Standardization and diagnostic utility of the Frontal Assessment Battery for healthy people and patients with dementia in the Chilean population

Fabrissio Grandi^{1,2,3,4}[®], David Martínez-Pernía^{1,2,5}[®], Mario Parra⁶[®], Loreto Olavarria^{2,3}[®], David Huepe⁵[®], Patricia Alegria⁷[®], Álvaro Aliaga⁸[®], Patricia Lillo^{1,9,10}[®], Carolina Delgado^{11,12}[®], Marcela Tenorio^{4,13}[®], Ricardo Rosas¹⁴[®], Oscar López^{15,16}[®], James Becker^{15,16,17}[®], Andrea Slachevsky^{1,2,3,18}[®]

ABSTRACT. The Frontal Assessment Battery (FAB) is a screening test that measures executive functions. Although this instrument has been validated in several countries, its diagnostic utility in a Chilean population has not been studied yet. **Objectives:** This study aimed to (1) adapt FAB in a Chilean population; (2) study the psychometric properties of the FAB in a Chilean population; (3) assess the sociodemographic influence in the performance of the FAB in a sample of healthy controls (HC); and (4) develop normative data for this healthy group. **Methods:** A HC (n=344) and a group of patients with dementia (n=156) were assessed with the Chilean version of FAB. **Results:** FAB showed good internal consistency (Cronbach's alpha=0.79) and acceptable validity based on the relationship with other variables. Factor analysis showed the unidimensionality of the instrument. Significant differences were found in the total FAB value between the HC and dementia groups. With the matched sample, the established

¹Gerosciences Center for Brain Health and Metabolism, Santiago, Chile.

²Universidad de Chile, Faculty of Medicine, Hospital del Salvador, Memory and Neuropsychiatric Clinic, Neurology Department, Santiago, Chile.

³Universidad de Chile, Faculty of Medicine, Neuropsychology and Clinical Neuroscience Laboratory, Physiopathology Department, Neuroscience and East Neuroscience Departments, Santiago, Chile.

⁴Universidad de los Andes, School of Psychology, Santiago, Chile.

⁵Universidad Adolfo Ibañez, School of Psychology, Center for Social and Cognitive Neuroscience, Santiago, Chile.

⁶University of Strathclyde, School of Psychological Sciences and Health, Glasgow, Scotland.

⁷Clínica Alemana, Physical Medicine and Rehabilitation Service, Santiago, Chile.

⁸Diego Portales Universidad, School of Psychology, Santiago, Chile.

⁹Universidad de Chile, Faculty of Medicine, South Neuroscience Department, Santiago, Chile.

¹⁰Complejo Hospitalario San José, Neurology Unit, Santiago, Chile.

¹¹Universidad de Chile, School of Medicine, Department of Neuroscience, Santiago, Chile.

¹²Universidad de Chile, Hospital Clínico, Department of Neurology and Neurosurgery, Healthy Brain Unit, Santiago, Chile.

¹³Millennium Institute for Caregiving Research, Santiago, Chile.

¹⁴Pontificia Universidad Católica de Chile, Center for the Development of Inclusion Technologies, Santiago, Chile.

¹⁵University of Pittsburgh, Department of Psychiatry, Pittsburgh, PA, USA.

¹⁶University of Pittsburgh, Department of Neurology, Pittsburgh, PA, USA.

¹⁷University of Pittsburgh, Department of Psychology, Pittsburgh, PA, USA.

¹⁸Universidad del Desarrollo, Clínica Alemana, Department of Medicine, Neurology Unit, Santiago, Chile.

Correspondence: Andrea Slachevsky; Email: andrea.slachevsky@uchile.cl

Disclosure: The authors report no conflicts of interest.

Funding: ANID/FONDAP/15150012; ANID/FONDEF/ID18/10113; ANID/Fondecyt/1191726, 1210176, and 1210195 and MULTI-PARTNER CONSORTIUM TO EXPAND DEMENTIA RESEARCH IN LATIN AMERICA [ReDLat, supported by the National Institutes of Health, National Institutes of Aging (P30 AG066468), Alzheimer's Association (SG-20-725707), Tau Consortium, and Global Brain Health Institute] and Alzheimer's Association GBHI ALZ UK-20-639295.

Received on June 18, 2021; Received in its final form on September 06, 2021; Accepted on September 17, 2021.

CC BY

This study was conducted by the Memory and Neuropsychiatric Clinic, Neurology Department, Hospital del Salvador and Faculty of Medicine, Universidad de Chile, Santiago, Chile.

cutoff point was 13.5, showing a sensitivity of 80.8% and a specificity of 90.4%. Regression analysis showed that education and age significantly predicted FAB performance in the healthy group. Finally, normative data are provided. **Conclusions:** This study shows that FAB is a useful tool to discriminate between healthy people and people with dementia. However, further studies are needed to explore the capacity of the instrument to characterize the dysexecutive syndrome in people with dementia in the Chilean population.

Keywords: Executive Function; Dementia; Mental Status and Dementia Tests; Neurodegenerative Diseases.

PADRONIZAÇÃO E UTILIDADE DIAGNÓSTICA DO FRONTAL BATERIA DE AVALIAÇÃO PARA PESSOAS SAUDÁVEIS E PACIENTES COM DEMÊNCIA NA POPULAÇÃO CHILENA

RESUMO. A Bateria de Avaliação Frontal (FAB) é um teste de rastreio que mede as funções executivas. Embora esse instrumento tenha sido validado em vários países, sua utilidade diagnóstica em uma população chilena ainda não foi estudada. **Objetivos:** (1) Adaptar a FAB para uma população chilena; (2) estudar as propriedades psicométricas da FAB em uma população chilena; (3) avaliar a influência sociodemográfica no desempenho da FAB em uma amostra de controles saudáveis; e (4) desenvolver dados normativos para este último grupo. **Métodos:** Um grupo controle saudável (n=344) e um grupo de pacientes com demência (n=156) foram avaliados com a versão chilena da FAB. **Resultados:** A FAB apresentou boa consistência interna (alfa de Cronbach=0,79) e validade aceitável com base na relação com outras variáveis. A análise fatorial mostrou a unidimensionalidade do instrumento. Diferenças significativas foram encontradas no valor total da FAB entre os grupos controle saudável e demência. Com a amostra pareada, o ponto de corte estabelecido foi de 13,5, que apresentou sensibilidade de 80,8% e especificidade de 90,4%. A análise de regressão mostrou que a escolaridade e a idade predisseram significativamente o desempenho da FAB no grupo saudável. Finalmente, os dados normativos são fornecidos. **Conclusões:** O presente estudo mostrou que a FAB é uma ferramenta útil para discriminar entre pessoas saudáveis e aquelas com demência. No entanto, mais estudos são necessários para explorar a capacidade do instrumento para caracterizar a síndrome disexecutiva em pessoas com demência na população chilena.

Palavras-chave: Função Executiva; Demência; Testes de Estado Mental e Demência; Doenças Neurodegenerativas.

INTRODUCTION

he executive function (EF) comprises a wide f L range of cognitive processes and behavioral competencies, including reasoning, problem-solving, planning, sequencing, resistance to interference, multitasking, cognitive flexibility, and the capacity to deal with novelty, among others¹. These processes mainly depend on neural circuits involving the prefrontal cortex, the basal ganglia, the parietal cortex, the cerebellum, and the thalamus². Assessing EF can be helpful in the diagnosis and prognosis of many brain disorders and other neuropsychiatric conditions, such as vascular cognitive impairment, frontotemporal dementia, parkinsonian disorders, and schizophrenia³. Along with the comprehensive neuropsychological evaluation of executive dysfunction, brief screening tools that are easy and quick to administer and contribute to determining whether a person presents with executive impairments and, accordingly, improving the quality of preliminary diagnostic workup are used^{4,5}. In this context, the Frontal Assessment Battery (FAB) was devised as a rapid bedside screening of frontal functions. The FAB comprises six subtests that assess different domains of EF⁵. Each subset explores a specific cognitive or behavioral domain related to the functions of frontal lobes, including conceptualization, mental flexibility, motor programming, sensitivity to interference, inhibitory control, and environmental autonomy. The global performance on these six subtests gives a composite score that summarizes the severity of the dysexecutive syndrome⁶. The FAB has good correlations with other executive measures such as the Wisconsin Card Sorting Test (WCST) (number of perseverative errors: rho=0.68; and number of criteria: rho=0.77) as well as measures of general cognitive functioning (Mattis Dementia Rating Scale) (rho=0.82)⁵.

Since its first publication, the FAB has been adapted to diverse languages and cultures, including Brazil⁷, Korea⁸, Japan⁹, Italy¹⁰, Germany¹¹, France^{5,12}, China¹³, Portugal¹⁴, Spain¹⁵, Turkey¹⁶, Taiwan¹⁷, and Persia¹⁸. Several studies have reported that the FAB has presented adequate reliability and validity.

The diagnostic utility of the FAB has been reported in patients with Alzheimer's disease⁸, amyotrophic lateral sclerosis¹⁹, frontotemporal dementia¹², and in small study of patients with stroke¹³. Age, education, and race influence the performance in executive tests^{16,20,21}. Although some empirical work has been done on FAB in Latin America^{7,21}, there has not been yet any studies in this region that provide normative data in Spanish. More studies are needed in Spanish-speaking Latin America and the Caribbean (LAC) countries to support its use in clinical practice.

Therefore, our aims were to (a) adapt FAB in a Chilean population; (b) study the psychometric properties of the FAB in this population (healthy people and people with dementia); (c) assess the influence of sociodemographic variables in the performance of the FAB in the healthy controls (HC); and (d) develop normative data in this healthy group.

METHODS

Participants

This normative study involved 344 HC (194 women and 150 men). All of them were native Spanish speakers (Chilean), lived in the community, and met the following inclusion criteria: (a) with at least a minimal writing capacity (correct writing regardless of orthographic errors due to low education); (b) scores >24/30 on the Mini-Mental State Examination (MMSE)²⁰ (c); scores <5 on the Geriatric Depression Scale²² (d); scores <51 in the Zung Anxiety Scale²³. Subjects were excluded if they had current major psychiatric diseases including alcohol or drug abuse, were taking psychoactive drugs, had history of brain injury (e.g., stroke, dementia, or any other neurological illness detected on a semi-structured clinical interview), or had a severe sensory deficit (loss of vision and/or hearing) that could impede neuropsychological evaluation.

They were recruited through a variety of advertisements at citizen activity centers and workplaces. Participation was voluntary, and the participants did not receive any compensation for their contribution to the study. This study was approved by the Comité de Ética of the Servicio Metropolitano Oriente, Santiago, Chile. Written informed consent was obtained from all the participants.

The clinical sample included 156 patients with dementia syndromes (83 women and 73 men) (Table 1). All patients were evaluated in the Cognitive Neurology

 Table 1. Demographic and neuropsychological characteristics of healthy controls and patients.

	Control (n=344)	Dementia (n=156)	p-value
Age ^a	55.35±18.096	74.2±7.626	<0.001
Years of education ^a	12.62±4.565	11.42±4.7	<0.01
MMSE ^a	28.53±1.578	19.68±5.631	<0.001
FAB total ^a	16.07±1.751	10.06±3.865	<0.01
FAB subtest 1ª	2.32±0.685	1.5±1.002	<0.01
FAB subtest 2 ^a	2.73±0.530	1.56±1.013	<0.01
FAB subtest 3ª	2.53±0.755	1.42±0.993	<0.01
FAB subtest 4 ^a	2.86±0.402	1.82±1.187	<0.01
FAB subtest 5 ^a	2.63±0.729	1.26±1.133	<0.01
FAB subtest 6ª	2.98±0.178	2.48±0.963	<0.01

^aResults are expressed in mean±standard deviation; MMSE: Mini-Mental State Examination; FAB: Frontal Assessment Battery. and Dementia Unit (UNCD) at the Department of Neurology, Hospital del Salvador in Santiago, Chile. The UNCD receives patients with suspected dementia from primary care facilities. A diagnosis was made by a neurologist based on the *DSM-IV-TR* criteria for dementia using multidisciplinary approach (neurological, neuropsychological, laboratory, and neuroimaging data). There were 115 patients with Alzheimer's disease, 17 with frontotemporal dementia behavioral variant, 6 with Lewy body dementia, 3 with vascular dementia, 2 with mixed dementia, 1 with semantic dementia, 1 with progressive supranuclear Palsy, 1 with alcoholic dementia, and 10 with dementia of unknown etiology.

Instruments and procedure

All participants were initially assessed with the MMSE. The adaptation of the FAB to Spanish was achieved by two translations from English to Spanish based on the original FAB, followed by two back-translations from Spanish to English that were reviewed with one of the authors' original FAB. The forward- and back-translations were performed independently by different individuals, in each case by one bilingual expert in the field of dementia and by one bilingual layperson. The Chilean version of the FAB (FAB-Ch) can be found in the Supplementary material (Appendix 1). It maintains the structure and number of items of the original English version and is grouped into six sections: conceptualization, mental flexibility, motor programming, sensitivity to interference, inhibitory control, and environmental autonomy. The original lexical fluency task with letter "S" in the English version was changed to lexical fluency with letter "A" because the number of words starting with A in Spanish is higher than those starting with letter S. Each subtest is scored from 3 (high score) to 0 (low score). The maximum score is 18 points.

Statistical analyses

All analyses were conducted using IBM SPSS Statistics 25 for Microsoft Windows (IBM Corp., Armonk, NY, USA). Descriptive and comparative analyses were performed using Student's *t*-tests to compare between the two groups. Regarding psychometric aspects, reliability was explored via internal consistency of the instrument with Cronbach's alpha.

Evidence of validity based on the relationship with other variables was evaluated by assessing the association between the performance on the FAB-Ch and MMSE. We also studied the correlation between our instrument and two EFs tests: (a) number of sorts in WCST and (b) categorical fluency, collected in the Chilean-Argentine version of the ACE-III test²⁴ using Pearson`s correlation. In case of categorical fluency, the participant must identify the names of animals.

The diagnostic utility was determined using the Receiver Operating Characteristic (ROC) analysis to calculate sensitivity and specificity values. The first analysis was carried out with the complete sample, and the second analysis included the matched sample in the variables age and education. The influence of sociodemographic variables in the HC was also studied using linear regression. Finally, we present mean and standard deviation (SD) of the total FAB-Ch scores stratified by age and education, as well as scores in the single subtests of this instrument.

RESULTS

Sociodemographic variables

Demographic and neuropsychological data of the sample are presented in Table 1. The HC was younger [t(502)=-12.485, p<0.001] and showed more years of education [t(502)=-2.639, p<0.01] than the patients group. People with dementia performed significantly worse than the HC on the MMSE [analysis of covariance (ANCOVA) covaried by age and years of education: F(1,499)=579.60; p<0.001]. In the case of the HC, the proportion of women was 53.2% and that of men was 45.8%, while dementia patients showed a proportion of 56.4% for women and 43.6% for men. This last case is probably associated with epidemiological variables. Alternatively, to control the effect of demographic variables on the difference between HC and patients group, we performed an analysis in a subsample of participants matched by age and education level. The outcome was very similar to that obtained from total data (Table 2).

Performance on Chilean version of the Frontal Assessment Battery: total score and subtests

Significant differences were found in the total FAB-Ch values between the HC and the dementia group (Table 1). Regarding the scores obtained in the subtests that make up the FAB-Ch, significant differences were again found between the two study groups in the domains of "conceptualization," "mental flexibility," "motor programming," "sensitivity to interference," "inhibitory control," and "environmental autonomy." Additionally, a multivariate analysis of covariance (MANCOVA) was conducted to compare results across subtests of the FAB-Ch by diagnosis category controlling for age, sex, and years of education. Performance differed significantly between the two groups for each subtest of the FAB-Ch [Wilks' lambda=0.488, F(6,487)=77.352; p<0.001]. Both

HC and patients group differed in each of the subtests: Conceptualization [F(1,492)=91.176, p<0.001]; mental flexibility [F(1,492)=198.732, p<0.001]; motor programming [F(1,492)=141.212, p<0.001]; sensitivity to interference [F(1,492)=171.490, p<0.001]; inhibitory control [F(1,492)=148.245, p<0.001]; and environmental autonomy [F(1,492)=60.176, p<0.001].

Psychometric properties

Reliability

The Cronbach's alpha for the FAB-Ch considering the 6 subscales and calculated for all 500 subjects was α =0.797, which shows a good reliability of the instrument. Cronbach's values of the six subtests suggest that all items positively contributed to the overall reliability.

Validity based on the relationship with other variables The FAB-Ch showed a statistically significant association with the MMSE (Pearson's r=0.83; p<0.001, n=499) and other measures of EF (number of sorts in WCST: r=0.678, p<0.001, n=413; and category fluency: r=0.71, p<0.001, n=493) collected in the ACE-III test, so we have a high validity based on the relationship with other variables²⁵.

Structure of the Chilean version of the Frontal Assessment Battery

The six subscales of the FAB-Ch were subjected to an Exploratory Factorial Analysis in order to obtain its

Table 2. Demographic and neuropsychological characteristics of healthy controls and patients matched by age and education level.

	Control (n=122)	Dementia (n=118)	p-value
Ageª	72.25±6.891	70.84±6.73	0.11
Years of education ^a	fears of education ^a 11.63±4.857 1		0.117
MMSE ^a	28.29±1.639	20.06±5.307	<0.001
FAB total ^a	15.77±1.875	10.35±3.925	<0.001
FAB subtest 1ª	2.36±0.739	1.57±0.977	<0.001
FAB subtest 2ª	2.66±0.625	1.55±1.038	<0.001
FAB subtest 3ª	2.43±0.862	1.5±0.988	<0.001
FAB subtest 4ª	2.86±0.44	1.88±1.176	<0.001
FAB subtest 5ª	2.42±0.87	1.28±1.136	<0.001
FAB subtest 6ª	2.99±0.091	2.54±0.905	<0.001

^aResults are expressed in mean±standard deviation; MMSE: Mini-Mental State Examination; FAB: Frontal Assessment Battery.

factorial structure. We used Kaiser's criterion (eigenvalues>1.0) and the extraction method was by principal axis factoring. The factors were then orthogonally rotated using a varimax rotation. The Kaiser–Meyer–Olkin test for sampling adequacy was 0.85, which indicates that factor analysis is appropriate. Bartlett's test of sphericity reached statistical significance (χ^2 =805.95, p<0.001), supporting the factorability of the correlation matrix. The results showed that the FAB-Ch has a unidimensional structure. The explained variance was 41%, and the factorial loadings were mostly above 0.5.

Utility of the Chilean version of the Frontal Assessment Batteryto classify patient and healthy controls

The results of the ROC curve analysis for the FAB are shown in Table 3 and Figure 1. The area under the curve (AUC) for the FAB was 0.92 (95% confidence interval: 0.89–0.95), indicating an overall high diagnostic usefulness of the test²⁶. The optimal balance between sensitivity

Table 3. Sensitivity and specificity of the Chilean version of the Frontal Assessment Battery for the discrimination of dementia patients (n=156) and healthy controls (n=344) in the complete sample.

Cutoff point	Sensitivity	Specificity
-1.00	0.000	1.000
1.00	0.006	1.000
2.50	0.026	1.000
3.50	0.038	1.000
4.50	0.064	1.000
5.50	0.141	1.000
6.50	0.212	1.000
7.50	0.282	1.000
8.50	0.359	1.000
9.50	0.429	1.000
10.50	0.532	0.991
11.50	0.596	0.985
12.50	0.718	0.959
13.50	0.808	0.904
14.50	0.853	0.805
15.50	0.910	0.689
16.50	0.968	0.520
17.50	0.987	0.218
19.00	1.000	0.000

and specificity for the FAB was obtained with a cutoff point of 13.5 (sensitivity=80.8%, specificity=90.4%).

Finally, we evaluated the sample matched by age and education level. The optimal balance between sensitivity and specificity for the FAB was again obtained with a cutoff point of 13.5 (Table 4 and Figure 2).

Influence of sociodemographic variables in the healthy controls

Multiple regression analysis was used to test whether sociodemographic variables (i.e., gender, age, and years of education) significantly predicted FAB-Ch performance in the normative sample. The results of the regression indicated these predictors explained 34.9% of the variance [r²=0.349, F(3,344)=60.796, p<0.001]. Both education [β =0.569, t(344)=12.831, p<0.01] and age [β =-0.127, t(344)=-0.127, p<0.01] significantly predicted FAB-Ch score. Based on this analysis, we calculated an FAB-Ch predicted value for each patient using the formula: 13.977-0.012×age (years)+0.218×education (years). We then subtracted the patient's actual score on the FAB-Ch score from the predicted score. The mean difference between FAB-Ch observed score (10.06±3.86) and the FAB-Ch predicted score (15.79±1.03) was -5.73 (SD=3.67). This value is significantly different from zero [t(155)=-19.53, p<0.001].

Normative data in the healthy control group

We created a table of normative values based only on age and education. Table 5 shows the normative data for total scores for the FAB-Ch in the HC group.

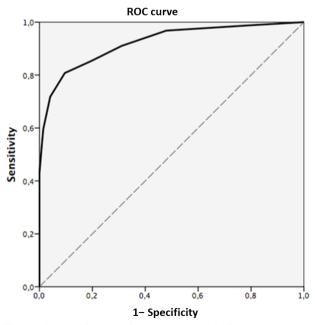


Figure 1. Receiver Operating Characteristic curve for Frontal Assessment Battery score in healthy controls and dementia patients. **Table 4.** Sensitivity and specificity of the Chilean version of the Frontal Assessment Battery on healthy controls (n=122) and patients (n=128) matched by age and education level.

Cutoff point	Sensitivity	Specificity
-1.00	0.000	1.000
1.00	0.008	1.000
2.50	0.025	1.000
3.50	0.042	1.000
4.50	0.051	1.000
5.50	0.110	1.000
6.50	0.195	1.000
7.50	0.263	1.000
8.50	0.339	1.000
9.50	0.407	1.000
10.50	0.517	0.992
11.50	0.568	0.975
12.50	0.686	0.926
13.50	0.780	0.852
14.50	0.822	0.762
15.50	0.890	0.664
16.50	0.958	0.434
17.50	0.983	0.164
19.00	1.000	0.000

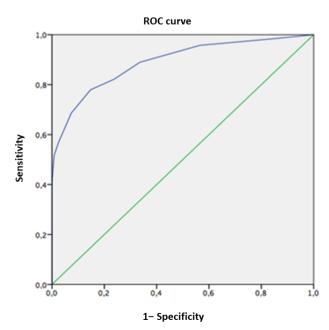


Figure 2. Receiver Operating Characteristic curve for Frontal Assessment Battery score in healthy controls and dementia patients matched by age and education level.

DISCUSSION

This report describes the standardization of the FAB-Ch in a Chilean sample of an HC and patients with dementia syndrome. We provide psychometric evidence and normative data of this instrument.

Regarding psychometric properties, the FAB-Ch has strong evidence of reliability based on internal consistency, similar to the data reported in previous studies^{5,18}. The Chilean version has a high correlation with two measures of EFs: the WCST and categorical fluency, which provides an acceptable validity based on the relationship with other variables. However, it is important to highlight that categorical fluency is not a pure executive test and depends also on language and semantic memory²⁷⁻²⁹. Nevertheless, Junquera et al.³⁰ showed that the executive component of this instrument significantly predicted conversion to dementia (1 year later) in patients with mild cognitive impairment who presented a dysexecutive phenotype, independently of impairment at baseline. This result is consistent with other studies with different populations showing that the FAB-Ch has appropriate convergent validity for testing frontal lobe function^{13,17}.

We also found that the FAB-Ch strongly correlated with the MMSE, which is a measure of global cognitive function, which is different from previous results^{5,13,31,32}. These results are unexpected since the MMSE does not formally evaluate EFs³³. One possible explanation for this finding is the interaction between education and MMSE performance, with the former being associated with the FAB-Ch¹³. An alternative explanation is that FAB is sensitive to the disease progression, making it useful to monitor the clinical course of dementing diseases.

The factor analysis identified a single factor explaining most of the variance of the FAB-Ch, similar to previous findings³⁴. The optimal balance between sensitivity and specificity for the FAB-Ch was obtained with a cutoff point of 13.5, highlighting that this test can discriminate between HC and people with dementia syndrome.

Performance on the FAB-Ch is explained by education and age, while gender does not contribute to performance. Cognitive aging is associated with a mild decline in $EF^{35,36}$, and education affects performance on executive tests^{37,38}. Our results are consistent with previous data on the effect of sociodemographic factors on the FAB-Ch^{7,8,10,11,34,39,40}.

The availability of a normative sample including people with a wide range of educational levels is essential for using FAB-Ch in clinical practice, especially in countries like Chile, where the range of educational levels in the populations is very heterogeneous wide⁸.

Age	n	Education	Mean	SD	Median	Maximum	Minimum
20–49	30	0–8	15.00	1.23	15.00	17.00	13.00
	29	9–12	15.86	1.83	16.00	18.00	10.00
	66	13 or +	17.11	1.14	17.00	18.00	13.00
50–69	30	0–8	14.17	1.76	14.00	17.00	10.00
	25	9–12	16.04	1.67	16.00	18.00	12.00
	63	13 or +	16.79	1.25	17.00	18.00	13.00
70–89	26	0–8	14.00	1.72	14.00	17.00	10.00
	22	9–12	16.14	1.32	16.00	18.00	13.00
	53	13 or +	16.74	1.26	17.00	18.00	12.00

Table 5. Normative data for the Frontal Assessment Battery score total.

SD: standard deviation.

Several limitations warrant consideration in generalizing our observations. First, although we have participants of different ages and educational ranges, the variability of the data is small, which could impact on the relative position of an individual concerning standard scores. Therefore, assessors interpreting FAB-Ch scores should always review the overall distribution of scores on this test and consider the raw score obtained by the individual, which could be especially important when, for example, trying to determine if a person's score is far outside the normal range⁴¹. Second, the main limitation of our study is that we only provide indirect evidence of the ability of the FAB-Ch to detect a dysexecutive syndrome (validity based on the relationship with other variables). We did not provide specific evidence of the utility of the FAB-Ch in the diagnosis of a dysexecutive syndrome. In this way, it is important to note that as has been highlighted for other screening instruments, FAB-Ch cannot lead to the specific diagnosis of the type of dementia, such as Alzheimer's disease or frontotemporal dementia⁴. The aim of FAB-Ch is to establish the presence and degree of severity in a specific domain (not the type of diagnosis)⁴. Emphasizing this limitation is particularly important since executive dysfunction is present in many dementia syndromes (e.g., Lewy body dementia, vascular dementia, frontotemporal dementia, and Alzheimer's disease)⁴²⁻⁴⁴.

In this study, we did not consider types of dementia in the analysis as its aim was to investigate the sensitivity and specificity of this screening tool relative to FC. In this line, we do not have measures of the level of severity of dementia from the point of view of functionality or a global level of severity of dementia such as the Global Deterioration Scale (GDS). However, we have the MMSE, a cognitive screening test that is widely used as a measure of cognitive severity, which can reduce this limitation

In addition, illiterate subjects were excluded, and only 13 participants of the HC have 4 years of education or below. Therefore, our norms have limited use for people with low educational level, who are still an important percentage of the Latin American population⁴⁵. More studies are needed to establish norms in subjects with a low socioeconomic status. This study included only Chilean subjects, consequently limiting our data to other Spanish-speaking countries. Yet, recent normative data for 10 Spanish-language neuropsychological tests in 11 Latin American countries suggest that most of the differences in test performance are explained by age and educational factors. Inter-country factors only account for a small proportion of variance⁴⁶. Finally, the inclusion of HC whose performances are 24 or higher on the MMSE could be criticized as too strict. However, as 95.9% of our sample has more than 4 years of education, this criterion ensured the inclusion of healthy subjects without cognitive impairment⁴⁷.

In conclusion, the main results of our study are (a) the FAB-Ch is an instrument with strong evidence of reliability and validity based on international standard, (b) an adequate diagnosis utility for dementia, (c) the effect of aging and level of education on FAB-Ch performances, and (d) the availability of normative data for the FAB-Ch, improving the usefulness of this instrument in clinical settings. In addition to other tests such as the MMSE, the administration of the FAB-Ch allows a more comprehensive evaluation in the diagnosis process of dementia. Future studies need to address if FAB-Ch presents good diagnostic utility to show the degree of executive dysfunction and its contribution in the differential diagnosis of types dementia.

ACKNOWLEDGMENTS

Study data were partially collected and managed using RedCap electronic data capture tools hosted at Faculty of Medicine, University of Chile, GERO, Memory and Neuropsychiatry Clinic, Faculty of Medicine Hospital del Salvador, University of Chile, and German Clinic.

Authors' contributions. FG: conceptualization, data curation, formal analysis, investigation, methodology, project administration, and writing – original draft. DM, PA, AA, PL, CD, and RR: investigation, supervision, and validation. MP: conceptualization, formal analysis, investigation, methodology, supervision, validation,

and writing – review & editing. LO: conceptualization, data curation, investigation, and writing – review & editing. DH: investigation, methodology, supervision, and validation. MT: conceptualization, formal analysis, investigation, methodology, methodology, supervision, validation, writing – original draft, and writing – review & editing. OL: formal analysis, investigation, methodology, supervision, and validation. JB: conceptualization, investigation, methodology, and supervision. AS: conceptualization, data curation, formal analysis, funding acquisition, investigation, methodology, project administration, supervision, validation, writing – original draft, and writing – review & editing.

REFERENCES

- Ferguson HJ, Brunsdon VEA, Bradford EEF. The developmental trajectories of executive function from adolescence to old age. Sci Rep. 2021;11(1):1382. https://doi.org/10.1038/s41598-020-80866-1
- Hwang K, Bruss J, Tranel D, Boes AD. Network localization of executive function deficits in patients with focal thalamic lesions. J Cogn Neurosci. 2020;32(12):2303-19. https://doi.org/10.1162/jocn_a_01628
- Orellana G, Slachevsky A. Executive functioning in schizophrenia. Front Psychiatry. 2013;4:35. https://doi.org/10.3389/fpsyt.2013.00035
- Della Sala S, Morris RG. When no more research is needed (without further reflection). Cortex. 2020;123:A1. https://doi.org/10.1016/j.cortex.2019.12.018
- Dubois B, Slachevsky A, Litvan I, Pillon B. The FAB: a Frontal Assessment Battery at bedside. Neurology. 2000;55(11):1621-6. https://doi. org/10.1212/wnl.55.11.1621
- Coen RF, McCarroll K, Casey M, McNulty H, Laird E, Molloy AM, et al. The Frontal Assessment Battery: Normative Performance in a Large Sample of Older Community-Dwelling Hospital Outpatient or General Practitioner Attenders. J Geriatr Psychiatry Neurol. 2016;29(6):338-43. https://doi. org/10.1177/0891988716666381
- Beato R, Amaral-Carvalho V, Guimarães HC, Tumas V, Souza CP, Oliveira GN de, et al. Frontal assessment battery in a Brazilian sample of healthy controls: normative data. Arq Neuro-Psiquiatr. 2012;70(4):278-80. https:// doi.org/10.1590/s0004-282x2012005000009
- Kim TH, Huh Y, Choe JY, Jeong JW, Park JH, Lee SB, et al. Korean version of frontal assessment battery: psychometric properties and normative data. Dement Geriatr Cogn Disord. 2010;29(4):363-70. https:// doi.org/10.1159/000297523
- Kugo A, Terada S, Ata T, Ido Y, Kado Y, Ishihara T, et al. Japanese version of the Frontal Assessment Battery for dementia. Psychiatry Res. 2007;153(1):69-75. https://doi.org/10.1016/j.psychres.2006.04.004
- Appollonio I, Leone M, Isella V, Piamarta F, Consoli T, Villa ML, et al. The Frontal Assessment Battery (FAB): normative values in an Italian population sample. Neurol Sci Off J Ital Neurol Soc Ital Soc Clin Neurophysiol. 2005;26(2):108-16. https://doi.org/10.1007/s10072-005-0443-4
- Benke T, Karner E, Delazer M. FAB-D: German version of the Frontal Assessment Battery. J Neurol. 2013;260(8):2066-72. https://doi. org/10.1007/s00415-013-6929-8
- Slachevsky A, Villalpando JM, Sarazin M, Hahn-Barma V, Pillon B, Dubois B. Frontal assessment battery and differential diagnosis of frontotemporal dementia and Alzheimer disease. Arch Neurol. 2004;61(7):1104-7. https:// doi.org/10.1001/archneur.61.7.1104
- Mok VCT, Wong A, Yim P, Fu M, Lam WWM, Hui AC, et al. The validity and reliability of chinese frontal assessment battery in evaluating executive dysfunction among Chinese patients with small subcortical infarct. Alzheimer Dis Assoc Disord. 2004;18(2):68-74. https://doi.org/10.1097/01. wad.0000126617.54783.7
- Lima CF, Meireles LP, Fonseca R, Castro SL, Garrett C. The Frontal Assessment Battery (FAB) in Parkinson's disease and correlations with formal measures of executive functioning. J Neurol. 2008;255(11):1756-61. https://doi.org/10.1007/s00415-008-0024-6

- Rodriguez del Alamo A, Catalán Alonso MJ, Carrasco Marín L. FAB: a preliminar Spanish application of the frontal assessment battery to 11 groups of patients. Rev Neurol. 2003;36(7):605-8. PMID: 12666037
- Kudiaki C, Aslan A. Executive functions in a Turkish sample: associations with demographic variables and normative data. Appl Neuropsychol. 2008;15(3):194-204. https://doi. org/10.1080/09084280802324416
- Wang T-L, Hung Y-H, Yang C-C. Psychometric properties of the Taiwanese (Traditional Chinese) version of the Frontal Assessment Battery: a preliminary study. Appl Neuropsychol Adult. 2016;23(1):11-20. https:// doi.org/10.1080/23279095.2014.995792
- Asaad S, Ashrafi F, Omidbeigi M, Nasiri Z, Pakdaman H, Amini-Harandi A. Persian version of frontal assessment battery: Correlations with formal measures of executive functioning and providing normative data for Persian population. Iran J Neurol. 2016;15(1):16-22. PMID: 27141272
- Barulli MR, Fontana A, Panza F, Copetti M, Bruno S, Tursi M, et al. Frontal assessment battery for detecting executive dysfunction in amyotrophic lateral sclerosis without dementia: a retrospective observational study. BMJ Open. 2015;5(9):e007069. https://doi.org/10.1136/ bmjopen-2014-007069
- Tombaugh TN, McIntyre NJ. The Mini-Mental State Examination: A comprehensive review. J Am Geriatr Soc. 1992;40(9):922-35. https:// doi.org/10.1111/j.1532-5415.1992.tb01992.x
- Aschiero MB, Aguilar MJG, Grasso L. Influencia de variables sociodemográficas en el rendimiento de la Frontal Assessment Battery en adultos mayores Argentinos. Rev Iberoam Psicol. 2019;12(2):115-24. https://doi. org/10.33881/2027-1786.rip.12211
- Yesavage JA, Brink TL, Rose TL, Lum O, Huang V, Adey M, et al. Development and validation of a geriatric depression screening scale: a preliminary report. J Psychiatr Res. 1983 de 1982;17(1):37-49. https:// doi.org/10.1016/0022-3956(82)90033-4
- Zung WW. A rating instrument for anxiety disorders. Psychosom J Consult Liaison Psychiatry. 1971;12(6):371-9. https://doi.org/10.1016/S0033-3182(71)71479-0
- Bruno D, Slachevsky A, Fiorentino N, Rueda DS, Bruno G, Tagle AR, et al. Argentinian/Chilean validation of the Spanish-language version of Addenbrooke's Cognitive Examination III for diagnosing dementia. Neurología. 2020;35(2):82-8. https://doi.org/10.1016/j.nrleng.2019.03.009
- Carlson KD, Herdman AO. Understanding the Impact of Convergent Validity on Research Results. Organ Res Methods. 2012;15(1):17-32. https://doi.org/10.1177/1094428110392383
- Polo TC, Miot HA. Use of ROC curves in clinical and experimental studies. J Vasc Bras. 2020;19:e20200186. https://doi.org/10.1590/1677-5449.200186
- Cosentino S, Scarmeas N, Albert SM, Stern Y. Verbal fluency predicts mortality in Alzheimer disease. Cogn Behav Neurol. 2006;19(3):123-9. https://doi.org/10.1097/01.wnn.0000213912.87642.3d
- Shao Z, Janse E, Visser K, Meyer AS. What do verbal fluency tasks measure? Predictors of verbal fluency performance in older adults. Front Psychol. 2014;5:772. https://doi.org/10.3389/fpsyg.2014.00772

- Amunts J, Camilleri JA, Eickhoff SB, Heim S, Weis S. Executive functions predict verbal fluency scores in healthy participants. Sci Rep. 2020;10:11141. https://doi.org/10.1038/s41598-020-65525-9
- Junquera A, García-Zamora E, Olazarán J, Parra MA, Fernández-Guinea S. Role of executive functions in the conversion from mild cognitive impairment to dementia. J Alzheimers Dis. 2020;77(2):641-53. https://doi. org/10.3233/JAD-200586
- Beato RG, Nitrini R, Formigoni AP, Caramelli P. Brazilian version of the Frontal Assessment Battery (FAB): Preliminary data on administration to healthy elderly. Dement Neuropsychol. 2007;1(1):59-65. https://doi. org/10.1590/S1980-57642008DN10100010
- Lipton AM, Ohman KA, Womack KB, Hynan LS, Ninman ET, Lacritz LH. Subscores of the FAB differentiate frontotemporal lobar degeneration from AD. Neurology. 2005;65(5):726-31. https://doi.org/10.1212/01. wnl.0000174437.73416.7b
- Kahokehr A, Siegert RJ, Weatherall M. The frequency of executive cognitive impairment in elderly rehabilitation inpatients. J Geriatr Psychiatry Neurol. 2004;17(2):68-72. https://doi.org/10.1177/0891988704264536
- Graver CJ, Hajek CA, Bieliauskas LA. An investigation of impaired scores on the frontal assessment battery in a VA population. Appl Neuropsychol. 2011;18(4):278-83. https://doi.org/10.1080/09084282.2011.595454
- Gerstorf D, Siedlecki KL, Tucker-Drob EM, Salthouse TA. Executive dysfunctions across adulthood: measurement properties and correlates of the DEX self-report questionnaire. Neuropsychol Dev Cogn B Aging Neuropsychol Cogn. 2008;15(4):424-45. https://doi. org/10.1080/13825580701640374
- Sathouse TA, Atkinson TM, Berish DE. Executive functioning as a potential mediator of age-related cognitive decline in normal adults. J Exp Psychol Gen. 2003;132(4):566-94. https://doi.org/10.1037/0096-3445.132.4.566
- Ostrosky-Solis F, Ardila A, Rosselli M, Lopez-Arango G, Uriel-Mendoza V. Neuropsychological Test Performance in Illiterate Subjects. Arch Clin Neuropsychol. 1998;13(7):645-60. https://doi.org/10.1093/arclin/13.7.645
- Tripathi R, Kumar K, Bharath S, Marimuthu P, Varghese M. Age, education and gender effects on neuropsychological functions in healthy Indian older adults. Dement Neuropsychol. 2014;8(2):148-54. https://doi.org/10.1590/ S1980-57642014DN82000010

- Cunha PJ, Nicastri S, de Andrade AG, Bolla KI. The frontal assessment battery (FAB) reveals neurocognitive dysfunction in substance-dependent individuals in distinct executive domains: Abstract reasoning, motor programming, and cognitive flexibility. Addict Behav. 2010;35(10):875-81. https://doi.org/10.1016/j.addbeh.2010.05.005
- Iavarone A, Ronga B, Pellegrino L, Loré E, Vitaliano S, Galeone F, et al. The Frontal Assessment Battery (FAB): normative data from an Italian sample and performances of patients with Alzheimer's disease and frontotemporal dementia. Funct Neurol. 2004;19(3):191-5. PMID: 15595714
- Suhr JA, Angers K. Neuropsychological Testing and Assessment. In: Sellbom M, Suhr JA, editors. The Cambridge Handbook of Clinical Assessment and Diagnosis. Cambridge: Cambridge University Press; 2019. p. 191–207. (Cambridge Handbooks in Psychology).
- Amanzio M, Bartoli M, Cipriani GE, Palermo S. Executive Dysfunction and Reduced Self-Awareness in Patients With Neurological Disorders. A Mini-Review. Front Psychol. 2020;11:1697. https://doi.org/10.3389/ fpsyg.2020.01697
- D'Onofrio G, Panza F, Sancarlo D, Addante F, Solfrizzi V, Cantarini C, et al. Executive dysfunction detected with the Frontal Assessment Battery in Alzheimer's disease versus vascular dementia. J Alzheimers Dis. 2018;62(2):699-711. https://doi.org/10.3233/JAD-170365
- Firbank M, Kobeleva X, Cherry G, Killen A, Gallagher P, Burn DJ, et al. Neural correlates of attention-executive dysfunction in lewy body dementia and Alzheimer's disease. Hum Brain Mapp. 2016;37(3):1254-70. https:// doi.org/10.1002/hbm.23100
- Arango-Lasprilla JC, Rivera D, Garza MT, Saracho CP, Rodríguez W, Rodríguez-Agudelo Y, et al. Hopkins Verbal Learning Test- Revised: Normative data for the Latin American Spanish speaking adult population. NeuroRehabilitation. 2015;37(4):699-718. https://doi.org/10.3233/NRE-151286
- Guàrdia-Olmos J, Peró-Cebollero M, Rivera D, Arango-Lasprilla JC. Methodology for the development of normative data for ten Spanish-language neuropsychological tests in eleven Latin American countries. NeuroRehabilitation. 2015;37(4):493-9. https://doi.org/10.3233/NRE-151277
- Gonzalez J, Aguilar L, Oporto S, Araneda L, Vásquez M, von Bernhardi R. Normalización del "Mini-Mental State Examination" según edad y educación, para la población de Santiago de Chile. Rev Memoriza. 2009;3:23-34.

APPENDIX 1. CHILEAN'S VERSION OF THE FAB

1. Semejanzas (conceptualización).

"¿En qué se parecen...?"

a. Un plátano y una naranja.

- b. Una mesa y una silla
- c. Un tulipán, una rosa y una margarita.

Ayudar al paciente en caso de fracaso total "no se parecen" o parcial "los 2 tienen cáscara" en el primer ítem, no en los siguientes. Sólo las respuestas de categoría (frutas, muebles, flores) se consideran correctas. Puntaje: 3 correctas=3; 2 correctas=2; 1 correcta=1; ninguna correcta=0.

2. Fluidez léxica (flexibilidad mental).

"Diga todas las palabras que pueda (por ejemplo animales, plantas y objetos, pero no nombres propios ni apellidos) que comiencen con A". Si no responde en los primeros 5 segundos decirle "por ejemplo, árbol". Si se detiene por más de 10 segundos, insista "cualquier palabra que empiece con A". Tiempo: 60 segundos. Las repeticiones, derivaciones (árbol, arbolito), nombres propios y apellidos no se cuentan.

Puntaje: 10 o más palabras=3; 6 a 9=2; 3 a 5=1; menos de 3=0.

3. Secuencias motoras (programación).

"Mire con atención lo que hago"; el examinador frente al paciente realiza 3 veces la prueba de Luria (golpear con nudillo, canto y palma) con su mano izquierda. *"Con su mano derecha haga lo mismo que yo, primero juntos, después solo".* El examinador hace la serie 3 veces con el paciente y le dice *"ahora haga lo mismo Ud. solo"*.

Puntaje: 6 series consecutivas correctas=3; 3 a 5 series correctas=2; no lo hace solo, pero sí 3 series consecutivas con el examinador=1; no logra ni siquiera imitar 3 veces=0.

4. Instrucciones conflictivas (sensibilidad a la interferencia).

"Cuando yo golpeo 1 vez, debe golpear 2 veces"; para asegurar que comprendió las instrucciones, se hace una serie de 3 ensayos: 1-1-1. *"Cuando yo golpeo 2 veces, debe golpear una";* para asegurar que comprendió las instrucciones, se hace una serie de 2-2-2. El examinador realiza la siguiente serie: 1-1-2-1-2-2-1-1-2.

Puntaje: sin errores=3; 1 o 2 errores=2; más de 2 errores=1; si golpea igual que el examinador al menos 4 veces consecutivas=0.

5. Go-no Go (control inhibitorio).

"Cuando yo golpeo 1 vez, debe golpear 1 vez"; para asegurar que comprendió la instrucción, se hace una serie de 3 ensayos: 1-1-1. *"Cuando yo golpeo 2 veces, no debe golpear";* para asegurar que comprendió la instrucción, se hace una serie de 3 ensayos: 2-2-2. El examinador realiza la siguiente serie: 1-1-2-1-2-2-2-1-1-2.

Puntaje: sin errores=3; 1 o 2 errores=2; más de 2 errores=1; golpea igual que el examinador al menos 4 veces seguidas=0.

6. Conducta de prehensión (autonomía del ambiente).

El examinador se sienta frente al paciente, que tiene las manos sobre sus rodillas, con las palmas hacia arriba. El examinador acerca lentamente sus manos hasta tocar las del paciente para ver si se las toma espontáneamente. Si lo hace, dice *"ahora, no me tome las manos"* y vuelve a tocárselas.

Puntaje: no le toma las manos=3; duda o pregunta qué tiene que hacer=2; las toma sin vacilar=1; las toma aún después de decirle que no lo haga=0.

Puntaje:

Semejanzas=3 -2 -1 -0
 Fluencia lexical=3 -2 -1 -0
 Secuencias motoras=3 -2 -1 -0
 Instrucciones conflictivas=3 -2 -1 -0
 Go-no go=3 -2 -1 -0
 Conducta de prehension=3 -2 -1 -0
 Total= /18