

**Appendix 1** to Sehatzadeh S, Daskalakis ZJ, Yap B, et al. Unilateral and bilateral repetitive transcranial magnetic stimulation for treatment-resistant depression: a meta-analysis of randomized controlled trials over 2 decades. *J Psychiatry Neurosci* 2019.

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## Supplemental Data

**Table S1: Quality Assessment for the Body of Evidence According to the GRADE System for Rating the Quality of Evidence (GRADE): Randomized Controlled Trials of Repetitive Transcranial Magnetic Stimulation Versus Sham**

| Number of Studies                  | Design | Risk of Bias            | Inconsistency          | Indirectness           | Imprecision            | Publication Bias       | GRADE    |
|------------------------------------|--------|-------------------------|------------------------|------------------------|------------------------|------------------------|----------|
| <b>Unilateral rTMS Versus Sham</b> |        |                         |                        |                        |                        |                        |          |
| <b>WMD</b>                         |        |                         |                        |                        |                        |                        |          |
| 18                                 | RCT    | Serious limitations*    | No serious limitations | No serious limitations | No serious limitations | No serious limitations | Moderate |
| <b>Remission Rate</b>              |        |                         |                        |                        |                        |                        |          |
| 13                                 | RCT    | Serious limitations*    | No serious limitations | No serious limitations | No serious limitations | No serious limitations | Moderate |
| <b>Response Rate</b>               |        |                         |                        |                        |                        |                        |          |
| 17                                 | RCT    | Serious limitations*    | No serious limitations | No serious limitations | No serious limitations | No serious limitations | Moderate |
| <b>Bilateral rTMS Versus Sham</b>  |        |                         |                        |                        |                        |                        |          |
| <b>WMD</b>                         |        |                         |                        |                        |                        |                        |          |
| 4                                  | RCT    | No serious limitations† | No serious limitations | No serious limitations | No serious limitations | No serious limitations | High     |
| <b>Remission Rate</b>              |        |                         |                        |                        |                        |                        |          |
| 6                                  | RCT    | No serious limitations† | No serious limitations | No serious limitations | No serious limitations | No serious limitations | High     |
| <b>Response Rate</b>               |        |                         |                        |                        |                        |                        |          |
| 7                                  | RCT    | No serious limitations† | No serious limitations | No serious limitations | No serious limitations | No serious limitations | High     |

RCT = randomized controlled trial; rTMS = repetitive transcranial magnetic stimulation; WMD = weighted mean difference.

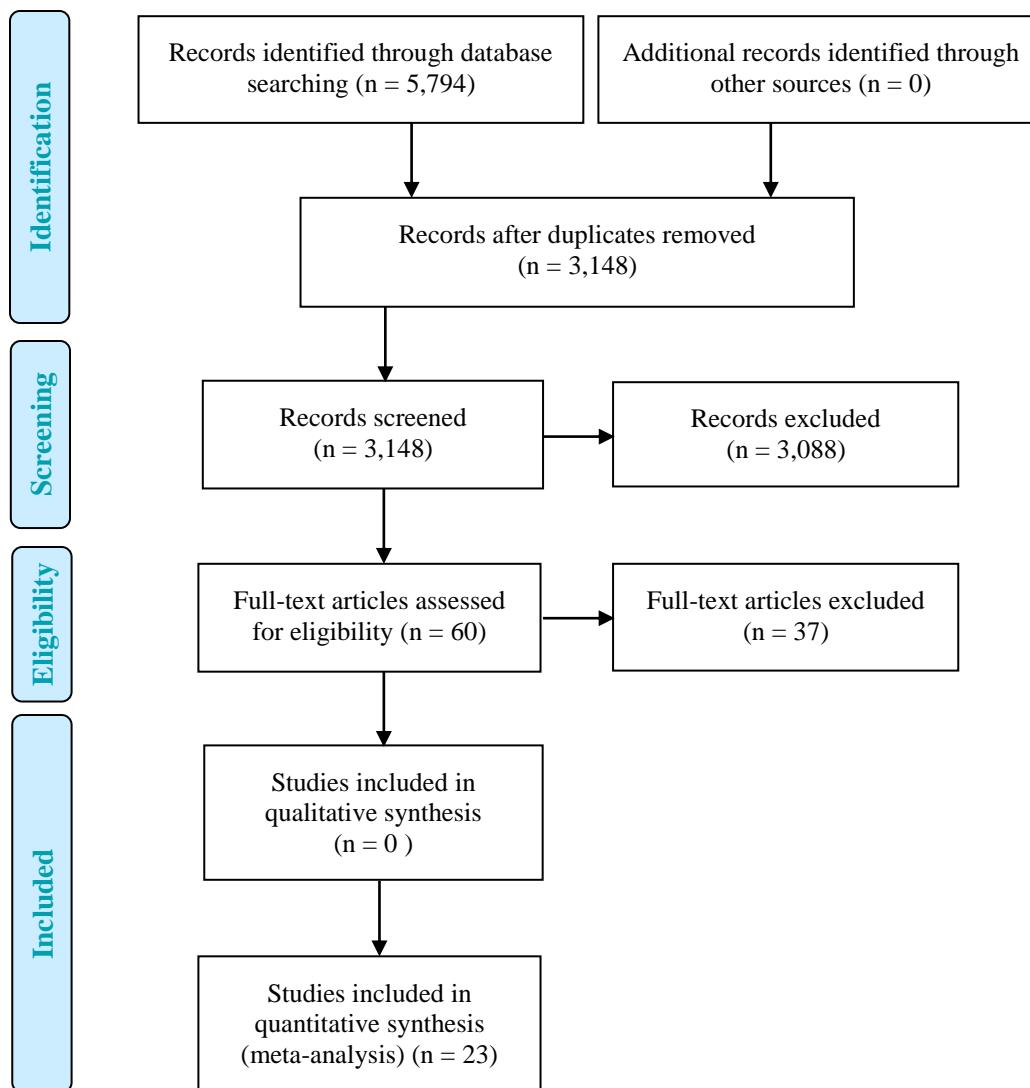
\*Lack of allocation concealment in most studies.

†Allocation concealment performed in 3 studies.

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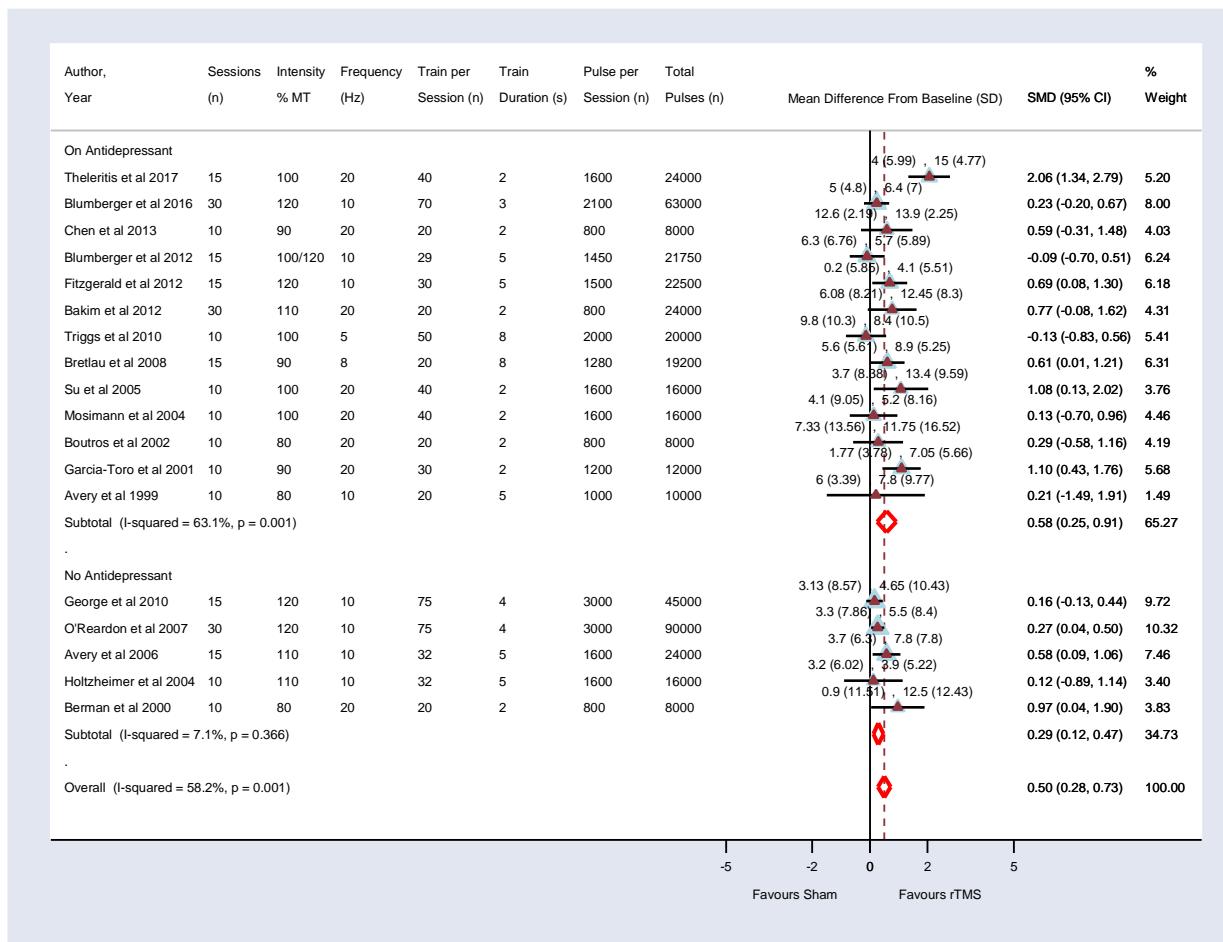


**Figure S1: Flow Diagram for the Preferred Reporting Items for Systematic Reviews and Meta-analyses (PRISMA)**

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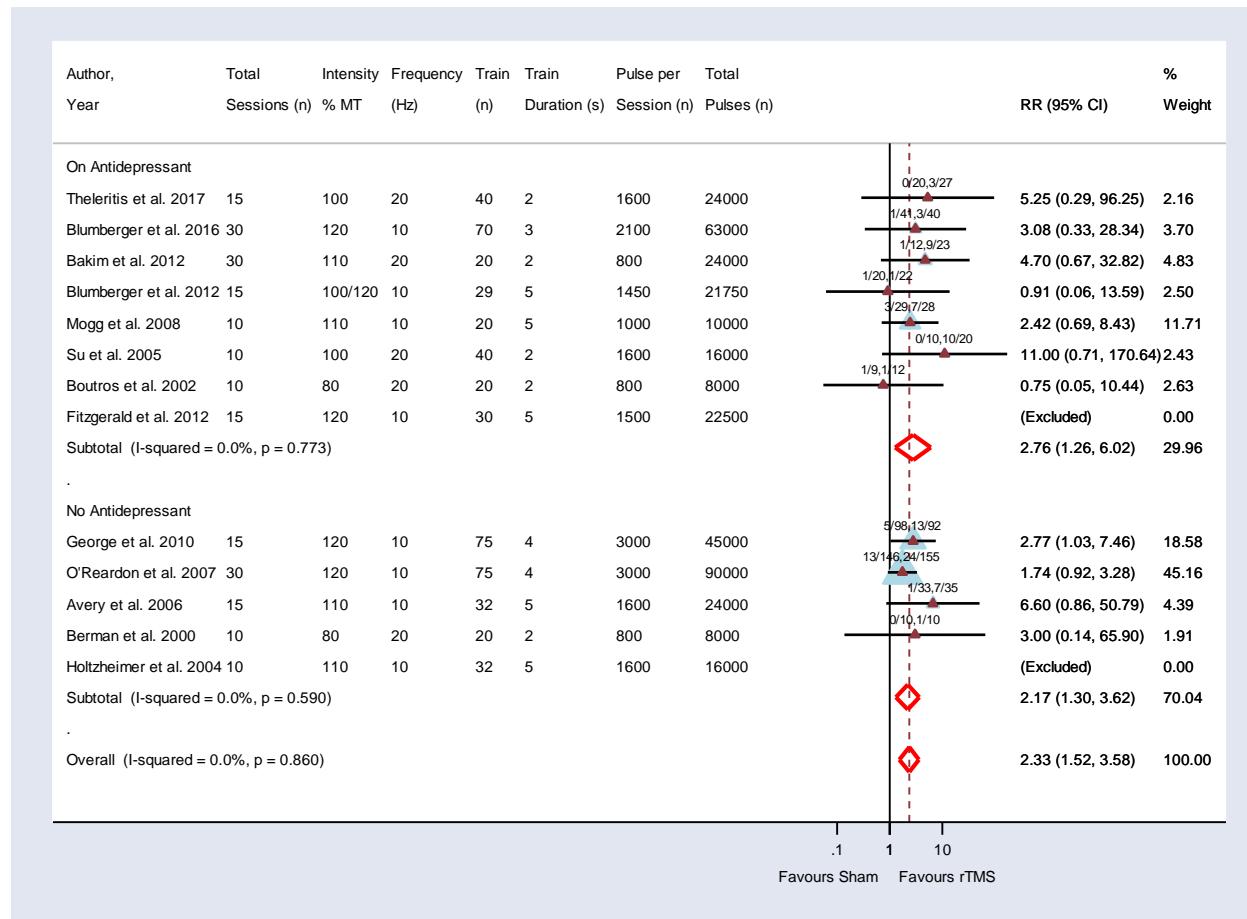
**Figure S2: Standardized Mean Difference in Depression Scores: Unilateral High Frequency Repetitive Transcranial Magnetic Stimulation Versus Sham**

CI = confidence interval; Hz = hertz; MT = motor threshold; rTMS = repetitive transcranial magnetic stimulation; SD = standard deviation; SMD = standardized mean difference.

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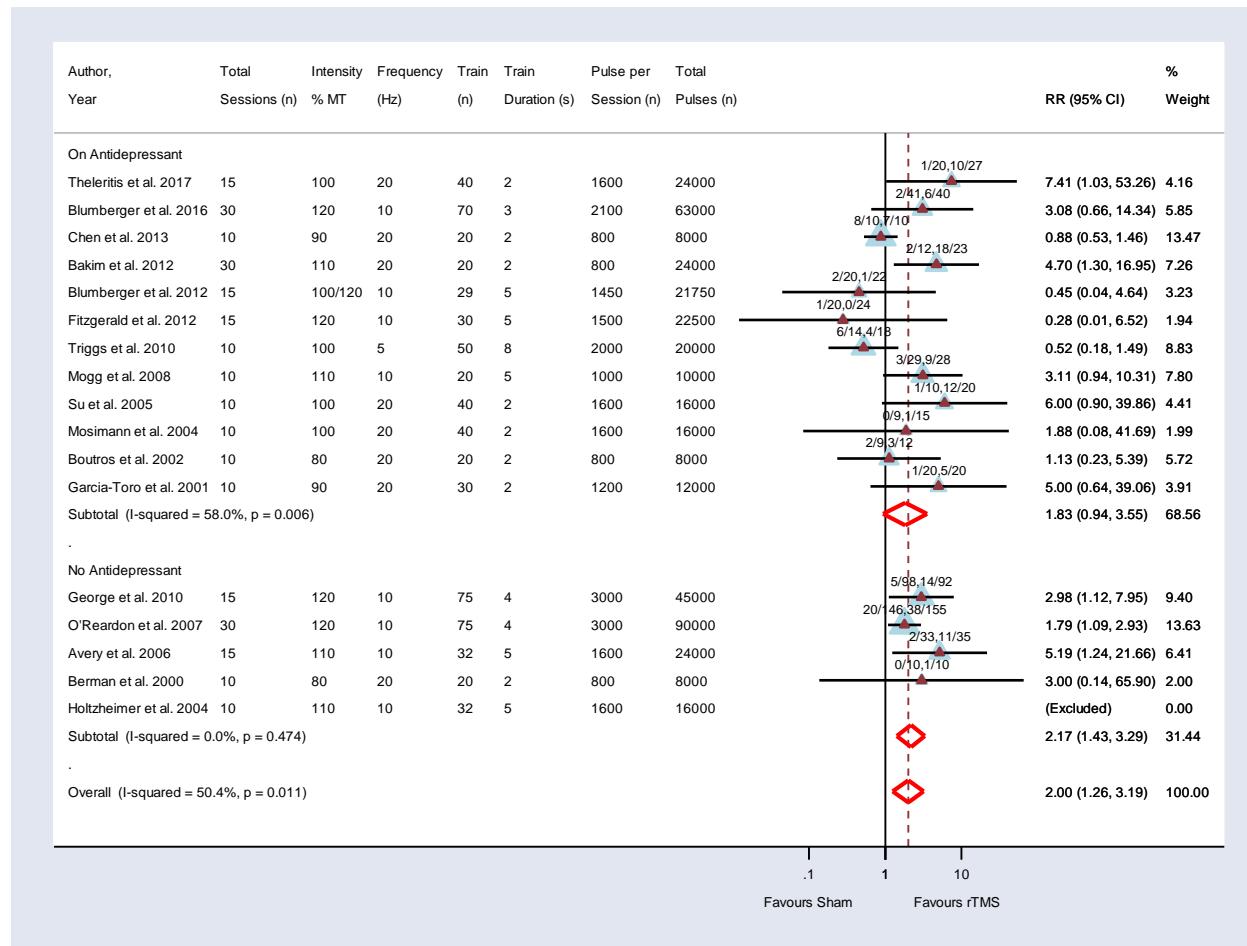
**Figure S3: Remission Rates and Rate Ratios: Unilateral High Frequency Repetitive Transcranial Magnetic Stimulation Versus Sham**

CI = confidence interval; Hz = hertz; MT = motor threshold; RR = rate ratio; rTMS = repetitive transcranial magnetic stimulation.

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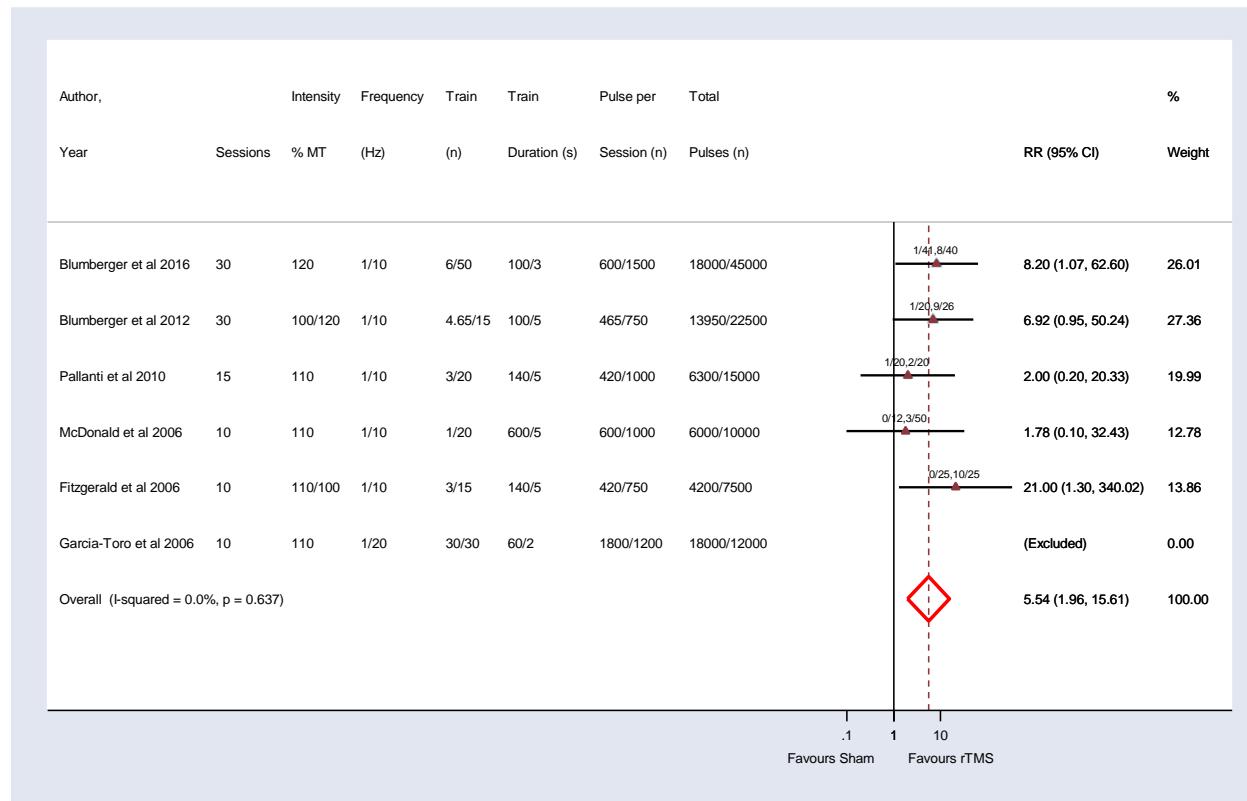
**Figure S4: Response Rates and Rate Ratios: Unilateral High Frequency Repetitive Transcranial Magnetic Stimulation Versus Sham**

CI = confidence interval; Hz = hertz; MT = motor threshold; RR = rate ratio; rTMS = repetitive transcranial magnetic stimulation.

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**Figure S5: Remission Rates and Rate Ratios: Bilateral Sequential Repetitive Transcranial Magnetic Stimulation Versus Sham**

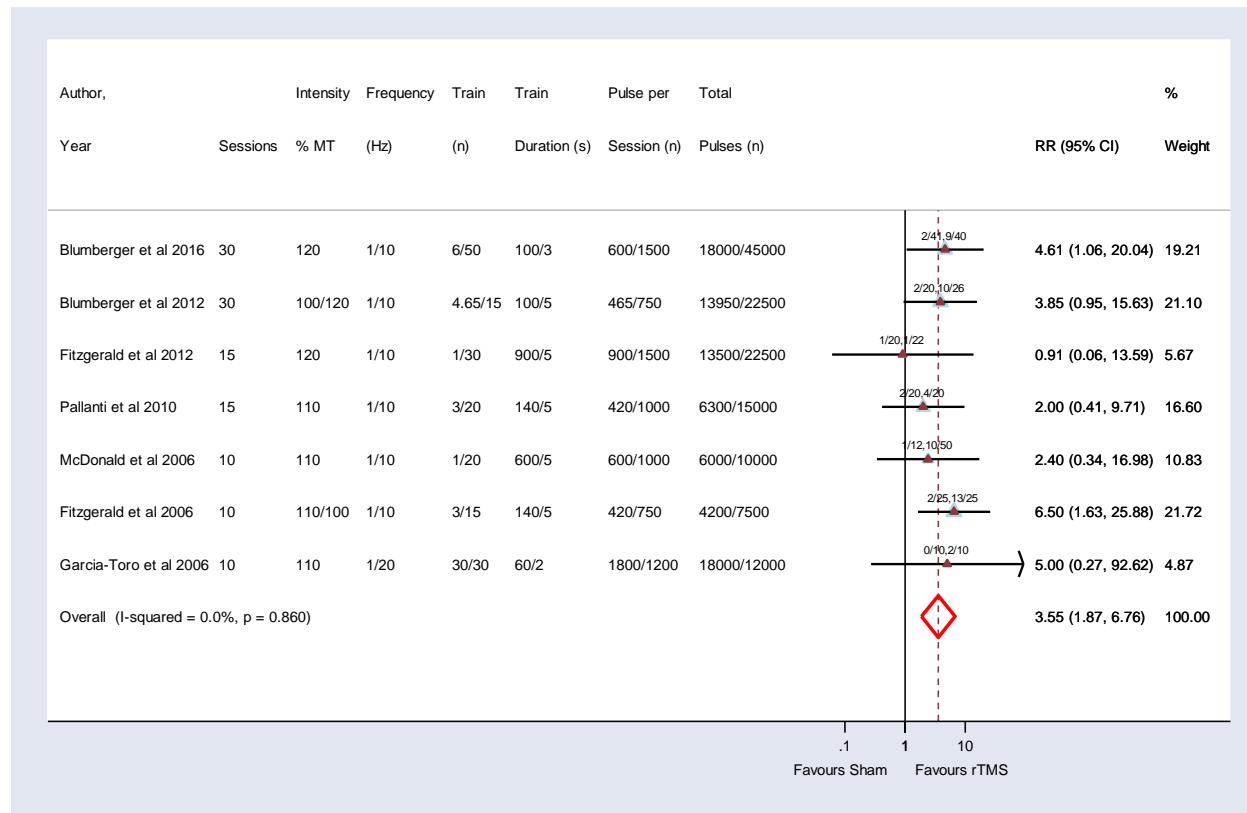
*Note:* Numbers separated by slash relate to the technical parameters applied to the right and left DLPFC stimulation (right/left).

CI = confidence interval; Hz = hertz; MT = motor threshold; RR = rate ratio; rTMS = repetitive transcranial magnetic stimulation.

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**Figure S6: Response Rates and Rate Ratios: Bilateral Sequential Repetitive Transcranial Magnetic Stimulation Versus Sham**

*Note:* Numbers separated by / relate to the technical parameters applied to the right and left DLPFC stimulation (right/left).

CI = confidence interval; Hz = hertz; MT = motor threshold; RR = rate ratio; rTMS = repetitive transcranial magnetic stimulation.

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## **Literature Search – Repetitive Transcranial Magnetic Stimulation**

**Search date:** April 03, 2017

**Database:** EBM Reviews - Cochrane Central Register of Controlled Trials <February 2017>, EBM Reviews - Cochrane Database of Systematic Reviews <2005 to March 29, 2017>, EBM Reviews - Health Technology Assessment <4th Quarter 2016>, EBM Reviews - NHS Economic Evaluation Database <1st Quarter 2016>, Embase <1980 to 2017 Week 14>, Ovid MEDLINE(R) Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily and Ovid MEDLINE(R) <1946 to Present>, PsycINFO <1967 to March Week 4 2017>

### **Search Strategy:**

---

- 1 Depression/ (432736)
- 2 exp Depressive Disorder/ (500483)
- 3 (depressi\* or dysthymic or melancholia or TRD or "involutional psychos\*" or paraphrenia).ti,ab. (1011971)
- 4 or/1-3 (1207712)
- 5 Transcranial Magnetic Stimulation/ (35458)
- 6 (((transcranial or trans-cranial) adj2 magnetic adj2 stimulation\*) or rtms or tms).mp. (55077)
- 7 or/5-6 (55077)
- 8 4 and 7 (7994)
- 9 8 use ppez,coch,cctr,clhta,cleed (2591)
- 10 Depression/ (432736)
- 11 Major Depression/ (161223)
- 12 (depressi\* or dysthymic or melancholia or TRD or "involutional psychos\*" or paraphrenia).ti,ab. (1011971)
- 13 or/10-12 (1157605)
- 14 Transcranial Magnetic Stimulation/ (35458)
- 15 (((transcranial or trans-cranial) adj2 magnetic adj2 stimulation\*) or rtms or tms).ti,ab. (47284)
- 16 or/14-15 (54084)
- 17 13 and 16 (7755)
- 18 17 use emez (3789)
- 19 exp Major Depression/ (168247)
- 20 (depressi\* or dysthymic or melancholia or TRD or "involutional psychos\*" or paraphrenia).ti,ab. (1011971)
- 21 or/19-20 (1028669)

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- 22 exp Transcranial Magnetic Stimulation/ (35458)
- 23 (((transcranial or trans-cranial) adj2 magnetic adj2 stimulation\*) or rtms or tms).mp. (55077)
- 24 or/22-23 (55077)
- 25 21 and 24 (7332)
- 26 25 use psyb (1527)
- 27 9 or 18 or 26 (7907)
- 28 limit 27 to english language [Limit not valid in CDSR; records were retained] (7258)
- 29 limit 28 to yr="2014 -Current" (2356)
- 30 29 use ppez (525)
- 31 29 use emez (1239)
- 32 29 use psyb (401)
- 33 29 use cctr (167)
- 34 29 use coch (4)
- 35 29 use clhta (19)
- 36 29 use cleed (1)
- 37 remove duplicates from 29 (1407)

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## Literature Search – Repetitive Transcranial Magnetic Stimulation

**Search date:** Nov 20, 2014

**Databases searched:** Ovid MEDLINE/In-Process, Embase, EBM Databases, PsycINFO

**Limits:** 1994-current; English

**Databases:** EBM Reviews - Cochrane Database of Systematic Reviews 2005 to October 2014, EBM Reviews - Database of Abstracts of Reviews of Effects 4th Quarter 2014, EBM Reviews - Cochrane Central Register of Controlled Trials October 2014, EBM Reviews - Cochrane Methodology Register 3rd Quarter 2012, EBM Reviews - Health Technology Assessment 4th Quarter 2014, EBM Reviews - NHS Economic Evaluation Database 4th Quarter 2014, Ovid MEDLINE(R) Daily Update November 19, 2014, Ovid MEDLINE(R) In-Process & Other Non-Indexed Citations and Ovid MEDLINE(R) 1946 to Present, Embase 1974 to 2014 November 19

### Search Strategy:

| #  | Searches  | Results |
|----|---|---------|
| 1  | Depression/   | 341444  |
| 2  | exp Depressive Disorder/ use prmz,acp,cctr,coch,clcmr,dare,clhta,cleed                                    | 92220   |
| 3  | Major Depression/ use oemezd  | 37851   |
| 4  | Treatment Resistant Depression/ use oemezd  | 742     |
| 5  | (depressi* or dysthymic or melancholia or TRD or "involutional psychos*" or paraphrenia).ti,ab.           | 642939  |
| 6  | or/1-5  | 783031  |
| 7  | Transcranial Magnetic Stimulation/  | 22067   |
| 8  | ((transcranial or trans-cranial) adj2 magnetic adj2 stimulation*) or rtms or tms).mp.                     | 36404   |
| 9  | or/7-8  | 36404   |
| 10 | 6 and 9   | 4766    |
| 11 | limit 10 to yr="1994 -Current" [Limit not valid in DARE; records were retained]                           | 4743    |
| 12 | limit 11 to english language [Limit not valid in CDSR,ACP Journal Club,DARE,CLCMR; records were retained] | 4305    |
| 13 | remove duplicates from 12   | 2734    |

**Database:** PsycINFO <1987 to November Week 3 2014>

**Search Strategy:**

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| # | Searches  | Results |
|---|---|---------|
| 1 | exp Major Depression/   | 93059   |
| 2 | (depressi* or dysthymic or melancholia or TRD or "involutional psychos*" or paraphrenia).ti,ab. | 180727  |
| 3 | or/1-2  | 186314  |
| 4 | exp Transcranial Magnetic Stimulation/  | 4565    |
| 5 | ((transcranial or trans-cranial) adj2 magnetic adj2 stimulation*) or rtms or tms).mp.           | 6312    |
| 6 | or/4-5  | 6312    |
| 7 | 3 and 6   | 1182    |
| 8 | limit 7 to (english language and yr="1994 -Current")  | 1081    |

#### **HEED**

depressi\* OR dysthymic OR melancholia OR TRD OR psychos\* OR paraphrenia =all data  
AND  
transcranial OR trans-cranial OR rtms OR tms =all data

5 results