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Meta-Analysis of Cognitive Rehabilitation Interventions in Veterans and Service Members with Traumatic Brain Injuries

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Abstract

Main objective: Cognitive difficulties are some of the most frequently experienced symptoms following mild-to-moderate traumatic brain injuries (TBIs). There is meta-analytic evidence that cognitive rehabilitation improves cognitive functioning after TBI in non-Veteran populations, but not specifically within the Veteran and Service Member (V/SM) population. The purpose of the current meta-analysis was to examine the effect of cognitive rehabilitation interventions for V/SMs with history of mild-to-moderate TBI.

Design and main measures: This meta-analysis was preregistered with PROSPERO (CRD42021262902) and used the Preferred Reporting Items for Systematic Reviews and Meta-analyses (PRISMA) checklist for reporting guidelines. Inclusion criteria required studies to

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Competing interests:

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The authors have no competing interests to declare.

have (1) randomized controlled trials; (2) used adult participants (age 18 or older) who were US Veterans or active-duty Service Members who had a history of mild-to-moderate TBI; (3) cognitive rehabilitation treatments designed to improve cognition and/or everyday functioning; (4) used objective neuropsychological testing as a primary outcome measure; and (5) been published in English. At least two reviewers independently screened all identified abstracts and full-text articles and coded demographic and effect size data. The final search was run on 2/24/2023 using four databases (PubMed, PsycInfo, Web of Science, and Google Scholar). Study quality and bias were examined using the revised Cochrane risk-of-bias tool for randomized trials.

Results: We identified eight articles meeting full criteria (total participant*s*=564, 97% of whom had a history of mild TBI). Compared to control groups, participants showed a small, but significant, improvement in overall objective neuropsychological functioning after cognitive rehabilitation interventions. Interventions focusing on teaching strategies had a larger effect size than did those focusing on drill-and-practice for both objective neuropsychological test performance as well as performance-based measures of functional capacity.

Conclusion: There is evidence of cognitive improvement in V/SMs with TBI histories after participation in cognitive rehabilitation. Clinician-administered interventions focusing on teaching strategies may yield the greatest cognitive improvement in this population.

Keywords

traumatic brain injury; cognitive training; cognitive remediation

Meta-Analysis of Cognitive Rehabilitation Interventions in Veterans and Service Members

Traumatic brain injuries (TBIs) have been labelled the "signature injury" of post-9/11 Veterans and military Service Members (V/SMs). Between 2000 and 2019, more than 400,000 active-duty SMs sustained a TBI, with the majority (82.8%) categorized as mild injuries and 11% categorized as moderate¹. A random representative sample of post-9/11 Veterans found 17.3% met criteria for TBI acquired during military service². Although most symptoms of mild TBI usually resolve within 90 days in civilian populations³, post-concussive symptoms may persist longer in military populations and often interfere with optimal functioning^{4,5}. While there is more variability in recovery from moderate TBI, between 48–75% of participants with moderate TBI having favorable outcomes on the Glasgow Outcome Scale -Extended and 32% reported no disability^{6,7}.

TBIs can have a significant impact on both individuals and family members/caregivers, resulting in increased impairment in daily activities, depression, anxiety, social isolation, and decreased quality of life⁸. In addition, TBIs are costly to healthcare systems (e.g., for Veterans, presence of TBI confers three times higher healthcare costs²). Common symptoms after mild to moderate TBIs include headaches, changes in mood, and cognitive symptoms⁹. Cognitive dysfunction appears to result in higher healthcare utilization, as individuals with cognitive impairment require three times as many hospitalizations as those without cognitive impairment¹⁰. As such, it is important to know the most efficacious treatments for cognitive

impairments among Veterans and SMs with a history of mild-to-moderate TBI, as well as any moderating factors of treatment response.

There is meta-analytic evidence for successful post-TBI cognitive rehabilitation in the general population, including attentional-based skills training¹¹, memory skills training¹², and problem-solving training¹³ with post-treatment improvements observed on neuropsychological test performance and subjective cognitive symptoms. However, it is not well understood if these post-intervention improvements translate into meaningful changes in everyday functioning or how long they last¹³. It is also not clear if these interventions are efficacious in V/SMs, whose TBIs often occur in the context of psychological trauma and who may have higher rates of comorbidities¹⁴. One previous review of cognitive rehabilitation treatments in V/SM populations found support for cognitive rehabilitation¹⁵. The present study provides an update and expansion to this review by performing a meta-analysis of cognitive rehabilitation for V/SMs with a history of mild-to-moderate TBI and incorporating analyses of study quality. We examined changes in performance on both neuropsychological tests and functional measures. Additionally, we built on previous research by examining effects on everyday functioning, and when possible, durability of the treatment effects. Finally, we examined moderating factors (e.g., type of treatment, treatment length, age) through subgroup analyses and meta-regression.

Methods

This meta-analysis was preregistered with PROSPERO (CRD42021262902) and used the Preferred Reporting Items for Systematic Reviews and Meta-analyses (PRISMA) checklist for reporting guidelines¹⁶ (see Supplemental Digital Content for PRISMA checklist). All deviations from the preregistration are explicitly noted in the Supplemental Digital Content.

We developed sets of keywords related to the following elements: (A) cognitive rehabilitation and other behavioral/neuropsychological interventions; (B) traumatic brain injury/acquired brain injury; and (C) Veteran/military populations. Preliminary searches were conducted in several databases to gauge the precision of the search, scan article metadata for additional relevant keywords, and refine final inclusion and exclusion criteria. The final Boolean search string was: (("Cognitive training" or "Cognitive strategy training" or "Cognitive skills training" or "Cognitive rehabilitation" or "Cognitive remediation" or "Cognitive intervention" or "Memory training" or "Attention training" or "Executive function training" or "Problem solving therapy" or "Neurorehabilitation" or "Deal injury" or "concussion" or "postconcussive syndrome" or "post-concussive syndrome")) AND (Veteran or military or army or navy or "air force" or "marine corps" or "service member" or "active duty").

The final inclusion criteria required studies to have (1) randomized controlled trials; (2) used adult participants (age 18 or older) who were US Veterans or active-duty Service Members who had a history of mild-to-moderate TBI; (3) cognitive rehabilitation treatments designed to improve cognition and/or everyday functioning; (4) used objective neuropsychological testing as a primary outcome measure; and (5) been published in English. Samples of

mixed populations (e.g., including civilians, mixed etiologies of cognitive impairment) were used if TBI or Veteran/Service Member groups were reported separately. Due to the small number of studies in this area, we included three studies of samples with mixed mild and moderate TBI severity; in these studies, a minority of participants had moderate TBI (25%). Exclusion criteria included (1) severe TBI, (2) non-military population, and (3) trials using only self-report outcomes. We excluded studies of severe TBI due to differences in mechanisms, symptom trajectory, prognosis, and treatment needs, resulting in minimal overlap in the rehabilitation literature^{17–21}.

Data screening, extraction, and coding

At least two reviewers (T.A., S.P., C.H.) independently screened all identified abstracts and full-text articles, with discrepancies resolved through discussion. At least two reviewers independently extracted demographic and effect size data from articles meeting full criteria (T.A., B.E., C.H.). We extracted all available summary effect size data; when multiple effect sizes were reported, we preferentially used raw mean values and standard deviations. When available, we preferentially used or calculated pre-post change scores for both groups, rather than using only post-intervention scores. When studies reported insufficient information to determine study eligibility or calculate an effect size, we contacted authors for information. Studies were reviewed for possible overlapping samples; we used the study with the largest sample size or most comprehensive reporting of neuropsychological outcome measures.

Additionally, we coded studies for the following information: type of neuropsychological test domain (e.g., attention, memory), measures of functional capacity and self-reported everyday functioning, and whether the intervention focused on teaching strategies (strategy-based interventions) or drill-and-practice approaches to cognitive rehabilitation. We coded each neuropsychological test as described in Strauss et al²² (see Table 1 for included neuropsychological tests). Strategy-based interventions were defined as interventions teaching strategy use with the goal of improving daily functioning even in the absence of improvement in cognitive functioning. Drill-and-practice interventions were defined as interventions with the goal of strengthening or restoring the impaired skills to improve cognitive functioning through the use of repeated drills or cognitive excercises^{23,24} (see Supplemental Digital Content for full definitions).

Statistical analysis

We entered raw or standardized scores for all groups in a study (e.g., intervention group vs. control group) directly into Comprehensive Meta-Analysis 2.0 (CMA) using Hedges and Olkin's random-effects model to calculate the overall effect size for both the primary and subgroup analyses. For studies with multiple outcomes, a meta-analysis for the results of the individual study was conducted to give one effect size based on recommendations provided by Borenstein et al.²⁵. We considered the mean effect sizes as significant if p<0.05 or if the 95% confidence interval (CI) did not include zero; if discrepancies occurred, we used the 95% CI. All effect sizes were transformed into Cohen's *d* for the analyses, with the classification of small (*d*=0.2), medium (*d*=0.5), and large (*d*=0.8) effects based on Cohen's recommendations²⁶. Moderator analyses (meta-regression) were used if there are at least ten

studies per moderator category²⁵. Exploratory moderator analyses were conducted if there were at least eight studies without statistically significant heterogeneity between studies²⁷.

To estimate how unpublished null results could lower the effect sizes, we used Rosenthal's failsafe N-analysis, which estimates how many missing studies with statistically insignificant results are needed to reduce the statistical significance to nonsignificant in the meta-analysis²⁵. Additionally, we report a power analysis to determine whether there were enough studies to power both the primary and subgroup analyses used in the metaanalysis²⁸. We also visually inspected funnel plots and performed a trim and fill analysis for outlier studies among both the primary analysis as well as the subgroup analyses. We assessed heterogeneity using estimates of Q, τ^2 , and I². Three studies included a minority of moderate TBI severity participants (6%, 21%, and 25%)^{29,30,36}. One of the three studies included mild, moderate, and severe TBI, but the primary author of this study provided data with only the mild and moderate severity participants³⁰.

Results:

Search and sample characteristics

The final searches were run on 9/8/2022 and 2/24/2023 using keywords related to cognitive rehabilitation interventions, traumatic brain injuries, and Veteran or military populations (see Supplemental Digital Content for full search structure, databases used, and preliminary search methods), with reference treeing (i.e., searching articles pulled for full text screening and examining their references and cited by lists) completed on 2/24/2023. We screened the titles and abstracts of 636 unique articles (see Figure 1). After initial screening, we examined 88 articles.

Following full-text screening, we identified 8 articles meeting full criteria (total participants=564; intervention=303, control=261; see Table 2 for included studies and descriptions)²⁹⁻³⁶. All included articles were peer reviewed (i.e., no preprints or unpublished works met all inclusion criteria). The sample size ranged from 17-119 (median n=40.5). The average age of study participants was 36.7 (SD=6.8; see Table 3 for demographic information for all included studies); intervention and control participants did not differ in age. Average education did not differ between intervention and control participants. On average, participants had 14.2 years of education (SD=1.2). Between 81–100% (mean 88.3%, SD 11.4) of participants were male. Limited racial and ethnic information were reported by the majority of studies, limiting available information for the meta-analytic sample. Using data collected by more than one study, on average 65.4% (SD=9.7) of participants identified as White, 15.5% (SD=2.5) African American, 7% (SD=5.0) Other, and 21.5% (SD=12.5) reported Hispanic ethnicity. The average length of time since TBI was 6 years (M=71.8 months; SD=52.0 months, range 5-189 months). There was limited information on pre-intervention cognitive treatments, which were only reported in two studies. One study reported 14% of the sample had previous cognitive rehabilitation treatment, and one study reported that 24% of the sample had previous TBI rehabilitation treatment and 12% were currently in TBI rehabilitation treatment (see Table 2).

Intervention lengths ranged from 4–15 weeks (M=9.5; SD=3.7). Four studies used a strategy-based approach, three studies used a drill-and-practice approach, and one study had three intervention conditions (one drill-and-practice, which we included with the other drill-and-practice interventions, and two combinations of strategy-based and drill-and-practice approaches, which we considered separately). Sufficient information was provided to use pre-post change scores for seven studies. We ran a post-hoc sensitivity analysis comparing results with and without the study with only post-intervention scores, which revealed minimal differences (see Supplemental Digital Content for sensitivity analysis results). Sensitivity analysis did not reveal a difference when including these studies compared to using only samples with mild injury severity (see Supplemental Digital Content). Agreement between coders was greater than 96% and κ =0.85 for all aspects of the screening and coding process; there was 100% agreement for full article inclusion and article coding after discussion.

Risk of Bias assessment

Two reviewers (T.A., C.H.) independently coded for study quality using the revised Cochrane Risk-of-Bias Tool for Randomized Trials, second edition³⁷ (RoB 2), as well as additional indicators of study quality (see Supplemental Digital Content).

Results of the Risk of Bias

Overall, there was low concern for study bias (see Supplemental Digital Content). Two studies had baseline differences between the intervention and control group, probably due to small sample sizes.

Overall analysis

Compared to control groups, participants showed a small, but statistically significant, improvement in objective neuropsychological functioning after cognitive rehabilitation interventions (k=8, d=0.22, [95% CI (0.01, 0.43)], p=0.04; see Figure 2), and small, but not statistically significant, effect on performance-based measure of functional capacity (k=4, d=0.16, [95% CI (-0.48, 0.81)], p=0.62). There was no evidence of significant heterogeneity between studies for the primary analysis (Q(7)=8.14; p=0.32; $f^2 = 14.03$). We found no evidence of publication bias (see Supplemental Digital Content), though only one additional study with null findings would be needed for the improvement in objective neuropsychological testing to no longer be significant.

Cognitive domains

There were significant effects on memory (k=6, d=0.42, [95% CI (0.13, 0.70)], p=0.01) and executive functioning (k=6, d=0.26, [95% CI (0.01, 0.51)], p=0.04), but not on attention (k=7, d=0.12, [95% CI (-0.12, 0.35)], p=0.33; see Figure 2). Data in other domains (language, visuospatial) were not sufficient to examine due to being included in only one study.

Strategy-based interventions

Studies focusing on teaching strategies had a small, statistically significant effect on objective neuropsychological performance (k=4, d=0.37, [95% CI (0.08, 0.67)], p=0.01), and a moderate-to-large effect on performance-based measures of functional capacity (k=2, d=0.72, [95% CI (0.03, 1.07)], p<0.01). There was no evidence of significant heterogeneity (Q(3)=2.38, p=0.50, $l^2<0.01$). We found no evidence of publication bias (see Supplemental Digital Content). Three missing null studies would be needed for the statistically significant finding of improvement in objective neuropsychological performance to no longer be significant.

Drill-and-practice interventions

Studies using a drill-and-practice approach had a negligible effect on objective neuropsychological test performance that was not statistically significant (k=4, d=0.10, [95% CI (-0.26, 0.46)], p=0.59). Small (non-significant) improvements on measures of functional capacity favored the control groups (k=2, d=-0.45, [95% CI (-1.39, 0.44)], p=0.32). There was no evidence of significant heterogeneity (Q(3)=3.95, p=0.27, $\hat{P}<0.01$). We found no evidence of publication bias (see Supplemental Digital Content).

Mixed interventions

One study used three different treatment groups, with two using both drill-and-practice and strategy-based elements. There was a small, not statistically significant effect of these mixed interventions on neuropsychological performance (k=1 but with two groups with different treatments, d=0.30, [95% CI (-0.38, 0.45)], p=0.88).

Types of control group

Four studies used active control groups and four studies used non-active control groups (wait list control, treatment as usual/usual care). There was greater improvement in interventions using active conditions as control groups (d=0.37, [95% CI (0.08, 0.67)], p=0.01) than studies using non-active control groups (not statistically significant; d=0.11, [95% CI (-0.19, 0.40)], p=0.48).

Exploratory meta-regression

We used meta-regression to examine the relationship between neuropsychological outcomes and participant demographic factors (age, education, time since TBI, and presence of comorbid PTSD) as covariates in four independent models with neuropsychological test performance change scores as the outcome. There was no significant effect for percentage of the sample with comorbid PTSD (*b*=–0.004, [95% CI (–0.01, 0.01)], SE(0.01), *p*=0.42); age (*b*=0.01, [95% CI (–0.04, 0.07)], SE(0.03), *p*=0.64); education (*b*=0.03, [95% CI (–0.39, 0.46)], SE(0.22), *p*=0.88); or time since TBI (*b*=–0.002, [95% CI (–0.01, 0.01)], SE(0.01), *p*=0.59). There was also no relationship between length of the intervention and neuropsychological test performance (*b*=0.01, [95% CI (–0.06, 0.88)], SE(0.04), *p*=0.70).

Durability of treatment effects

Four studies included post-intervention follow-up visits to measure durability of treatment effects, with three studies repeating objective measures after a twelve week no-contact/no training period. When limiting analysis to the studies with sufficient data, treatment effects on overall neuropsychological test performance at 10- or 12-week follow-up (d=0.45, [95% CI (0.01, 0.90)], p=0.04) were similar to treatment effects immediately post-treatment ((d=0.40, [95% CI (0.33, 0.77)], p=0.03).

Discussion

TBIs are a prevalent concern for V/SM populations and there is a need to identify efficacious treatments. The present meta-analysis examined the effects of cognitive rehabilitation in Veterans with history of mild-to-moderate TBI. Compared to control participants, we found evidence of small effect size improvements for cognitive rehabilitation on objective neuropsychological performance, with small effect size improvements on memory and executive functioning tests, but no significant change in attention performance. Interventions using strategy-based approaches yielded larger effects than drill-and-practice interventions. We found the effect of the active intervention was larger in studies using active control groups. This finding was unexpected, as more robust control conditions are typically associated with lower effect sizes³⁸. One possible reason may be inconsistency of participant blinding of active control conditions, due to difficulty in providing an active control condition that is not easily identifiable to the study participants as the control condition. Another reason is that many of the inactive control conditions, particularly treatment as usual conditions, consisted of a high level of clinical contact and specialty appointments. There was no effect of length of the intervention on neuropsychological test performance, nor did individual factors (age, education, time since TBI, presence of comorbid PTSD) moderate outcomes, although this finding may be due to minimal statistical power and limited variability in the studies. In the studies that included followup assessments, participants maintained treatment gains in global neuropsychological performance three months post-intervention. Although this finding merits replication, these studies provide preliminary evidence of sustained benefit of the interventions on objective neuropsychological test performance. It should be noted that 97% of the participants included in this meta-analysis had a history of mild TBI, so these findings may not generalize to individuals with more severe TBIs.

Findings from this analysis are comparable with a recent meta-analysis of cognitive rehabilitation in non-Veteran populations, which found a small treatment effect (d = 0.30) for cognitive rehabilitation treatments for acquired brain injuries (e.g., TBI and stroke), with a smaller and statistically nonsignificant effect for studies only examining participants with TBI³⁹. The larger effect size seen in our analysis is somewhat surprising, as many pharmacological and psychotherapeutic trials find lower treatment gains in Veteran populations compared to civilian populations⁴⁰. However, Veterans receive their care in a very different healthcare system, and their injuries may have been more likely to be witnessed, resulting in earlier specialized care and rehabilitation.

Subgroup analyses found strategy-based treatments to have a small, statistically significant effect on objective neuropsychological performance (d = 0.37), and a large, statistically significant effect on performance-based measures of functional capacity (d = 0.72). There were no significant effects for either objective neuropsychological test performance, or performance-based measures of functional capacity for drill-and-practice interventions. These findings are important, as The Institute of Medicine's report on TBI encourages interventions to focus on functional outcomes as many decontextualized treatments do not translate into increased daily functioning⁴¹.

These findings are also consistent with the best practice guidelines recommended by Cicerone et al.⁴², based on the evaluation of 491 studies of cognitive rehabilitation after TBI or stroke, as well as the 2023 INCOG 2.0 guidelines for cognitive rehabilitation treatments following brain injuries⁴³. In the most recent edition of Cicerone's living review/practice guidelines, drill-and-practice, computer-assisted programs are reported to have emerging efficacy, but current practice guidelines state these programs should be managed by a rehabilitation clinician, rather than solely computer-delivered⁴². While the INCOG 2.0 guidelines include both drill-and-practice treatments and strategy training, the guidelines recommend that drill-and-practice treatments should focus on real-world activities⁴³. They recommend teaching internal compensatory strategies for mild-to-moderate memory deficits, training in external compensatory strategies for more severe impairment⁴⁴, and metacognitive strategy use for mild-to-moderate attention deficits⁴⁵. Computer-based training without a therapist was not recommended. As there are benefits to both drill-andpractice and computer-based programs (e.g., greater flexibility in adapting the program or having the program adjust to participant abilities, easily scalable, reduced costs due to in-home and self-administered treatments), their recommendation of the use of drilland-practice treatments that focus on real world activities, strategy development and use, facilitated by a TBI-experienced clinician, may increase the efficacy of these programs, particularly for functional capacity in Veterans. Further research can explore the benefit of interventions utilizing both strategy training and drill-and-practice on cognitive domains.

There are several strengths to the current analysis. First, while there was a wide range of types of interventions, intervention lengths, and varying amounts of comorbid mental health concerns, there was a low amount of systematic heterogeneity between studies. As such, we believe there can be greater confidence in the findings of this study. Second, all studies used normed neuropsychological tests, and the age and demographic factors of the study participants included in this study are well matched to the normative samples of these tests.

Limitations

There were also limitations in both the primary studies included in the analysis, as well as in our statistical analyses. The studies included in this analysis used different neuropsychological batteries, with few studies measuring multiple domains, thus precluding further analysis at the domain level. Additionally, mild TBIs are frequently comorbid with other mental health concerns, including PTSD, depression, and anxiety. These conditions were inconsistently measured and described in the primary studies. There was limited information on the previous treatment experiences of participants, with only two studies

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providing information on previous cognitive rehabilitation, and only one study describing previous or concurrent mood treatment. Future meta-analyses will benefit from primary studies providing details on their sample's previous treatment history.

There are also some limitations in our analysis due to lower power of meta-regressions, as well as the restricted age and education range in the primary studies. Although the recommended number of studies sufficient for meta-regressions typically vary between 10–25, there is some evidence that eight studies may provide sufficient information in the absence of significant heterogeneity²⁷. It is possible we were unable to detect whether age or education moderated treatment response due to limited range of these variables in the primary studies. Additionally, we likely were underpowered to detect an effect with only eight studies. Future meta-analyses with additional studies and greater between-study variability will be able to evaluate the moderating effects of these treatments. The average age of participants in the included studies (35.6 years) was also lower than the average age of Veterans reporting TBI (49.9 years)^{46,47}. As such, our findings may not apply to older Veterans. However, there are also advantages to our restricted age range, in that there is a low possibility of cognitive impairments due to age-related decline or dementia rather than secondary to TBI.

Based on the results of this meta-analysis, we conclude that clinician-administered cognitive rehabilitation interventions with a focus on teaching strategies produce greatest cognitive improvement in V/SMs with a history of mild-to-moderate TBI. As many of these treatments are transdiagnostic and symptom-based, rather than etiology specific, further research will benefit from examining the effect of cognitive rehabilitation treatments in Veterans with non-TBI causes of cognitive impairment. As other types of treatments are studied, such as neuromodulation or psychopharmacology, next steps will include comparison of these treatments as monotherapy and combination therapy.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Data availability statement:

The data that supports the findings of this study are available in the supplemental digit content of this article and available upon reasonable request from the first author.

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*This number was marked ineligible by automation tools and then each record was manually reviewed by the first author to check the record was ineligible.



Figure 2.

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Table 1.

Neuropsychological test domains

Domain	Tests coded in meta-analysis	N (%) of studies
Attention	Integrated Visual and Auditory Continual Performance Test, Paced Auditory Serial Addition Test, Trail Making Test-A, WAIS-III Auditory Consonant Trigrams *, WAIS-III Letter Number Sequencing *, WAIS-IV Digit Span, WAIS-IV Letter Number Sequencing, WAIS-IV Symbol Search	8 (100%)
Memory	Brief Visual Memory Test-Revised [*] , California Verbal Learning Test-2 nd edition, Hopkins Verbal Learning Test-Revised, Memory for Intentions Screening Test, RBANS Immediate and Delayed Memory, Rey Auditory Verbal Learning Test [*] , Ruff Light Trails Test	7 (88%)
Executive Functions	Controlled Oral Word Association Test/FAS, D-KEFS Color-Word Inhibition, D-FEKS Trail Making, EXAMINER: Flanker and Set-shifting tests *, Verbal Fluency/FAS, Trail Making Test-B, Wisconsin Card Sorting Test, 64 card	7 (88%)
Language	RBANS Language	1 (13%)
Visual Perception	RBANS Visuospatial	1 (13%)
Performance-based functional capacity	Goal Processing Scale, performance subtests, Timed Instrumental Activities of Daily Living, UCSD Performance-Based Skills Assessment, Brief Version, Virtual Reality Tactical Driving Quotient, Virtual Reality Operational Driving Quotient	4 (50%)
*		

Used in a composite and subtest scores are unavailable.

** Functional based capacity examples were not provided by Strauss et al., and instead based on recent literature reviews of the topic. Note: WAIS-III = Wechsler Adult Intelligence Scale-3rd edition, WAIS-IV = Wechsler Adult Intelligence Scale-4th edition, RBANS =Repeatable Battery for the Assessment of Neurological Status, D-KEFS = Delis-Keplin Executive Function System.

Cohen's d (95% CI)	<i>d</i> = -0.07 (-0.78, 0.44)	d = -0.03 (-0.62, 0.56)	<i>d</i> =0.10 (-0.49, 0.69)	$\begin{array}{l} d=-1.63 \ (-2.77, \ -0.49) \\ d=-2.56 \ (-3.89, \ -1.23) \\ d=-0.69 \ (-1.71, \ 0.32) \\ d=0.81 \ (-0.22, \ 1.84) \\ d=0.81 \ (-0.22, \ 1.84) \\ d=0.92, \ 1.06) \\ d=0.91 \ (-0.92, \ 1.06) \\ d=0.13 \ (-1.12, \ 0.86) \\ d=0.13 \ (-1.12, \ 0.81) \\ d=-0.18 \ (-1.17, \ 0.81) \\ d=-0.19 \ (-1.39, \ 1.01) \\ d=-0.19 \ (-1.39, \ 1.01) \\ d=-0.36 \ (-0.82, \ -0.31) \end{array}$	$ \begin{aligned} d &= 0.33 \ (-0.07, \ 0.72) \\ d &= 0.19 \ (-0.21, \ 0.58 \\ d &= 0.03 \ (-0.21, \ 0.58 \\ d &= 0.13 \ (-0.12, \ 0.67) \\ d &= 0.13 \ (-0.25, \ 0.53) \\ d &= 0.13 \ (-0.25, \ 0.52) \\ d &= 0.13 \ (-0.27, \ 0.52) \\ d &= 0.73 \ (0.09, \ 1.12) \end{aligned} $
Outcomes included in meta-analyses	PASAT	PASAT	PASAT	TMT-A WAIS-IV Coding WAIS-IV DS WAIS-IV DS CVLT-II Trials 1–5 CVLT-II LDFR CVLT-II LDFR CVLT-II LDFR CVWAT Letters COWAT Letters COWAT Animals TMT-B VR TDQ VR ODQ	WAIS-IV DS WAIS-IV PSI CVLT-II Trials 1–5 CVLT-II LDFR CVLT-II LDFR D-KEFS C-WI D-KEFS TM Switching WCST-64 Total Errors
Strategy-based or Drill-and- practice	Drill-and- practice	Mixed (clinician administered)	Mixed (clinician administered)	Drill-and- practice	Strategy-based (clinician administered)
Length/intensity of treatment	10 hours of in- clinic, computerized CR treatment per week throughout the 6-week treatment trial	5 hours of individual therapy sessions, 2 hours group therapy sessions, 3 hours of weekly computer-based weekly computer-based total) per week for 6 weeks	4 hours of individual therapy sessions, 3 hours group therapy sessions, 3 hours of CBT, mindfulness and computer-based training (10 hours total) per week for 6 weeks	Six 90-minute sessions (9 hours total) conducted over a four-week period	60–75 minute individually sessions delivered weekly for 12 weeks
Treatment Description	A: Commercially available brain fitness or brain training computer games targeting attentional processes and general cognitive activation	B: Individual and group therapy (cognitive rehabilitation) and 3 hours of weekly computer-based training targeting attentional processes and general cognitive activation	C: Individual and group therapy (cognitive rehabilitation, mindfulness, and CBT) and 3 hours of weekly CBT, mindfulness, and computer-based training targeting attentional processes and general cognitive activation	Virtual driving practice, including component cognitive skills pertinent to driving (e.g., dual processing, response inhibition, working memory), and performing memory), and performing composite driving skills (e.g., road hazards, yielding)	All standard components and structure of CPT with elements from CogSMART including compensatory cognitive strategies for attention (e.g., breaks, self-talk, environmental adjustments), memory (e.g., prospective memory (e.g., prospective memory (e.g., calendar use, routines, linking tasks, automatic places), and
Sample characteristics	Intervention Sample size: A: 30 B: 30 C: 32 Control Sample size: 34 Total Sample size: 126 Average Age: 31.3 Education: 35% >12 of education	% mate: 100% Hacechtnicity:77% White, 23% other, 29% Haspanic ethnicity TBI severity: 100% mild TBI* % previous treatment: Not provided		Intervention Sample size: 11 Control Sample size: 6 Total Sample size: 17 Average Age: 51.7 Education: 16.9 years % male: 65% % Mhite, 18% African American, 5% Latino, 12% Other TBI severity: 75% mild, 25% mod Previous treatment: Not provided	Intervention Sample size: 51 Control Sample size: 49 Total Sample size: 100 Average Age: 34.4 Education: 13.7 years % male: 98% Race/ethnicity: 47% White, 13% African American, 9% Asian, 23% Hispanic/Latino, 4% Native Hawaiian/Pacific Islander, 2% Native American/Native Alaskan S% Other
Study	Cooper et al., 2017			Ettenhofer et al., 2019**	Jak et al., 2019

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Table 2.

Demographic and treatment information for All studies included in the meta-analysis

	Sample characteristics TBI severity: 94% mild, 6% mod.	Treatment Description executive functioning (e.g.,	Length/intensity of treatment	Strategy-based or Drill-and- practice	Outcomes included in meta-analyses	Cohen's <i>d</i> (95% CI)
	Preventy. 74% mut, 0% mor. Previous treatment: 14% had prior cognitive rehabilitation	executive turbuouning (v.g., goal setting, problem solving)				
et	Intervention Sample size: 41 Control Sample size: 42 Average Age: 33.8 Average Age: 33.8 Education: 14.4 years % male: 81% % male: 81% TBI severity: 100% white TBI severity: 100% mild* Previous treatment: 24% had previous TBI treatment, 12% were currently in TBI treatment	Brain plasticity-based computerized cognitive training (BrainHQ) program targeting speed/ accuracy of information processing	5 hours of self-delivered training per week for 13 weeks	Drill-and- practice	Cognitive Composite TIADL	d = 0.43 (-0.07, 9.2) $d = -0.04 (-0.53, 0.45)$
t al.,	Intervention Sample size: 21 Control Sample size: 24 Average Age: 32.9 Education: 12.9 years % male: 100% Race/ethnicity: Not provided TBI severity: 100% mild* Previous treatment: Not provided	Interactive metronome therapy: Patient executes various repeated movements in time to a beat, while a computer provides precision feedback on performance targeting planning, sequencing, and processing information	15 hours over approximately 7 weeks (length ranged from 5–17 weeks)	Drill-and- practice	IVA-CPT RBANS Attention WAIS-IV Coding WAIS-IV DS WAIS-IV LNS WAIS-IV LNS WAIS-IV LNS WAIS-IV LNS RBANS Imm. Memory RBANS Delayed Memory D-KEFS TM D-KEFS TM RBANS Language RBANS Visuospatial	$\begin{array}{c} d=0.03 \ (-0.55, 0.6) \\ d=0.60 \ (0.20, 1.2) \\ d=0.46 \ (-0.13, 1.05) \\ d=0.19 \ (-0.22, 0.60) \\ d=0.13 \ (-0.25, 0.83) \\ d=0.31 \ (-0.28, 0.83) \\ d=0.31 \ (-0.28, 0.83) \\ d=0.17 \ (0.06, 1.25) \\ d=0.72 \ (0.06, 1.25) \\ d=0.17 \ (0.05, 0.25) \\ d=0.17 \ (0.05, 0.25) \\ d=0.06 \ (-0.22, 0.64) \\ d=0.06 \ (-0.52, 0.64) \\ \end{array}$
ic et	Intervention Sample size: 21 Control Sample size: 19 Total Sample size: 40 Average Age: 45.3 Aucation: 14.4 years % male: 88% male: 88% Race/ethnicity: 67% White TBI severity: 100% mild* Previous treatment: Not provided	Cognitive rehabilitation training that targets executive control functions and metacognitive goal management strategies.	2 hours of group training per week, 3 hours of individual sessions (total) over 5 weeks	Strategy-based (clinician administered)	WM composite Sus. att. composite Recall composite Delayed composite Inhibition composite MF composite GPS Overall Performance Score.	$ \begin{array}{l} d=0.48 \ (0.19, \ 0.63) \\ d=0.77 \ (0.17, \ 1.4) \\ d=0.14 \ (-0.44, \ 0.72) \\ d=0.14 \ (-0.4, \ 0.75) \\ d=0.06 \ (-0.92, \ 0.67) \\ d=0.09 \ (-0.14) \ 0.67) \\ d=0.09 \ (-0.11, \ 1.12) \\ d=0.09 \ (0.01, \ 1.12) \end{array} $
h et	Intervention Sample size: 50 Control Sample size: 69 Total Sample size: 119 Average Age: 35.1 Aucation: 13.7 years % male: 95% Raec/ethnicity: 100% mild* Previous treatment: Not provided	Group-based compensatory cognitive rehabilitation treatment emphasizing compensatory strategies in prospective memory, attention, learning and memory, and executive functioning	2 hours of group training per week for 10 weeks	Strategy-based (clinician administered)	WAIS-IV DS WAIS-IV Coding HVLT-Total Recall HVLT Retention DKEFS TF DKEFS PF UPSA-B	$ \begin{array}{l} d=0.37 \ (-0.5, 0.80) \\ d=0.03 \ (-0.39, 0.45) \\ d=0.04 \ (0.14, 0.87) \\ d=0.44 \ (0.13, 0.87) \\ d=0.06 \ (-0.37, 0.48) \\ d=0.07 \ (-0.35, 0.49) \\ d=0.44 \ (0.01, 0.04) \\ d=0.58 \ (0.06, 0.90) \end{array} $
/ et	Intervention Sample size: 16 Control Sample size: 18 Total Sample size: 34	Manualized, 12-week, multimodal compensatory cognitive training intervention	2 hours per week (1- hour CogSMART and 1- hour standard supported	Strategy-based (clinician administered)	WAIS-III DS CVLT-II Trials 1–5 CVLT-II LDFR	d = -0.45 (-1.12, 0.23) d = -0.08 (-0.75, 0.59) d = -0.09 (-0.77, 0.58)

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Study	Sample characteristics	Treatment Description	Length/intensity of treatment	Strategy-based or Drill-and- practice	Outcomes included in meta-analyses	Cohen's d (95% CI)
	Average Age: 32.0 Education: 13.6 years % male: 94% Race/ethnicity: 58.6% White, 35.4% Hispanic TBI severity: 79% mild, 21% mod Previous treatment: Not provided	emphasizing habit learning and compensatory strategies in prospective memory, attention, learning and memory, and executive functioning	employment) for 12 weeks		MIST Summary MIST 24-hour D-KEFS LF D-KEFS CF D-KEFS CF D-KEFS CS WCST-64	$ \begin{aligned} d &= -0.08 \; (-0.74, \; 0.61) \\ d &= 0.73 \; (0.04, \; 1.43) \\ d &= 0.28 \; (-0.41, \; 0.95) \\ d &= 0.28 \; (-0.41, \; 0.95) \\ d &= 0.16 \; (-0.51, \; 0.84) \\ d &= -0.30 \; (-0.97, \; 0.38) \end{aligned} $

Note: CI = confidence interval, TBI = traumatic brain injury, PASAT = Paced Auditory Serial Addition Test (PASAT), TMT = Trail Making Test, WAIS-IV = Wechsler Adult Intelligence Scale-4th edition,

DS = Digit Span, SS = Symbol Search, COWAT = Controlled Oral Word Association Test, CVLT-II = California Verbal Learning Test-2nd edition, SDFR = Short delay free recall, LDFR = Long delay free Making, C-W1 = Color-Word Inhibition, WCST-64 Wisconsin Card Sorting Test – 64 items, RAVLT = Rey Auditory Verbal Learning Test, DR = Delayed Recall, RULT = Ruff Light Trails Test, WMS-III recall, VR = Virtual Reality, TDQ = Tactical Driving Quotient, ODQ = Operational Driving Quotient, PSI = Processing Speed Index, DKEFS = Delis-Kaplan Executive Functioning System, TM = Trail

Intelligence Scale-3rd edition, ACT = Auditory Consonant Trigrams, DF = Design Fluency, VF = Verbal Fluency, DVT = Digital Vigilance Test, HVLFR = Hopkins Verbal Memory Test-Revised, BVMT-R = Wechsler Memory Scale, 3rd edition, TIADL = Timed Instrumental Activities of Daily Living, RBANS = Repeatable Battery for the Assessment of Neurological Status, IVA-CPT = Integrated Visual and Auditory Continuous Performance Test, LNS = letter number sequencing, UPSA-B = University of San Diego (UCSD) Performance-Based Skills Assessment, Brief Version, WAIS-III = Wechsler Adult = Brief Visual Memory Test-Revised, GPS = The Goal Processing Scale, LF = Letter Fluency, CF = Category Fluency, CS = Switching Fluency, MIST = Memory for Intentions Screening Test.

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	Intervention	Control	Total [*]
			All studies:
Number of participants	303	261	564
Average Age (mean, SD)	35.6 (6.2)	36.1 (8.4)	36.7 (6.8)
Average Education (mean, SD)	14.2 (1.8)	14.0 (0.6)	14.2(1.2)
Average % Male (mean, SD)			88.3% (11.4)
Average racial demographics **			65.4% (9.7) White, 15.5% (2.5) African American ^{**} , 9% (0) Asian ^{**} , 4% (0) Native Hawaiian/Pacific Islander ^{**} , 2% (0) Native American/Native Alaskan ^{**} , 7% (5) Other ^{**}
Average ethnic demographics **			21.1% (12.5) Hispanic Ethnicity
Length of intervention in weeks (mean, SD), range			9.5(3.7), Range 4–15
			Strategy-based
Number of participants	106	115	221
Average Age (mean, SD)	33.2 (2.7)	34.4 (0.4)	36.3 (4.2)
Average Education (mean, SD)	13.4 (0.3)	13.9 (0.2)	13.9 (0.3)
Average % Male (mean, SD)			93.8% (3.63)
Average racial demographics **			63.9% (3.8) White, 13% (0) African American **, 9% (0) Asian ** 4% (0) Native Hawaiian/Pacific Islander **, 2% (0) Native American/Native Alaskan **,
Average ethnic demographics **			29.2% (6.2) Hispanic Ethnicity
Length of intervention in weeks (mean, SD), range		•	9.8 (2.9), Range 5–12
			Drill-and-Practice
Number of participants	149	105	254
Average Age (mean, SD)	37.5 (7.4)	37.4 (10.9)	37.1 (8.6)
Average Education (mean, SD)	15.0 (2.2)	14.2 (0.8)	14.7 (1.7)
Average % Male (mean, SD)			86.5 (14.6)
Average racial demographics **			73% (5.7) White, 18% (0) African American ** , 17.5% (5.5) Other **
Average ethnic demographics **			17% (12) Hispanic Ethnicity
Length of intervention in weeks (mean, SD), range			9.3 (4.4), Range 4–15

* Some studies only gave overall information for demographic factors. As such, the total score differs somewhat from the intervention and control group only information

** Detailed demographics are available by study in Table 2. Racial/ethnic categories with two stars indicate the information was reported in <4 studies.

*** Demographic information for the mixed and restorative interventions are reported together as only one study included a mixed intervention (Cooper et al., 2007), and then demographic information was not reported separately.