

NKR24 - PICO6 - Schizophrenia: Cognitive remediation

Characteristics of studies

Characteristics of included studies

Belucci 2002

Methods	
Participants	
Interventions	
Outcomes	
Identification	
Notes	

Risk of bias table

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

Burda 1994

Methods	
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Risk of bias table

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

d'Amato 2011

Methods	<p>Study design: Randomized controlled trial Study grouping: Parallel group Open Label: Cluster RCT: YES</p>
Participants	<p>Baseline Characteristics TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 32.2 (6.0) ● Sex (male %): 76.3 ● Length of illness (years), mean (sd): 8.1 (4.5) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 33.4 (6.9) ● Sex (male %): 74.4 ● Length of illness (years), mean (sd): 8.7 (6.6) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>Included criteria: Patients were eligible if they met the DSM IV criteria for schizophrenia, were clinically stabilized without any modification of their medication for at least one month, spoke French fluently and were aged between 18 and 40 years. Excluded criteria: Exclusion criteria were past or present neurological disorders or substance dependence or abuse, pregnancy, and not being able to give informed consent.</p>
Interventions	<p>Intervention Characteristics TAU</p> <ul style="list-style-type: none"> ● Description: standard treatment only <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: The therapy consisted of 14 individual two-hour sessions over a 7-week period providing supportive, graduated training and practice in selecting, executing, and monitoring cognitive operations. CRT was conducted by a psychologist on a computer with a special input panel (joystick and ergonomic pads) using RehaCom® software package (SCHUHFRIED, GmbH). Four procedures have been chosen from amongst the nineteen different procedures available in RehaCom®, to train four cognitive functions involved in different stages of the

	information processing: attention/concentration, working memory, logic, and executive functions.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Global cognition score, Z score ● Social functioning ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Days at hospital ● QoL ● Symptoms, total score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Verbal learning and memory, RAVLT, learning (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Verbal learning and memory, other scale ● Working memory, other. <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Syptomatic relapse ● Symptomatic remission
Identification	<p>Sponsorship source: Funding for this study was provided by PHRC 2005 and had no further role in the study design; in the collection, analysis and interpretation of data; in the writing of the report; and in the decision to submit the paper for publication.</p> <p>Country: France</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Thierry d'Amato</p> <p>Institution: Université de Lyon, Lyon, F-69003, France</p> <p>Email: thierry.damato@ch-le-vinatier.fr</p> <p>Address: CH le Vinatier, Service Pr. d'Amato, 95 Boulevard Pinel, 69677 Bron cedex, France</p>
Notes	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics:</p> <p>Pretreatment:</p> <p>Continuous outcomes:</p> <p><i>Jesper Østrup Rasmussen Scales:</i> Verbal Working memory: Auditory Number Sequencing (ANS). The participants are presented with clusters of numbers (e.g. 936) of increasing length (from 2 digits to a maximum of 8 digits). They are asked to tell the tester the numbers in order, from lowest to highest. A key measure was the maximal span recalled (high=better) Verbal learning and memory: Word List Memory test (WLM). 16 words are presented auditorily by the computer to the subject who must then recall as many as possible. Key measures were the total recall on first trial (high=better) En del præsenteres kun som Cohens d: Impact of CRT on all clinical and neuropsychological assessments was calculated using Cohen's d effect size. We reported a large effect of CRT on verbal learning (1.56) and a medium effect size on verbal memory (0.52), and working memory (0.41)</p> <p><i>Elisabeth Ginnerup-Nielsen</i> Quality of life was assessed by the self-report quality of life for people with schizophrenia (SQoL) higher=better Verbal Working memory: Auditory Number Sequencing (ANS). The participants are presented with clusters of numbers (e.g. 936) of increasing length (from 2 digits to a maximum of 8 digits). They are asked to tell the tester the numbers in order, from lowest to highest. A key measure was the maximal span recalled; higher=better Verbal learning and memory: Word List Memory test (WLM) higher=better</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Quote: "Seventy-seven patients were randomised to either the active group (CRT patients, standard treatment and cognitive remediation program, n = 39)" Comment: unclear how
Allocation concealment (selection bias)	Unclear risk	Comment: Not described
Blinding of participants and personnel (performance bias)	High risk	Comment: Probably not possible. Not described
Blinding of outcome assessment (detection bias)	Low risk	Quote: "The assessors were blind to the participants' assigned randomisation and had no other role in the project that would undermine the blinding."
Incomplete outcome data (attrition bias)	Low risk	Comment: No it analysis but apparently no dropouts
Selective reporting (reporting bias)	Unclear risk	Comment: Det er ikke helt de samme outcomes de agiver de vil måle, som de resultater de præsenterer. Desuden er resultaterne meget kortfattet præsenteret.
Other bias	Low risk	

Dickinson 2010

Methods	<p>Study design: Randomized controlled trial</p> <p>Study grouping: Parallel group</p> <p>Open Label:</p> <p>Cluster RCT:</p>
Participants	<p>Baseline Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 48.5 (8.8) ● Sex (male %): 75 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 46.9 (6.6) ● Sex (male %): 65.7 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): <p>Included criteria: Participation was open to individuals diagnosed with schizophrenia or schizoaffective disorder. Diagnoses used the Structured</p>

	<p>Clinical Interview for DSM-IV (SCID), information from the participants' mental health care providers, and medical records. Eligible individuals were 21 to 60 years old, clinically stable on regimens or second-generation or low-dose first-generation antipsychotics, without a history of significant brain trauma, neurological disorder, or substance dependence within the previous 3 months, and without physical limitations precluding effective use of computer-based exercises.</p> <p>Excluded criteria:</p>
Interventions	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● <i>Description:</i> This condition was designed to control for nonspecific treatment effects. It specified an equal number of one-on-one computer sessions with the same trainers who conducted the remediation sessions. It offered supportive trainer interactions and matched experience with computers and varied computer activities. Control activities were selected for game-like properties and low cognitive demand. Participants in this condition did not receive problem-solving training or guided practice on the exercises used in the remediation condition. The control sessions were also videotaped and reviewed in supervision meetings. <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● <i>Description:</i> Training was organized in three phases. During the first phase, trainers introduced a simple and general problem-solving approach, which was reinforced consistently through all phases of training. The participants were prompted to identify the challenges in each exercise, articulate a plan to address them, implement the plan, monitor its effectiveness, and adjust their strategy as needed. Computer-assisted cognitive remediation shaped these problem-solving techniques through extensive practice. Master's-level trainers guided participants at an individualized pace through a varied curriculum of engaging, educational computer exercises, selected to gradually enhance processing speed, attention, working memory, episodic memory, and executive functioning, i.e., reasoning and problem solving (30) (Figure 1). Time in individual sessions was split; practice of cognitive exercises (roughly two-thirds of each session) alternated with trainer prompts, queries and feedback, and strategy review. The training sessions were videotaped and reviewed in a weekly supervision meeting to promote consistency across different participants and trainers and allow adjustments to individual participant needs. We sought to complete three remediation sessions per week, with a maximum of 15 weeks allowed for completion of the 36-session training program.
Outcomes	<p><i>Continuous:</i></p> <ul style="list-style-type: none"> ● Social functioning (Maryland Assessment of social competence) Z-score ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, other scale ● QoL ● Global cognition score, ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, (see note). higher=better ● Verbal learning and memory, RAVLT, learning (high=better) ● symptoms BPRS <p><i>Dichotomous:</i></p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	<p>Sponsorship source: Funded by NIMH grant MH-67764 and the VA Rehabilitation Research and Development Service.</p> <p>Country: USA</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Dwight Dickinson</p> <p>Institution: Mental Illness Research, Education and Clinical Center, Veterans Integrated Services Network 5, Baltimore VA Medical Center; and the Department of Psychiatry, University of Maryland School of Medicine, Baltimore</p> <p>Email: dwight.dickinson@va.gov</p> <p>Address:</p>
Notes	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics:</p> <p>Pretreatment:</p> <p>Continuous outcomes:</p> <p><i>Elisabeth Ginnerup-Nielsen</i> Working memory measured by: N-back paradigm (35); accuracy on the 1- and 2-back conditions WAIS-III (36) letter-number sequencing subtest. number correct. Forstår det som higher=better Working memory - intervention eot: -0.04 control group eot: -0.13 (Maryland Assessment of social competence): Svært at finde beskrivelse af scoring står opgivet som en Z score</p> <p><i>Jesper Østrup Rasmussen</i> The FU results is not included, because the period was only 3 mo (The participants were assessed before treatment, immediately after treatment, and 3 months after treatment). Scales: Symptoms: BPRS, total (Low=better) Working memory: WAIS-111 (High=better). Global Cognition: SCoRS (interviewer global rating was significantly correlated with the BACS composite score, the UPSA total score, and the ILSI total score. Higher ratings reflect a greater degree of impairment).</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Comment: What is adaptive urn? Only randomised
Allocation concealment (selection bias)	Unclear risk	Comment: Not described.
Blinding of participants and personnel (performance bias)	Low risk	Comment: The participants were not informed that they were assigned to "treatment" or "control" conditions; rather, individuals in both groups were told that the aim of the study was to determine whether participation in a "computer activities program" improved thinking skills. Personnel not relevant in this type of intervention
Blinding of outcome assessment (detection bias)	Low risk	Quote: "The cognitive assessors (who also administered the functional measures) were research assistants, trained and supervised by a neuropsychologist (0.0.). The assessors were blind to the participants' assigned condition and had no other role in the project that would undermine blinding."
Incomplete outcome data (attrition bias)	Low risk	Quote: "Of the 35 participants who engaged in the remediation condition, 30 (85.7%) completed at least 30 remediation sessions (mean=32.2)." Comment: Dropout: Control: 25.9% Intervention: 17,64
Selective reporting (reporting bias)	Low risk	Quote: "ClinicalTrials.gov identifiers: NCT00295048, NCT00261794." Comment: Outcome from protocol assessed

Other bias	Low risk
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Farreny 2012

Methods	Study design: Study grouping: Open Label: Cluster RCT:
Participants	Baseline Characteristics TAU <ul style="list-style-type: none"> ● Age, mean (sd): ● Sex (male %): 67.9 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): Cognitive remediation <ul style="list-style-type: none"> ● Age, mean (sd): ● Sex (male %): 67.6 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): Included criteria: 1) they had a diagnosis of schizophrenia or schizoaffective disorder and more than 2 years' illness duration; 2) they had finished primary studies or they were able to successfully complete a reading comprehension task used for 13-year-old students; 3) if they had a Mini Mental State Examination score of 24 or more and a Global Assessment of Functioning score between 40 and 70. Excluded criteria: 1) they were suffering acute illness exacerbation; 2) they had intellectual disability or any neurological disorder; 3) they were participating in social skills training, cognitive remediation or any other psychological intervention differing from usual care; 4) they had had a switch of antipsychotic drug the month before the trial or during the 40 week study period; 5) and/or a diagnosis of alcohol or drug dependence within 6 months prior to inclusion.
Interventions	Intervention Characteristics TAU <ul style="list-style-type: none"> ● Description: Cognitive remediation <ul style="list-style-type: none"> ● Description: REPYFLEC CR is a strategy-based training that targets executive function and metacognition. It is carried out using paper and pencil and a blackboard (required to develop some of the tasks, explanations, examples, etc.); in a group format (4–6 participants), over 4 months twice a week and consisting of 32 sessions lasting 1 h. We developed a Spanish manual where training is described session by session; incorporating the materials for developing sessions, some theoretical points and bibliography for therapists. Working contents are divided into two main areas: Problem Solving (PS) and Cognitive Flexibility (CF).
Outcomes	Continuous: <ul style="list-style-type: none"> ● Social functioning SFS higher=better ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, other scale ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other. ● Verbal learning and memory, RAVLT, learning (high=better) Dichotomous: <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	Sponsorship source: This research was supported by Fundació La Caixa and Instituto de Salud Carlos III Country: Spain Setting: Comments: Authors name: Aida Farreny Institution: Parc Sanitari Sant Joan de Déu, Fundació Sant Joan de Déu, Centro de Investigación Biomédica en Red de Salud Mental (CIBERSAM), Sant Boi de Llobregat, Barcelona, Spain Email: afarreny@pssjd.org Address:
Notes	Identification: Participants: Study design: Baseline characteristics: <i>Jesper Østrup Rasmussen</i> Some of baseline characteristics only for the total sample: Some 89% of participants had a diagnosis of schizophrenia (n=54), principally paranoid-type (n=35); and the remaining 11% (n=7) of schizoaffective disorder. The average age was 40.6 years (SD: 7.6) and average illness duration was 17.5 years (SD: 8.9). Intervention characteristics: Pretreatment: Continuous outcomes: <i>Elisabeth Ginnerup-Nielsen</i> Some Wechsler Memory Scale-III (WMS-III) (Wechsler, 2004) subscales were selected with the aim of assessing verbal and visual memory (Texts I and II, and Scenes I and II) Social functioning: We used the Spanish validation (Torres and Olivares, 2005) of the Social Functioning Scale (SFS) (Birchwood et al., 1990) for measuring social behavior and relationships, autonomy, employment-occupation and leisure. Raw scoring was used for each subscale and for total score (min. 0–max. 223) with a higher score indicating a better result <i>Jesper Østrup Rasmussen</i> Length of intervention: 4 mo, FU 6 mo (40 weeks after start of treatment). Scales: Social functioning: Higher standardised scores indicate better functioning (range 55 to 135) Memory: WMS-III (Some Wechsler Memory Scale-III (WMS-III) (Wechsler, 2004) subscales were selected with the aim of assessing verbal and visual memory (Texts I and II, and Scenes I and II). Raw scores were used.) (high=better) Dichotomous outcomes: Adverse outcomes:

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Quote: "The participants were assigned to the experimental and control groups through a randomization procedure once the baseline assessments had been performed." Comment: Hmm still unclear which procedure. 34/28 in the two group is relatively skewed. At the same time there were no significant differences between groups
Allocation concealment (selection bias)	Unclear risk	Comment: Not described.
Blinding of participants and personnel (performance bias)	High risk	Quote: "metacognition. It is carried out using paper and pencil and a blackboard (required to develop some of the tasks, explanations, examples, etc.); in a group format (4-6 participants)," Comment: probably not possible to blind
Blinding of outcome assessment (detection bias)	Low risk	Quote: "psychologists carrying out the assessment were blinded to the treatment until the conclusion of the study."
Incomplete outcome data (attrition bias)	Unclear risk	no details
Selective reporting (reporting bias)	Low risk	Quote: "This clinical trial is registered at ClinicalTrials.gov: NCT01279070 and was approved by the Parc Sanitari Sant Joan de Déu Ethics Committee."
Other bias	Low risk	

Garrido 2013

Methods	<p>Study design: Randomized controlled trial</p> <p>Study grouping: Parallel group</p> <p>Open Label:</p> <p>Cluster RCT:</p>
Participants	<p>Baseline Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 33.21 (6.89) ● Sex (male %): 76 ● Length of illness (years), mean (sd): 10.68 (6.66) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): 64.41 (10.62) <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 33.37 (8.32) ● Sex (male %): 71 ● Length of illness (years), mean (sd): 11.84 (8.23) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): 63.58 (10.99) <p>Included criteria: DSM-IV criteria for schizophrenia disorder confirmed by Structured Clinical Interview for DSM-IV Axis I Disorders (SCID-I; First et al., 1997); age between 18 and 55 years; estimated IQ of 85 or above according to the Vocabulary subtest on the Wechsler Adult Intelligence Scale-III (WAIS-III); patients were considered sufficiently stable if they had a Global Assessment of Functioning score (GAF; Endicott et al., 1976) of 40 or higher and they maintained a stable dose and type of psychiatric medication for at least one month prior to inclusion.</p> <p>Excluded criteria: a score of 6 or higher (severe or extremely severe) on any item of PANSS-P Positive Syndrome Scale, Spanish version (Peralta and Cuesta, 1994); absence of cognitive impairment confirmed by neurocognitive assessment (when raw scores were less than 1 standard deviations of the mean score obtained from the respective normative data in their corresponding manual, they were considered non-impaired); current substance abuse or drug dependence in the last year, defined by the Structured Clinical Interview for DSM-IV (SCID-I; First et al., 1997); traumatic brain injury or history of neurological illness; electroconvulsive therapy in the previous 12 months; psychiatric comorbidity and plan to change medication during the trial.</p>
Interventions	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Description: The active control condition consisted of watching videos for 60 min on a computer with the same features as the therapy condition and led by the same staff who conducted the CACR. At the end of each session, participants had to answer five multiple choice questions referring to different points in the video. Patients were also instructed to write down what had impressed them or what they had liked best about the documentary. <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: 48 sessions of computer-assisted cognitive remediation (CACR) Participants received either CACR therapy or active control within two weeks of randomization (Fig. 1). The intervention was conducted over six months and comprised 48 sessions in both cases. The sessions lasted 60 min and were held twice a week.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning ● Working memory, WAIS III ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, CVLT long term recall ● QoL gls-total ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other. ● Verbal learning and memory, RAVLT, learning (high=better) ● Verbal learning and memory, CVLT short term recall <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	<p>Sponsorship source: This study was supported by the "Fundació La Marató TV-3" (012810).</p> <p>Country: Spain</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Gemma Garrido</p> <p>Institution: Department of Mental Health, Consorci Sanitari de Terrassa, Terrassa (Barcelona), Spain</p> <p>Email: ggarrido@cst.cat</p> <p>Address: Department of Mental Health, Consorci Sanitari de Terrassa, Terrassa, Barcelona, Spain</p>

Notes	<p>Identification: Participants: Study design: Baseline characteristics: Intervention characteristics: Pretreatment: Continuous outcomes: <i>Elisabeth Ginnerup-Nielsen</i> WAIS III = Letter-Number sequencing subtest higher =better The Heinrichs-Carpenter Quality of Life Scale (QLS; Heinrichs et al., 1984) was administered to assess the two secondary outcomes. The QLS assesses overall quality of life and functioning on 21 items rated from 0 to 6 (higher scores reflecting better quality of life). Verbal learning was assessed with California Verbal Learning Test (CVLT; Delis et al., 2000), taking into account the number of words recalled in short-term and long-term free recall. <i>Jesper Østrup Rasmussen</i> Scales: Working memory: WAIS III. QoL: The QLS assesses overall quality of life and functioning on 21 items rated from 0 to 6 (higher scores reflecting better quality of life). Verbal learning and memory: CVLT (taking into account the number of words recalled in short-term and long-term free recall. (Kan sammenlignes med RAVLT learning og delayed)) Short-term free recall Intervention: 11.42 (3.55) Control 11.00 (2.47) Long-term free recall Intervention: 11.65 (3.25) Control: 11.61 (2.52) Dichotomous outcomes: Adverse outcomes:</p>
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Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Participants were randomly assigned using computer-generated random numbers after baseline assessment. Randomization was independently conducted by V.V. who took no part in the implementation of assignments or clinical and neuropsychological assessments. After baseline assessment, patients were assigned to the CACR group or active control group."
Allocation concealment (selection bias)	Unclear risk	Comment: Not described. probably done but unclear
Blinding of participants and personnel (performance bias)	Low risk	Quote: "Both patients and relatives were blinded to the group allocation, and the raters were blind to treatment condition."
Blinding of outcome assessment (detection bias)	Low risk	Comment: Both patients and relatives were blinded to the group allocation, and the raters were blind to treatment condition.
Incomplete outcome data (attrition bias)	High risk	Comment: Big difference in dropout rates: Intervention: 18,4% Control: 37,9%
Selective reporting (reporting bias)	Low risk	Quote: "The trial registration number is NCT01598220." Comment: All outcome from protocol assessed
Other bias	Low risk	

Gharaeipour 2012

Methods	<p>Study design: Randomized controlled trial Study grouping: Parallel group Open Label: Cluster RCT:</p>
Participants	<p>Baseline Characteristics TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 27.62 (5.66) ● Sex (male %): 66.67 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): 14.86 (5.99) ● Level of functioning (GAF, GAS) at baseline, mean (sd): <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 29.81 (7.61) ● Sex (male %): 76.20 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): 15.71 (3.20) ● Level of functioning (GAF, GAS) at baseline, mean (sd): <p>Included criteria: Schizophrenia, as determined by the Structured Clinical Interview for DSM-IV, signed informed consent, age over 20 years, and being fluent in speaking, reading, and writing Farsi. Excluded criteria: Auditory or visual impairment, evidence of mental retardation, history of traumatic brain injury, presence or history of any neurologic illness, and substance abuse or dependence.</p>
Interventions	<p>Intervention Characteristics TAU</p> <ul style="list-style-type: none"> ● Description: <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: The program was made up of educational, experiential (trying out strategies) and reflective (group discussion) components. Patients were engaged in approximately 40 h of neurocognitive exercises that provided equal practice time in areas of attention and concentration learning and memory, and executive functions. Sessions were conducted in a seminar format. At the start of each week the clinical team provided a preview of topics to be presented. The group facilitator usually had a handout and presented information on a whiteboard. Participants took notes, asked questions, shared experiences relating to the topic and offered explanations to each other. Participants were taught about various compensatory strategies relevant to their needs, and they were given the opportunity to practice these strategies. They were also encouraged to relate the tasks carried out within the exercise group to everyday activities.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, other scale ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, ACT higher=better ● Verbal learning and memory, RAVLT, learning (high=better) <p>Dichotomous:</p>

	<ul style="list-style-type: none"> ● Symptomatic remission ● Syptomatic relapse
Identification	<p>Sponsorship source: None to declare Country: Iran Setting: Comments: Authors name: Manouchehr Gharaeipour Institution: Tehran University of Medical Sciences, Department of Psychology, Hazrat Rasoul Akram Hospital, Niayesh, Satar Khan, Tehran, Iran Email: m-gharaeipour@farabi.tums.ac.ir Address:</p>
Notes	<p>Identification: Participants: Study design: Baseline characteristics: Intervention characteristics: <i>Elisabeth Ginnerup-Nielsen</i> Pretreatment: Continuous outcomes: <i>Elisabeth Ginnerup-Nielsen</i> Rey Auditory Verbal Learning Test (RAVLT; Lezak, 2012) assesses verbal learning and verbal memory. Auditory Consonant Trigrams (ACT; Stuss et al., 1987) assesses verbal working memory. higher = better Global cognition score: Computed z scores of the all cognitive measures and summed those z scores separately for the baseline and posttreatment assessments <i>Jesper Østrup Rasmussen</i> Working memory: ACT (The number of letters correctly remembered after 5 second delay interval was tallied. The maximum score was 15 - High=better) Dichotomous outcomes: Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Comment: Unclear how random Only "randomised". Quote: "were randomly assigned to the cognitive remediation condition (n = 21) or the control condition (n = 21), independent of the assessors."
Allocation concealment (selection bias)	Unclear risk	Quote: "Randomization was independently conducted by a Bachelor's level research assistant who was not involved in study treatments." Comment: But could still foresee allocation
Blinding of participants and personnel (performance bias)	High risk	Comment: probably not blinded Not possible.
Blinding of outcome assessment (detection bias)	Low risk	Quote: "Testing on measures occurred prior to random group assignment (baseline) and after a two month intervention (posttreatment). All tests and scales were administered, scored and interpreted by two trained raters with Master's degrees in clinical psychology who were blind to treatment conditions."
Incomplete outcome data (attrition bias)	Low risk	Quote: "All participants in both groups completed the program and no one dropped out of the study."
Selective reporting (reporting bias)	Low risk	Comment: no trial protocol. But described outcomes seem assessed
Other bias	Low risk	

Hadas-Lidor 2001

Methods	
Participants	
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Risk of bias table

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

Lecardeur 2009

Methods	<p>Study design: Study grouping: Open Label: Cluster RCT:</p>
Participants	<p>Baseline Characteristics TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 40.5 (8.9) ● Sex (male %): 87.5 ● Length of illness (years), mean (sd): 12.5 (7.8) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 45 (5.9)

	<ul style="list-style-type: none"> ● Sex (male %): 75 ● Length of illness (years), mean (sd): 20.7 (4.3) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>Included criteria: Individuals were eligible if aged between 18 and 55 years, were fluent in French, met DSM-IV diagnostic criteria (American Psychiatric Association, 1994) for schizophrenia (n=15), schizoaffective disorder (n=8) or delusional disorder (n=1). Participants were stabilized according to their psychiatrist.</p> <p>Excluded criteria: Change in treatment over 2 months prior to the start of the study, meeting criteria for past neurological disorders or substance dependence, and not being able to give informed consent.</p>
Interventions	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Description: <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: Mental Flexibility Therapy (MFT) This therapy was centered on the remediation of mental flexibility. The exercises were oriented towards social situations and activities of daily living. We adopted a maximum of modalities by which mental flexibility could be solicited (language, hearing, touch, etc.). Both therapies were short, with 9 sessions of approximately 1 h, 2 sessions/week (i.e. each therapy lasted 5 weeks). This short format was selected since the duration of CRT was not related to the effect on cognitive performances in a recent meta-analysis (McGurk et al., 2007). This rhythm ensured the maintenance of acquisitions and improved performances. Sessions lasting 1 h took into account tiredness and attentional disturbances typical in schizophrenia patients. Several patients reported that this duration was adjusted to the rhythm of their daily living activities. A group format was chosen to create a stimulating environment that favoured interactions, communications, transfer of knowledge and experiences between participants. We used various media for the exercises proposed to patients.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, PANSS total score ● Verbal learning and memory, other scale ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other. ● Verbal learning and memory, RAVLT, learning (high=better) <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	<p>Sponsorship source: Funding for the study was provided by Fonds de la recherche en santé du Québec (FRSQ) to MCL. LL was supported by the Chaire de Schizophrénie Eli Lilly de l'Université de Montréal (Québec, Canada).</p> <p>Country: Canada</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Laurent Lecardeur</p> <p>Institution: Pavillon Albert-Prévost, Hôpital du Sacré-Coeur de Montréal, Montréal, Québec, Canada</p> <p>Email: lecardeur@cyceron.fr</p> <p>Address:</p>
Notes	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics:</p> <p>Pretreatment:</p> <p>Continuous outcomes:</p> <p>Jesper Østrup Rasmussen Scales: Symptoms: PANSS (low=better)</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Quote: "Group allocation was pseudo-randomized according to patient availability. Neither cognitive deficits nor symptoms were taken into account when patients were assigned to a group."
Allocation concealment (selection bias)	Unclear risk	Comment: Not described
Blinding of participants and personnel (performance bias)	High risk	Comment: Not possible.
Blinding of outcome assessment (detection bias)	High risk	Quote: "scales. Assessors were not blind to group assignment."
Incomplete outcome data (attrition bias)	Unclear risk	Comment: 20 % dropout rates no itt
Selective reporting (reporting bias)	Low risk	
Other bias	Low risk	

Lindenmayer 2008

Methods	<p>Study design: Randomized controlled trial</p> <p>Study grouping: Parallel group</p> <p>Open Label:</p> <p>Cluster RCT:</p>
Participants	<p>Baseline Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 43.33 (8.65) ● Sex (male %): 88

	<ul style="list-style-type: none"> ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 85 <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 43.58 (10.34) ● Sex (male %): 91 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 82 <p>Included criteria: a DSM-IV (14) chart diagnosis of schizophrenia, schizoaffective disorder, or bipolar disorder; absence of psychiatric history of mental retardation, brain injury, or neurological disorder; stable use of medication for at least three months without plans for changing medication; and proficiency in English.</p> <p>Excluded criteria:</p>
Interventions	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Description: The computerized control condition controlled for staff time and computer exposure, which consisted of three weekly, one-hour computer sessions. Similar to the group assigned to cognitive remediation, the groups comprised six to eight patients and were supervised by two or three hospital staff. Patients were assigned to a computer station and given instruction on using the mouse. Patients were then able to play computer games <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: The cognitive remediation program consisted of 24 hours of computerized practice over a 12-week period and a weekly discussion group to facilitate transfer of cognitive skills to daily activities. All study participants had access to the broad range of treatments available to inpatients at Manhattan Psychiatric Center while involved in the trial. These treatments included pharmacological treatment and management, and psychosocial group interventions administered in a required 20-hours-per-week "treatment mall" program. This program included aggression management; mental illness and chemical abuse interventions; social skills training and preparation for community living; and a comprehensive educational program that teaches patients about their medication, mental illness, and healthy lifestyles. Both groups attended the same mix of small group programs, both in terms of number of groups and type of groups.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning ● Working memory, WAIS (note) ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, other ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other ● Verbal learning and memory, RAVLT, learning (high=better) <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	<p>Sponsorship source: Not stated.</p> <p>Country: USA</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Jean-Pierre Lindenmayer</p> <p>Institution: Department of Psychiatry, New York University, New York City</p> <p>Email: lindenmayer@nki.rfmh.org</p> <p>Address:</p>
Notes	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics:</p> <p>Pretreatment:</p> <p>Continuous outcomes:</p> <p><i>Elisabeth Ginnerup-Nielsen</i> Global cognition: Her er brugt composite z-score - Overall cognitive function, psychomotor speed, and verbal learning WAIS er brugt: Digit span (number correct) (Forward) Possible scores range from 0 to 12, with higher scores indicating improvement in verbal working memory from the Wechsler Adult Intelligence Scale-Revised. Resultater Backwards: Intervention: Mean (sd) 4.5 (2.3) (N=42) Control: 3.9 (2.0) N=33</p> <p><i>Jesper Østrup Rasmussen</i> Scales: Verbal learning and memory: RAVLT learning: Possible scores range from 0 to 75 with higher scores indicating higher verbal learning memory. Verbal learning and memory: RAVLT delayed: Possible scores range from 0 to 15, with higher scores indicating increased delayed recall. Working memory: WAIS-R, backward: Possible scores range from 0 to 12, with higher scores indicating improvement in verbal working memory from the Wechsler Adult Intelligence Scale-Revised. Global cognition: WRAT, z score:</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Randomization was conducted with a computer-generated randomization sequence (www.randomization.com) conducted by the study coordinator."
Allocation concealment (selection bias)	Unclear risk	Comment: Not described
Blinding of participants and personnel (performance bias)	High risk	Comment: Not possible. Efforts have been done to make interventions equal but some parts of the intervention are difficult to blind. Also to treating therapists..
Blinding of outcome assessment (detection bias)	Unclear risk	Quote: "Interviewers were blind to the condition to which the patient had been assigned during the treatment phase." Comment: and: Work measures were collected by the vocational rehabilitation director (not blind to the conditions) during the 12-month follow-up and included cumulative hours, weeks worked, and wages earned.

Incomplete outcome data (attrition bias)	Low risk	Quote: "Third, intent-to-treat analyses were conducted to" Comment: Low dropout rates and ITT Quote: "Forty-five were randomized to cognitive remediation and 40 to the control condition. Seventy-two patients completed the entire trial of 24 hours"
Selective reporting (reporting bias)	Unclear risk	Quote: "Positive and Negative Syndrome Scale (PANSS) covering the prior week of functioning (21). Symptoms were assessed at baseline and at six-week, 12-week, six-month, and 12-month follow-ups." Comment: 6 and 12 months not described anywhere in the methods section it says that PANSS will be reported at FU, can't find those answers.
Other bias	Low risk	

Lu 2012

Methods	<p>Study design: Study grouping: Open Label: Cluster RCT:</p>
Participants	<p>Baseline Characteristics TAU <ul style="list-style-type: none"> ● Age, mean (sd): 38 (9) ● Sex (male %): 58.7 ● Length of illness (years), mean (sd): 24 (8) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 100 Cognitive remediation <ul style="list-style-type: none"> ● Age, mean (sd): 37 (8) ● Sex (male %): 63.5 ● Length of illness (years), mean (sd): 23 (9) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 100 <p>Included criteria: schizophrenia (based on the third edition of the Chinese Classification and Diagnostic Criteria of Mental Disorders[5]), had a duration of illness of at least five years, were clinically stable at the time of enrollment (i.e., total score of Positive and Negative Syndrome Scale[6] [PANSS] <60 or a drop in the total PANSS score of more than 50% after initial treatment of acute symptoms), were 18-65 years of age, and were receiving maintenance treatment with a combination of clozapine and risperidone (the most common combined treatment regimen for chronic patients in our setting).</p> <p>Excluded criteria: Excluded patients included those with co-morbid mental retardation, perceptual disorders, organic brain disease, serious medical disorders, impulse control problems, or severe depressive or anxiety symptoms, and those who were pregnant or lactating.</p> </p>
Interventions	<p>Intervention Characteristics TAU <ul style="list-style-type: none"> ● Description: The TAU group received routine occupational and recreational therapy and general mental health education including instructor-led music therapy, dance therapy and physical exercises as well as psychological counseling during daily ward rounds. This treatment also lasted 45 minutes per day and was provided five days per week for three months. The four therapists who provided CRT were different from the clinicians who provided TAU. Cognitive remediation <ul style="list-style-type: none"> ● Description: The CRT employed in this study used a Chinese version of the CRT manual[3,4] developed by Ann Delahunty and Rodney Morice. CRT is a comprehensive neuropsychological training method targeting various major cognitive deficits of schizophrenia that combines verbal reinforcement, errorless learning, individualized guidance and other cognitive therapeutic techniques. The cognitive functioning of patients in three primary areas—cognitive flexibility, working memory and planning—is improved as they complete cognitive tasks of increasing difficulty under the guidance of four trained therapists.[7-10] Training sessions lasted for 45 minutes each and were repeated five days a week for three months. </p>
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning (SSSI) ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, other scale ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other. ● Verbal learning and memory, RAVLT, learning (high=better) <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	<p>Sponsorship source: The study was supported by the Third People's Hospital of Lanzhou Municipality. Country: China Setting: Comments: Authors name: Hongbo LU Institution: The Third People's Hospital of Lanzhou Municipality, Lanzhou, Gansu Province, China Email: hongbolv0308@126.com Address:</p>
Notes	<p>Identification: Participants: Study design: Baseline characteristics: Intervention characteristics: Pretreatment: Continuous outcomes: Elisabeth Ginnerup-Nielsen The Wisconsin Card Sorting Test[11] (WCST) was used to evaluate before versus after changes in cognitive function. Men denne rapporterer 5 forskellige outcomes hhv. 1) Total number of completed cards. 2) Number of cards correctly categorized 3) Number of perseverative errors. 4) Number of random errors 5) Number of categories completed CRT -mean (sd) 973.0 (12.2) 25.8 (4.5) 25.0 (4.0) 30.7 (6.3)</p>

	<p>3.5 (1.1) N=60Control: mean (sd) 79.3 (11.5) 22.5 (2.6) 27.5 (3.2) 36.0 (5.9) 3.0 (1.2) (N=62) the Scale of Social Skills of chronic schizophrenialnpatients[12] (SSSI) was used to assess social functioning, <i>Jesper ØStrup Rasmussen Scales: Socialfunctioning: SSSI</i> (an interviewer-rated questionnaire developed in China specifically for chronic psychiatric inpatients with 10 items (score on 0-2 point Likert scales) that have good internal consistency (alpha=0.89), test-retest reliability (rs=0.97), and inter-rater reliability (rs=0.98);[12] the total score (range 0-20) is used in the current study, with lower scores representing better functioning.Jeg har ikke medtaget Wisconsin sorting card test, da den beskrives som at måle executive functions, og ikke hukommelse (wikipedia). Dichotomous outcomes: Adverse outcomes:</p>
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Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Comment: unclear how
Allocation concealment (selection bias)	Unclear risk	Comment: Not described.
Blinding of participants and personnel (performance bias)	High risk	Comment: Not possible.
Blinding of outcome assessment (detection bias)	High risk	Quote: "Two psychologists were trained to administer these instruments; their inter- rater reliability for the three instruments was assessed by comparing their results for 15 patients (ICC for WCST=0.80, ICC for SSSI=0.87, and ICC for ITAQ=0.92). These psychologists were not blind to the treatment status of the patients they evaluated."
Incomplete outcome data (attrition bias)	Low risk	Quote: "Four patients dropped out during the study leaving 60 in the CRT group and 62 in the TAU group in the final analysis." Comment: Low dropout rates.
Selective reporting (reporting bias)	Low risk	
Other bias	Low risk	

Man 2012

Methods	<p>Study design: Randomized controlled trial Study grouping: Parallel group Open Label: Cluster RCT:</p>
Participants	<p>Baseline Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 35.1 (10.2) ● Sex (male %): 70 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 34.9 (8.5) ● Sex (male %): 56 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>therapist-administered int. (TA)</p> <ul style="list-style-type: none"> ● Age, mean (sd): 41.6 (7.7) ● Sex (male %): 61 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>Included criteria: schizophreniaaged 18 to 55 years who were mentally stable and calm and had a basic attention span of at least 3 minutes Excluded criteria: (1) impaired physicalfunctions inhibiting the operation of a keyboard or mouse,(2) visual impairment such as blindness, partial blindnessand other visual problems, (3) other neurological problems such as epilepsy, (4) pre- and post-morbid mental retardationof severe or moderate grades, (5) previous training of similarcomputerised programmes, or (6) a deviation quotient of<85 in the Test of Nonverbal Intelligence version III (TONI-3),7 which is a language-free intelligence test that measuresabstract/figural problem-solving ability.</p>
Interventions	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Description: <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: The 12-session CAEL and TA programmes weredeveloped based on the work scenario of a conveniencestore worker involving four major tasks: stock keeping,cleansing, food servicing, and cashiering. Five principlesof errorless learning were applied: (1) the to-be-learnedtask was broken down into components, (2) training beganon simple tasks and proceeded gradually to more difficultones, (3) high levels of success were maintained at eachstage with use of aids and abundant positive reinforcement,(4) each component was over-learned through repetitive,successful practice until performed nearly automatically,and (5) the learned components were recombined, addingone component at a time, until the task was trained entirely. <p>therapist-administered int. (TA)</p> <ul style="list-style-type: none"> ● Description: The TA programme was produced by print-screening thescenes of CAEL to form an administration handbook for each session. Thus the two programmes were of similarcontent and structure, but different in the mode of delivery.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning CWPP higher=better ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, Cognistat ● QoL ● Global cognition score higher=better ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other.

	<ul style="list-style-type: none"> ● Verbal learning and memory, RAVLT, learning (high=better) <p><i>Dichotomous:</i></p> <ul style="list-style-type: none"> ● Symptomatic remission ● Syptomtomatic relapse
Identification	<p>Sponsorship source: The study was supported by the Health and Health Services Research Fund, Food and Health Bureau, Hong Kong SAR Government (#05060231).</p> <p>Country: Hong Kong</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: DWK Man</p> <p>Institution: Department of Rehabilitation Sciences, The Hong Kong Polytechnic University</p> <p>Email: David.Man@polyu.edu.hk</p> <p>Address:</p>
Notes	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics: <i>Jesper Østrup Rasmussen</i> The computerassisted intervention was chosen, because most of the included studies are computerassisted.</p> <p>Pretreatment:</p> <p>Continuous outcomes: <i>Elisabeth Ginnerup-Nielsen</i> Til Global cognition score er brugt: Vocational Cognitive Rating Scale Memory er baseret på Neurobehavioral Cognitive Status Examination memory subscale (cognistat) Til social functioning er brugt Chinese Work Personality Profile Social skills subscale <i>Jesper Østrup Rasmussen</i> Length of intervention: 12 weeks. FU 3 month - not included (4 mo cutoff). scales: Memory: The Neurobehavioral Cognitive Status Examination (NCSE) Chinese version, 9 which is a standardised examination of global cognitive function. It assesses multiple domains of cognitive functioning, namely: orientation, attention, language, construction, memory, calculation and reasoning. It was used to detect changes in the global cognitive functioning of the subjects after training. (High=better) OBS KUN learning IKKE memory - subgruppeanalyse. Jeg har ikke medtaget vocational skalaen, eller work profile skalaen, tænker de er for minded på arbejde. Jeg har ikke medtaget Wisconsin, obs om det evt. er et mål for learning.</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Comment: Not clear how
Allocation concealment (selection bias)	Unclear risk	Comment: Not described
Blinding of participants and personnel (performance bias)	Unclear risk	Quote: "follow-up. The patients and assessors did not know the expected results of the training programmes." Comment: Treating Personnel not blinded. And patient knew if they were in control group. At the same time some effort has been done to eliminate bias.
Blinding of outcome assessment (detection bias)	Low risk	Comment: assessed by independent raters
Incomplete outcome data (attrition bias)	Unclear risk	Quote: "commencement. Of 90 subjects with schizophrenia aged 18 to 55 years who were mentally stable and calm and had a basic attention span of at least 3 minutes, 80 completed the study and 10 dropped out (owing to early discharge, incomplete training or data set)." Comment: No itt and relatively small but skewed dropout 7/30, 3/10 and 0/30
Selective reporting (reporting bias)	Low risk	Comment: No trial protocol. All described outcomes assessed.
Other bias	Low risk	

Medalia 1998

Methods	
Participants	
Interventions	
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Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	no details
Allocation concealment (selection bias)	Unclear risk	no details
Blinding of participants and personnel (performance bias)	High risk	
Blinding of outcome assessment (detection bias)	Low risk	An outside consultant who was blind to group status, a psychologist trained in rating videotaped psychiatric interviews with the BPRS, viewed each taped interview and provided a BPRS rating.
Incomplete outcome data (attrition bias)	Low risk	
Selective reporting (reporting bias)	Low risk	
Other bias	Low risk	

Poletti 2010

Methods	<p>Study design: Randomized controlled trial</p> <p>Study grouping: Parallel group</p> <p>Open Label:</p> <p>Cluster RCT:</p>
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<p>Participants</p>	<p>Baseline Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 34.69 (7.63) ● Sex (male %): 61.9 ● Length of illness (years), mean (sd): 9.88 (6.24) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 100 <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 34.00 (9.87) ● Sex (male %): 58.6 ● Length of illness (years), mean (sd): 10.17 (8.24) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 100 <p>Included criteria: To be included, patients had to satisfy DSM-IV criteria for schizophrenia and the following conditions: 1. To have been treated with a stable dose of the same antipsychotic therapy for at least 6 months, and be responsive (30% or more response based on PANSS scores) and clinically stabilised. 2. To have participated in a rehabilitation programme, including both cognitive-behavioural and psychosocial programmes, three hours a day, three times a week, for three months.</p> <p>Excluded criteria:</p>
<p>Interventions</p>	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Description: SRT + PBO. The control condition consisted of one hour a week of computer-aided non-domain-specific activity and two extra hours a week of SRT (patients were randomly assigned to one of the non-cognitive groups previously described), for a period of 12 weeks. Subjects completed a total of 36 hours. <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: SRT + CRT. The experimental condition consisted of three 1-hour sessions a week of domain-specific computer-aided exercises, for a period of 12 weeks. This gave a total of 36 hours. Sets of exercises were individually created for each patient on the basis of the quality of baseline performances at neuropsychological assessment: for each poor performance, a domain-specific exercise was included, while for each good performance a non-domain-specific exercise was added. The computer-aided training employed the Cogpack Software (Marker, 1987-2007). This computer programme includes different neurocognitive exercises that can be divided into domain-specific exercises, aimed at training specific cognitive areas among the ones known to be impaired in schizophrenia (verbal memory, verbal fluency, psychomotor speed and coordination, executive function, working memory, attention) and non-domain-specific exercises, that do not focus on one specific function but require the use of several functions at a time and engage functions such as culture, language and simple calculations skills.
<p>Outcomes</p>	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, mean of 5 trials ● QoL (QLS) higher=better ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, digit seq. higher=better. ● Verbal learning and memory, RAVLT, learning (high=better) <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
<p>Identification</p>	<p>Sponsorship source: This work was supported by the Italian Ministry of University and Scientific Research, grant number 2001064198.</p> <p>Country: Italy</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Sara Poletti</p> <p>Institution: C.E.R.M.A.C. (Centro di Eccellenza Risonanza Magnetica ad Alto Campo), University Vita-Salute San Raffaele, Milan, Italy</p> <p>Email: poletti.sara@hsr.it</p> <p>Address:</p>
<p>Notes</p>	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics:</p> <p>Pretreatment:</p> <p>Continuous outcomes:</p> <p><i>Elisabeth Ginnerup-Nielsen</i> Verbal and working memory based on BACS Verbal memory - words recall (mean of 5 trials) Working memory ((digit sequencing) n correct sequences) QLS: Daily functioning was assessed by the Quality of Life Scale (QLS; Heinrichs, Hanlon, & Carpenter, 1984), a semi-structured interview made up of 21 items that evaluates three different areas of social functioning:</p> <p><i>Jesper Østrup Rasmussen</i> Scales: Working memory: BACS (high = better) Verbal memory (ikke learning - obs subgruppe): BACS (high=better) QoL: Quality of Life Scale (high=better)</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "programme. Following admission to the study, computer-generated random number tables allocated each patient to one of the following conditions:"
Allocation concealment (selection bias)	Unclear risk	Comment: Not described.
Blinding of participants and personnel (performance bias)	High risk	Quote: "The rehabilitation therapist was blind to the randomisation and neuropsychological assessment." Comment: Blind personnel but patients probably not

Blinding of outcome assessment (detection bias)	Low risk	Quote: "1987), administered by trained psychiatrists who were blind to treatment randomisation and neuropsychological testing."
Incomplete outcome data (attrition bias)	High risk	Quote: "36 patients in the SRT (standard rehabilitation treatment) + PBO (placebo) group and 50 in the SRT + CRT group (Cavallaro et al., 2009); 13 subjects dropped out from the SRT + PBO group and 11 from the SRT + CRT group." Comment: Large skewed dropout and no ITT Dropouts: Intervention: 22%TAU: 36%
Selective reporting (reporting bias)	Low risk	Comment: No trial protocol but all outcome reported seems assessed
Other bias	Low risk	

Rass 2012

Methods	<p>Study design: Randomized controlled trial Study grouping: Parallel group Open Label: Cluster RCT:</p>
Participants	<p>Baseline Characteristics TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 43.9 (8.9) ● Sex (male %): 90 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 100 <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 37.2 (12.5) ● Sex (male %): 58.8 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 100 <p>Included criteria: Axis-I diagnosis of schizophrenia or schizoaffective disorder(SZ) was obtained by Structured Clinical Interview for DSM-IV (SCID-I:First et al., 2001), clinical observations, and chart review. Inclusion criteria for all participants were: 1) age between 18 and 50 years;2) no history of electroconvulsive therapy; 3) no history of neurological illness; 4) no current alcohol or drug dependence (DSM-IV criteria) as ascertained by administration of the SCID sections on substance use disorders; 5) no hearing impairments on audiometry;6) verbal I.Q. above 70; 7) visual acuity (with correction) of 20/30 or better; and 8) no alcohol use in the 24 h prior to testing. Excluded criteria:</p>
Interventions	<p>Intervention Characteristics TAU</p> <ul style="list-style-type: none"> ● Description: The TAU participants came in only for assessments. <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: participants completed assigned tasks for 2 h, including breaks, two days per week for ten weeks, a treatment schedule consistent with other studies that showed positive outcomes. The CR group completed a cognitive training regimen using software that applies adaptive algorithms to continuously adjust the demands of each task according to performance (Mahncke et al., 2006). Participants completed auditory exercises described previously by Fisher et al. (2009a) and visual exercises. The visual module aims to improve the speed and accuracy of visual processing, to facilitate perception, to improve visual memory, and to reduce response time.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, HVLT delayed higher=better ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other. ● Verbal learning and memory, RAVLT, learning (high=better) ● Verbal learning and memory HVLT Total higher=better <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	<p>Sponsorship source: We are grateful for the support from NIMH RO1 MH62150 (BFO), NIMH R21MH091774, and IUSM/CTR, NIH/NCRR grant number RR025761 to BFO; NIMH R01MH074983 to WPH; NARSAD (ARB) and NIDA T32 DA024628-01 (OR). Country: USA Setting: Comments: Authors name: Olga Rass Institution: Department of Psychological and Brain Sciences, Indiana University, 1101 East 10th Street, Bloomington, IN 47405, USA Email: rasso@indiana.edu Address:</p>
Notes	<p>Identification: Participants: Study design: Baseline characteristics: Intervention characteristics: Pretreatment: Continuous outcomes: <i>Elisabeth Ginnerup-Nielsen</i> Verbal memory function was assessed using the Letter-Number Sequencing subtest of the Wechsler Adult Intelligence Scale (WAIS-III; Wechsler, 1997) and the Hopkins Verbal Learning Test (HVLT; Brandt and Benedict, 2001). Forstær det som at WAIS ligger under HVLT?? <i>Jesper Østrup Rasmussen</i> Length of intervention: 10 weeks, FU after 20 weeks (only 2,5 mo FU, our cut off is 4 mo, not included) Scales: Verbal learning: HVLT (high=better) Verbal memory (delayed): HVLT (high=better) Global cognition: ? Dichotomous outcomes: Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Comment: Participants were randomly assigned using a random number table to cognitive remediation treatment (CR) or an active control study arm.
Allocation concealment (selection bias)	Unclear risk	no details
Blinding of participants and personnel (performance bias)	High risk	Comment: Not possible
Blinding of outcome assessment (detection bias)	Low risk	Comment: Assessments were completed at baseline, five weeks, ten-weeks, and twenty weeks follow-up by staff blind to treatment condition.
Incomplete outcome data (attrition bias)	Low risk	Comment: No it and 4/21 and 0/10 dropout
Selective reporting (reporting bias)	Unclear risk	Quote: "WAIS-III Spatial Span forward and backward assessed visual working memory" Comment: Mentioned as an outcome but not assessed? No trial protocol
Other bias	Unclear risk	no details

Royer 2012

Methods	<p>Study design:</p> <p>Study grouping:</p> <p>Open Label:</p> <p>Cluster RCT:</p>
Participants	<p>Baseline Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 35.5 (9.0) ● Sex (male %): ● Length of illness (years), mean (sd): 11.8 (8.6) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 31.0 (7.6) ● Sex (male %): ● Length of illness (years), mean (sd): 10.6 (7.8) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): <p>Included criteria: DSM-IV diagnosis of schizophrenia more than 70 Intelligence quotient (IQ; Wechsler Adult Intelligence Scale [WAIS-R]; Britton & Savage, 1966), cognitive deficiency in at least one attention TEA test (score below the second percentile; Zimmermann & Fimm, 1994), and/or on memory (score below 2 standard deviations [SD] in the Grober & Buschke test, 1987), and/or on executive functions (score below the fifth percentile; Rousset & Godefroy, 2008).</p> <p>Excluded criteria: mental retardation (IQ B70), traumatic brain injury, presence or history of any neurological condition, and criteria for substance abuse or dependence.</p>
Interventions	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Description: <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: The training programme was carried out for 6 hours/week for 6 months. Each 2 hour training session comprised paper and pencil exercises for 100 min in groups of six to eight persons, and individual training for 20 min with computer exercises. The cognitive training programme consisted of a series of exercises of increasing complexity. The first 6 weeks focused on attention, the next 3 weeks on language, the next 8 weeks introduced working and long-term memory, and the final 7 weeks focused on planning and problem solving. A psychologist assisted the patients in implementing strategies (i.e., compensation approach) in order to find a way adapted to the patient deficit to perform the exercises. Computerised exercises. A computerised training programme (REHACOM† software; Schuhfried Company) was used. The exercises were repetitive (i.e., restitution approach) and consisted of five 10-min sessions with tasks designed to train vigilance, divided attention, reaction time, visuomotor and visuoconstruction skills. The level of difficulty increased with achievement, every 90 s. If the patient was not successful, another task was proposed.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other. ● Verbal learning and memory, RAVLT, learning (high=better) ● long-term verbal memory delayed Groebe dFR16 higher=better <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	<p>Sponsorship source: Funding for this study was provided by a grant of the French ministry of health.</p> <p>Country: France</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Aurélie Royer</p> <p>Institution: Department of Psychiatry, University Hospital, Saint-Etienne, France</p> <p>Email: aureroyer@hotmail.com</p> <p>Address:</p>

Notes	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics:</p> <p>Pretreatment:</p> <p>Continuous outcomes: <i>Elisabeth Ginnerup-Nielsen</i> working memory: backward span test (WAIS-R; Wechsler, 1981), two-back test (Zimmermann & Fimm, 1994) long-term verbal memory: the two parallel forms of Grober and Buschke test (1987); <i>Jesper Østrup Rasmussen</i> Scales: working memory: WAIS-R, backward span test (high=better) long-term verbal memory: the two parallel forms of Grober and Buschke test (d/FR) (high=better)</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>
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Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Comment: they were randomly allocated to one of two groups (CRT or TAU) using a centralised randomisation procedure.
Allocation concealment (selection bias)	Unclear risk	no details
Blinding of participants and personnel (performance bias)	High risk	Comment: Not possible.
Blinding of outcome assessment (detection bias)	Unclear risk	Quote: "Cognitive functions were assessed by trained neuropsychologists" Comment: Nothing said about blinding of assessors
Incomplete outcome data (attrition bias)	High risk	Comment: 3/28 patients dropped out in the CRT group and 8/18 in the TAU group. No it analysis dropout rates: Intervention: 9,7% Control: 33.3
Selective reporting (reporting bias)	Low risk	Comment: No trial protocol .. But all outcome seems reported
Other bias	Low risk	

Sanchez 2014

Methods	<p>Study design:</p> <p>Study grouping:</p> <p>Open Label:</p> <p>Cluster RCT:</p>
Participants	<p>Baseline Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 36.92 (10.5) ● Sex (male %): 77.1 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): 43.33 (16.3) ● Schizophrenia, Schizoaffective, schizophreniform (%): 100 <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 33.60 (9.4) ● Sex (male %): 75 ● Length of illness (years), mean (sd): ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): 38.88 (13.5) ● Schizophrenia, Schizoaffective, schizophreniform (%): 100 <p>Included criteria: Diagnostic criteria for schizophrenia according to the American Psychiatric Association's Diagnostic and Statistical Manual of Mental Disorders, Fourth Edition, Text Revision (DSM-IV-TR).</p> <p>Excluded criteria: -evidence of alcohol or drug abuse in the last 30 days;- previous history of a significant lack of consciousness;- mental retardation; and- relevant neurological or medical conditions.</p>
Interventions	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Description: The control group received standard treatment and participated in group activities including drawing, reading the daily news, and constructing objects using different materials (such as paper or wood). These activities were accomplished in a group format and with the same frequency as the implementation of REHACOP. <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Description: REHACOP group attended 90-min sessions at least 3 days per week over 3 months. REHACOP is a structured program based on paper-pencil tasks and uses the principles of restoration, compensation, and optimization. Training procedures gradually increase the level of cognitive effort and demand. REHACOP trains patients in traditionally impaired cognitive domains such as attention, memory, processing speed, language, and executive functioning.
Outcomes	<p>Continuous:</p> <ul style="list-style-type: none"> ● Social functioning WHODAS lower=better ● Working memory, WAIS ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, PANSS total score ● Verbal learning and memory, HVLT ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other. ● Verbal learning and memory, RAVLT, learning (high=better) <p>Dichotomous:</p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse

Identification	<p>Sponsorship source: Health Department of the Basque Government(2010111136, 2011111102); Educational and ScienceDepartment of the Basque Government (BFI09.123).</p> <p>Country: Spain</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Pedro Sánchez</p> <p>Institution: Refractory Psychosis Unit, Hospital Psiquiátrico de Alava, Vitoria, Spain</p> <p>Email: nojeda@deusto.es</p> <p>Address:</p>
Notes	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics:</p> <p>Pretreatment:</p> <p>Continuous outcomes: <i>Elisabeth Ginnerup-Nielsen</i> Functional disability was assessed with the GlobalAssessment of Functioning (GAF) scale,42 the CGIscale,43 and the Disability Assessment Schedule scalefrom World Health Organization (DAS-WHO).44The 4 functional disability characteristic indicators offered bythe DAS-WHO were analyzed.The evaluation of cognitive functioning included tests toassess processing speed, working memory, verbal learningand memory, verbal fluency, and executive functioning.All cognitive measures were converted into Z-scores, higher=better Er der en en ved en Z-score??For learning and verbal memory, authors includedlearning and long-term recall from the Hopkins VerbalLearning TestWorking memory (Cronbach's $\alpha = .73$) was assessed using Digit Forwardand Digit Backwards from WAIS-III <i>Jesper Østrup Rasmussen</i> scales:Verbal learning and memory: HVLT (high=better)Working memory: Digit Backwards from WAIS-III (high=better)Symptoms; PANNS (Low=better)Socialfunctioning: WHO-DAS, social competence subscale (low=better)</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Assignment to the program was performed using a computer-generated random- ization list."
Allocation concealment (selection bias)	Unclear risk	Comment: Not described.
Blinding of participants and personnel (performance bias)	High risk	Comment: Not possible
Blinding of outcome assessment (detection bias)	Low risk	Comment: All raters were blind to thetreatment condition and had no other role in the projectthat would undermine the blinding.
Incomplete outcome data (attrition bias)	Low risk	Comment: 2/38 and 6/54 5% in intervention group 11 % in control. Relatively small but unequal dropout. ITT should have been doneLow dropout rates.
Selective reporting (reporting bias)	Low risk	Comment: No trial protocol but all outcome seems relevant and reported
Other bias	Low risk	

Silverstein 2005

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Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	no details
Allocation concealment (selection bias)	Unclear risk	no details
Blinding of participants and personnel (performance bias)	High risk	
Blinding of outcome assessment (detection bias)	High risk	Raters were not blind to treatment condition.
Incomplete outcome data (attrition bias)	Unclear risk	no information
Selective reporting (reporting bias)	Low risk	
Other bias	Low risk	

Tan 2013

Methods	<p>Study design: Randomized controlled trial</p> <p>Study grouping: Parallel group</p> <p>Open Label:</p> <p>Cluster RCT:</p>
Participants	<p>Baseline Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● Age, mean (sd): 36.80 (10.08) ● Sex (male %): 55.9 ● Length of illness (years), mean (sd): 11.96 (8.87) ● Length of illness (month), mean (sd): ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 100 <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● Age, mean (sd): 32.70 (10.86) ● Sex (male %): 58.3 ● Length of illness (years), mean (sd): 9.28 (6.85) ● Length of illness (month), mean (sd):

	<ul style="list-style-type: none"> ● Level of functioning (GAF, GAS) at baseline, mean (sd): ● Schizophrenia, Schizoaffective, schizofreniform (%): 100 <p>Included criteria: Diagnostic and Statistical Manual of Mental Disorders, Fourth Edition (DSM-IV) diagnoses of schizophrenia or schizoaffective disorder, as certified by a psychiatrist. Global Assessment of Functioning score of above 30.</p> <p>Excluded criteria: patients with known neurological, cardiovascular and respiratory diseases as well as developmental disabilities were excluded.</p>
Interventions	<p>Intervention Characteristics</p> <p>TAU</p> <ul style="list-style-type: none"> ● <i>Description:</i> The PE programme was adapted from the Structured Exercise Programme implemented by the Centre for Psychiatric Rehabilitation at Boston University (Hutchinson et al., 2005). To match the treatment intensity and duration of CR, the PE programme was also carried out for 5 hours (three sessions) per week over 12 weeks. The PE programme consisted of exercises in the gymnasium, as well as physical-based counselling <p>Cognitive remediation</p> <ul style="list-style-type: none"> ● <i>Description:</i> Computer-assisted cognitive exercises. The computer-assisted cognitive exercises were conducted for up to 5 hours (three sessions) each week for 12 weeks. In addition to the computer exercises, participants received cognitive-based counselling fortnightly.
Outcomes	<p><i>Continuous:</i></p> <ul style="list-style-type: none"> ● Social functioning ● Working memory, WAIS digit span higher=better ● Verbal learning and memory, WLM (high=better) ● Verbal learning and memory, RAVLT, total (high=better) ● Days at hospital ● Symptoms, total score ● Verbal learning and memory, other scale ● QoL ● Global cognition score, Z score ● Verbal learning and memory, RAVLT, delayed (high=better) ● Working memory, other. ● Verbal learning and memory, RAVLT, learning (high=better) <p><i>Dichotomous:</i></p> <ul style="list-style-type: none"> ● Symptomatic remission ● Symptomatic relapse
Identification	<p>Sponsorship source: This research was partially funded by a grant of S\$10,000 from the Institute of Mental Health Research Department. Study number: 175/2006.</p> <p>Country: Singapore</p> <p>Setting:</p> <p>Comments:</p> <p>Authors name: Bhing-Leet Tan</p> <p>Institution: Occupational Therapy Department, Institute of Mental Health, Singapore</p> <p>Email: Bhing_Leet_TAN@imh.com.sg</p> <p>Address:</p>
Notes	<p>Identification:</p> <p>Participants:</p> <p>Study design:</p> <p>Baseline characteristics:</p> <p>Intervention characteristics:</p> <p>Pretreatment:</p> <p>Continuous outcomes:</p> <p><i>Elisabeth Ginnerup-Nielsen</i> QOL assessed by: The World Health Organization Quality of Life Scale-Brief (WHOQOL-Brief) QOL og PANSS (symptoms) kun opgivet via F-score. Lastly, the Wechsler Adult Intelligence Scale (WAIS)-Digit Span Forward and Backward was administered as a test of attention and working memory. I SKEMA ER BRUGT BACKWARDS</p> <p><i>Jesper Østrup Rasmussen</i> Scales: Working memory: WAIS, Digit Span Backward (high=better) Verbal memory and learning: RAVLT (high=better)</p> <p>Dichotomous outcomes:</p> <p>Adverse outcomes:</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Comment: During randomisation, a biostatistician independent to the study generated a random allocation of treatment using the computer.
Allocation concealment (selection bias)	Low risk	Comment: This sequence was placed in numbered sealed envelopes and given to co-investigators who recruited the participants.
Blinding of participants and personnel (performance bias)	Low risk	Comment: Not described exactly what was done, but since both groups received an intervention, there was likely a similar "placebo effect" in both groups. In addition, all therapists were told that CR and PE were interventions likely to yield benefits to participants and that the efficacy of both interventions was under investigation in this study. Hence, all therapists and participants were informed that the topic of the research study was 'The effects of CR and PE on functional outcomes among people with schizophrenia'. They were not told that the CR was the treatment that was being researched and that PE was the placebo treatment.
Blinding of outcome assessment (detection bias)	Unclear risk	Quote: "These two measurements were administered at baseline and upon completion of the 12-week treatment." Comment: The recruiting therapists, vocational training/day rehabilitation therapists as well as the therapists involved in conducting CR and PE were not involved in administering the neurocognitive and Rockport Walking tests. Probably not blinded
Incomplete outcome data (attrition bias)	High risk	Comment: 10/36 and 10/34 dropped out. Even if it was done this is a relatively large dropout. High dropout rates: Intervention: 30.6% Control: 38.2%
Selective reporting (reporting bias)	High risk	Comment: No trial protocol and WHO-QOL and PANSS unclearly reported
Other bias	Low risk	

Twamley 2008

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Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	no details
Allocation concealment (selection bias)	Unclear risk	no details
Blinding of participants and personnel (performance bias)	High risk	
Blinding of outcome assessment (detection bias)	Low risk	Examiners were trained to a high level of interrater reliability (ICC > .90) and were blind to group assignment.
Incomplete outcome data (attrition bias)	High risk	Thirty-eight participants completed a baseline assessment and at least one follow-up evaluation. Three participants dropped out of the SP group, and 11 dropped out of the CT group. Of the CT drop-outs, 7 dropped out after randomization but before the group started, and four attended one or two CT sessions but did not return. One found the group setting too anxiety-provoking, one felt he had no cognitive problems, one relapsed on alcohol and stopped coming to the clinic entirely, and one discontinued his antipsychotic medication and had a symptom exacerbation. The 14 drop-outs did not differ statistically from the 38 completers on age, education, gender, ethnicity, diagnosis, duration of illness, antipsychotic dose, positive and negative symptom severity, or premorbid IQ.
Selective reporting (reporting bias)	Low risk	
Other bias	Low risk	

Van der Gaag 2002

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Risk of bias table

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

Velligan 2000

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Risk of bias table

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

Velligan 2002

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Risk of bias table

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

Velligan 2008A

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Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	no details
Allocation concealment (selection bias)	Unclear risk	no details
Blinding of participants and personnel (performance bias)	High risk	
Blinding of outcome assessment (detection bias)	Low risk	The SOFAS score was based upon all information obtained during several hours of assessments conducted by blinded raters. In an effort to maintain treatment blinds, all subjects and collaterals were asked at the beginning of each assessment neither to divulge information about any visits made by staff of the research project nor to refer to any items they may have received as part of the study. If blinds were broken, alternative raters blind to group assignment completed the remaining assessments.
Incomplete outcome data (attrition bias)	Low risk	By the end of 24 months, 31%, 37% and 17% of participants dropped out of TAU, GES, and CAT respectively
Selective reporting (reporting bias)	High risk	No raw estimates are reported, only effect sizes (Results of a mixed effects regression model examining treatment group differences over time...)
Other bias	Low risk	

Velligan 2008B

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Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	no details
Allocation concealment (selection bias)	Unclear risk	no details
Blinding of participants and personnel (performance bias)	High risk	
Blinding of outcome assessment (detection bias)	Low risk	In an effort to maintain treatment blinds, all subjects and collaterals were asked at the beginning of each assessment neither to divulge information about any visits made by staff of the research project nor to refer to any items they may have received as part of the study. If blinds were broken, alternative raters blind to group assignment completed the remaining assessments.
Incomplete outcome data (attrition bias)	Low risk	105 were randomized. Of these 105, there were 95 subjects with baseline and follow-up data for data analyse
Selective reporting (reporting bias)	High risk	some outcomes insufficiently reported (e.g. only ES)
Other bias	Low risk	

Wykes 2007

Methods	
Participants	
Interventions	
Outcomes	
Identification	
Notes	

Risk of bias table

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

Wykes 2007A

Methods	
Participants	
Interventions	
Outcomes	
Identification	
Notes	

Risk of bias table

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	participants randomized to group by an independent trial statistician following a baseline assessment. Block randomization was used with CRT and control treatment being assigned randomly to 4 patients each within blocks of 8.
Allocation concealment (selection bias)	Low risk	
Blinding of participants and personnel (performance bias)	High risk	
Blinding of outcome assessment (detection bias)	Unclear risk	Symptom and quality of life assessments were assessed by an independent rater who was blind to group allocation. Self-report assessments (cognition and self-esteem) and informant ratings (social behaviour) were collected by a research assistant who was not blind to group allocation.
Incomplete outcome data (attrition bias)	Low risk	Low dropout rates.
Selective reporting (reporting bias)	Low risk	
Other bias	Low risk	

Footnotes

Characteristics of excluded studies

Alwi 2010

Reason for exclusion	Wrong setting
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Arango Lasprilla 2012

Reason for exclusion	Wrong comparator
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Bell 2009

Reason for exclusion	Wrong comparator
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Bor 2011

Reason for exclusion	Data indgår i et andet studie
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Bowie 2012

Reason for exclusion	Wrong comparator
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Bucci 2013

Reason for exclusion	Wrong comparator
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Burton 2011

Reason for exclusion	Wrong intervention
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Cavallaro 2009

Reason for exclusion	Wrong comparator
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Cella 2014

Reason for exclusion	Wrong outcomes
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Dang 2014

Reason for exclusion	Wrong outcomes
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Eack 2010

Reason for exclusion	Wrong outcomes
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Eack 2010a

Reason for exclusion	Wrong outcomes
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Eack 2013

Reason for exclusion	Wrong outcomes
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Farreny 2013

Reason for exclusion	Konference abstract uden data
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Farreny 2013a

Reason for exclusion	Data indgår i et andet studie
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Franck 2013

Reason for exclusion	Wrong comparator
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Hodge 2010

Reason for exclusion	Doublet. Allerede inkluderet studie
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Kidd 2014

Reason for exclusion	Wrong patient population
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Klingberg 2012

Reason for exclusion	Wrong comparator
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Kurtz 2009

Reason for exclusion	Wrong comparator
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Lalova 2013

Reason for exclusion	Wrong comparator
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Lewandowski 2011

Reason for exclusion	Wrong comparator
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Lindenmayer 2008a

Reason for exclusion	Doublet. Allerede inkluderet studie
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Lindenmayer 2013

Reason for exclusion	Wrong comparator
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Linke 2013

Reason for exclusion	Konference abstract uden data
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Matsui 2009

Reason for exclusion	Wrong intervention
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McGurk 2008

Reason for exclusion	Wrong setting
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McGurk 2008a

Reason for exclusion	Wrong comparator
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McGurk 2013

Reason for exclusion	Konference abstract uden data
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Pontes 2013

Reason for exclusion	Pilot study, ikke europæisk sample, meget lille sample
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Reeder 2013

Reason for exclusion	Konference abstract uden data
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Ross 2011

Reason for exclusion	Wrong intervention
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Sartory 2008

Reason for exclusion	Referat af Wykes 2007 studie
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Schmidt 2011

Reason for exclusion	Wrong setting
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Surti 2011

Reason for exclusion	Konference abstract uden data
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Twamley 2010

Reason for exclusion	Konference abstract uden data
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Twamley 2011

Reason for exclusion	Konference abstract uden data
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Twamley 2011a

Reason for exclusion	Wrong study design
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Twamley 2012

Reason for exclusion	Dublet. Allerede inkluderet studie
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Vesterager 2010

Reason for exclusion	Konference abstract uden data
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Vinogradov 2013

Reason for exclusion	Konference abstract uden data
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Vinogradov 2014

Reason for exclusion	Konference abstract uden data
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Vita 2011

Reason for exclusion	Wrong intervention
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Footnotes

Characteristics of studies awaiting classification

Footnotes

Characteristics of ongoing studies

Footnotes

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Data and analyses

1 TAU vs Cognitive remediation

Outcome or Subgroup	Studies	Participants	Statistical Method	Effect Estimate
1.1 Global cognition score (Z score, final score), end of treatment	2	118	Std. Mean Difference (IV, Random, 95% CI)	-0.28 [-0.70, 0.13]
1.2 Social function End of treatment	6	479	Std. Mean Difference (IV, Random, 95% CI)	-0.56 [-0.96, -0.16]
1.2.2 SBS (higher=worse) End of treatment	1	79	Std. Mean Difference (IV, Random, 95% CI)	-0.17 [-0.62, 0.27]
1.2.3 SFS (higher=better)	1	57	Std. Mean Difference (IV, Random, 95% CI)	-0.15 [-0.67, 0.37]
1.2.4 SSSI (higher=better)	1	122	Std. Mean Difference (IV, Random, 95% CI)	-0.31 [-0.67, 0.05]
1.2.5 WHODAS (higher=worse)	1	84	Std. Mean Difference (IV, Random, 95% CI)	-0.35 [-0.79, 0.09]
1.2.6 SoFAS (higher=better)	2	137	Std. Mean Difference (IV, Random, 95% CI)	-1.24 [-1.70, -0.78]
1.3 Social functioning, FU	4	261	Std. Mean Difference (IV, Random, 95% CI)	-0.26 [-0.51, -0.01]
1.3.2 SBS (higher=worse) FU	1	76	Std. Mean Difference (IV, Random, 95% CI)	0.04 [-0.42, 0.49]
1.3.3 SFS (higher=better)	1	47	Std. Mean Difference (IV, Random, 95% CI)	-0.20 [-0.78, 0.38]
1.3.4 SoFAS (higher=better)	2	138	Std. Mean Difference (IV, Random, 95% CI)	-0.45 [-0.79, -0.11]
1.4 Working memory, end of treatment	9	574	Std. Mean Difference (IV, Random, 95% CI)	-0.66 [-1.04, -0.27]
1.4.2 ANS (higher=better)	1	77	Std. Mean Difference (IV, Random, 95% CI)	-0.21 [-0.66, 0.24]
1.4.3 WAIS digit span backward (higher=better)	1	70	Std. Mean Difference (IV, Random, 95% CI)	-0.47 [-0.95, 0.01]
1.4.4 WAIS III digit span backwards (higher=better)	2	130	Std. Mean Difference (IV, Random, 95% CI)	-1.33 [-2.60, -0.07]
1.4.5 ACT (higher=better)	1	42	Std. Mean Difference (IV, Random, 95% CI)	-1.00 [-1.65, -0.35]
1.4.6 BACS (higher=better)	1	100	Std. Mean Difference (IV, Random, 95% CI)	-0.33 [-0.73, 0.07]
1.4.7 WAIS III letter-number seq. (higher=better)	2	80	Std. Mean Difference (IV, Random, 95% CI)	-0.55 [-1.12, 0.03]
1.4.8 WAIS-R digit span backwards (higher=better)	1	75	Std. Mean Difference (IV, Random, 95% CI)	-0.27 [-0.73, 0.19]

1.5 Verbal learning and memory, Total	2	97	Std. Mean Difference (IV, Random, 95% CI)	-0.50 [-2.37, 1.37]
1.5.1 HVLt total (higher=better) End of treatment (final value)	1	27	Std. Mean Difference (IV, Random, 95% CI)	0.48 [-0.32, 1.27]
1.5.2 RAVLT total (higher=better) End of treatment (final value)	1	70	Std. Mean Difference (IV, Random, 95% CI)	-1.43 [-1.96, -0.90]
1.6 Verbal learning	6	330	Std. Mean Difference (IV, Random, 95% CI)	-0.23 [-0.55, 0.09]
1.6.1 RAVLT, learning (higher=better)End of treatment (final value)	2	116	Std. Mean Difference (IV, Random, 95% CI)	-0.54 [-1.05, -0.03]
1.6.2 CVLT short term recal (higher=better)End of treatment (final value)	1	49	Std. Mean Difference (IV, Random, 95% CI)	-0.13 [-0.71, 0.45]
1.6.3 WLM (first recall) (higher=better) End of treatment (final value)	1	77	Std. Mean Difference (IV, Random, 95% CI)	-0.44 [-0.90, 0.01]
1.6.4 WMS ST (higher=better)	1	57	Std. Mean Difference (IV, Random, 95% CI)	0.35 [-0.18, 0.87]
1.6.5 HVLt, learning (higher=better) End of treatment	1	31	Std. Mean Difference (IV, Random, 95% CI)	0.04 [-0.69, 0.78]
1.7 Verbal memory	10	578	Std. Mean Difference (IV, Random, 95% CI)	-0.34 [-0.71, 0.04]
1.7.1 CVLT long tern recall (higher =better) End of treatment (final value)	1	49	Std. Mean Difference (IV, Random, 95% CI)	-0.01 [-0.59, 0.57]
1.7.2 HVLt delayed (higher=better) End of treatment (final value)	1	27	Std. Mean Difference (IV, Random, 95% CI)	0.50 [-0.30, 1.29]
1.7.3 RAVLT delayed (higher=better)End of treatment (final value)	3	188	Std. Mean Difference (IV, Random, 95% CI)	-0.78 [-1.74, 0.17]
1.7.4 Cognistat (higher=better) End of treatment (final value)	1	80	Std. Mean Difference (IV, Random, 95% CI)	-0.04 [-0.49, 0.42]
1.7.5 Groebe DfR16 (higher=better)End of treatment (final value)	1	46	Std. Mean Difference (IV, Random, 95% CI)	-0.95 [-1.58, -0.33]
1.7.6 BACS verbal memory, (high = better)End of treatment (final value)	1	100	Std. Mean Difference (IV, Random, 95% CI)	-0.32 [-0.72, 0.08]
1.7.7 WMS LT (higher=better)	1	57	Std. Mean Difference (IV, Random, 95% CI)	0.27 [-0.25, 0.79]
1.7.8 HVLt-R percent retained	1	31	Std. Mean Difference (IV, Random, 95% CI)	-0.37 [-1.11, 0.37]
1.8 Symptoms, end of treatment	6	367	Std. Mean Difference (IV, Random, 95% CI)	-0.12 [-0.32, 0.08]
1.8.1 PANSS (higher=worse) End of treatment	5	306	Std. Mean Difference (IV, Random, 95% CI)	-0.18 [-0.40, 0.04]
1.8.2 BPRS (higher=worse)	1	61	Std. Mean Difference (IV, Random, 95% CI)	0.19 [-0.32, 0.70]
1.9 QoL, end of treatment	4	257	Std. Mean Difference (IV, Random, 95% CI)	-0.85 [-2.03, 0.34]
1.9.1 QOLI (higher=better)	1	31	Std. Mean Difference (IV, Random, 95% CI)	0.14 [-0.60, 0.87]
1.9.2 QLS (higher=better)	2	149	Std. Mean Difference (IV, Random, 95% CI)	-1.62 [-3.45, 0.20]
1.9.3 SQoL (higher=better)	1	77	Std. Mean Difference (IV, Random, 95% CI)	-0.25 [-0.70, 0.20]
1.10 Days at hospital	0	0	Mean Difference (IV, Fixed, 95% CI)	Not estimable

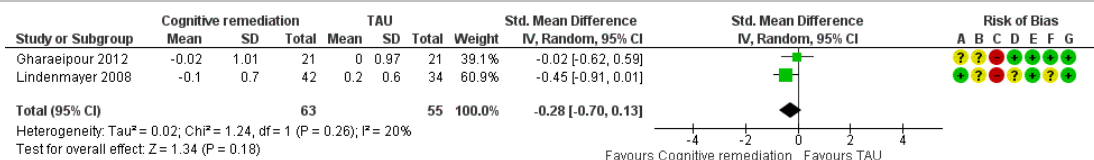
2 TAU vs Cognitive remediation original data

Outcome or Subgroup	Studies	Participants	Statistical Method	Effect Estimate
2.1 Global cognition score (Z score, final score), end of treatment	3	145	Std. Mean Difference (IV, Random, 95% CI)	0.02 [-0.62, 0.66]
2.2 Social functioning, end of treatment	3	263	Std. Mean Difference (IV, Random, 95% CI)	-0.29 [-0.53, -0.04]
2.2.1 SFS (higher=better)	1	57	Std. Mean Difference (IV, Random, 95% CI)	-0.15 [-0.68, 0.37]
2.2.2 SSSI (higher=worse)	1	122	Std. Mean Difference (IV, Random, 95% CI)	-0.31 [-0.66, 0.05]
2.2.3 WHODAS (higher=worse)	1	84	Std. Mean Difference (IV, Random, 95% CI)	-0.35 [-0.79, 0.08]
2.3 Social functioning SFS (higher=better), longest FU	1	47	Mean Difference (IV, Random, 95% CI)	-0.20 [-9.24, 8.84]
2.4 Social function End of treatment	4		Std. Mean Difference (IV, Random, 95% CI)	-0.26 [-0.47, -0.05]
2.4.1 LSP (high=worse) end of treatment	0		Std. Mean Difference (IV, Random, 95% CI)	Not estimable
2.4.2 SBS (higher=worse) End og treatment	1		Std. Mean Difference (IV, Random, 95% CI)	-0.17 [-0.62, 0.27]
2.4.3 SFS (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.15 [-0.67, 0.37]
2.4.4 SSSI (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.31 [-0.67, 0.05]
2.4.5 WHODAS (higher=worse)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.35 [-0.79, 0.09]
2.5 Social functioning, FU	2		Std. Mean Difference (IV, Random, 95% CI)	-0.05 [-0.41, 0.30]
2.5.1 LSP (higher=worse) FU	0		Std. Mean Difference (IV, Random, 95% CI)	Not estimable
2.5.2 SBS (higher=worse) FU	1		Std. Mean Difference (IV, Random, 95% CI)	0.04 [-0.42, 0.49]
2.5.3 SFS (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.20 [-0.78, 0.38]
2.6 Working memory	9	574	Std. Mean Difference (IV, Random, 95% CI)	-0.61 [-0.99, -0.23]
2.6.1 ANS (higher=better) End of treatment (final value)	1	77	Std. Mean Difference (IV, Random, 95% CI)	-0.21 [-0.66, 0.24]
2.6.2 WAIS digit span backward (Higher =better) End of treatment (final value)	1	70	Std. Mean Difference (IV, Random, 95% CI)	-0.47 [-0.95, 0.00]
2.6.3 WAIS III digit span backwards End of treatment (final value)	2	130	Std. Mean Difference (IV, Random, 95% CI)	-1.33 [-2.60, -0.07]
2.6.4 ACT (higher=better) End of treatment (final value)	1	42	Std. Mean Difference (IV, Random, 95% CI)	-1.01 [-1.66, -0.37]
2.6.5 BACS (digit seq.) (higher=better) End of treatment (final value)	1	100	Std. Mean Difference (IV, Random, 95% CI)	-0.33 [-0.72, 0.07]

2.6.6 WAIS III (letter-number seq. subtest) (higher=better)End of treatment (final value)	2	80	Std. Mean Difference (IV, Random, 95% CI)	-0.32 [-0.93, 0.29]
2.6.7 WAIS-R digit span backwards (higher=better) End of treatment (final value)	1	75	Std. Mean Difference (IV, Random, 95% CI)	-0.27 [-0.73, 0.18]
2.7 Working memory, end of treatment	10		Std. Mean Difference (IV, Random, 95% CI)	-0.60 [-0.93, -0.28]
2.7.1 working memory (fra NICE)	2		Std. Mean Difference (IV, Random, 95% CI)	-0.33 [-0.72, 0.06]
2.7.2 ANS (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.21 [-0.66, 0.24]
2.7.3 WAIS digit span backward (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.47 [-0.95, 0.01]
2.7.4 WAIS III digit span backwards (higher=better)	2		Std. Mean Difference (IV, Random, 95% CI)	-1.33 [-2.60, -0.07]
2.7.5 ACT (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-1.00 [-1.65, -0.35]
2.7.6 BACS (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.33 [-0.73, 0.07]
2.7.7 WAIS III letter-number seq. (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.59 [-1.18, 0.00]
2.7.8 WAIS-R digit span backwards (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.27 [-0.73, 0.19]
2.8 Verbal learning and memory, Total	2	97	Std. Mean Difference (IV, Random, 95% CI)	-0.50 [-2.37, 1.37]
2.8.1 HVLT total (higher=better) End of treatment (final value)	1	27	Std. Mean Difference (IV, Random, 95% CI)	0.48 [-0.32, 1.27]
2.8.2 RAVLT total (higher=better) End of treatment (final value)	1	70	Std. Mean Difference (IV, Random, 95% CI)	-1.43 [-1.96, -0.90]
2.9 Verbal learning	5	299	Std. Mean Difference (IV, Random, 95% CI)	-0.27 [-0.63, 0.09]
2.9.1 RAVLT, learning (higher=better)End of treatment (final value)	2	116	Std. Mean Difference (IV, Random, 95% CI)	-0.54 [-1.05, -0.03]
2.9.2 CVLT short term recal (higher=better)End of treatment (final value)	1	49	Std. Mean Difference (IV, Random, 95% CI)	-0.13 [-0.71, 0.45]
2.9.3 WLM (first recall) (higher=better) End of treatment (final value)	1	77	Std. Mean Difference (IV, Random, 95% CI)	-0.44 [-0.90, 0.01]
2.9.4 WMS ST (higher=better)	1	57	Std. Mean Difference (IV, Random, 95% CI)	0.35 [-0.18, 0.87]
2.10 Verbal memory	10	631	Std. Mean Difference (IV, Random, 95% CI)	-0.73 [-1.32, -0.14]
2.10.1 CVLT long term recall (higher =better) End of treatment (final value)	1	49	Std. Mean Difference (IV, Random, 95% CI)	-0.01 [-0.59, 0.57]
2.10.2 HVLT delayed (higher=better) End of treatment (final value)	2	111	Std. Mean Difference (IV, Random, 95% CI)	-0.71 [-3.04, 1.62]
2.10.3 RAVLT delayed (higher=better)End of treatment (final value)	3	188	Std. Mean Difference (IV, Random, 95% CI)	-0.78 [-1.74, 0.17]
2.10.4 Cognistat (higher=better) End of treatment (final value)	1	80	Std. Mean Difference (IV, Random, 95% CI)	-2.52 [-3.13, -1.92]
2.10.5 Groebe DFR16 (higher=better)End of treatment (final value)	1	46	Std. Mean Difference (IV, Random, 95% CI)	-0.95 [-1.58, -0.33]
2.10.6 BACS verbal memory, (high = better)End of treatment (final value)	1	100	Std. Mean Difference (IV, Random, 95% CI)	-0.32 [-0.72, 0.08]
2.10.7 WMS LT (higher=better)	1	57	Std. Mean Difference (IV, Random, 95% CI)	0.27 [-0.25, 0.79]
2.11 Verbal learning and memory (fra NICE)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.33 [-1.01, 0.36]
2.12 Symptoms, End of treatment	3	202	Std. Mean Difference (IV, Random, 95% CI)	-0.02 [-0.37, 0.34]
2.12.1 PANSS (higher=worse)	2	141	Std. Mean Difference (IV, Random, 95% CI)	-0.12 [-0.65, 0.40]
2.12.2 BPRS (higher=worse)	1	61	Std. Mean Difference (IV, Random, 95% CI)	0.19 [-0.31, 0.70]
2.13 Symptoms, end of treatment	6		Std. Mean Difference (IV, Random, 95% CI)	-0.12 [-0.32, 0.08]
2.13.1 PANSS (higher=worse) End of treatment	5		Std. Mean Difference (IV, Random, 95% CI)	-0.18 [-0.40, 0.04]
2.13.2 BPRS (higher=worse)	1		Std. Mean Difference (IV, Random, 95% CI)	0.19 [-0.32, 0.70]
2.14 QoL, end of treatment	3	226	Std. Mean Difference (IV, Random, 95% CI)	-1.16 [-2.58, 0.26]
2.14.1 SQoL (higher=better)	1	77	Std. Mean Difference (IV, Random, 95% CI)	-0.25 [-0.70, 0.20]
2.14.2 QLS (higher=better)	2	149	Std. Mean Difference (IV, Random, 95% CI)	-1.63 [-3.45, 0.20]
2.15 QoL, end of treatment	4		Std. Mean Difference (IV, Random, 95% CI)	-0.85 [-2.03, 0.34]
2.15.1 QOLI (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	0.14 [-0.60, 0.87]
2.15.2 QLS (higher=better)	2		Std. Mean Difference (IV, Random, 95% CI)	-1.62 [-3.45, 0.20]
2.15.3 SQoL (higher=better)	1		Std. Mean Difference (IV, Random, 95% CI)	-0.25 [-0.70, 0.20]
2.16 Days at hospital	0	0	Mean Difference (IV, Fixed, 95% CI)	Not estimable
2.17 New Outcome	0	0	Odds Ratio (M-H, Fixed, 95% CI)	Not estimable
2.18 Verbal learning, change from baseline	1	84	Mean Difference (IV, Random, 95% CI)	-0.47 [-0.58, -0.36]
2.18.1 HVLT (higher=better), end of treatment	1	84	Mean Difference (IV, Random, 95% CI)	-0.47 [-0.58, -0.36]

Figures

Figure 1 (Analysis 1.1)

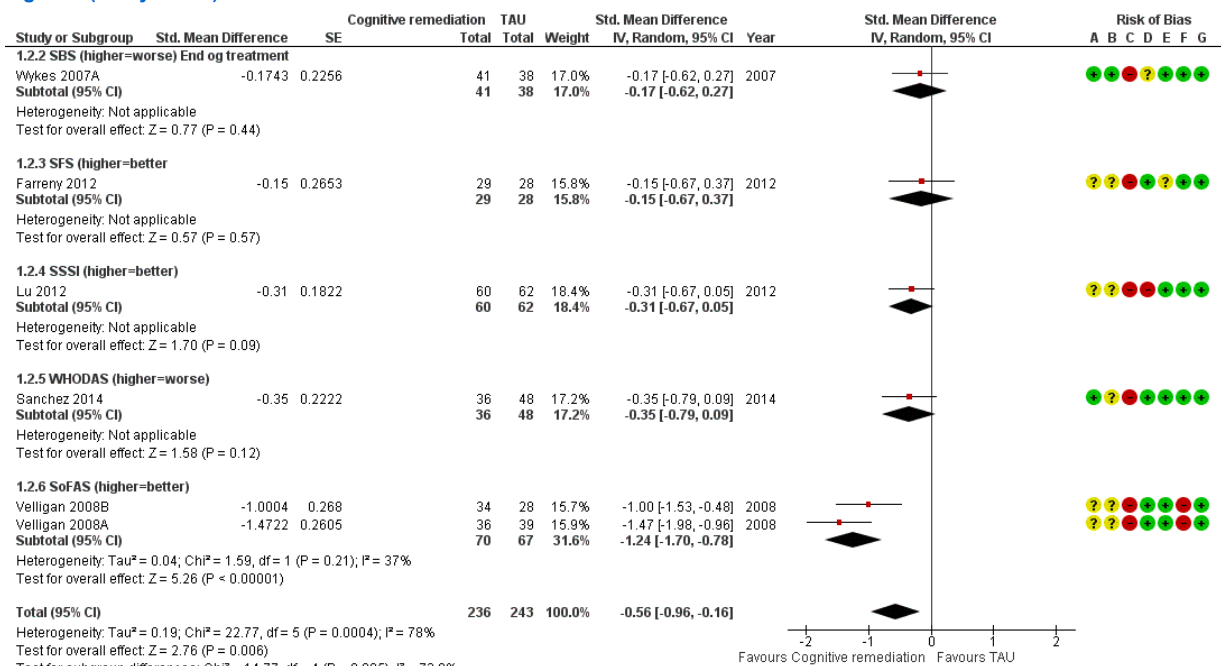


Risk of bias legend

- (A) Random sequence generation (selection bias)
- (B) Allocation concealment (selection bias)
- (C) Blinding of participants and personnel (performance bias)
- (D) Blinding of outcome assessment (detection bias)
- (E) Incomplete outcome data (attrition bias)
- (F) Selective reporting (reporting bias)
- (G) Other bias

Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.1 Global cognition score (Z score, final score), end of treatment.

Figure 2 (Analysis 1.2)

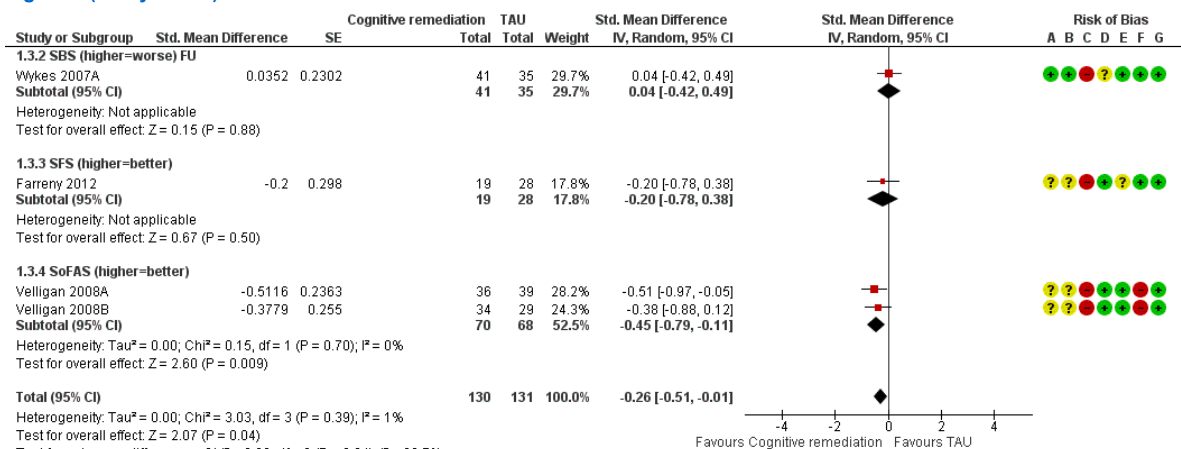


Risk of bias legend

- (A) Random sequence generation (selection bias)
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- (D) Blinding of outcome assessment (detection bias)
- (E) Incomplete outcome data (attrition bias)
- (F) Selective reporting (reporting bias)
- (G) Other bias

Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.2 Social function End of treatment.

Figure 3 (Analysis 1.3)

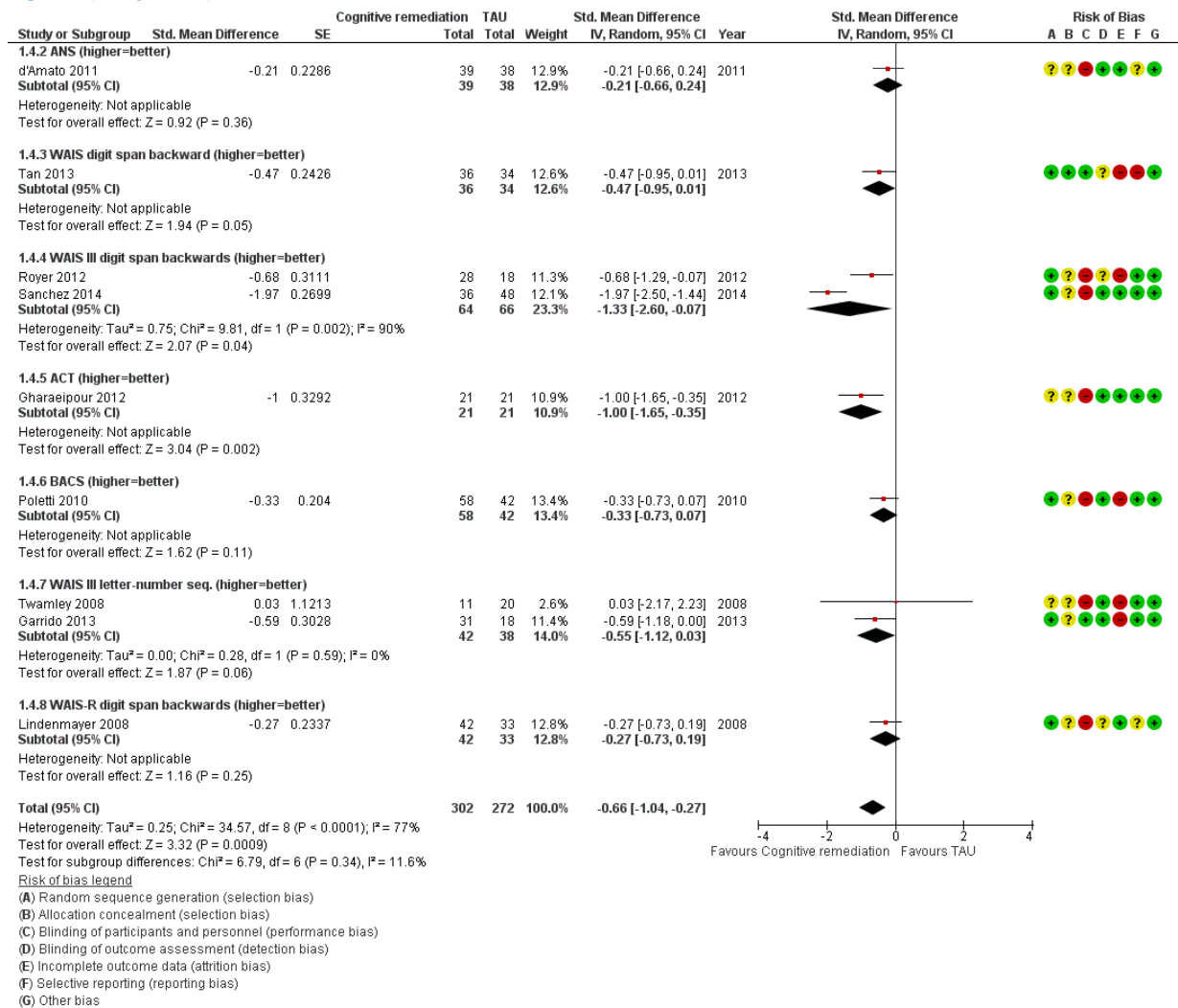


Risk of bias legend

- (A) Random sequence generation (selection bias)
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- (C) Blinding of participants and personnel (performance bias)
- (D) Blinding of outcome assessment (detection bias)
- (E) Incomplete outcome data (attrition bias)
- (F) Selective reporting (reporting bias)
- (G) Other bias

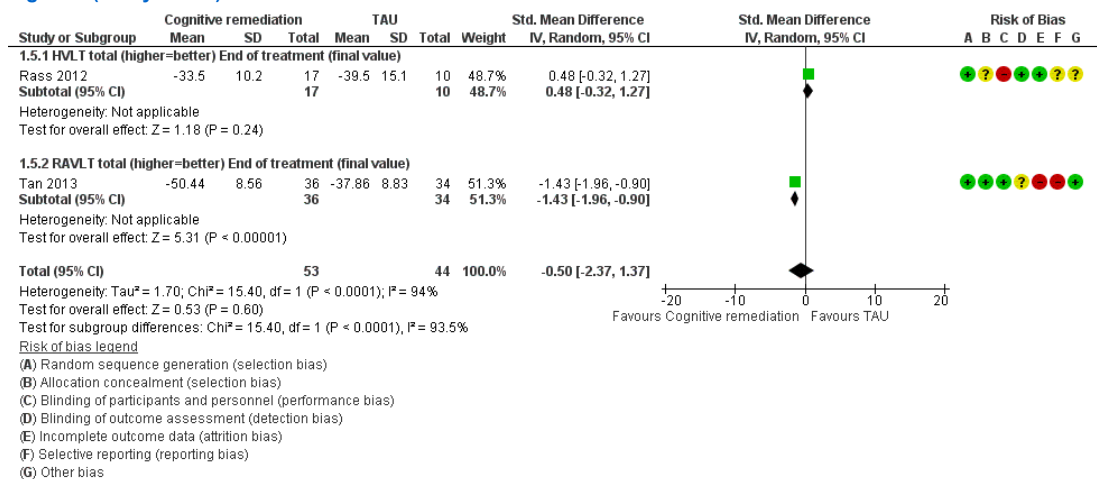
Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.3 Social functioning, FU.

Figure 4 (Analysis 1.4)



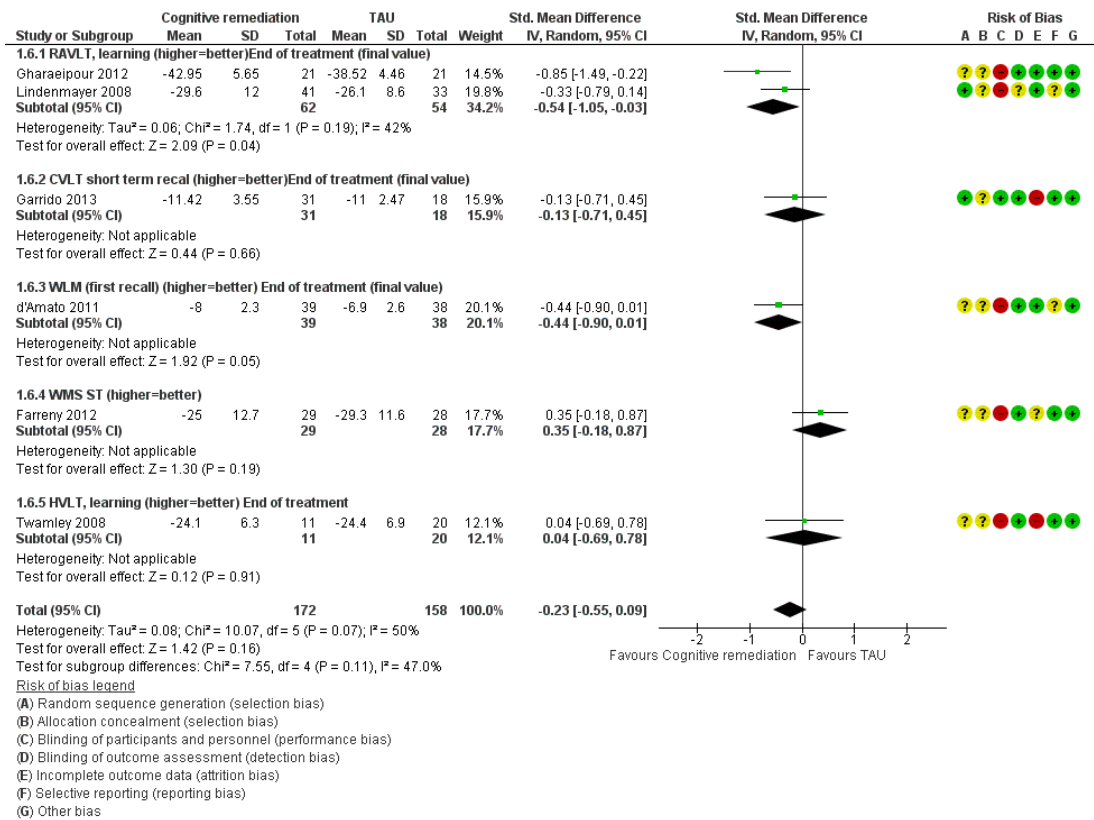
Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.4 Working memory, end of treatment.

Figure 5 (Analysis 1.5)



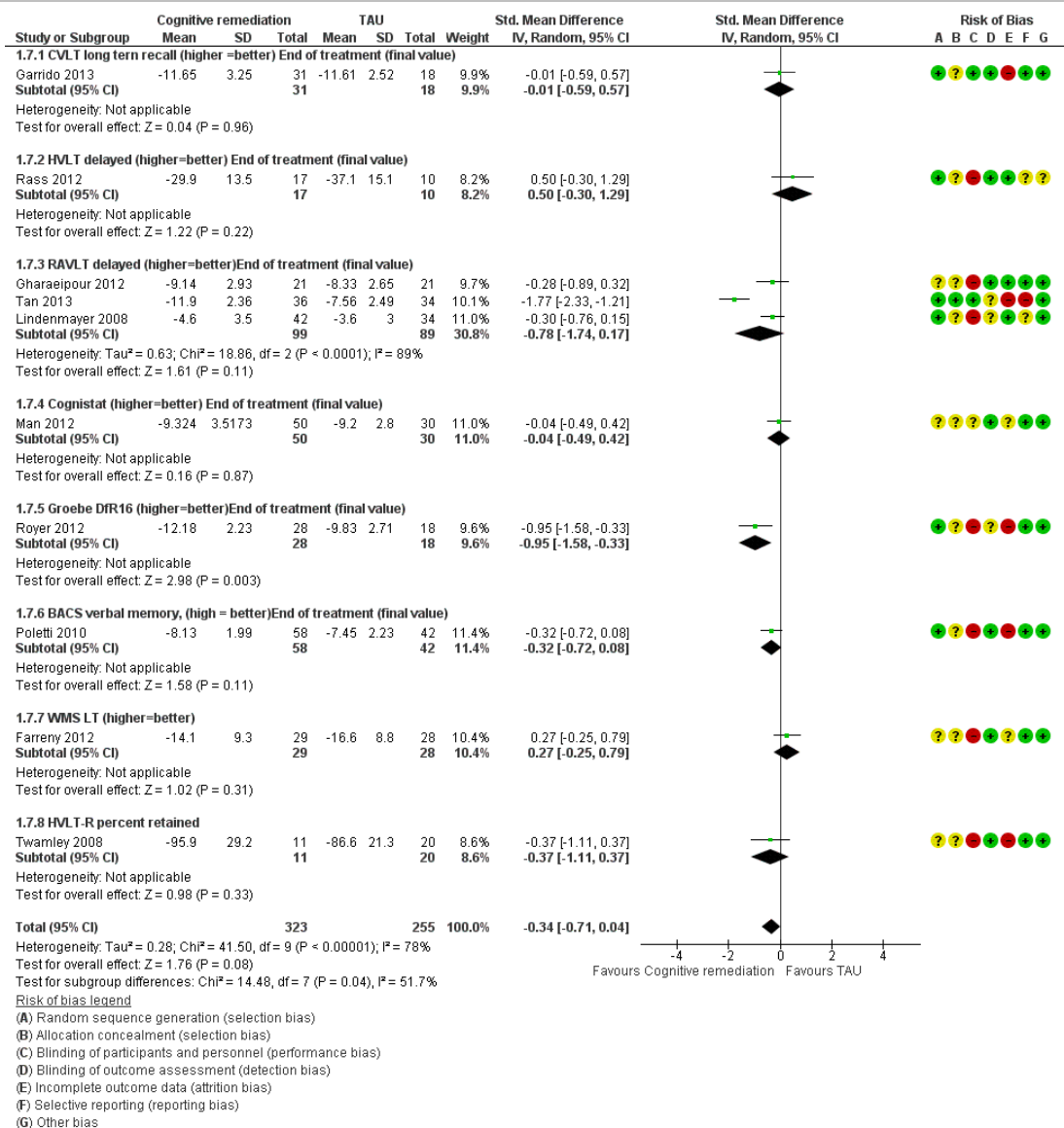
Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.5 Verbal learning and memory, Total.

Figure 6 (Analysis 1.6)



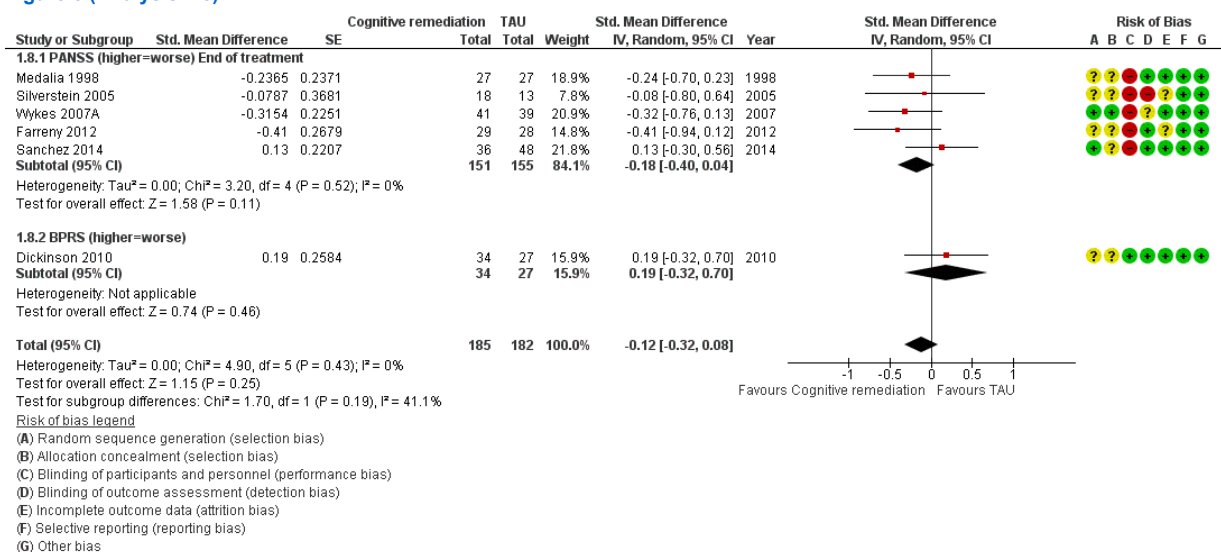
Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.6 Verbal learning.

Figure 7 (Analysis 1.7)



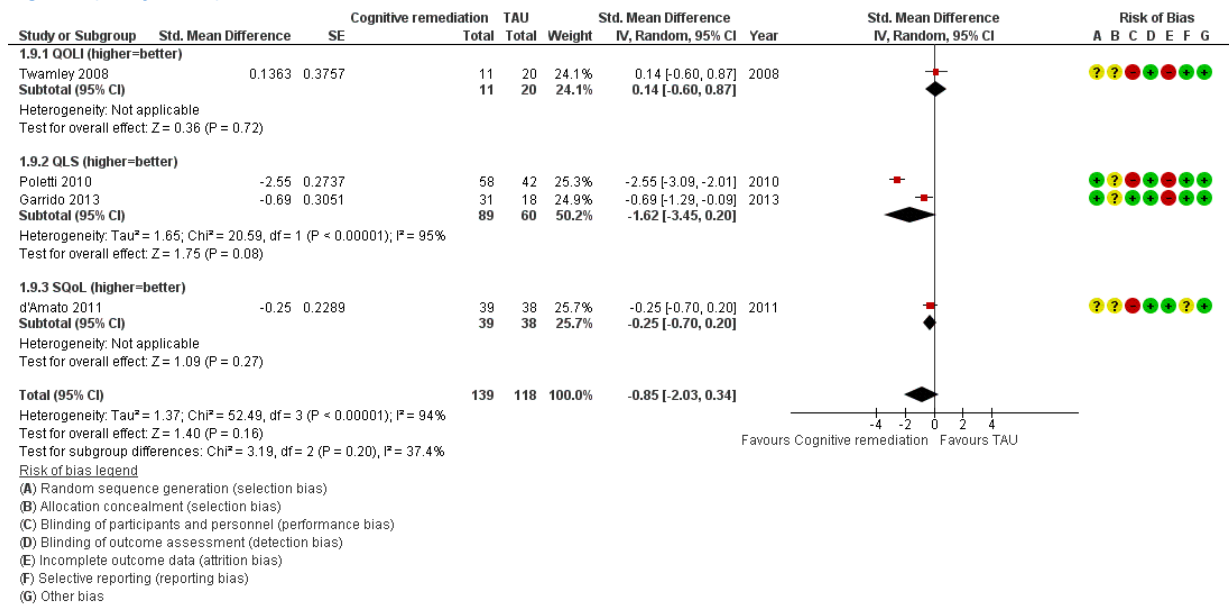
Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.7 Verbal memory.

Figure 8 (Analysis 1.8)



Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.8 Symptoms, end of treatment.

Figure 9 (Analysis 1.9)



Forest plot of comparison: 1 TAU vs Cognitive remediation, outcome: 1.9 QoL, end of treatment.