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Boston cognitive assessment (BOCA) — a comprehensive self-administered smartphone- and computer-based at-home test for longitudinal tracking of cognitive performance

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Abstract

Longitudinal cognitive testing is essential for developing novel preventive interventions for dementia and Alzheimer's disease; however, the few available tools have significant practice effect and depend on an external evaluator. We developed a self-administered 10-min at-home test intended for longitudinal cognitive monitoring, Boston Cognitive Assessment or BOCA. The goal of this project was to validate BOCA. BOCA uses randomly selected non-repeating tasks to minimize practice effects. BOCA evaluates eight cognitive domains: 1) Memory/Immediate Recall, 2) Combinatorial Language Comprehension/Prefrontal Synthesis, 3) Visuospatial Reasoning/Mental rotation, 4) Executive function/Clock Test, 5) Attention, 6) Mental math, 7) Orientation, and 8) Memory/Delayed Recall. BOCA was administered to patients with cognitive impairment (n = 50) and age- and education-matched controls (n = 50). Test scores were significantly different between patients and controls (p < 0.001) suggesting good discriminative ability. The Cronbach's alpha was 0.87 implying good internal consistency. BOCA demonstrated strong correlation with Montreal Cognitive Assessment (MoCA) (R = 0.90, p < 0.001). The study revealed strong (R = 0.94, p < 0.001) test-retest reliability of the total BOCA score one week after participants' initial administration. The practice effect tested by daily BOCA administration over 10 days was insignificant ($\beta = 0.03$, p = 0.68). The effect of the screen size tested by BOCA administration on a large computer screen and re-administration of the BOCA to the same participant on a smartphone was insignificant $(\beta = 0.82, p = 0.17; positive \beta)$ indicates greater score on a smartphone). BOCA has the potential to reduce the cost and improve the quality of longitudinal cognitive tracking essential for testing novel interventions designed to reduce or reverse cognitive aging. BOCA is available online gratis at www.bocatest.org.

Keywords: Cognitive testing, Mild cognitive impairment, Dementia, Cognition, Telehealth

Introduction

Many treatable health conditions (e.g., sleep disorders, hypertension, diabetes, heart failure, hypothyroid), deficiencies (e.g., vitamin B12, tryptophan), as well as lack of movement and social interactions can affect memory and thinking [1–3]. Longitudinal monitoring of cognitive



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health can help clinicians assess if an underlying condition is causing cognitive decline and guide timely therapeutic interventions [4]. In addition, longitudinal monitoring is essential for testing novel interventions designed to reduce or reverse cognitive aging [5]. Standard cognitive assessments are not suited for weekly/ monthly cognitive evaluations. First, they ubiquitously rely on trained professionals. While this approach has a high sensitivity and specificity for the detection of dementia, it is time and resource intensive. Second, the number of variations of standard tests is often limited resulting in practice effects [6]. Therefore, there is a clear need for a self-administered cognitive test that can be repeated periodically and is resistant to practice effects [7]. Such a test could be performed at home or in the clinic by using randomly selected non-repeating tasks to minimize practice effects.

In the last two decades, the availability of computerized cognitive testing with diagnostic accuracy comparable to traditional pen-and-paper neuropsychological testing have improved significantly [8]. The National Institutes of Health Toolbox Cognition Battery [9], the Cognitive Stability Index [10], CogState [11], BrainCheck [12], Neurotrack [13], CNS Vitals [14] can replace existing paper screening tests like Montreal Cognitive Assessment (MoCA), but they still require a trained evaluator administering the test to a patient. At home approaches have also been developed and validated (e.g., the Computer Assessment of Mild Cognitive Impairment [15], COGselftest [16], and MicroCog [11]). These tests showed good neuropsychological parameters, but were primarily designed for single use cases and are not validated for longitudinal cognitive tracking [8]. To the best of our knowledge, to date, only one test has been specifically designed for at-home longitudinal cognitive monitoring. The Brain on Track self-administered web-based test for longitudinal cognitive assessment was developed in 2014, in Portugal [17]. However, the *Brain on Track* was not available in English as of 2022. Therefore, we aimed to develop an online self-administered test for longitudinal cognitive assessment that could be used at home on multiple devices including smartphones, tablets, and computers.

Previously, BOCA has been validated against and shown strong correlations (r=0.80, p<0.01) with the Telephone Interview for Cognitive Status (TICS) [18]. BOCA demonstrated good internal consistency (Cronbach's alpha=0.79), adequate content validity, and strong (r=0.89, p<0.001) test-retest reliability of the total BOCA score one week after participants' initial administration. This study aimed to assess the convergent validity of the BOCA with the MoCA test in a larger group of participants, and provide further evidence of the BOCA's validity and reliability.

Methods

Boston Cognitive Assessment or BOCA is a 10-min, self-administered online test that uses randomly selected non-repeating tasks to minimize practice effects. BOCA includes eight subscales: Memory/Immediate Recall, Memory/Delayed Recall, Executive function/Clock Test, Visuospatial Reasoning/Mental rotation, Attention, Mental math, Language/Prefrontal Synthesis, and Orientation (Table 1). The maximum total score is 30, with higher scores indicating better cognitive performance.

Principles for test development

As a self-administered test, BOCA was designed to be user-friendly and self-explanatory. The testing within each subscale is preceded by a *training session* (Table 1). The goal of a training session is to familiarize participants with both the task and the answering protocol in each subscale. Training sessions are unscored and are repeated until the participant finds the correct answer. During training sessions, users are provided with the "Correct/Incorrect" feedback. Once a user has confirmed his/her ability to answer a question correctly in a given subscale, they are able to proceed to scored questions of that subscale.

Assessment in each subscale is accomplished by a set of questions with gradually increasing level of difficulty. For example, assessment in the Mental Math subscale has four levels of difficulty: in the first level, the participant is expected to add two single-digit numbers (e.g., 7+6=?); in the second level, one-single-digit number and one two-digit number (e.g., 7+16=?); in the third level, they must add two, two-digit numbers (e.g., 17+16=?); and in the fourth level, they must subtract a two-digit number from another two-digit number (e.g., 37-16=?) (Table 1).

Each level begins with a set of announced instructions and has a limited duration of 45 s, excluding the tests' instructions. As BOCA is intended to be used repeatedly, randomly selected non-repeating tasks are used to minimize practice effects.

BOCA was programmed to work on all devices including smartphones and tablets and in all browsers. BOCA is light on data usage transmitted over the web and can be administered over slower 3G connections. The sound check at the beginning of the test ensures that users clearly hear the instructions. Furthermore, users are asked to find a solution in their mind and to avoid using pen and paper.

Memory / immediate recall

The names of 5 animals are announced verbally. After a short pause, 16 buttons are displayed indicating the names of the 5 announced animals amidst 11 distractor animals. Participants are expected to select the Vyshedskiy et al. BMC Neurology (2022) 22:92 Page 3 of 12

Table 1 BOCA subscales, example questions, and scoring

Subscale	Subscale description	Max. Score
1. Memory / Immediate Recall	The names of 5 animals are announced verbally. After a short pause, 16 buttons are displayed indicating the names of the 5 announced animals and 11 other random animals. The participant is expected to select the 5 announced animals. This subscale includes three attempts scored as follows: all five animals selected correctly on the 1st attempt: 2 points; all five animals selected correctly on the 2nd attempt: 1 point; otherwise, zero point.	2
2. Language / Prefrontal Synthesis	Questions are announced verbally and the participant is expected to select the answer by pressing a picture on the screen.	
	Training: Integration of one modifier. E.g., 'select the blue square,' select the green triangle.'	0
	Level 1: Integration of one modifier. E.g., 'select the blue square,' select the green triangle.'	1
	Level 2: Integration of two modifiers. E.g., 'select the large blue square,' select the small green triangle.'	
	Level 3: Spatial prepositions <i>on top of</i> and <i>under.</i> E.g., 'select the square on top of the circle,'select the circle under the triangle.'	1
	Level 4: Two objects integration. E.g., 'If the tiger was eaten by the lion, who is still alive?'; 'If the boy was overtaken by the girl, who won?'	1
	Level 5: Three objects integration. E.g., 'The girl is taller than the boy. The monkey is taller than the girl. Who is the shortest?'	1
3. Visuospatial Reasoning / Mental rotation	The participant is expected to select the object, that, when rotated, is identical to the object on top.	
	Training: easy	0
	Level 1: easy	1
	Level 2: moderate	1
	Level 3: challenging	1
4. Executive function / Clock test	The participant is expected to calculate the time difference between the two clocks.	
	Training: easy	0
	Level 1: easy	2
	Level 2: moderate	1
	Level 3: challenging	1
5. Attention	The participant is instructed to click the announced digits in forward and backward order.	
	Training: Click the 4 digits in the order that you hear them	0
	Level 1: Click the 4 digits in the order that you hear them	1
	Level 2: Click the 5 digits in the order that you hear them	1
	Level 3: Click the 3 digits in the backward order	1
	Level 4: Click the 4 digits in the backward order	1
6. Mental math	The participant performs mental math by adding or subtracting two numbers.	
	Training: Single-digit number addition. E.g., $7+6=?$	0
	Level 1: Single-digit number addition. E.g., $7+6=?$	1
	Level 2: One-digit number plus two-digit number. E.g., $7 + 16 = ?$	1
	Level 3: Two-digit number plus two-digit number. E.g., $17 + 16 = ?$	1
	Level 4: Two-digit number minus two-digit number. E.g., 37–16=?	1
7. Orientation	The participant is expected to select today's month, year, and day of the week.	
	Level 1: What month is it today?	1
	Level 2: What year is it today?	1
	Level 3: What day of the week is it today?	1
8. Memory / Delayed Recall	The participant is expected to select the five animals named at the beginning of the test. The score equals to the number of correctly named animals.	5
Total maximum score		30

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5 announced animals. This subscale includes three attempts scored as follows: all five animals selected correctly on the 1st attempt: 2 points; all five animals selected correctly on the 2nd attempt: 1 point; otherwise, zero points are given. With 16 answer choices, the probability of selecting five correct animals by chance is 0.02%; four correct animals 1.3%; three correct animals 13%; two correct animals 38%; one correct animal 38%; and zero correct animal 11%.

Language / prefrontal synthesis

Combinatorial language comprehension and prefrontal synthesis subscale has five levels of difficulty. In level 1, participants are expected to integrate one modifier (colors: red, blue and green) with a noun (geometrical figures: square, triangle, and circle) and click on the corresponding picture on the screen. With 9 answer choices, the probability of selecting the correct answer by chance is 11%.

In level 2, participants are expected to integrate two modifiers (color and size) with a noun (geometrical figures: square, triangle, and circle) and click on the corresponding picture on the screen (e.g., 'select the large blue square,' 'select the small green triangle'). As in level 1, since the three colors, two sizes, and the three geometrical figures are equally familiar, the difficulty is standardized across all possible assessments. With 18 answer choices, the probability of selecting the correct answer by chance is 5.5%.

In level 3, participants are expected to integrate spatial prepositions (on top of or under) with geometrical figures (square, triangle, and circle) and click on the corresponding picture (e.g., 'select the square on top of the circle,' select the circle under the triangle'). In line with previous levels, since the two spatial prepositions and the three geometrical figures are equally familiar, the difficulty is standardized across all possible tasks. With 12 answer choices, the probability of selecting the correct answer by chance is 8.3%.

In level 4, participants are expected to mentally combine two objects as instructed by a semantically-reversible sentence. Sentences in which swapping the subject and the object result in a new meaning are called semantically-reversible sentences. By contrast, in a nonreversible-sentence (e.g., "The boy writes a letter") swapping the subject and the object results in a sentence with no real meaning ("The letter writes a boy"). With 2 answer choices, the probability of selecting the correct answer by chance is 50%.

In level 5, the participant is expected to mentally combine three objects as instructed by a semantically-reversible sentence (e.g., "The boy is taller than the girl. The monkey is taller than the boy. Who is the shortest?" In

this case the girl is the shortest.). With 3 answer choices, the probability of selecting the correct answer by chance is 33.3%.

Correct answers in all levels are scored as one, incorrect answers are scored as zero.

Visuospatial reasoning / mental rotation

The mental rotation subscale has three difficulty levels. In each level, participants are expected to select an object, that, when rotated, is identical to the object on top. At each level, the task is selected randomly from the pool of 120 shapes. To ensure standardization across assessments, the pool of 120 shapes were carefully selected to be equal in the number of visual features. With 3 answer choices in each difficulty level, the probability of selecting the correct answer by chance is 33.3%. Correct answers in all levels are scored as one, incorrect answers are scored as zero.

Executive function / clock test

In this subscale, participants are expected to calculate the time difference between two analog clocks. The test has three difficulty levels. In level 1, both minute hands are on the hour and both hour-hands are between 1 and 6 (the difference between hour-hands is no greater than 4 h)

In level 2, both minute hands are on the hour, whereas one hour-hand is between 1 and 6 and the other is between 7 and 11.

In level 3, one minute-hand is on the hour and the other is on half-hour, whilst one hour-hand is between 1 and 6 and the other is between 7 and 11. Participants respond by separately dialing hours and minutes (the minute-dial uses 5-min steps). With 12h choices and 12min choices on each dial there are 144 answers, the probability of selecting the correct answer by chance is 0.7%.

Correct answers in levels 2 and 3 are scored as one and in level 1 as two, incorrect answers are scored as zero.

Attention

Attention testing in BOCA has four levels of difficulty. In level 1, participants are expected to remember and click the four digits in the forward order. The numbers (zero to 9) are randomly drawn from a pre-recorded list. To ensure a consistent difficulty across tasks, mathematical rules were incorporated into the algorithm, for example, the algorithm avoids consecutive numbers 0,1,2,3. With 10 answer choices (zero to 9), the probability of clicking the four digits in the correct order by chance is 0.02%.

In level 2, participants are expected to remember and click five digits in the forward order. With 10 answer choices, the probability of clicking the five digits in the correct order by chance is 0.003%.

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In level 3, participants are expected to remember and click three digits in the backward order. With 10 answer choices, the probability of clicking the three digits in the correct order by chance is 0.14%.

In level 4, participants are expected to remember and click four digits in the backward order. With 10 answer choices, the probability of clicking the four digits in the correct order by chance is 0.02%.

Correct answers in all levels are scored as one, incorrect answers are scored as zero.

Mental math

Mental math testing in BOCA has four levels of difficulty. In level 1, participants are expected to add two single-digit numbers (e.g., 7+6=?). To ensure standardization across assessments, only numbers between 6 and 9 are used in this task.

In level 2, participants are expected to add one single-digit number and one two-digit number (e.g., 7+16=?). To ensure standardization across assessments, the one-digit number is between 6 and 9 and the two-digit number is between 16 and 19.

In level 3, participants are expected to add two two-digit numbers (e.g., 17+16=?). To ensure standardization across assessments, the first two-digit number is between 16 and 19 and the second two-digit number is between 26 and 29.

In level 4, participants are expected to subtract a two-digit number from another two-digit number (e.g., 37–16=?). To ensure standardization across assessments, the first two-digit number is between 31 and 35 and the second two-digit number is between 16 and 19.

With 21 answer choices (3 rows of 7 digits in each row), the probability of clicking the correct digit by chance is 4.8%. Correct answers in all levels are scored as one, incorrect answers are scored as zero.

Orientation

Orientation is tested in BOCA in 3 levels. In level 1, participants are expected to select the current month. With 12 answer choices, the probability of clicking the correct month by chance is 8.3%.

In level 2, participants are expected to select the current year. With 21 answer choices, the probability of clicking the correct year by chance is 4.8%.

In level 3, participants are expected to select the current day of the week. With 7 answer choices, the probability of clicking the correct day of the week by chance is 14.3%.

Correct answers in all levels are scored as one, incorrect answers are scored as zero.

Memory / delayed recall

In the delayed recall subscale participants are expected to select the five animals named at the beginning of the BOCA test in the subscale 1. The delayed recall score equals to the number of correctly named animals. With 16 answer choices, the probability of selecting 5 correct animals by chance is 0.02%; the probability of selecting 4 correct animals by chance is 1.3%; the probability of selecting 3 correct animals by chance is 13%; the probability of selecting 2 correct animals by chance is 38%; the probability of selecting 1 correct animal by chance is 38%; and the probability of selecting no correct animal by chance is 11%.

Montreal cognitive assessment (MoCA)

BOCA was compared to Montreal Cognitive Assessment (MoCA), a golden standard pen-and-paper test of global cognition [19]. The MoCA assesses several cognitive domains: 1) Memory/ Delayed Recall is assessed by learning of five nouns and recalling them 5 min later (5 points); 2) Visuospatial abilities are assessed using a number/letter connection exercise (1 point), a threedimensional cube copy (1 point), and a clock-drawing task (3 points); 3) Multiple aspects of executive functions are assessed using an alternation trail-making task (1 point), a phonemic fluency task (1 point), and a two-item verbal abstraction task (2 points). 3) Naming is assessed using a three-item naming task with animals (lion, camel, rhinoceros; 3 points); 4) Language is assessed using a repetition of two syntactically complex sentences (2) points); 5) Abstract thinking is assessed by asking a patient to describe the similarity between two objects (2 points); 6) Attention is evaluated using a sustained attention task (target detection using tapping; 1 point) and digit repetition forward and backward (1 point each); 7) Arithmetic is assessed by a serial subtraction task (3 points); 8) Temporal and spatial orientation is evaluated by asking the subject for the date and the city in which the test is occurring (6 points). The maximum score is 30 points with higher scores indicating better cognitive performance.

Participants

Participants were invited to the study online and recruited from the geriatric and memory clinics in eastern Massachusetts and Washington, DC. Each participants in the study gave their informed consent prior to their inclusion. In participants with cognitive impairment caregivers gave their informed consent.

The overall inclusion criteria for participants was \geq 25 years of age. Cognitive impairment was defined as the presence of subjective cognitive complaints over

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a period of at least 6 months reported by the patient or family members and MoCA score of less than 25. No participant were excluded.

Participants completed BOCA at home. They were not supervised during completion of BOCA.

Statistical analysis

To assess the correlation between BOCA and MoCA test scores, Pearson correlation was utilized with the corresponding p-value determined by the Student's t-distribution table. The conditions for independent two samples t-test were assessed via the Shapiro-Wilk test for normal distribution. All subtests' scores met the normality assumption with p > 0.05. The significance level alpha = 0.05 was utilized in concluding if the correlation was significant. Sample estimates are reported as the Pearson correlation coefficient denoted by 'R'. Confidence Intervals (CI) are indicated at 95%.

Cronbach's alpha was obtained to determine internal consistency of the BOCA test. It was computed through a two-way mixed effect model, with assumptions that a group of multiple participants is randomly selected from a population. Using this model, the participants and the subscale scores are considered sources of random effects. Subscale scores were determined through pooled scaling. Labels for interpretation of correlational strength were as follows: 0.1–0.3 indicated a small or weak association, 0.31–0.5 indicated a medium or moderate association, and 0.51–1.0 indicated a large or strong association [20].

For test-retest analysis, the Pearson correlation was used to measure the strength of the linear relationship between the first BOCA exam and the second BOCA exam. Pearson correlation coefficient is given as the quotient of the standardized form of covariance of the test-retest total scores and the standard deviation of each trial under the normal distribution assumption.

Data security

The data in transit is encrypted using SSL. SSL stands for Secure Sockets Layer, a security protocol that creates an encrypted link between a web server and a web browser. SSL certificates secure online transactions and keep all information private and secure. The test data are stored in the secure cloud database in a reputable cloud provider.

Results

Differences in the score between patients and controls

Cognitive impairment was defined as the presence of subjective cognitive complaints over a period of at least 6 months reported by the patient or family members and MoCA score of less than 25. Participants with cognitive impairment ("patients", n=50) were matched

Table 2 Characteristics of participants. The data shown as Mean (SD)

	Controls (N=50)	Patients (N = 50)	P - Value
Age	70.6 (13.1)	70.0 (14.2)	0.82
Age range (years)	26-95	25-93	n/a
Education (years)	16.2 (2.3)	16.1 (2.9)	0.94
Males	28%	36%	0.14

Table 3 MoCA performance in patients and controls

MoCA Subscales	Controls	Patients	P - Value
Visuospatial	4.33 (0.88)	2.73 (1.72)	< 0.001
Naming	2.96 (0.20)	2.67 (0.63)	0.003
Attention 1 and 2	1.96 (0.20)	1.59 (0.67)	< 0.001
Attention 3	2.86 (0.41)	1.78 (1.28)	< 0.001
Orientation	5.84 (0.37)	4.45 (1.68)	< 0.001
Delayed Recall	3.33 (1.41)	0.59 (1.04)	< 0.001
Language	2.61 (0.53)	1.96 (0.94)	< 0.001
Abstraction	1.90 (0.47)	1.47 (0.79)	0.001
MoCA Total Score	26.80 (1.88)	18.16 (5.63)	< 0.001

to controls (n=50) based on education obtained and age. Accordingly, there were no significant differences between patients and controls regarding age and education (Table 2).

All participants completed the BOCA and MoCA. The average total MoCA score was 26.80 (95% Confidence Interval: 26.26; 27.34) in controls and 18.16 (CI: 16.56; 19.75) in patients. The MoCA total score was statistically different between controls and patients (t (98) = 10.25, p < 0.001), Table 3. All MoCA subscale scores were also significantly different among controls and patients (Table 3).

The average total BOCA score was 27.30 (CI: 26.67; 27.94) in controls and 18.28 (CI: 16.35; 20.21) in patients. The BOCA total score was statistically different between controls and patients (t (98) = 8.911; p < 0.001). All BOCA subscale scores were also significantly different between the controls and patients, Table 4. There was strong positive statistically significant correlation between the BOCA total score and the MOCA total score with R = 0.90 (CI: 0.86, 0.93), p < 0.001.

Linear regression was computed for patients and controls to assess the relationship between the total BOCA score (independent variable) and their age (dependent variable), Fig. 1. As expected, there was a mild negative correlation between the total BOCA score and age in both patients: R = -0.12 (CI: -0.38, 0.15), p = 0.39; and controls: R = -0.29 (CI: -0.52, -0.01), p = 0.04.

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Table 4 BOCA performance in patients and controls

BOCA Subscales	Controls	Patients	P - value
Memory/Immediate Recall	1.64 (0.76)	0.54 (0.09)	< 0.001
Language	4.62 (0.40)	3.82 (1.41)	< 0.001
Mental Rotation	2.58 (1.16)	1.74 (1.02)	< 0.001
Clock Test	3.64 (0.65)	2.28 (1.82)	< 0.001
Attention	3.64 (0.27)	2.30 (1.03)	< 0.001
Mental Math	3.66 (0.63)	2.56 (1.57)	< 0.001
Orientation	2.96 (0.36)	2.22 (0.86)	< 0.001
Memory/Delayed Recall	4.44 (0.84)	2.82 (1.40)	< 0.001
BOCA Total Score	27.30 (2.16)	18.28 (7.34)	< 0.001

Linear regression was also computed for patients and controls to assess the relationship between the total BOCA score and their number of years of education, Fig. 2. There was a small insignificant correlation between the total BOCA score and years of education in both patients: R = -0.13 (CI: -0.39, 0.16), p = 0.37; and controls: R = 0.15 (CI: -0.13, 0.41), p = 0.28.

BOCA reliability analysis

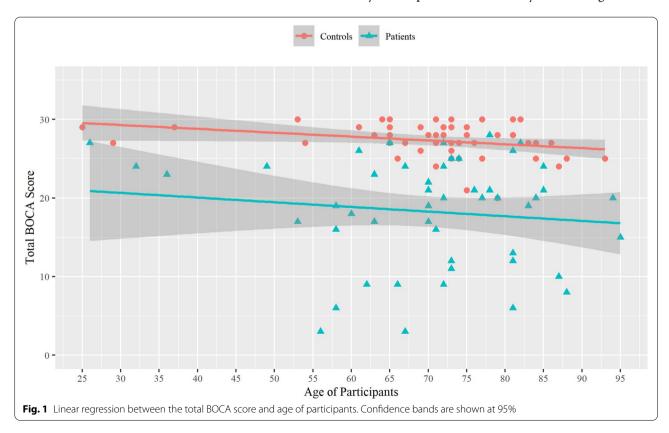
The correlation matrix for BOCA subscales was obtained via the Pearson method, Table 5. The strongest correlations were between the Orientation and the Attention

subscales (R=0.62, t(98)=7.72, p<0.001),the Orientation and the Clock Test subscales (R=0.57, t(98)=6.56; p<0.001), and the Immediate Memory and the Attention subscales (R=0.55, t(98)=6.49; p<0.001). All but one bivariate correlation was significant at the 0.01 level. The only insignificant correlation at the 0.01 level was between the Delayed Recall and the Language subscales (R=0.24, t(98)=2.48, p=0.015).

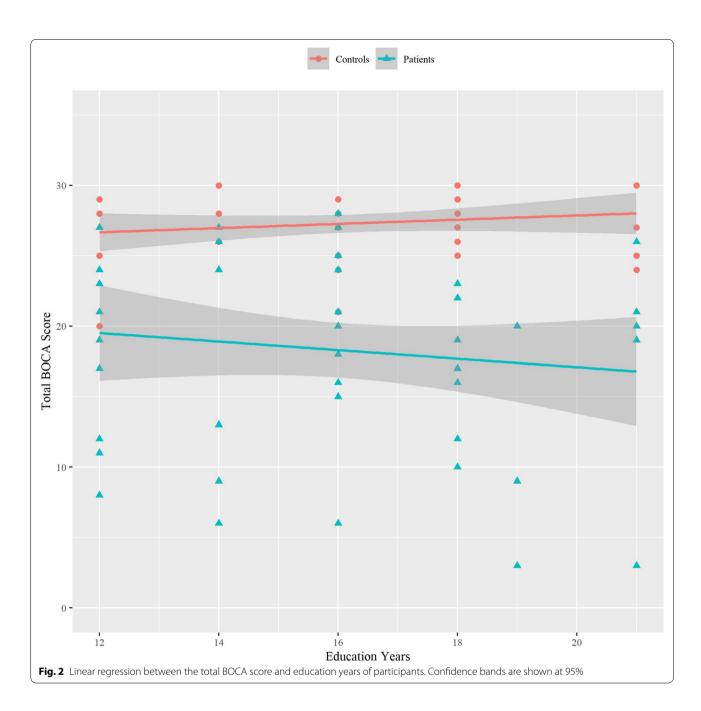
Internal consistency of the eight BOCA subscales was assessed using Cronbach's alpha. Results indicated good internal consistency $\alpha = 0.87$ (CI: 0.81, 0.90), p < 0.001. Item-total correlation (ITC) was evaluated using Pearson's product moment coefficient between each subscale score and the total score, Table 6. All subscales demonstrated high (>0.5) ITC. The highest ITC for the Orientation subscale was 0.72 and the lowest ITC for the Mental Rotation subscale was 0.501.

One subscale at a time was then removed and the Cronbach's alpha was re-calculated for the remaining 7 subscales, Table 6. For the purposes of this test, all subscale scores were standardized. The resulting Cronbach's Alpha were all positive, demonstrated high (>0.83) internal consistency, and remained stable.

The Kaiser-Meyer-Olkin measure of sampling adequacy indicated that the strength of the relationships among subscales was high (KMO=0.886) thus factor analysis was possible. Factor analysis of the eight BOCA



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subscales yielded one factor with an eigenvalue of 4.14 accounting for 51.76% of the variance in the data. Factor two had an eigenvalue of 0.801 and accounted for 10.01% of the variance. The remaining six factors had eigenvalues below 0.8. Based on the low amount of variance explained by the second factor, results of the Scree Plot, and moderate to strong positive correlations between all eight BOCA subscales and the first factor, only the first factor was retained. This factor was subsequently identified as global cognitive functioning and encompassed all of the eight subscales.

The BOCA test-retest reliability was evaluated by calculating a Pearson correlation coefficient between the first administration of the BOCA and the re-administration of the BOCA to the same participants approximately one-week later (93 participants), Table 7. The one-week test-retest correlation coefficient for the BOCA total score was $R\!=\!0.94$ (CI: 0.91 0.96), $p\!<\!0.001$, revealing excellent BOCA long-term stability. As shown in Table 7, each of the BOCA subscales showed significant test-retest correlations one week after participants initial completion of the test.

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Table 5 BOCA subscales Pearson correlation matrix

BOCA Subscale	Immediate Recall	Language	Mental Rotation	Clock Test	Attention	Mental Math	Orientation	Delayed Recall
Memory / Immediate Recall	-							
Language	0.41**	-						
Mental Rotation	0.34**	0.3**	=					
Clock Test	0.42**	0.45**	0.36**	-				
Attention	0.55**	0.43**	0.33**	0.48**	-			
Mental Math	0.44**	0.42**	0.45**	0.53**	0.49**	_		
Orientation	0.50**	0.44**	0.41**	0.57**	0.62**	0.49**	_	
Memory / Delayed Recall	0.52**	0.24*	0.39**	0.49**	0.50**	0.35**	0.53**	_

^{*}p < 0.05, **p < 0.01

Practice effect was assessed by administering BOCA daily to 10 participants and analyzed through a linear mixed effect model, where the trial number was modeled as a fixed effect, participants as random effects, and test score as the dependent variable. An interaction term was added to account for the effect of each trial nested within each individual participant. The effect of the trial number on the BOCA total score was insignificant (β =0.03, SE=0.07, p=0.68), Fig. 3. The fixed effect intercept of the trial number had β =19.3, SE=2.5, p<0.0001. The effect of the trial number on individual BOCA subscales was also insignificant, Table 8.

BOCA performance on small screens was evaluated by administration of the BOCA on a smartphone and re-administration of the BOCA to the same participant on a large screen device (a computer or an iPad; 22 participants). To avoid bias, the order of administration was varied randomly between participants: a computer/iPad was used first in 55% participants and a smartphone was used first in 45% participants. A linear mixed effect model, where the screen size was modeled as a fixed effect, participants as random effects, and test score as the dependent variable. The effect of the screen size on the BOCA total score was insignificant ($\beta = 0.82$, SE = 0.57, p = 0.17; positive β indicates greater score on a smartphone), Table 9. In 50% of participants the BOCA total score was slightly greater on a smartphone, in 27% the BOCA total score was slightly greater on a computer, and in 22% of participants the BOCA total score did not change between a computer and a smartphone. The effect of the screen size in individual BOCA subscales was also insignificant with some subscales on average slightly better on a computer and some on a smartphone, Table 9.

Discussion

This manuscript discusses the implementation of the Boston Cognitive Assessment or BOCA, a self-administered at-home test used for cognitive screening and

longitudinal monitoring. The manuscript shows the results of BOCA's validation procedure in 50 participants with cognitive impairment (patients) and 50 controls matched by education obtained and age. Test scores were significantly different between patients and controls (p < 0.001) suggesting good discriminative ability. Internal consistency of BOCA was high (Cronbach's alpha = 0.87) and BOCA exhibited excellent test-retest reliability

Table 6 Factor analysis for BOCA

BOCA Subscales	Item-Total Correlation	Cronbach's Alpha if Item Deleted
Memory/Immediate Recall	0.63	0.84
Language	0.51	0.85
Mental Rotation	0.50	0.85
Clock Test	0.66	0.83
Attention	0.68	0.83
Mental Math	0.63	0.84
Orientation	0.72	0.83
Memory/Delayed Recall	0.59	0.85

Table 7 Test-Retest Reliability (N = 93)

BOCA Subscales	BOCA 1	BOCA 2	Pearson Correlation
Memory/Immediate Recall	1.11 (0.96)	1.23 (0.93)	0.68**
Language	4.32 (0.92)	4.10 (1.12)	0.50**
Mental Rotation	2.04 (1.12)	2.24 (0.96)	0.52**
Clock Test	3.06 (1.31)	3.13 (1.32)	0.58**
Attention	2.84 (1.41)	2.84 (1.44)	0.82**
Mental Math	2.90 (1.40)	2.92 (1.38)	0.84**
Orientation	2.60 (0.81)	2.57 (0.91)	0.53**
Memory/Delayed Recall	3.59 (1.66)	3.71 (1.59)	0.70**
Total Score	22.52 (7.52)	22.75 (7.74)	0.94**

^{**}p < 0.01

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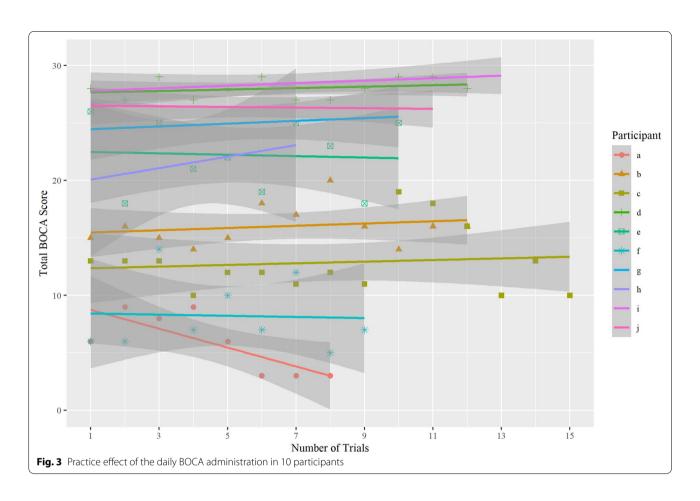


Table 8 Practice effect analysis

BOCA Subscales	Beta (β)	Standard Error	T Statistic (β/SE)	P - Value
Memory/Immediate Recall	0.01	0.02	0.93	0.36
Language/PFS	-0.02	0.03	-0.66	0.52
Mental Rotation	0.01	0.02	0.56	0.58
Clock Test	0.02	0.02	0.65	0.53
Attention	0.01	0.02	0.75	0.46
Mental Math	0	0.03	-0.19	0.85
Orientation	0.01	0.02	0.58	0.58
Memory/Delayed Recall	0	0.03	-0.26	0.80
Total Score	0.03	0.07	0.42	0.68

(R=0.94). In further support of the BOCA's validity, exploratory factor analysis yielded a single factor explaining a plurality of the variance in participants scores. This factor was felt to reflect global cognitive functioning, and suggests the test's internal structure matches the construct intended to be measured. Finally, practice effect, assessed by administering BOCA daily for 10 days, was insignificant ($\beta=0.03$, SE=0.08, p=0.74) confirming the absence of learning even when BOCA is used daily.

These results agree with previously reported observations comparing BOCA to the Telephone Interview for Cognitive Status (TICS) that demonstrated BOCA's ability to differentiate patients from controls (p<0.001), good BOCA's internal consistency (Cronbach's alpha=0.79), strong correlations with the TICS (r=0.80, p<0.01), and strong (r=0.89, p<0.001) test-retest reliability of the total BOCA score one week after participants' initial administration [18]. Furthermore, this study reconfirmed

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Table 9 Comparison between BOCA administration on a large screen (computer or iPad) and a small screen smartphone (77% responded on an iPhone; 23% responded on an Android smartphone). Positive β indicates greater score on a smartphone

BOCA Subscales	Beta (β)	Standard Error (SE)	T-Statistic (β/SE)	P - Value
Memory/Immediate Recall	0.36	0.19	1.89	0.07
Language/PFS	-0.18	0.19	-0.94	0.36
Clock Test	0.41	0.28	1.48	0.15
Mental Rotation	-0.23	0.16	-1.42	0.17
Attention	0.14	0.27	0.51	0.61
Mental Math	0.23	0.20	1.16	0.26
Orientation	0.18	0.14	1.28	0.21
Memory/Delayed Recall	-0.09	0.16	-0.57	0.58
Total Score	0.82	0.57	1.44	0.17

the factor structure of the BOCA in a novel sample, and demonstrated that education does not significantly influence patient scores, which is a common limitation of other global screening instruments. Although not included in this study, previous evaluations of the BOCA found it to possess equal or better sensitivity (79%) and specificity (90%) to mild cognitive impairment relative to other global screening tests [18].

BOCA was designed to be a self-administered and selfexplanatory test. BOCA experience starts with receiving an email or SMS with a link to the test. Clicking on the link, automatically opens the browser with BOCA introductory page or the BOCA app on an iPhone. The mandatory sound check ensures that patients can hear the instructions. The testing in each subscale is preceded by a training session in order to familiarize participants with both the task and the answering protocol. Training sessions are unscored and are repeated until the participant finds the correct answer. Once a user has confirmed his/ her ability to answer correctly in a given subscale, they proceed to scored questions of that subscale. The testing in each subscale is accomplished by a set of questions which gradually increase in difficulty. Randomly selected non-repeating tasks are used to minimize practice effects.

BOCA achieves high sensitivity to cognitive impairment by using eight orthogonal subscales: Memory/Immediate recall, Memory/Delayed recall, Executive function/Click test, Visuospatial reasoning/Mental rotation, Attention, Mental math, Language/Prefrontal synthesis, and Orientation, Table 1. In selecting BOCA subscales, we strived to cover as many neurologically distinct cognitive processes as possible. The variety of cognitive tasks had to be balanced against the test duration. The maximum total score is 30, with higher scores indicating better cognitive performance. The resulting BOCA takes an average $11.0\pm1.8\,\mathrm{min}$ to complete and therefore satisfies our preset duration specification.

There was no difference between results of the BOCA administered on a smartphone and on a computer or iPad (Table 9) and no participant complained about illegible buttons or test stimuli. This is not surprising as all instructions in BOCA are auditory and therefore do not depend on the screen size; no reading is required except the names and numbers on the buttons. Additionally, BOCA uses big buttons that are several times bigger than buttons commonly found in a smartphone browser. The smallest smartphone used by participants was iPhone SE (41% of participants) with a screen size of 4.7-in.

BOCA advantages and limitations

The BOCA test shares potential advantages with the other computerized cognitive tests: convenience and cost-effectiveness, reduction of the examiner bias, reduction of performance anxiety, automatic recording and storing of the results, and progress tracking [8, 17]. The main criticisms of at-home cognitive tests relate to the presence of potential technical difficulties faced by older adults [8]. To address this concern, BOCA uses unscored training sessions in each subscale, large unambiguous buttons, and an internet browser to present the test. No installation of any files is required and the test can be taken on any device, including smartphones and tablets. All participants in this trial have taken the test at home with minimal help from their caretakers, suggesting good usability in the at-home setting. As the number of adults using smartphones increases, the lack of familiarity should decrease as well.

However, unsupervised testing at home also creates the potential issue of patient non-compliance. For example, patients could record names of animals on paper or use a calculator in the math subscale. In the future, it may be possible to add eye tracking and face-movement recognition to avoid this potential issue.

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Computerized tests performed at home also have potential technological hurdles, such as different hardware and internet speed of the patients' computers. Using an internet browser to present the test (instead of locally installed software) guarantees homogeneity in the different platforms and simplifies the procedure for patients. Once patients receive an email or SMS from their provider, they click on the link, automatically opening the browser with the test. Furthermore, BOCA preloads all of the data needed for each subscale before its initiation, resulting in independence from internet speed. A slower 3G network can affect the waiting time for the test to start, but not the duration of the test.

High internal consistency (Cronbach's alpha=0.87), excellent test-retest reliability, good discriminative ability, and the absence of practice effect validate BOCA as an effective cognitive test for longitudinal clinical use. BOCA is the first self-administered at-home test intended for cognitive monitoring. It has the potential to revolutionize cognitive tracking essential for testing novel interventions designed to reduce or reverse cognitive aging. Additionally, the test can be used to assess the effect of anesthesia, long-term effect of cancer drugs, COVID fog, and other conditions known to affect cognition. BOCA is available online gratis at www.bocatest.org

Acknowledgments

We wish to thank Dr. C. Prather for insightful comments on this manuscript.

Authors' contributions

Study design: AV, MK, IP. Developed the BOCA test: AV. Implementation of the BOCA test: AV, AS, DD, LdT, SO. Acquisition of data: EF, PJ, MA, SB, AG, VM, SM, IP. Analysis of data: RN. Drafting the manuscript: AV. Manuscript revision for important intellectual content: AV, RN, DG, MK, PJ. All authors have read and approved the manuscript.

Funding

This research received no specific grant from any funding agency in the public, commercial, or not-for-profit sectors.

Availability of data and materials

Data are available from the corresponding author upon reasonable request. The data are not publicly available due to them containing information that could compromise research participant privacy.

Declarations

Ethics approval and consent to participate

The research has been approved by the George Washington University Committee on Human Research, USA, Institutional Review Board (IRB). All methods were performed in accordance with the relevant guidelines and regulations. All participants in the study gave their informed consent prior to their inclusion. In participants with cognitive impairment the caregiver's informed consent was also requested.

Consent for publication

Not Applicable.

Competing interests

The authors declare no conflict of interest.

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Received: 26 July 2021 Accepted: 8 March 2022 Published online: 15 March 2022

References

- Buratti L, et al. Obstructive sleep apnea syndrome: an emerging risk factor for dementia. CNS Neurol Disord Drug Targets. 2016;15:678–82.
- Bakulski KM, et al. Heavy metals exposure and Alzheimer's disease and related dementias. J Alzheimers Dis. 2020;76(4):1215-42.
- Iuliano E, et al. Physical exercise for prevention of dementia (EPD) study: background, design and methods. BMC Public Health. 2019;19:1–9.
- Kim J, et al. Tracking cognitive decline in amnestic mild cognitive impairment and early-stage Alzheimer dementia: mini-mental state examination versus neuropsychological battery. Dement Geriatr Cogn Disord. 2017:44:105–17.
- Foster PP, et al. Roadmap for interventions preventing cognitive aging. Front Aging Neurosci. 2019;11:268.
- Cooley SA, et al. Longitudinal change in performance on the Montreal cognitive assessment in older adults. Clin Neuropsychol. 2015;29:824–35.
- Ruano L, et al. Tracking cognitive performance in the general population and in patients with mild cognitive impairment with a self-applied computerized test (brain on track). J Alzheimers Dis. 2019;71:541–8.
- 8. Zygouris S, Tsolaki M. Computerized cognitive testing for older adults: a review. Am J Alzheimers Dis Demen. 2015;30:13–28.
- Weintraub S, et al. The cognition battery of the NIH toolbox for assessment of neurological and behavioral function: validation in an adult sample. J Int Neuropsychol Soc. 2014;20:567.
- Erlanger DM, et al. Development and validation of a web-based screening tool for monitoring cognitive status. J Head Trauma Rehabil. 2002;17:458–76.
- Green RC, Green J, Harrison JM, Kutner MH. Screening for cognitive impairment in older individuals: validation study of a computer-based test. Arch Neurol. 1994;51:779–86.
- Groppell S, et al. A rapid, mobile neurocognitive screening test to aid in identifying cognitive impairment and dementia (BrainCheck): cohort study. JMIR Aging. 2019;2:e12615.
- 13. Hamermesh RG, Kind L, Knoop C-I. Neurotrack and the Alzheimer's Puzzle; 2016.
- Gualtieri CT, Johnson LG. Reliability and validity of a computerized neurocognitive test battery, CNS Vital Signs. Arch Clin Neuropsychol. 2006;21:623–43.
- Saxton J, et al. Computer assessment of mild cognitive impairment. Postgrad Med. 2009;121:177–85.
- Dougherty JH Jr, et al. The computerized self test (CST): an interactive, internet accessible cognitive screening test for dementia. J Alzheimers Dis. 2010;20:185–95.
- Ruano L, et al. Development of a self-administered web-based test for longitudinal cognitive assessment. Sci Rep. 2016;6:19114.
- Gold D, et al. The Boston cognitive assessment: psychometric foundations of a self-administered measure of global cognition. Clin Neuropsychol. 2021;1–18.
- Nasreddine ZS, et al. The Montreal cognitive assessment, MoCA: a brief screening tool for mild cognitive impairment. J Am Geriatr Soc. 2005;53:695–9.
- 20. Laerd statistics. Pearson product moment correlation. Statistical tutorials and software guides. https://statistics.laerd.com/statistical-guides/pears on-correlation-coefficient-statistical-guide-2.php. (2021).

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