic review of studies on the effectiveness of rTMS, found improvement in post-stroke aphasia with or without conventional rehabilitation [27], Ren et al., in their meta-analysis also reported a positive effect on language recovery in patients with post-stroke aphasia [28], but Gadenz et al, in a systematic review of randomized controlled trials, reported uncertain benefits in neurologic disorders related to communication; the authors analysed 3 studies about dysphagia 1 about dysarthria in Parkinson's disease and 1 about linguistic deficits in Alzheimer's disease, which could underestimate rTMS effects [29].

Most of studies included, stimulated the contralateral Broca's area (only Szaflarski et al, stimulated ipsilateral using excitatory intermittent theta-burst stimulation) this on base of theories about post-stroke aphasia recover, which proposed that brain may use two strategies to recover speech-relevant regions (the structural repair of primarily speech-relevant regions or the activation of compensatory areas) [30]. Karbe et al., found that brain recruit right-hemispheric regions for speech processing when the left-hemispheric centers were permanently impaired as compensatory mechanisms. On this way, Thiel et al., analysed the potential to compensate damage of left-hemisphere by other brain zones using functional neuroimaging and transcranial magnetic stimulation in patients with brain tumors as a model; they found that time is factor which determines successful integration of right-brain [31], But post-stroke recovery is also based on plastic changes in the central nervous system and activation of perilesional zones and reactivation of impaired zones and this could be done by rTMS [32]. The use of rTMS in contralesional brain zones is explained by the inhibition of

Table 3: Effect of repetitive transcranial magnetic stimulation in the treatment of aphasia post- stroke according to the literature review.

Study	Outcome measurement	Δ Measure overall P value	Listening	Speaking	Reading	Writing	Naming	Compression	Expression	Repetition	Description
Hara et al. [15]	SLTA	CL df:5.9p<0.01 IL df:6.6p<0.01	CL df:0.9 p>0.05 IL df:1.6 p<0.05	CL df:1.7 p<0.01 IL df:2.8 p<0.01	CL df:0.7 p<0.01 IL df:0.5 p>0.05	CL df:1.4 p<0.05 IL df:1.6 p<0.01	No	No	No	No	No
Rubi-fessen et al. [16]	AAT FIM Anelt A	Real df:6.20 p<.001 Sham df:3.26 p<.001	No	No	No	Real df:4.7 p<.001 Sham df:1.87 p=0.007	Real df:6.47 p<.001 Sham df:2.07 p=0.057	Real df:4.40 p<.001 Sham df:1.60 p=0.045	Real df:0.53p=0.013 Sham df:0.27 p=0.052	Real df:2.93 p=0.005 Sham df:2.34p<.001	No
Li Y et al. [19]	BNT, BADE, CCAT, AAT	No	No	No	No	No	SMD:0.51(0.16-0.86)	SMD:0.31 (-0.14-0.75)	No	SMD: 0.31 (-0.04-0.65)	No
Otal et al. [42]	AAT, BDAE, CPNT PNT	No	No	No	No	No	SMD: 0.51 p=0.0003	No	No	No	No
Yoon et al. [21]	K-WAB	No	No	No	No	No	Real, df=17 p=0.039	No	No	Real df=12.2 p=0.042	No
Wang et al. [20]	CCAT	SYN= Post1 df: 1.6 p<0.05	No	No	No	No	SYN= Post1 df:20% p<0.05 Post2 df:26.1% p<0.05	No	SYN= Post1df:2 p<0.05	No	SYN= Post1 df:1.3 p<0.05
Tsai et al. [17]	CCAT	Post1 df:1.2 p<0.001 Post2 df:1.7 p<0.05	No	No	No	No	Post1 df:1.2 p<0.05	No	Post1 df:1.4 p<0.01 Post2 df:1.5 p<0.05	Post1df: 0.9 p<0.05 Post2 df: 1 p<0.05	Post1df:0.8 p<0.05 Post2 df:1.4 p<0.05
Khedr et al. [18]	HSS ASRS	df=1.6 p= 0.004	No	No	No	No	df=2.1 p=0.01	df=1.7 p=0.04	No	df=1.5 p=0.002	No
Barwood et al. [22]	BNT BDAE	df=4.9 p< 0.05	No	No	No	No	df=3.9 p< 0.05	No	No	df= 4.01 p< 0.05	No
Seniow et al. [23]	BDAE	No	No	No	No	No	Anterior-follow up Real df=64.9 Sham df=40.4 p=0.03	No	No	Posterior-post Real df=5.2 Sham df=2.1 p=0.02 Severe-follow up Real df=7.7 Sham df=3.5 p=0.016	No
Thiel et al. [44]	AAT	df=16.2 p=0.003	No	No	No	No	df=5.2 p=0.002	No	No	No	No
Waldowski et al. [45]	CPNT ASRS BDAE	No	No	No	No	No	df=13 p=0.016	No	No	No	No
Szaflarski et al. [46]	SFT COWAT BNT BDAE	No	No	No	No	No	No	No	No	No	No
Weiduschat et al. [47]	AAT	df=19.8 p=0.002	No	No	No	No	No	No	No	No	No
Barwood et al. [43]	BNT BDAE	df=10 p< 0.01	No	No	No	No	df=10 p< 0.01	No	No	No	No

Abbreviations: CL: Contralateral; IL: Ipsilateral; df: Difference From Baseline To Posterior Assessment; No: no Statistical Significance; SMD: Standard Mean Difference; BNT: Boston Naming Test; SLTA: The Standard Language Test of Aphasia; AAT: Aachen Aphasia Test; FIM: The Functional Independence Measure; Anelt A-scale Amsterdam-Nijmegen Everyday Language Test; BNT: Boston Naming Test; BDAE: Boston Diagnostic Aphasia Examination; CCAT: Criteria Cognitive Aptitude Test; AAT: Aachen Aphasia Test; PNT: Philadelphia Naming Test; K-WAB: Korean-version of the Western Aphasia Battery; ASRS: Aphasia Severity Rating Scale; HSS: High Sensation Seeking Test; CPNT: Computerized Picture Naming Test; SFT: Semantic Fluency Test; COWAT: Controlled Oral Word Association Test

over-activation in the Broca's homologous to the lesion that lead to inhibition on perilesional brain zones [33]. One of the first works on rTMS in aphasia post-stroke was done by Naeser et al., they used a 1 Hz 1200 pulses at 90% of motor threshold in anterior part of Broca's homologue, this area was identified on each patient by MRI scan; they reported improve in picture naming at 2 months post-rTMS [34], and this due to improve in inter hemispheric modulation of semantic procession for picture naming [32]; author remarks the possibly modulating the distributed bi-hemispheric language network for naming; however, is possible that patient's improved naming scores are related to their having learned the test items, this is one of the issues in neuropsychological test re-test assessment.

From our study results, some questions emerged; first, is the test scores the best way to measure aphasia? Second, post-stroke aphasia population have to be measured also as Health-related quality of life? Changing in naming outcomes is enough to improve quality of life? Third, what are the most useful marginal improvements to communicate?

First, there is a lack of consensus about the most suitable aphasia scale, but the use of Boston test is widely accepted for clinical trials [14]. Second, the recommendation for future clinical trials is the inclusion of Health-related quality of life (HRQL) and also the inclusion of people with aphasia and report separately to determine the main predictors of HRQL [35]. The improvements on some sub-test scores are insufficient to improve basic communication skills in real life. Third, it could be possible that marginal changes in speaking, listening, writing and reading made the real difference for patients. We considered that more investigations could be useful in this field.

As a weakness of this study we identify the heterogeneity of the scales used by the studies to measure outcomes in aphasic patients; compare studies results by scale changes from baseline results to post stimulation results, is a task for this kind of studies. Changes in mean were reported for the majority of studies included in the systematic review; we avoid measuring the magnitude of its median changes for results, our focus was reporting differences on scales results. As a weakness we also identified the differences in inclusion and exclusion criteria, following of patients included, time post-stroke, use of SLT and other coadjutants in stroke. The only one study included in the systematic review that showed improve on speaking, reading and writing skills, was the Hara et al., study, on it the SLTA scale was used for measure study outcomes-no other study used it. The SLTA was developed for classification of severity of aphasia of Japanese speakers in 1975; this test includes specific tasks and assessments considered complicated, but focused on daily communications skills [36]. There is a lack of studies published that included assessment only for this scale-different to the Japanese language-is impossible to assume that speaking, reading and writing skills improve in Japanese population with rTMS by this study along. Some authors propose an ecological approach aimed at restoring the patient ability to communicate in daily context, for this approach propose of new alternatives for help them is a validate alternative.

Wang et al., include synchronous verbal training during rTMS; they investigate the efficacy with a simultaneous picture-

naming activity and demonstrated a superior language for verbal expression also [20]. This study showed change from baseline assessment to post 1 and post 2, for conversation and expression there were changes in all of groups, but exist doubt regard to changes from baseline assessment and evaluations until 90 days' post-stroke with or without any intervention. Lazar et al., study, reported that patients improved by approximately 70% of the maximum potential recovery with language therapy; it was considered as standard care from this results, and alternatives of standard care that could be useful [37]. From this study, authors consider that coadjutants to speech therapy could be used and it could be added value to standard care (SLT) [24]. Thus many gap data come from these results, severely of aphasia, the time and the quality of speech therapy; this was other issues found in studies included in our study.

Hills in her study outline about targets of therapy and important aspects of recovery; she recalls for evaluations of speech articulation, reading writing and grammatical skills. The purpose of these results is improved communications assessments, and guide for toward communication enhancement programs and others (book clubs, social organizations) [24]. For an inclusion of a holistic outcome measure in post-stroke aphasic patients, the inclusion of different assessment close related to aphasia may be included as emotional distress, activity limitations, social factors, quality of life, cognitive function, and so on. From this perspective authors designed alternative measure tools, one of them is aphasia quality of life scale-39 (SAQOL), this stroke-specific scale incorporates patients' views about their own health perception. Hilari et al., tested the feasibility and psychometric evaluation of this scale: 87% were able to complete the SAQOL, and derivate a version that identified 4 sub domains (physical, psychosocial, communication and energy) to Resume it [35]. Many other authors have evaluated quality of life in stroke survivors. The overall results are a decrease in QL that depends on demographic factors, comorbidity, disability and psychosocial factors [38,39]. Frances et al., measured outcomes related to quality of life in patients with post-stroke and aphasia. They used modified outcome measure included the stroke impact scale, the 36-item from medical outcomes, reintegration to normal living scale and activity card sort. They concluded with recommendation for future investigations and practitioners the inclusion of outcomes that assess self-reported quality of life, as an understanding of consequences of stroke [40]. There are movements towards self-reported outcomes, patient preferences and approaches of a holistic measure of results in clinical practice to focus new treatments or fitting current alternatives; from this view patient reported outcomes (PRO) could help to capture patient perspective -determine which health outcome are relevant from them [41-47].

CONCLUSION

Our study shows benefits measured by aphasia scales with rTMS; in overall and sub-test scores there were improve from baseline to post-evaluations, sub-test scores showed improve mainly in naming, and less for other sub-test; neither study reported improve in basic communication skills. Our results support the use of rTMS in patients with aphasia post-stroke; chronic and early. We considered that minimal improvements in

patients with aphasia are relevant in a condition that is difficult and considered a task-due to the complexity of it. We also considered that more research has to be done focused on quality of life changes with rTMS as a holistic assessment from patient perspective and incorporate it in the health system if it shows benefits for daily life of patients with aphasia post-stroke.

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REFERENCES

1. Feigin VL, Forouzanfar MH, Krishnamurthi R, Mensah GA, Connor M, Bennett DA, et al. Global and regional burden of stroke during 1990-2010: findings from the Global Burden of Disease Study 2010. *Lancet*. 2014; 383: 245-254.
2. Engelter ST, Gostynski M, Papa S, Frei M, Born C, Ajdacic-Gross V, et al. Epidemiology of aphasia attributable to first ischemic stroke: incidence, severity, fluency, etiology, and thrombolysis. *Stroke*. 2006; 37: 1379-1384.
3. Stroke Association, stroke statistics. Information guide. London, 2016.
4. Herrmann M, Bartels C, Wallesch CW. Depression in acute and chronic aphasia: symptoms, pathoanatomical-clinical correlations and functional implications. *J Neurol Neurosurg Psychiatry*. 1993; 56: 672-678.
5. Laska AC, Mårtensson B, Kahan T, von Arbin M, Murray V. Recognition of depression in aphasic stroke patients. *Cerebrovasc Dis*. 2007; 24: 74-79.
6. Laska AC, Hellblom A, Murray V, Kahan T, Von Arbin M. Aphasia in acute stroke and relation to outcome. *J Intern Med*. 2001; 249: 413-422.
7. Kauhanen ML, Korpelainen JT, Hiltunen P, Määttä R, Mononen H, Brusin E, et al. Aphasia, depression, and non-verbal cognitive impairment in ischaemic stroke. *Cerebrovasc Dis*. 2000; 10: 455-461.
8. Davidson B, Howe T, Worrall L, Hickson L, Togher L. Social participation for older people with aphasia: the impact of communication disability on friendships. *Top Stroke Rehabil*. 2008; 15: 325-340.
9. Lyon JG. Communication Use and Participation in Life for Adults with Aphasia in Natural Settings The Scope of the Problem. *AJSLP*. 1992; 1: 7-14.
10. Madeline Cruice, Linda Worrall, Louise Hickson, Robert Murison. Finding a focus for quality of life with aphasia: Social and emotional health, and psychological well-being. *Aphasiology*. 2003; 17: 333-353.
11. Fama ME, Turkeltaub PE. Treatment of poststroke aphasia: current practice and new directions. *Semin Neurol*. 2014; 34: 504-513.
12. Sinanović O, Mrkonjić Z, Zukić S, Vidović M, Imamović K. Post-stroke language disorders. *Acta Clin Croat*. 2011; 50: 79-94.
13. Pedersen PM, Vinter K, Olsen TS. Aphasia after stroke: type, severity and prognosis. The Copenhagen aphasia study. *Cerebrovasc Dis*. 2004; 17: 35-43.
14. Berthier ML. Poststroke aphasia: epidemiology, pathophysiology and treatment. *Drugs Aging*. 2005; 22: 163-182.
15. Hara T, Abo M, Kobayashi K, Watanabe M, Kakuda W, Senoo A. Effects of low-frequency repetitive transcranial magnetic stimulation combined with intensive speech therapy on cerebral blood flow in post-stroke aphasia. *Transl Stroke Res*. 2015; 6: 365-374.
16. Rubi-Fessen I, Hartmann A, Huber W, Fimm B, Rommel T, Thiel A, et al. Add-on Effects of Repetitive Transcranial Magnetic Stimulation on Subacute Aphasia Therapy: Enhanced Improvement of Functional Communication and Basic Linguistic Skills. A Randomized Controlled Study. *Arch Phys Med Rehabil*. 2015; 96: 1935-1944.
17. Tsai PY, Wang CP, Ko JS, Chung YM, Chang YW, Wang JX. The Persistent and Broadly Modulating Effect of Inhibitory rTMS in Nonfluent Aphasic Patients a Sham-Controlled, Double-Blind Study. *Neurorehabil Neural Repair*. 2014; 28: 779-787.
18. Khedr EM, Abo El-Fetoh N, Ali AM, El-Hammady DH, Khalifa H, Atta H, et al. Dual-hemisphere repetitive transcranial magnetic stimulation for rehabilitation of poststroke aphasia a randomized, double-Blind clinical trial. *Neurorehabil Neural Repair*. 2014. 28: 740-750.
19. Li Y, Qu Y, Yuan M, Du T. Low-frequency repetitive transcranial magnetic stimulation for patients with aphasia after stroke: A meta-analysis. *J Rehabil Med*. 2015; 47: 675-681.
20. Wang CP, Hsieh CY, Tsai PY, Wang CT, Lin FG, Chan RC. Efficacy of synchronous verbal training during repetitive transcranial magnetic stimulation in patients with chronic aphasia. *Stroke*. 2014; 45: 3656-3662.
21. Yoon TH, Han SJ, Yoon TS, Kim JS, Yi TI. Therapeutic effect of repetitive magnetic stimulation combined with speech and language therapy in post-stroke non-fluent aphasia. *NeuroRehabilitation*. 2015; 36: 107-114.
22. Barwood CH, Murdoch BE, Whelan BM, Lloyd D, Riek S, O' Sullivan JD, et al. Improved language performance subsequent to low-frequency rTMS in patients with chronic non-fluent aphasia post-stroke. *Eur J Neurol*. 2011; 18: 935-943.
23. Seniów J, Waldowski K, Leśniak M, Iwański S, Czepiel W, Członkowska A. Transcranial magnetic stimulation combined with speech and language training in early aphasia rehabilitation: a randomized double-blind controlled pilot study. *Top Stroke Rehabil*. 2013; 20: 250-261.
24. Hillis AE. The 'standard' for poststroke aphasia recovery. *Stroke*. 2010; 41: 1316-1317.
25. Bortnik KE, Boone KB, Wen J, Lu P, Mitrushina M, Razani J, et al. Survey results regarding use of the Boston Naming Test: Houston, we have a problem. *J Clin Exp Neuropsychol*. 2013; 35: 857-866.
26. Camara WJ, Nathan JS, Puente AE. Psychological test usage: Implications in professional psychology. *Professional Psychology: Research and Practice*, 2000; 31: 141-154.
27. Wong IS, Tsang HW. A review on the effectiveness of repetitive transcranial magnetic stimulation (rTMS) on post-stroke aphasia. *Rev Neurosci*. 2013; 24: 105-114.
28. Ren CL, Zhang GF, Xia N, Jin CH, Zhang XH, Hao JF, et al. Effect of low-frequency rTMS on aphasia in stroke patients: a meta-analysis of randomized controlled trials. *PLoS One*. 2014; 9: e102557.
29. Gadenz CD, Moreira Tde C, Capobianco DM, Cassol M. Effects of Repetitive Transcranial Magnetic Stimulation in the Rehabilitation of Communication and Deglutition Disorders: Systematic Review of Randomized Controlled Trials. *Folia Phoniatr Logop*. 2015; 67: 97-105.
30. Karbe H, Thiel A, Weber-Luxenburger G, Herholz K, Kessler J, Heiss WD. Brain plasticity in poststroke aphasia: what is the contribution of the right hemisphere? *Brain Lang*. 1998; 64: 215-230.
31. Thiel A, Habedank B, Herholz K, Kessler J, Winhuisen L, Haupt WF, et al. From the left to the right: How the brain compensates progressive loss of language function. *Brain Lang*. 2006; 98: 57-65.

32. Lefaucheur JP. Stroke recovery can be enhanced by using repetitive transcranial magnetic stimulation (rTMS). *Neurophysiol Clin.* 2006; 36: 105-115.
33. Connor LT, DeShazo Braby T, Snyder AZ, Lewis C, Blasi V, Corbetta M. Cerebellar activity switches hemispheres with cerebral recovery in aphasia. *Neuropsychologia.* 2006; 44: 171-177.
34. Naeser MA, Martin PI, Nicholas M, Baker EH, Seekins H, Kobayashi M, et al. Improved picture naming in chronic aphasia after TMS to part of right Broca's area: an open-protocol study. *Brain Lang.* 2005; 93: 95-105.
35. Hilari K, Needle JJ, Harrison KL. What are the important factors in health-related quality of life for people with aphasia? A systematic review. *Arch Phys Med Rehabil.* 2012; 93: S86-S95.
36. Tamura A, Shichijo F, Matsumoto K. A study on simplification of the Standard Language Test of Aphasia (SLTA). *Tokushima J Exp Med.* 1996; 43: 39-46.
37. Lazar RM, Minzer B, Antonello D, Festa JR, Krakauer JW, Marshall RS. Improvement in aphasia scores after stroke is well predicted by initial severity. *Stroke.* 2010; 41: 1485-1488.
38. Carod-Artal FJ. Determining quality of life in stroke survivors. *Expert Rev Pharmacoecon Outcomes Res.* 2012; 12: 199-211.
39. Hilari K, Byng S. Health-related quality of life in people with severe aphasia. *Int J Lang Commun Disord.* 2009; 44: 193-205.
40. Tucker FM, Edwards DF, Mathews LK, Baum CM, Connor LT. Modifying health outcome measures for people with aphasia. *Am J Occup Ther.* 2012; 66: 42-50.
41. Devlin NJ, Parkin D, Browne J. Patient-reported outcome measures in the NHS: new methods for analysing and reporting EQ-5D data. *Health Econ.* 2010; 19: 886-905.
42. Otal B, Olma MC, Flöel A, Wellwood I. Inhibitory non-invasive brain stimulation to homologous language regions as an adjunct to speech and language therapy in post-stroke aphasia: a meta-analysis. *Front Hum Neurosci.* 2015; 9: 236.
43. Barwood CH, Murdoch BE, Riek S, O'Sullivan JD, Wong A, Lloyd D, et al. Long term language recovery subsequent to low frequency rTMS in chronic non-fluent aphasia. *NeuroRehabilitation.* 2013; 32: 915-928.
44. Thiel A, Hartmann A, Rubi-Fessen I, Anglade C, Kracht L, Weiduschat N, et al. Effects of noninvasive brain stimulation on language networks and recovery in early poststroke aphasia. *Stroke.* 2013; 44: 2240-6.
45. Waldowski K, Seniów J, Leśniak M, Iwański S, Członkowska A. Effect of low-frequency repetitive transcranial magnetic stimulation on naming abilities in early-stroke aphasic patients: a prospective, randomized, double-blind sham-controlled study. *Scientific World Journal.* 2012; 2012: 518568.
46. Szaflarski JP, Vannest J, Wu SW, DiFrancesco MW, Banks C, Gilbert DL. Excitatory repetitive transcranial magnetic stimulation induces improvements in chronic post-stroke aphasia. *Med Sci Monit.* 2011; 17: CR132-139.
47. Weiduschat N, Thiel A, Rubi-Fessen I, Hartmann A, Kessler J, Merl P, et al. Effects of repetitive transcranial magnetic stimulation in aphasic stroke a randomized controlled pilot study. *Stroke.* 2011; 42: 409-415.

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