

Methodology for the development of normative data for ten Spanish-language neuropsychological tests in eleven Latin American countries

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en Cervell, Cognició i Conducta, Universitat de Barcelona, Espanya, Spain

^b*Faculty of Psychology and Education, University of Deusto, Bilbao, Spain*

^c*IKERBASQUE, Basque Foundation for Science, Bilbao, Spain*

Abstract.

BACKGROUND: Within the field of neuropsychology, there is a significant lack of normative data for individuals in Latin America.

OBJECTIVE: To describe the methodology utilized to obtain the data and create norms for 10 Spanish-language neuropsychological tests administered in 11 Latin-American countries in a sample of 3,977 healthy individuals between the ages 18 and 90.

METHOD: The same data manipulation process was applied to the data collected (regardless of the scale or country) using a regression-based procedure that takes into account sex, age, and educational influences on neuropsychological test scores.

CONCLUSIONS: Following this procedure, we were able to generate age, education, and sex (if relevant) based norms for each test in each of the 11 countries studied. These norms are presented in the 10 articles that comprise this special issue.

Keywords: Psychometric, norms, Latin America, neuropsychological tests

1. Introduction

Most psychological and neuropsychological tests are built using the Classical Test Theory (CTT), from this perspective norms are created using simple procedures like transformation in percentile scales, or T and D scores (Abad, Olea, Ponsoda, & García, 2011; Crawford, 2003). Within the field of neuropsychology, there

is a significant lack of norms for Spanish-speakers in Latin America. This article presents the methodology employed to standardize various neuropsychological tests administered in 11 Latin American countries.

There is extensive literature describing which demographic variables (e.g., sex, age, socio-economic status, and years of education) must be taken into account to generate appropriate norms for neuropsychological tests. It is widely accepted that the systematic error generated by both chronological age and education needs must be controlled for (Salthouse, 2000, 2001, 2009; Salthouse, Atkinson, & Berish, 2003). Therefore, there are several ways to control for age and education effects:

*Address for correspondence: Joan Guàrdia-Olmos, Department of Methodology of Behavioral Sciences, Faculty of Psychology, University of Barcelona, Passeig de la Vall d'Hebron, 171, 08035 Barcelona, Spain. Tel.: +34 933125099; Fax: +34 934021359; E-mail: jguardia@ub.edu.

1) A method that involves the adjustment of control systems for error correction, based on the generation of normative data for each reference group. In this procedure, it would be necessary to use the stratum system to identify normative groups based on classic standardization. Though this option is easy, it generates an enormous quantity of normative tables that would turn the final product into a measureless and useless contribution from the point of view of applied clinical research (Brooks, Strauss, Sherman, Iverson, & Slick, 2009; Kim et al., 2014; La Cour & Andersen, 2006; La Paglia et al., 2014; Mungas, Reed, Crane, Haan, & González, 2004; Peintinger & Klünemann, 2013; Salvadori, Poggesi, Pracucci, Inzitari, & Pantoni, 2015), 2) The score correction obtained from the use of partial regression coefficients from a linear model (Blesa et al., 2001; Villaseñor, Guàrdia, Jiménez, Rizo, & Però, 2010), and 3) The use of regression coefficients combined with the standard deviation of the residual to obtain normative data based on a percentile scale (Van der Elst, Dekker, Hurks, & Jolles, 2012; Van der Elst, Hurks, Wassenberg, Meijs, & Jolles, 2011; Van der Elst, Van Boxtel, Van Breukelen, & Jolles, 2006).

We had to choose a statistical approach to standardize the neuropsychological tests given the final resulting sample. We found an approach that would allow us to take into account a number of issues. For instance, the sample was obtained from a large number of countries and each country had slightly different distributions of age and educational levels. Furthermore, given that tests administered to these samples have different response ranges, as well as individual differences across countries, and well-known floor and ceiling effects made it more difficult to obtain normal distributions of acceptable symmetry values for our sample (Hartshorne & Germine, 2015; Schroeder & Salthouse, 2004). Therefore, we used the standardization procedure described by Van der Elst et al. (2006, 2011, 2012) in which regression coefficients are combined with the standard deviation of the residual.

Given the issues pertaining to this very heterogeneous sample of participants from 11 countries across Latin America, using regression coefficients combined with the standard deviation of the residual to obtain normative data based on a percentile scale seems to be the best procedure. Conventional norms tables will be generated in a quantity that is easy to handle and has greater clinical utility. Additionally, this procedure proposes correction systems that take real observed distributions into account instead of a series of arbitrary strata. In such a way they solve the ceiling and floor

effect problems. Finally, the procedure allows for a specific adjustment depending on the cohort studied (Van der Elst et al., 2006, 2011, 2012). Though this procedure does not radically solve the issues of asymmetric distributions, it does make it possible to take them into account, as the predictions of scale setting are based on the establishment of confidence intervals close to the average values (Russell, Russell, & Hill, 2005).

In light of all of the above, in this paper we will explain the methodology based on ordered regression coefficients and statistical models, with the intention to generate normative data and standardize different neuropsychological test in 11 Spanish-speaking countries from Latin America.

2. Method

2.1. Participants

The present study was conducted with a sample of 5,402 healthy individuals from 20 cities in 12 Spanish-speaking countries in Latin America (Argentina, Bolivia, Chile, Colombia, Cuba, El Salvador, Guatemala, Honduras, Mexico, Paraguay, Peru, and Puerto Rico). The results obtained from 1,425 Colombians have been published elsewhere (Arango & Rivera, 2015). Therefore, this article will present the procedures used to generate normative data from 3,977 healthy individuals from 14 cities for the remaining 11 countries.

The sample was designed taking into account the literacy level, the percentage of people with primary, secondary and tertiary studies, and the age distribution for each country. Thus, an empirical quota sampling was used, in order to have a statistical precision between 0.063 to 0.049 with a 95% confidence level estimated under the situation of maximum uncertainty ($\pi = 1 - \pi = 0.5$) (see Table 1). Due to the low number of participants across the three education levels, years of education was recoded into two groups (1 to 12 years and >12 years). These two groups differentiate between people with low and medium and those with a high level of education. A description of the sample differentiated by this codification of years of education and country of origin is shown in Table 2.

For all the countries, the inclusion and exclusion criteria were the same. The inclusion criteria were: a) were between 18 to 95 years of age, b) were born and currently lived in the country where the protocol was conducted, c) spoke Spanish as their

Table 1
Summary of the sampling designed and final sample obtained

Country	Age in years	Designed sample			Final sample		
		Years of education			Years of education		
		<6	6 to 12	>12	<6	6 to 12	>12
Argentina	<56	60	50	30	0	83	128
	56 a 75	60	50	30	1	49	40
	>75	60	40	20	3	12	4
Bolivia	<56	50	35	20	43	37	21
	56 a 75	50	35	20	40	39	18
	>75	50	30	10	37	30	9
Chile	<56	60	50	30	33	64	50
	56 a 75	60	50	30	37	58	20
	>75	60	40	20	29	20	9
Cuba	<56	60	50	30	0	109	39
	56 a 75	60	50	30	8	83	25
	>75	60	40	20	3	31	8
El Salvador	<56	50	35	20	34	37	22
	56 a 75	50	35	20	34	35	22
	>75	50	30	10	33	30	10
Guatemala	<56	50	35	20	15	48	41
	56 a 75	50	35	20	14	42	37
	>75	50	30	10	2	12	3
Honduras	<56	50	35	20	30	39	29
	56 a 75	50	35	20	30	26	14
	>75	50	30	10	8	7	1
Mexico	<56	225	225	100	170	262	154
	56 a 75	225	225	100	155	249	105
	>75	125	125	50	73	96	36
Paraguay	<56	50	35	20	13	71	35
	56 a 75	50	35	20	29	97	12
	>75	50	30	10	3	3	0
Peru	<56	60	50	30	4	46	117
	56 a 75	60	50	30	6	25	28
	>75	60	40	20	0	6	13
Puerto Rico	<56	40	35	20	4	65	85
	56 a 75	40	35	20	10	68	41
	>75	41	30	10	5	8	8

Table 2
Sample distribution by age, education and gender

	n Total	Age	Education		Gender	
		Mean (SD)	1 to 12 n (%)	>12 n (%)	Male n (%)	Female n (%)
Argentina	320	45.7 (19.5)	148 (46.3%)	172 (53.8%)	96 (30.0%)	224 (70.0%)
Bolivia	274	55.8 (22.0)	226 (82.5%)	48 (17.5%)	99 (36.1%)	175 (63.9%)
Chile	320	55.1 (19.6)	241 (75.3%)	79 (24.7%)	134 (41.9%)	186 (58.1%)
Cuba	306	53.0 (19.7)	234 (76.5%)	72 (23.5%)	142 (46.4%)	164 (53.6%)
El Salvador	257	56.0 (20.7)	203 (79.0%)	54 (21.0%)	100 (38.9%)	157 (61.1%)
Guatemala	214	53.2 (17.4)	133 (62.1%)	81 (37.9%)	95 (44.4%)	119 (55.6%)
Honduras	184	48.6 (18.8)	140 (76.1%)	44 (23.9%)	67 (36.4%)	117 (63.6%)
Mexico	1300	52.5 (20.5)	1005 (77.3%)	295 (22.7%)	431 (33.2%)	869 (66.8%)
Paraguay	263	53.0 (14.8)	216 (82.1%)	47 (17.9%)	101 (38.4%)	162 (61.6%)
Peru	245	43.4 (20.6)	87 (35.5%)	158 (64.5%)	87 (35.5%)	158 (64.5%)
Puerto Rico	294	50.9 (18.5)	160 (54.4%)	134 (45.6%)	126 (42.9%)	168 (57.1%)

native language, d) had completed at least one year of formal education, e) be able to read and write at the moment of evaluation. f) scored ≥ 23 on the Mini-Mental State Examination (MMSE, Folstein,

Folstein, & McHugh, 1975; Ostrosky-Solís, López-Arango, & Ardila, 2000; Villaseñor-Cabrera, Guàrdia-Olmos, Jiménez-Maldonado, Rizo-Curiel, & Peró-Cebollero, 2010), g) scored ≤ 4 on the Patient

Health Questionnaire–9 (PHQ-9, Kroenke, Spitzer, & Williams, 2001), and h) scored ≥ 90 on the Barthel Index (Mahoney & Barthel, 1965).

The exclusion criteria were: a) having a personal history of central nervous disease (stroke, epilepsy, multiple sclerosis, brain tumour, severe head trauma, etc.); b) having history of alcohol abuse or other psychotropic substances; c) having an active systemic disease or uncontrolled disease associated with cognitive impairment (diabetes mellitus, hypothyroidism, B12 vitamin deficit); d) having a history of psychiatric illness (major depression, bipolar disease, psychosis, etc.); e) having severe sensory deficits (vision and/or auditory loss) that could affect the administration of the test or the participants' performance in these test; f) Use of psychiatric or other drugs that could affect one's cognitive performance; and g) taking medications for chronic pain (e.g. Monoamine Oxidase Inhibitors – MAOI).

2.2. Instruments

A sociodemographic questionnaire was created for this project in order to collect data on age, years of education, sex, laterality (right, left handed, ambidextrous), residence zone (rural, urban), race (white, black, mestizo, indigenous), employment status (employed, un employed, student, retired, housekeeping), and marital status (single, married, cohabited, separated, divorced, widower). Then, all of the individuals were administered a comprehensive neuropsychological evaluation that was composed of the 10 neuropsychological tests:

1. Rey-Osterrieth Complex Figure test (ROCF; Rey, 2009).
2. Stroop colour and words test (Golden, 2010).
3. Modified Wisconsin card sorting test (M-WCST; Nelson, 1976; Schretlen, 2010).
4. Trial making test (TMT A-B; Reitan & Wolfson, 1985).
5. Brief test of attention (BTA; Schretlen, 1997).
6. Phonological and semantical verbal fluency test.
7. Boston naming test (Goodglas, Kaplan, & Barresi, 2005).
8. Symbol digit modalities test (SDMT; Smith, 2002).
9. Hopkins verbal learning test - Revised (HVLt-R; Benedict, Schretlen, Groninger, & Brandt, 1998).
10. Test of memory malingering (TOMM; Tombaugh, 2011).

2.3. Procedure

2.3.1. Training and administration procedure of the battery

The present study began November 1, 2012 with the participation of 15 institutions. Subsequently, the proposal that included the methodology and ethics was drafted and later delivered to the ethics committee of the University of Deusto (Bilbao, Spain). After approval, the publishing manuals, answer sheets and materials (booklets and encouragement cards) for each of the neuropsychological tests were bought. Spanish-language versions of instructions booklets were available for all instruments except for M-WCST and BTA. Spanish administration and instruction manuals for these instruments were created in collaborated with the publishers.

Likewise, tools and visual aids were created to achieve standard management process of the battery. These tools consisted of: a) a randomized list to determine the order of administration of the test for each participant in order to avoid order bias and cognitive conditioning. To do so, the function $f_x = RAND()$ was used in Microsoft Excel[®] and was set to take into account the interaction of the language test and verbal memory test; b) a framework for decision-making in the evaluation process; c) a template in Microsoft Excel[®] for entering information to limit bias input information. The template was designed using the configuration options: data validation = Customized (numeric variables), dropdown lists (categorical variables) and setting formats; d) examples showing the most frequent errors in the administration and scoring of each test, and e) a virtual folder with a security key for each city, administered by the study coordinator to track data entry.

Moreover, each coordinator selected a group of 6–12 undergraduates and/or graduate students, with which the coordinator reviewed the instructions and application and qualification of the tests. Once the group was ready, an online virtual training was performed through telemedicine platform VSee[®]. During the two-hour long training, the administration and scoring of the tests were reviewed and any doubts were resolved.

A pilot test with 40 protocols (these were excluded from the analysis of normative data) was conducted by analyzing data from the first two cases collected in each of the 20 centers in order to determine the adequate functioning for collection of information with the proposed design.

Data collection began March 2013 and ended August 2014. The protocol was administered in a single day and lasted about 70 minutes. Before starting the battery

administration, participants had to sign the informed consent.

Once the database was consolidated, we reviewed the data distribution of frequencies, comparing the values of various statistics and graphs, in order to check for correct processing and characteristics of distributions for each of the variables analyzed. This certified the database was generated correctly and the properties of the observed variables were known.

2.3.2. Normative procedure

An independent *t*-test by country was run for each neuropsychological measure to first determine whether there were any significant differences in test scores based on participant sex within each country. Since the sample size is very large, significant differences for gender in neuropsychological scores were needed with a medium or high effect size ($r > 0.3$, Cohen, 1988) in order to consider that differences in the scores between men and women are relevant. The effect size was obtained using the following expression:

$$r = \sqrt{\frac{t^2}{t^2 + v}}$$

when relevant, country sample sizes were stratified by sex when significant differences were found with medium or high effect sizes.

The lineal regression model was built using the raw (direct/unconverted) score in the scale being normed as the criterion variable. Participant age was included as a predictor variable, and years of education (dichotomized in two groups: 1 to 12 years coded as 0 and >12 years coded as 1) was also considered as a predictor variable for those cases where the differences in the raw scores between men and women met the above-mentioned criteria. All the regressions were run differentiating by country. If any predictive variable was not statistically significant, the linear regression analysis was repeated eliminating that predictive variable. For all linear regression models, the following were assessed: collinearity between the predictive variables, value of variance inflation factor (VIF) near to 1, the residuals normality using the Q-Q plot and the residuals histogram, the homoscedasticity as in the scatter plot of the standardized residuals and the predicted values, and finally, the existence of influent values by Cook's distance.

In order to obtain the exact percentile for an exact score, the procedure used by Van der Elst et al. (2006, 2011, and 2012) was followed. The steps to obtain the exact percentile were:

1. Obtain the predicted value using the regression equation obtained in the country of origin of the person where the original y_i was de score of each neuropsychological test:

$$\hat{y}_i = b_0 + b_1 \cdot Age_i + b_2 \cdot Educational\ level_i + b_3 \cdot Sex$$

2. Obtain the residual value:

$$e_i = y_i - \hat{y}_i$$

3. Standardize the residual obtained (transform to z score). To do this, it is necessary to divide the residual value by the standard deviation for the residual obtained in the fitted regression model.

$$z = \frac{e_i}{SD_e}$$

4. Search the exact probability associated with the z value using accumulated probability of the standardized normal distribution.

It is possible that some clinicians consider this procedure slow and expensive in time. For this reason, in the papers that the different normative data were presented for each scale, we present also tables with the approximated percentile. To do this, following the procedure used by Van der Elst et al. (2006, 2011, 2012), we used class mark age value starting with 20 years \pm 2 years until a class mark of 80 years old. For each class mark age and educational level (1: more than 12 years or 0:1 to 12 years) we apply the procedure explained in order to obtain the percentile. In this case we do not have the percentile for each age in years, we have the approximation for intervals of 5 years instead.

3. Limitations and future directions

Among the limitations encountered in the present study, there was an observed low symmetry in sample sizes between males and females for certain countries (e.g., Argentina). Additionally, discrepancy between designed and final sample was observed in certain countries more so than in others due to issues beyond researchers' control. For instance, in the case of Cuba, where education is obligatory and accessible to the population, it was extremely difficult to find participants with fewer than six years of education. Furthermore, the study used healthy participants. This resulted in floor and ceiling effects on some of the neuropsychological measures used. Additional limitations include lack of clinical sample(s) in the normative data, as well as lack of adequate representation of indigenous populations

from Latin America. Among inclusion criteria for this study was the requirement that participants must be able to read. Therefore illiterate individuals were not part of the final sample of this study. Directions for future work in this line of research may include creating normative data for illiterate individuals, as well as individuals with varying neurological disorders (e.g., dementias, TBI, multiple sclerosis, etc.) from Latin America.

4. Conclusions

The procedure used by Van der Elst et al. (2006, 2011, 2012) allows us to propose a convenient procedure to obtain sex, age, and education norms for 10 neuropsychological tests without the necessity to work with large samples. This procedure is, in fact, a simple procedure useful in clinical practice. Clinicians have two possibilities to obtain the percentile. The first one is to obtain the exact percentile using the procedure described and the information of the regression coefficients and the residual standard error provided in the different articles of each test. They also have tables corresponding to the standardized normal distribution or software that provide the cumulative probability to a z score in order to complete the process and obtain the exact percentile. The second one, for those clinicians who do not need to obtain the exact percentile, is to use the tables obtained for each test with the approximate percentile. These tables are very simple to use, and are similar to those tables that psychologists use in their regular practice. They only need to know the sex, age and educational level of the person assessed and, finally, the raw score in the test administered. With these values, it is necessary to search, in the table of their country, the sex if it is considered in the model, the educational level (1 to 12 and >12 years), the age group and the raw score, and then find the approximate percentile provided in the table. These tables will be including in each of the 10 manuscripts of the special issue.

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Conflict of interest

None.

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