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## CONCURRENT VALIDITY AND RELIABILITY OF AT-HOME TELENEUROPSYCHOLOGICAL EVALUATIONS AMONG PEOPLE WITH AND WITHOUT HIV

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### Abstract

**Objective:** To determine the reliability of teleneuropsychological (TNP) compared to in-person assessments (IPA) in people with HIV (PWH) and without HIV (HIV–).

**Methods:** Participants included 80 PWH ( $M_{age}=55.7$ ,  $SD_{age}=11.4$ ) and 23 HIV– ( $M_{age}=58.6$ ,  $SD_{age}=19.8$ ). Participants completed two comprehensive neuropsychological IPA before one TNP during the COVID-19 pandemic (March–December 2020). The neuropsychological tests included: Hopkins Verbal Learning Test-Revised (HVLTR Total and Delayed Recall), Controlled Oral Word Association Test (COWAT; FAS-English or PMR-Spanish), Animal Fluency, Verb Fluency, Wechsler Adult Intelligence Scale 3<sup>rd</sup> Edition (WAIS-III) Symbol Search and Letter Number Sequencing, Stroop Color and Word Test, Paced Auditory Serial Addition Test (Channel 1), and Boston Naming Test. Total raw scores and sub-scores were used in analyses. In the total sample and by HIV status, test-retest reliability and performance-level differences were evaluated between the two consecutive IPA (i.e., IPA1 and IPA2), and mean in-person scores (IPA-M), and TNP.

**Results:** There were statistically significant test-retest correlations between IPA1 and IPA2 ( $r$  or  $\rho = .603 - .883$ ,  $ps < .001$ ), and between IPA-M and TNP ( $r$  or  $\rho = .622 - .958$ ,  $ps < .001$ ). In the total sample, significantly lower test-retest scores were found between IPA-M and TNP on the COWAT (PMR), Stroop Color and Word Test, WAIS-III Letter Number Sequencing, and HVLTR Total Recall ( $ps < .05$ ). Results were similar in PWH only.

**Conclusions:** This study demonstrates reliability of TNP in PWH and HIV–. TNP assessments are a promising way to improve access to traditional neuropsychological services and maintain ongoing clinical research studies during the COVID-19 pandemic.

## Keywords

human immunodeficiency virus; COVID-19; validation study; neuropsychology; cognition; telehealth

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## Introduction

The use of telehealth (i.e., audio and videoconferencing to deliver healthcare) is rapidly growing, especially since the coronavirus disease (COVID-19) pandemic (Wosik et al., 2020). Within the field of neuropsychology, an advocacy team established by the Inter Organizational Practice Committee has been providing up-to-date recommendations and guidelines on the use of teleneuropsychological assessments (TNP) to clinicians (Bilder et al., 2020). Meanwhile, a host of training resources and virtual seminars/workshops have been disseminated across neuropsychological organizations, highlighting the rapid and immense need for knowledge within the context of neuropsychological research in this relatively new territory.

Telehealth allows for increased access to health care services, especially among persons with chronic illnesses and disabilities (Lillicrap, Hunter, & Goldswain, 2019). It decreases barriers to accessing appropriate care (e.g., lack of transportation, financial constraints, stigmatization) and can allow for patients needing specialty care to access providers despite geographical location (Gajarawala & Pelkowski, 2021; Moffatt & Eley, 2010; Speedie, Ferguson, Sanders, & Doarn, 2008). These advantages are especially pertinent given that already vulnerable populations (e.g., people with HIV; PWH) could be more susceptible to contracting COVID-19 and/or face more adverse health outcomes once infected (Mirzaei, McFarland, Karamouzian, & Sharifi, 2020).

Telehealth is generally well-accepted by patients, providers, and families (Parikh et al., 2013; Parsons et al., 2021; Shore, 2013), with patient reports of up to 98% satisfaction with videoconferencing and little concerns regarding privacy. Furthermore, some patients found videoconferencing more enjoyable and less anxiety-inducing in their naturalistic environment (Parikh et al., 2013). Despite the potential for decreased emotional connection compared to in-person assessments (IPA), videoconferencing has been found to provide similar personal interactions and possibly more frequent appointments (Bloem, Dorsey, & Okun, 2020; Marra, Hamlet, Bauer, & Bowers, 2020). With regard to TNP, patients and clinicians have found TNP evaluations acceptable and feasible during the COVID-19 pandemic and reported several favorable features, including saved travel time, reduced risk of COVID-19 exposure, and reduced concentration difficulties (Parsons et al., 2021).

Applications of TNP suggest a strong agreement between TNP and IPA across a variety of populations (e.g., older adults, patients with multiple sclerosis, cognitive impairment, psychiatric conditions, cerebrovascular accident) (Barcellos et al., 2021; Cullum, Hynan, Grosch, Parikh, & Weiner, 2014; Marra et al., 2020; Matchanova et al., 2020; Tailby et al., 2020; Wadsworth et al., 2018). Among cognitively impaired and non-impaired participants, scores on neuropsychological measures across domains (i.e., memory, attention, verbal fluency, language, executive function) between TNP (administered in-clinic) and

in-person conditions were highly concordant (Cullum et al., 2014). Furthermore, a recent systematic review (19 studies) of TNP validity concluded no significant effect related to video-administration of certain cognitive screeners (i.e., MMSE, MoCA), language tests (i.e., Boston Naming Test, Letter Fluency), attention/working memory tasks (i.e., Digit Span Total), and memory tests (Hopkins Verbal Learning Test-Revised) (Brearily et al., 2017; Marra et al., 2020). Majority of these studies conducted the TNP assessments over desktop/laptop (Marra et al., 2020). According to The Inter Organizational Practice Committee, Pearson recommends a display size of at least 9.75" on the patient side, although few studies have examined differences in TNP validity across different device types and sizes. Passell et al. (2021) found an association between device group and reaction time such that measures with more complex stimuli and responses (e.g., Trails A & B) were most affected by screen size. Together, these studies suggest that TNP is reliable and valid for various populations, and validity across various device types remains to be established.

To our knowledge, studies have yet to examine the reliability and comparability of remote TNP assessments versus traditional IPA among PWH. Previous TNP validations studies support the administration of TNP across multiple domains (e.g., memory, attention, executive function) that may be particularly relevant to assess among PWH (Becker et al., 1995; Heaton et al., 2015). Therefore, the cognitive effects of HIV may be amenable to TNP assessment.

Considering the COVID-19 pandemic, there is an immediate need for psychometrically-sound neuropsychological assessments that can be administered while patients are at-home, to maintain clinical neuropsychological care and ongoing research studies. However, several questions need to be answered before neuropsychologists can confidently use and interpret results from TNP evaluations. The current study will address two primary questions: How comparable are test scores obtained by TNP versus IPA? Is test reliability affected by mode of administration?

The specific purposes of this study are to (1) compare test-retest reliabilities between PWH and those without HIV (HIV-) at the participants' two most recent in-person assessments (IPA1 and IPA2), and between in-person and at-home TNP assessments, and (2) assess performance-level differences in neuropsychological assessment scores between IPA1 and IPA2, and between in-person and at-home TNP assessments. We hypothesized strong test-retest correlations and minimal performance-level differences in our samples of PWH and HIV-. We will additionally investigate the potential effects of technical aspects related to the remote assessment (e.g., participant device type, interruptions to testing) environment on raw scores at the TNP evaluation.

## Methods

### Participants

Participants included 80 PWH ( $M_{age}=58.7$ ,  $SD_{age}=11.0$ ) and 23 HIV- ( $M_{age}=61.9$ ,  $SD_{age}=16.7$ ) individuals who were enrolled in NIH-funded studies at the University of California, San Diego (UCSD) HIV Neurobehavioral Research Program (HNRP) from 2005-2020, demonstrated capacity to consent, and provided written informed consent.

All study procedures were approved by the UCSD Institutional Review Board and are in accordance with the Helsinki Declaration.

The current study is a secondary analysis of data from each participant's TNP evaluation, IPA1, and IPA2. Inclusion criteria were 1) age 18 years or older; 2) ability to provide informed consent; 3) negative urine toxicology for illicit drugs (excluding marijuana) or negative Breathalyzer test for alcohol on the day of the in-person study visits; 4) at least two completed IPAs; 5) greater than three months between the two prior IPAs; and 6) one remote TNP evaluation completed between March 2020-December 2020. Exclusion criteria at in-person and TNP evaluations were consistent among all parent studies and included 1) psychotic disorder diagnosis; 2) history of a non-HIV related neurological condition known to impact neurocognitive functioning (e.g., stroke, head injury with neurological complications; epilepsy); and 3) non-HIV related medical conditions associated with neurocognitive disorders. To determine language of test administration, participants were asked to self-report how well they spoke English and Spanish using a Likert-type scale (0=not well to 3=very well). Participants reporting equal scores for each language were tested in their preferred language (Mungas, Reed, Crane, Haan, & Gonzalez, 2004). Ninety-six participants were evaluated in English (93.2%). Given that each participant is their own control, we considered it appropriate to include the Spanish speakers in analyses.

### **In-Person Psychiatric and Neuromedical Evaluation**

The Composite International Diagnostic Interview (v2.1) was administered at the IPAs to assess current (i.e., past 12 months) and lifetime (i.e., >12 months ago) mood and substance use disorders (World Health Organization, 1997). The parent grants were funded before the publication of the DSM 5; therefore, diagnoses were made based on the DSM-IV criteria. HIV serostatus was determined using ELISA/Western blot by a CLIA-certified reference lab.

### **In-Person Neuropsychological Assessment**

Participants were administered a well-validated and comprehensive battery of neuropsychological assessments measuring seven cognitive domains: verbal fluency, executive function, processing speed, learning, delayed recall, working memory, and motor skills (Carey et al., 2004). This battery was designed in accordance with the international consensus conference recommendations (i.e., Frascati criteria) for HIV-associated neurocognitive disorder (Antinori et al., 2007; Heaton et al., 2010).

### **Teleneuropsychological Evaluation Setup**

TNP evaluations were conducted using HIPAA-compliant Zoom with both the examiner and participants using their personal devices in their respective home environments. To the extent possible, examiner setups were standardized. Examiner standardizations included use of a virtual private network; collection of participant responses with an iPad and stylus; secluded setting to minimize interruptions; computers with a camera and microphone. Because examiner screens are shared with the participant, all computer notifications were disabled. To protect participant privacy, Zoom meeting rooms were password-protected and examiners used headphones. All examiners operated from the same video-based platform; however, internet connection quality and computer hardware varied between examiners.

Because TNP evaluations were conducted in participants' naturalistic environments, participant standardizations were limited. HNRP schedulers recommended that participants find a private, quiet location in their homes, sit at a desk or table, wear headphones to improve audio quality and ensure confidentiality, and use a device with video capabilities for visual measures. Since June 2020, participants who did not have suitable home environments for TNP testing were provided the option of using a testing room (adhering to social distancing guidelines) at the HNRP for their TNP evaluation. Participants who connected by landline telephone received audio-only measures; participants who connected by tablet or personal computer received audio and visual measures; and participants who connected by smartphone received audio and visual measures, one of which needed to be adapted to conform to Zoom's non-adjustable mobile settings (i.e., Stroop Color and Word Test). Prior to testing, examiners administered a Remote Visit Questionnaire to assess participant testing environment (e.g., privacy, device used). After testing, examiners completed a second section of the Remote Visit Questionnaire to retrospectively capture the signal/connection quality during testing, audio quality, and interruptions during testing. Interruptions to testing were considered as any auditory or visual distraction that could influence neuropsychological performance (e.g., examinee's phone ringing, dog barking, family member speaking, garbage truck reversing).

### **Pre-Testing Sequence for the Remote Video Assessment**

Prior to beginning the TNP evaluation, participants received a brief introduction about the TNP procedure. To reduce the potential for distraction from self-view and video of the examiner, the video panel was minimized for participants only, leaving only test materials visible. Participants provided updated neurobehavioral and substance use histories for the interval between the prior IPA and the TNP visit. Examiners assessed substance use (e.g., alcohol, marijuana, methamphetamine) quantity since the previous IPA and lifetime quantity. Suspected intoxication at the TNP evaluation was indicated in behavioral notes.

Participants completed the Profile of Mood States (POMS), a 65-item self-report measure of mood (i.e., tension–anxiety, depression, anger–hostility, fatigue, confusion, vigor) over the previous seven days (McNair, Lorr, & Droppleman, 1981).

### **Teleneuropsychological Assessment Battery**

A comparison of the in-person and TNP batteries is presented in Table 1. Inclusion criteria for neurocognitive measures in the TNP battery were (a) brevity, (b) common use among HNRP studies, and (c) suitability for video-administration and response recording. Measures included Hopkins Verbal Learning Test-Revised (HVLTR Total and Delayed Recall) (Benedict, Schretlen, Groninger, & Brandt, 1998; Diaz-Santos et al., 2021), Controlled Oral Word Association Test (COWAT; FAS, PMR) and Category (Animal) Fluency (Borkowski, Benton, & Spreen, 1967; Marquine, Morlett Paredes, et al., 2021), Action (Verb) Fluency (Woods et al., 2005), Wechsler Adult Intelligence Scale 3<sup>rd</sup> Edition (WAIS-III) Symbol Search and Letter Number Sequencing, Stroop Color and Word Test (Gooding et al., 2021; Rivera Mindt et al., 2021; Stroop, 1935), Paced Auditory Serial Addition Test (PASAT - Channel 1) (Diehr, Heaton, Miller, & Grant, 1998; Gooding et al., 2021), and the 60-item

version of the Boston Naming Test (BNT; excluding item #48) (Kaplan, Goodglass, & Weintraub, 1983).

Individual test raw scores were converted into demographically-adjusted (i.e., age, sex, education, race/ethnicity) T-scores ( $M = 50$ ,  $SD = 10$  in healthy subjects) (Antinori et al., 2007; Cherner et al., 2021; Heaton, Miller, Taylor, & Grant, 2004; Heaton, Taylor, & Manly, 2003). Individual neuropsychological tests were considered impaired when T-scores  $< 40$  (Taylor & Heaton, 2001).

Given that neuropsychological assessments were designed to be administered in a face-to-face and in-person format, a few accommodations were made to facilitate TNP administration. First, considering internet and audio quality varies between participants, all TNP assessment instructions were presented both orally and visually. In the TNP format, verbal tasks (e.g., HVLTR), and tasks that rely on verbal responses to visually presented stimuli (e.g., BNT; visual presentation of stimuli via shared screen) are administered similarly to in-person. Three tasks that require visual stimuli or physical interaction with stimuli were reformatted (i.e., stimuli presentation via screen share instead of booklet, verbal response instead of motor) to be included in the video format. Comparisons between IPA and TNP administration of WAIS-III Symbol Search, Stroop Color and Word Test, and PASAT – Channel 1 are described in Supplementary Table 1.

### Statistical Analyses

HIV group differences on demographic characteristics were compared using independent  $t$ -tests and Chi-square statistics as appropriate. Raw scores were used for primary analyses. To examine test-retest reliability between IPA1 and IPA2, Pearson's correlation coefficients ( $r$ ) were used for normally distributed scores and Spearman's rho correlations for scores with skewed distributions. If raw scores on IPA1 and IPA2 were highly correlated ( $r$  or  $\rho > .500$ ,  $p < .05$ ), a mean in-person score was calculated for each neuropsychological test to represent average in-person performance (IPA-M) (Hemphill, 2003). Correlation coefficients were calculated for normally distributed scores and Spearman's rho correlations were used for scores with skewed distributions to examine test-retest reliability between IPA-M and TNP in the total sample and by HIV status. Paired  $t$ -tests were used to compare performance-level differences for normally distributed raw scores and Wilcoxon signed rank for skewed distributions between 1) IPA1 and IPA2; and 2) IPA-M and TNP in the total sample and by HIV status. Benjamini-Hochberg procedure was applied to correct for multiple comparisons using a false discovery rate of 0.05. Follow-up analyses using matched paired  $t$ -tests examined differences in T-scores between IPA2 and TNP to account for the effects of age, sex, education, and race/ethnicity.

Descriptive statistics from three of the Remote Visit Questionnaire items were calculated. Exploratory analyses were conducted to investigate the potential effects of technical aspects of the remote assessment environment on raw scores at the TNP evaluation. Linear regressions were used to examine the association between device type and change in raw scores from IPA2 to TNP (change score = TNP - IPA2). One-way analysis of variance (ANOVA) and Tukey's honest significant difference (HSD) tests were used to compare TNP raw scores between the four device types. A  $t$ -test was used to examine the effects



of examiner reported interruptions to testing on TNP raw scores. Statistical analyses were performed using JMP Pro version 14.0.0 (JMP®, Version <14.0.0>. SAS Institute Inc., Cary, NC, 1989-2007).

## Results

### Demographic Characteristics by HIV serostatus

Demographic characteristics at the TNP evaluation by HIV group are presented in Table 2. PWH had significantly fewer years of education and higher rates of lifetime substance use disorder ( $ps < .05$ ) than HIV-. The groups did not differ on age, sex, ethnicity, Wide Range Achievement Test 4 Reading, lifetime diagnosis Major Depressive Disorder, and mood ( $ps .09$ ). All participants completed two IPAs (days apart:  $M = 577$ ,  $SD = 716$ ;  $Mdn = 365$ ,  $IQR = 244-583$ ;  $range = 108-3970$ ) and one remote TNP evaluation (days apart from IPA2:  $M = 414$ ,  $SD = 238$ ;  $Mdn = 375$ ,  $IQR = 277-465$ ;  $range = 112-1655$ ). 49.5% of participants had re-tests more than one year apart between IPA1 and IPA2. 55.3% of participants had re-tests more than one year apart between IPA2 and TNP.

### Test-Retest Reliability of Neuropsychological Assessments

Results of correlation analyses are presented in Table 3. There were statistically significant correlations between IPA1 and IPA2 ( $r$  or  $\rho = .603 - .883$ ,  $mdn = .744$ ,  $ps < .001$ ) and between IPA-M and TNP ( $r$  or  $\rho = .622 - .958$ ,  $mdn = .801$ ,  $ps < .001$ ) across all neuropsychological assessment raw scores.

Correlations between IPA1 and IPA2 (Table 4) were statistically significant across neuropsychological assessment raw scores in PWH ( $r$  or  $\rho = .596 - .871$ ,  $mdn = .737$ ,  $ps < .001$ ) and HIV- groups ( $r$  or  $\rho = .556 - .943$ ,  $mdn = .826$ ,  $ps < .05$ ). Correlations between IPA-M and TNP (Table 5) were statistically significant across neuropsychological assessment raw scores in PWH ( $r$  or  $\rho = .631 - .960$ ,  $mdn = .820$ ,  $ps < .001$ ) except for the BNT ( $\rho = .593$ ,  $p = .122$ ), and in the HIV- group ( $r$  or  $\rho = .593 - .967$ ,  $mdn = .855$ ,  $ps < .05$ ). Correlations between IPA-M and TNP in the HIV- group for COWAT PMR and WAIS-III Letter Number Sequencing were not calculated due to small sample size ( $n = 5$ ).

### Performance-Level Differences between Neuropsychological Assessments

Results of matched paired  $t$ -tests or Wilcoxon Signed Rank test between IPA1 and IPA2 are presented in Table 6. There were no significant differences between raw scores on IPA1 and IPA2 ( $ps = .012$ ; Benjamini-Hochberg procedure cut off of 0.05 required). Results examining performance-level differences between IPA-M and TNP are presented in Table 7. There were significantly lower raw scores at TNP compared to IPA-M on COWAT (PMR) ( $t(7) = -3.7$ ,  $p = .007$ ), Stroop Word ( $t(49) = -6.1$ ,  $p < .001$ ), Stroop Color ( $t(48) = -3.9$ ,  $p < .001$ ), Stroop Incongruent ( $t(49) = -2.8$ ,  $p = .006$ ), WAIS-III Letter Number Sequencing ( $t(27) = -2.8$ ,  $p = .010$ ), and HVLTR Total Recall ( $t(81) = -3.6$ ,  $p < .001$ ). Cohen's effect size values for Stroop Color ( $d_z = .529$ ), Stroop Word ( $d_z = .387$ ), WAIS-III Letter Number Sequencing ( $d_z = .533$ ) and HVLTR Total Recall ( $d_z = .375$ ) suggest low to moderate practical significance. Effect size values for COWAT (PMR) ( $d_z = 1.11$ ) and Stroop Word ( $d_z = .867$ ) suggest high practical significance.



In PWH and the HIV– group, there were no significant differences between raw scores on IPA1 and IPA2, after correcting for multiple comparisons ( $ps > .05$ ). In PWH, there were significantly lower raw scores on TNP assessments compared to IPA-M on COWAT (PMR) ( $t(7) = -3.7, p = .007$ ), Stroop Word ( $t(38) = -4.6, p < .001$ ), Stroop Color ( $t(37) = -3.0, p = .004$ ), and HVLTR Total Recall ( $t(61) = -3.7, p < .001$ ) (Figure 1). In the HIV– group, there were lower raw scores on the TNP Stroop Word test compared to IPA-M ( $t(10) = -5.8, p < .001$ ) (Figure 2).

Follow-up analyses using matched paired  $t$ -tests examining differences in T-scores between IPA2 and TNP showed significantly lower T-scores at the TNP assessment on Stroop Word (TNP - IPA2 =  $-5.72$ ;  $t(57) = -5.86, p < .001$ ), Stroop Color (TNP - IPA2 =  $-2.7$ ;  $t(56) = -3.26, p = .002$ ), and HVLTR Total Recall (TNP - IPA2 =  $-3.65$ ;  $t(92) = -3.12, p = .002$ ). Neuropsychological test scores were considered impaired when T-score  $< 40$ . There were significantly more impaired scores in the TNP evaluation compared to IPA2 on Stroop Word (IPA2 = 18 (31%), TNP = 29 (50%);  $p < .001$ ), Stroop Color (IPA2 = 19 (33%), TNP = 22 (39%);  $p < .001$ ), and HVLTR Total Recall (IPA2 = 39 (42%), TNP = 57 (61%);  $p < .001$ ). On the Stroop Word test, 24% of participants went from an unimpaired to impaired score at the TNP evaluation and 5% went from impaired to unimpaired. On the Stroop Color test, 7% of participants went from an unimpaired to impaired score at the TNP evaluation and 2% went from impaired to unimpaired. On the HVLTR Total Recall, 26% of participants went from an unimpaired to impaired score at the TNP evaluation and 6% went from impaired to unimpaired.

Additional follow-up analyses were conducted to examine the potential effects of administration language. Seven participants were excluded that were tested in Spanish. Results of matched paired  $t$ -tests were consistent with the total sample, with observed differences in raw scores on Stroop Word, Stroop Color, Stroop Incongruent, WAIS-III Letter Number Sequencing, and HVLTR Total Recall ( $ps = .01$ ).

### Remote Visit Questionnaire Results

Device type, assessment type, and participant environment were evaluated from the Remote Visit Questionnaire. Of the total sample, 92 participants (89%) completed the Remote Visit Questionnaire. The most common device type used in the TNP evaluation was a smartphone (39%), followed by laptop/desktop (34%), tablet (14%), and traditional telephone (8%) (5% not documented). Audio only assessments were not limited to only via traditional telephone. Over 75% of the TNP evaluations were conducted using both video and audio. 21% of participants were interrupted at least once during the TNP evaluation (ex. “cathedral bells caused some disruption to participant’s attention span”). Results of a  $t$ -test comparing raw scores by participant interruption status revealed significant differences on the HVLTR Total Recall such that participants performed worse when there were interruptions during testing ( $M = 17.8$ ;  $SD = 5.1$ ) compared to no interruptions ( $M = 22.0$ ;  $SD = 6.1$ ;  $t(85) = -2.67, p = .009$ ). Additional follow-up analyses were conducted to examine the potential effects of interruptions to testing. Nineteen participants were excluded (16 PWH, 3 HIV–) that had interruptions to testing. Results of matched paired  $t$ -tests revealed differences in raw

scores on Stroop Word, Stroop Color, Stroop Incongruent, and HVLTR Total Recall ( $ps < .05$ ).

In the exploratory analyses, results revealed no significant association between device used at the TNP evaluation and change in performance from IPA2 to TNP. An ANOVA showed a significant omnibus difference across device type groups in TNP raw scores on Category Fluency – Animals ( $F(3,81) = 4.13, p = .008$ ). Follow-up pairwise comparisons showed poorer performance when administered via telephone compared to smartphone, tablet, and laptop/desktop ( $ps < .05$ ).

## Discussion

Results of this study add to a growing body of literature demonstrating that TNP assessments are reliable and valid across diverse populations and during the COVID-19 pandemic (Barcellos et al., 2021; Brearly et al., 2017; Cullum et al., 2014; Marra et al., 2020; Matchanova et al., 2020). Among our sample of PWH and HIV–, we established test-retest reliability between two IPAs that were approximately one year apart; and found significant and moderate to strong correlations between participants' IPA and TNP evaluations. Performance-level differences between IPA and TNP had variable effect sizes with small to moderate effect sizes for Stroop Color, Stroop Word, WAIS-III Letter Number sequencing, and HVLTR Total Recall; and large effect sizes for COWAT (PMR) and Stroop Word. Importantly, there was only a small sample of participants completed the COWAT (PMR). There were lower raw scores on the Stroop Color and Word Test, COWAT (PMR), WAIS-III Letter Number Sequencing, and HVLTR Total Recall at the TNP evaluation. Accounting for the effects of age, sex, education, and race/ethnicity, results indicated significant mean differences in T-scores on Stroop Word, Stroop Color, and HVLTR Total Recall with the greatest T-score point difference on the Stroop Word test (TNP - IPA2 =  $-5.72$ ). Across these three neuropsychological assessments, there were more participants that went from an unimpaired T-score at IPA2 to an impaired score at the TNP evaluation; however, several participants with impaired scores at IPA2 performed in the unimpaired range at the TNP evaluation. This could be attributable to practice effects, considering some participants had evaluations less than a year after their previous one; however, we might expect more participants to show improvement if there were significant practice effects (Dikmen, Heaton, Grant, & Temkin, 1999).

Despite statistical significance, differences in raw scores and T-scores between IPA-M and TNP on the COWAT (PMR), WAIS-III Letter Number Sequencing, and HVLTR Total Recall were minimal and possibly due to factors associated with COVID-19 (e.g., COVID-19 infection, stress, depression, social isolation), the TNP platform, or factors on the day of testing (e.g., pain, poor sleep, energy). (Hampshire et al., 2021; Suarez-Gonzalez, Rajagopalan, Livingston, & Alladi, 2021). The marginal difference in raw scores from the in-person to TNP evaluation observed on the HVLTR Total Recall (mean difference =  $-1.8$ ) is consistent with another study which found poorer performance on the HVLTR Total Recall in the video-based condition (mean difference =  $-2.11$ ) among mildly impaired stroke patients (Chapman, Gardner, Ponsford, Cadilhac, & Stolwyk, 2020). The authors attribute this possibly to mishearing words in the TNP condition or participant anxiety

with the TNP scenario. Results from our study indicate that interruptions to testing were significantly associated with worse performance on the HVLTR Total Recall and significant differences remained even after excluding participants that experienced interruptions. Therefore, differences observed on TNP tests could be attributable to other technical aspects of the TNP environment. For example, audio glitches may affect participant's understanding of task instructions, ability to clearly hear verbal stimuli, and adequate response collection by examiners (Gardner et al., 2021). These testing environment characteristics may be important to capture in TNP practice to understand whether a poor performance may reflect change in neurocognitive function or limitations of videoconferencing.

Few studies have investigated the reliability and validity of the Stroop Color and Word Test, or a similar response inhibition measure, in the TNP setting. One study among a pediatric sample (aged 6-20) found no significant performance-level differences on the DKEFS Color Word Interference Test between in-person and home-based TNP assessment administered via tablet or laptop (Harder et al., 2020). Another study among middle-to-older adults (aged 40-86) suggests moderate correlations between in-person and TNP assessments via desktop on the Stroop Color and Word Test (Zeghari et al., 2022). In the current study, more pronounced differences in raw scores observed across the Stroop Color and Word Test may be attributable to limitations of administering a time-bound visual neuropsychological assessment via videoconferencing. Particularly, lag-time in communication between an incorrect response and examiner feedback could limit the remaining time in the task for correct responses. Additionally, to be administered in the TNP modality, presentation of the stimuli was reformatted to balance the number of words on each slide (i.e., 60 words/slide, 2 slides) with the number of slide changes. Communication between the participant and examiner about changing slides could have also limited opportunity for participant correct responses. Considering differences in T-scores between IPA2 and TNP were greatest for Stroop Word and progressively decreased for the remaining subtests, it is also possible participants could benefit from more practice administration to better acclimate to video-administration of this test. On average, participants T-score on Stroop Word dropped six points at the TNP evaluation and 24% of participants went from an unimpaired score at IPA2 to an impaired score at TNP. To the degree that this difference is constant across individuals, and unrelated to their last in-person score, it may be possible to create and apply a time constant correction for this neuropsychological test.

Results of an exploratory analysis revealed no significant association between device used at TNP and change in performance from IPA2 to TNP, and comparability between TNP raw scores across device types, which may be suspected to account for differences on the Stroop Color and Word Tests. Despite these findings, several challenges remain in digital neuropsychology with regard to device type and characteristics including 1) variability in the perceptual, motor, and cognitive abilities needed for response behaviors; 2) variability in hardware and software between devices that may affect stimulus presentation and response latency; and 3) rapid changes in hardware, software, and device ownership which may affect tests and test norms (Germine, Reinecke, & Chaytor, 2019). These challenges are being rapidly investigated. Germine et al. (2019) found significant differences in response behavior on a digital trail making test with examinees (aged 18-35) taking less time to complete the task on an iPad compared to a personal computer, and more time on an iPhone.

Thus, the question around device type and screen size warrants reevaluation with a greater sample size and power to detect meaningful differences in the future, which could inform teleneuropsychological guidelines.

Among PWH, there were significant and strong correlations between IPA and TNP evaluations. Performance-level differences between IPA and TNP were minimal and consistent with findings from the total sample. Results suggest that TNP is a reliable alternative to IPA especially in the COVID-19 pandemic, but also more broadly when considering the health burden faced by this vulnerable population (Mirzaei et al., 2020). PWH are living longer and are more susceptible to age related neurodegenerative diseases and functional decline (Blackstone et al., 2012; Heaton et al., 2011; Wing, 2016). Neurocognitive and functional decline may limit the feasibility of attending in-person neuropsychological evaluations in aging PWH (Hearps, Schafer, High, & Landay, 2016). During the current COVID-19 pandemic, and especially among persons who are not vaccinated, PWH may be more fearful of going into the clinic for care or even COVID-19 testing services than the general population because of chronic immune impairment (Cooper, Woodward, Alom, & Harky, 2020; Fusco et al., 2020; Mirzaei et al., 2020). In addition to minimizing risk of additional infections, TNP evaluations have the potential to better maintain consistent access to care for PWH and provide benefits such as decreased time commitment, transportation expenses, and increase overall convenience (Gajarawala & Pelkowski, 2021; Moffatt & Eley, 2010; Speedie et al., 2008).

This study is unique from other studies examining TNP in that evaluations were typically completed at home rather than in a clinical space. Although clinical spaces provide certain testing environment standardizations (e.g., adequate internet connectivity, distraction-free), evidence suggests that cognitive performance in a naturalistic environment may be more aligned with actual cognitive functioning compared to in a clinic setting (Bloem et al., 2020; Moore et al., 2021; Rentz, 2016). It is important to note that TNP may not be feasible for all PWH, particularly those of the most vulnerable backgrounds (e.g., those experiencing homelessness, lower socioeconomic status, less acculturated, limited access to technology) as it introduces other potential barriers like resources to ascertain the necessary technology, need for reliable internet access, security concerns (Bilder et al., 2020; Mgbako et al., 2020). Although the HNRP does not currently provide technological devices (e.g., tablet) to participants for TNP testing, participants are provided the option of using a testing room at the HNRP for their evaluation.

There are several remaining questions that could not be addressed in this study: 1) Can published normative standards available for tests administered by IPA be used for those administered by TNP? Researchers may need to develop new normative data based on this modality or create adjustments for any differences derived from the TNP testing modality. 2) Can results of the same person using these two methods be compared to measure change or neurocognitive decline? Creating regression-based change scores would require a different set of visits than we used in the current study but can be helpful in determining significant change in neurocognitive performance. 3) Can results for different people in research studies be combined if some were administered the tests with IPA and some with TNP?

The current study is not without limitations. Our sample of PWH was relatively healthy which may have increased likelihood for test-retest reliability (R.K. Heaton et al., 2015). Due to the remote administration, motor tests (e.g., Grooved Pegboard) could not be administered. This may limit sensitivity in assessing some domains commonly impaired among PWH. Next, examiners did not explicitly ask participants not to write down information during the TNP testing. Although we detected significant differences between IPA-M and TNP on four neuropsychological assessments, there could be potential unmeasured confounders (e.g., distractibility, participant screen clarity, audio glitches) that may account for the differences. Audio glitches or disruptions when both the participant and examiner are speaking at the same time could have caused interference. While HNRP staff outline best practices for TNP evaluations, the lack of control over standardized testing environments in the TNP is a notable limitation. Furthermore, examinees were not asked to silence their device notifications during testing. While examiners noted this as an interruption to testing, we cannot fully rule out the potential impact of this distraction on test results. Results suggest that interruptions to testing were associated with worse performance on the HVLt-R Total Recall, despite no specific information about when the interruption occurred during testing. Thus, it may be beneficial to ask examinees to disable their device notifications during testing and for examiners to include standardized information about interruptions during testing. Since examiners do not directly assess substance use at the TNP and we were unable to conduct a urine toxicology test, we cannot confirm that examinees were toxicology negative. Considering there was small sample of Spanish speaking PWH that were administered the COWAT (PMR), results of significant differences between in-person and TNP may be interpreted with caution. Furthermore, there are other aspects of the Spanish-speaking sample that may make their neuropsychological evaluation particularly complex including socio-demographic, cultural, linguistic factors, and familiarity with telehealth and neuropsychological testing (Marquine, Rivera Mindt, et al., 2021). Future research may need to analyze the feasibility among Spanish-speaking PWH separately. The current study represents a secondary analysis of data from each participant's neuropsychological evaluations, not a randomized control trial to validate TNP assessment. The number of HNRP participants with a completed TNP evaluation is rapidly growing; therefore, follow-up analyses with a larger sample size and likely more statistical power will be beneficial. Future research may also investigate the role of emotional factors (e.g., depression, anxiety, financial instability, stress) on TNP performance during the COVID-19 pandemic, as well as whether performance on TNP assessments may adequately discriminate between impairment classifications.

Among our sample of PWH and HIV-, we provided evidence of test-retest reliability and performance-level comparability of our IPA and TNP. Considering the current COVID-19 pandemic, and possible additional pandemics in the future, there is an immediate need for reliable neuropsychological assessments that can be administered remotely, especially for PWH. TNP evaluation shows promise to improve access to neuropsychological services and maintain ongoing clinical research studies.

## Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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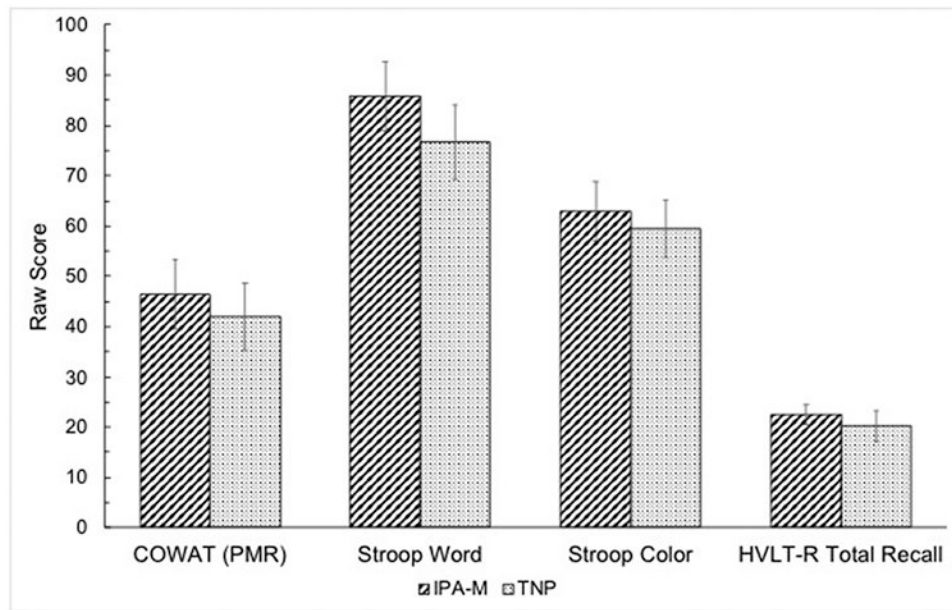


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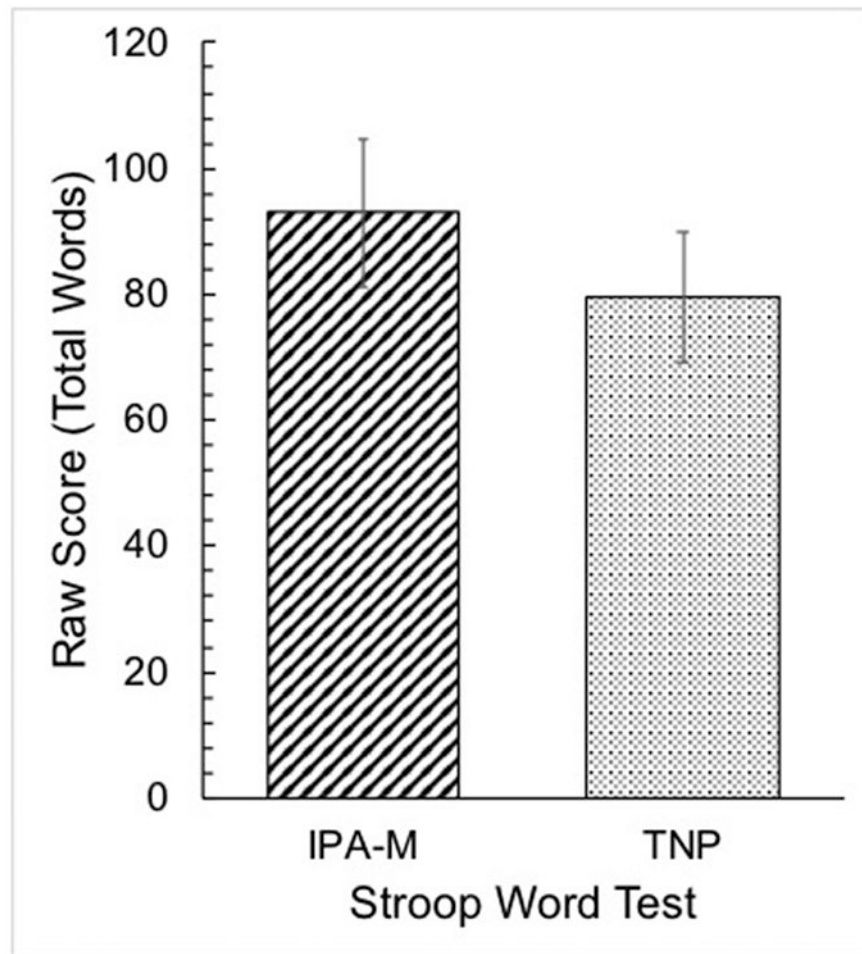


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**Figure 1.** Statistically significant performance-level differences between the mean in-person assessment scores (IPA-M) and the teleneuropsychological scores (TNP) in the HIV+ group



**Figure 2.** Statistically significant performance-level differences between the mean in-person assessment scores (IPA-M) and the teleneuropsychological scores (TNP) in the HIV– group

**Table 1.**

In-person battery versus teleneuropsychological battery

	In-Person	Teleneuropsych
<b>Verbal Fluency</b>		
Controlled Oral Word Fluency Test	X	X
Category Fluency (Animals)	X	X
Action (Verb) Fluency	X	X
<b>Executive Function</b>		
Trail Making Test, Part B	X	
Stroop Color and Word Test	X	X
Wisconsin Card Sorting Test (64-item)	X	
<b>Language</b>		
Boston Naming Test	X	X
<b>Working Memory</b>		
WAIS-III Letter-Number Sequencing	X <sup>a</sup>	X
Paced Auditory Serial Addition Test (Channel 1)	X	X
WMS-III Spatial Span	X <sup>a</sup>	
<b>Processing Speed</b>		
Trail Making Test, Part A	X	
WAIS-III Digit Symbol	X	
WAIS-III Symbol Search	X	X
<b>Learning and Memory</b>		
Hopkins Verbal Learning Test-Revised	X	X
Brief Visuospatial Memory Test-Revised	X	
<b>Motor Skills</b>		
Grooved Pegboard Test	X	

Note. WAIS-III = Wechsler Adult Intelligence Scale, Third Edition; WMS-III = Wechsler Memory Scale, Third Edition

<sup>a</sup>Not administered to every participant in-person

**Table 2.**

Participant demographic characteristics at the teleneuropsychological assessment by HIV serostatus

	<b>Total Sample N=103</b>	<b>PWH N=80</b>	<b>HIV- N=23</b>	<b>p-value</b>
Age (years)	59.4 (12.5)	58.7 (11.0)	61.9 (16.7)	.278
Education (years)	14.2 (2.8)	13.9 (2.8)	15.3 (2.5)	<b>.037</b>
Sex (male)	84 (81.6%)	66 (83%)	18 (78%)	.761
Ethnicity (Non-Hispanic White)	60 (58.2%)	45 (56%)	15 (65%)	.630
Testing Language (English)	96 (93.2%)	73 (91%)	23 (100%)	.344
WRAT4 Reading <sup>a</sup>	105.1 (12.9)	103.5 (11.3)	109.8 (16.0)	.093
Lifetime Major Depressive Disorder	56 (56%)	45 (57.7%)	11 (50%)	.628
Lifetime Substance Use Disorder	62 (62%)	54 (69.2%)	8 (36.4%)	<b>.007</b>
POMS Total Mood Disturbance <sup>b</sup>	59.4 (39.8)	60.6 (38.7)	55.5 (43.9)	.601
<b>HIV Disease Characteristics</b>				
History of AIDS	–	51 (64.6%)	–	–
Detectable plasma viral load <sup>cd</sup>	–	0 (0%)	–	–
Current CD4 count	–	643 [501, 850]	–	–
Nadir CD4 count	–	140 [49.3, 300.0]	–	–
Estimated years of HIV disease	–	24.0 [16.4, 30.7]	–	–
ARV Status (on cART)	–	75 (96.2%)	–	–

Note. WRAT4 Reading = Wide Range Achievement Test; POMS = Profile of Mood States; Values are presented as *M* (*SD*) or *Mdn* [*IQR*]. Bolded values indicate  $p < .05$ ; PWH = people with HIV

<sup>a</sup>N=62, administered at the first in-person visit only, not administered to Spanish-speakers

<sup>b</sup>N=92

<sup>c</sup>Defined as >50 copies/mL in plasma

<sup>d</sup>N=40

**Table 3.**

Correlation coefficients between the two most recent in-person assessments (IPA1 and IPA2) and the mean in-person assessment scores (IPA-M) and the teleneuropsychological scores (TNP)

	Correlations between IPA1 and IPA2		Correlations between IPA-M and TNP	
	Pearson's correlation coefficient or Spearman's rho	<i>p</i> -value	Pearson's correlation coefficient or Spearman's rho	<i>p</i> -value
<b>Verbal Fluency</b>				
COWAT (FAS)	.862	<.001	.880	<.001
Category Fluency - Animals	.732	<.001	.801	<.001
Action (Verb) Fluency	.791	<.001	.622	<.001
COWAT (PMR)	.829	.002	.958	<.001
<b>Executive Function</b>				
Stroop Word	.883	<.001	.743	<.001
Stroop Color	.876	<.001	.856	<.001
Stroop Incongruent	.744	<.001	.824	<.001
<b>Language</b>				
Boston Naming Test	.669	<.001	.763	<.001
<b>Working Memory</b>				
WAIS-III Letter Number Sequencing	.603	<.001	.652	<.001
PASAT (Channel 1)	.813	<.001	.805	<.001
<b>Processing Speed</b>				
WAIS-III Symbol Search	.742	<.001	.762	<.001
<b>Learning and Memory</b>				
HVLT-R Total Recall	.710	<.001	.696	<.001
HVLT-R Delayed Recall	.681	<.001	.838	<.001

Note. COWAT = Controlled Oral Word Fluency Test; WAIS-III = Weschler Adult Intelligence Scale, Third Edition; PASAT = Paced Auditory Serial Addition Test (Channel 1); HVLT-R = Hopkins Verbal Learning Test-Revised; Bolded values indicate  $p < .05$



**Table 4.**

Correlation coefficients between the two most recent in-person assessments (IPA1 and IPA2) by HIV status

	HIV+		HIV-	
	Pearson's correlation coefficient or Spearman's rho	<i>p</i> -value	Pearson's correlation coefficient or Spearman's rho	<i>p</i> -value
<b>Verbal Fluency</b>				
COWAT (FAS)	.835	<.001	.943	<.001
Category Fluency - Animals	.659	<.001	.891	<.001
Action (Verb) Fluency	.733	<.001	.886	<.001
COWAT (PMR)	.813	<.001	n=0	-
<b>Executive Function</b>				
Stroop Word	.871	<.001	.925	<.001
Stroop Color	.858	<.001	.918	<.001
Stroop Incongruent	.742	<.001	.789	<.001
<b>Language</b>				
Boston Naming Test	.678	<.001	.556	<b>.039</b>
<b>Working Memory</b>				
WAIS-III Letter Number Sequencing	.596	<.001	n=5	-
PASAT (Channel 1)	.788	<.001	.826	<.001
<b>Processing Speed</b>				
WAIS-III Symbol Search	.737	<.001	.760	<.001
<b>Learning and Memory</b>				
HVLT-R Total Recall	.678	<.001	.740	<.001
HVLT-R Delayed Recall	.676	<.001	.803	<.001

Note. COWAT = Controlled Oral Word Fluency Test; WAIS-III = Weschler Adult Intelligence Scale, Third Edition; PASAT = Paced Auditory Serial Addition Test (Channel 1); HVLT-R = Hopkins Verbal Learning Test-Revised; Bolded values indicate  $p < .05$

**Table 5.**

Correlation coefficients between the mean in-person assessment scores (IPA-M) and the teleneuropsychological scores (TNP) by HIV status

	HIV+		HIV-	
	Pearson's correlation coefficient or Spearman's rho	<i>p</i> -value	Pearson's correlation coefficient or Spearman's rho	<i>p</i> -value
<b>Verbal Fluency</b>				
COWAT (FAS)	.885	<.001	.855	<.001
Category Fluency - Animals	.820	<.001	.784	<.001
Action (Verb) Fluency	.932	<.001	.967	<.001
COWAT (PMR)	.960	<.001	n=0	-
<b>Executive Function</b>				
Stroop Word	.665	<.001	.926	<.001
Stroop Color	.845	<.001	.883	<.001
Stroop Incongruent	.792	<.001	.893	<.001
<b>Language</b>				
Boston Naming Test	.790	<.001	.593	.122
<b>Working Memory</b>				
WAIS-III Letter Number Sequencing <sup>a</sup>	.631	<.001	-.817	.184
PASAT (Channel 1)	.830	<.001	.607	<b>.048</b>
<b>Processing Speed</b>				
WAIS-III Symbol Search	.759	<.001	.836	<b>.001</b>
<b>Learning and Memory</b>				
HVLT-R Total Recall	.685	<.001	.714	<.001
HVLT-R Delayed Recall	.824	<.001	.856	<.001

Note. COWAT = Controlled Oral Word Fluency Test; WAIS-III = Wechsler Adult Intelligence Scale, Third Edition; PASAT = Paced Auditory Serial Addition Test (Channel 1); HVLT-R = Hopkins Verbal Learning Test-Revised; Bolded values indicate  $p < .05$

<sup>a</sup>  
n=4

**Table 6.**

Performance-level differences between the two most recent in-person assessments (IPA1 and IPA2)

	IPA1	IPA2	<i>p</i> -value	Cohens <i>d</i> <sub>z</sub>
<b>Verbal Fluency</b>				
COWAT (FAS) (n=69)	43.1 (14.6)	42.4 (14.5)	.461	.092
Category Fluency – Animals (n=84)	20.9 (6.4)	19.7 (6.0)	.012 <sup>a</sup>	.263
Action (Verb) Fluency (n=79)	15.5 (5.3)	15.7 (5.2)	.548	.059
COWAT (PMR) (n=7)	47.8 (13.9)	45.4 (13.7)	.421	.297
<b>Executive Function</b>				
Stroop Word (n=78)	87.7 (17.0)	87.1 (16.9)	.497	.073
Stroop Color (n=77)	63.6 (13.4)	61.8 (14.1)	.022 <sup>a</sup>	.262
Stroop Incongruent (n=77)	36.8 (10.7)	35.5 (13.4)	.211	.145
<b>Language</b>				
Boston Naming Test (n=56)	54.8 (5.8)	54.4 (8.6)	.864	.063
<b>Working Memory</b>				
WAIS-III Letter Number Sequencing (n=36)	8.9 (2.8)	8.9 (2.2)	1.00	0
PASAT (Channel 1) (n=76)	32.2 (11.7)	32.8 (12.6)	.642	.080
<b>Processing Speed</b>				
WAIS-III Symbol Search (n=86)	30.1 (8.2)	29.9 (9.0)	.803	.032
<b>Learning and Memory</b>				
HVLT-R Total Recall (n=87)	22.8 (6.3)	22.7 (6.4)	.911	.015
HVLT-R Delayed Recall (n=83)	7.2 (3.5)	7.3 (3.4)	.840	.036

Note. Values are presented as *M* (*SD*). COWAT = Controlled Oral Word Fluency Test; WAIS-III = Weschler Adult Intelligence Scale, Third Edition; PASAT = Paced Auditory Serial Addition Test (Channel 1); HVLT-R = Hopkins Verbal Learning Test-Revised; Bolded values indicate  $p < .05$

<sup>a</sup>Results considered not significant after applying Benjamini-Hochberg procedure (false discovery rate of 0.05)

**Table 7.**

Performance-level differences between the mean in-person assessment scores (IPA-M) and the teleneuropsychological scores (TNP)

	IPA-M	TNP	<i>p</i> -value	Cohens <i>d</i> <sub>z</sub>
<b>Verbal Fluency</b>				
COWAT (FAS) (n=68)	42.9 (14.0)	41.1 (14.0)	.036 <sup>a</sup>	.262
Category Fluency – Animals (n=83)	20.3 (5.8)	19.6 (6.0)	.112	.188
Action (Verb) Fluency (n=77)	15.6 (5.0)	14.8 (5.3)	.123	.178
COWAT (PMR) (n=7)	46.4 (14.1)	41.9 (13.4)	<b>.007</b>	1.11
<b>Executive Function</b>				
Stroop Word (n=50)	87.4 (16.3)	77.3 (16.2)	<b>&lt;.001</b>	.867
Stroop Color (n=49)	63.8 (14.6)	59.8 (12.9)	<b>&lt;.001</b>	.529
Stroop Incongruent (n=49)	36.6 (11.5)	33.8 (12.6)	<b>.006</b>	.387
<b>Language</b>				
Boston Naming Test (n=35)	55.7 (4.2)	55.2 (5.0)	.295	.154
<b>Working Memory</b>				
WAIS-III Letter Number Sequencing (n=28)	8.7 (2.0)	7.6 (2.7)	<b>.010</b>	.533
PASAT (Channel 1) (n=52)	34.1 (11.8)	32.5 (12.5)	.428	.210
<b>Processing Speed</b>				
WAIS-III Symbol Search (n=58)	31.2 (7.9)	29.9 (9.1)	.109	.218
<b>Learning and Memory</b>				
HVLT-R Total Recall (n=82)	22.9 (6.0)	21.1 (6.3)	<b>&lt;.001</b>	.375
HVLT-R Delayed Recall (n=77)	7.4 (3.2)	7.2 (3.2)	.591	.110

Note. Values are presented as *M* (*SD*). COWAT = Controlled Oral Word Fluency Test; WAIS-III = Weschler Adult Intelligence Scale, Third Edition; PASAT = Paced Auditory Serial Addition Test (Channel 1); HVLT-R = Hopkins Verbal Learning Test-Revised; Bolded values indicate  $p < .05$

<sup>a</sup>Results considered not significant after applying Benjamini-Hochberg procedure (false discovery rate of 0.05)