# Original Article



# Depression, Obstructive Sleep Apnea and Cognitive Impairment (DOC) Screen Completion Time Reflects Executive Function, Speed of Processing and Fluency

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ABSTRACT: Background: The depression, obstructive sleep apnea and cognitive impairment (DOC) screen assesses three post-stroke comorbidities, but additional information may be gained from the time to complete the screen. Cognitive screening completion time is rarely used as an outcome measure. Objective: To assess DOC screen completion time as a predictor of cognitive impairment in stroke/transient ischemic attack clinics. Methods: Consecutive English-speaking stroke prevention clinic patients consented to undergo screening and neuropsychological testing (n = 437). DOC screen scores and times were compared to scores on the NINDS-CSC battery using multiple linear regression (controlling for age, sex, education and stroke severity) and receiver operating characteristic (ROC) curve analysis. Results: Completion time for the DOC screen was  $3.8 \pm 1.3$  minutes. After accounting for covariates, the completion time was a significant predictor of the speed of processing ( $p = 0.002$ , 95% CI: -0.016 to -0.004), verbal fluency ( $p < 0.001$ , CI: -0.012 to -0.006) and executive function (p = 0.004, CI: −0.006 to −0.001), but not memory. Completion time above 5.5 minutes was associated with a high likelihood of impairment on executive and speed of processing tasks (likelihood ratios 3.9-5.2). Conclusions: DOC screen completion time is easy to collect in routine care. People needing over 5.5 minutes to be screened likely have deficits in executive functioning and speed of processing – areas commonly impaired, but challenging to screen for, after stroke. DOC screen time provides a simple, feasible approach to assess these underidentified cognitive impairments.

Résumé: Le temps de réalisation du test de dépistage de la triade DATC, reflet de la fonction exécutive, de la vitesse de traitement et de la fluidité verbale. Contexte : Le test de dépistage de la dépression, de l'apnée obstructive du sommeil et de troubles cognitifs (DATC) permet d'évaluer trois troubles comorbides post-AVC, mais le temps de réalisation du test lui-même pourrait fournir des renseignements additionnels. Toutefois, on utilise rarement la durée du test de dépistage de troubles cognitifs comme critère d'évaluation. Objectif : L'étude visait à évaluer la durée du dépistage de la triade DATC comme test prévisionnel de troubles cognitifs dans des centres de soins des AVC et des accidents ischémiques transitoires. Méthode : Des patients ( $n = 437$ ) consécutifs, de langue anglaise, inscrits dans des centres de prévention des AVC ont consenti à passer le test de dépistage ainsi que des tests neuropsychologiques. Les résultats obtenus au test de dépistage de la triade DATC et la durée des tâches ont été comparés aux résultats obtenus à la batterie de tests CSC du NINDS à l'aide de modèles de régression linéaire multiple (prise en considération de l'âge, du sexe, du degré d'instruction et du degré de gravité des AVC) et de l'analyse des courbes caractéristiques de la performance du test (ROC : en anglais). Résultats : Le temps de réalisation du test de dépistage de la triade DATC était de 3,8 ± 1,3 minutes. Après la prise en considération des covariables, la durée du test s'est révélée un facteur prévisionnel significatif de la vitesse de traitement ( $p = 0.002$ ; IC à 95 % : -0.016 à -0.004), de la fluidité verbale ( $p < 0.001$ ; IC : -0.012 à -0.006) et de la fonction cognitive ( $p = 0.004$ ; IC : -0,006 à -0,001), mais pas de la mémoire. Une durée de test supérieure à 5,5 minutes a été associée à des probabilités élevées de troubles de la fonction exécutive et de la vitesse des tâches de traitement (rapport de vraisemblance : 3,9-5,2). Conclusion : La durée du test de dépistage de la triade DATC est facile à consigner en milieu de soins usuels. Les personnes ayant besoin de plus de 5,5 minutes pour passer le test de dépistage connaissent probablement des troubles de la fonction exécutive et de la vitesse des tâches de traitement – sphères d'activité souvent perturbées

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mais difficiles à dépister – après un AVC. Le temps de dépistage de la triade DATC s'avère donc un moyen simple et facilement réalisable d'évaluation de ces troubles cognitifs souvent peu identifiés.

Keywords: Cognition; depression; executive function; sleep apnea; stroke

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### Introduction

Stroke is the leading cause of neurological disability in adults<sup>[1](#page-7-0)</sup> and survival after stroke is increasing. $2-4$  $2-4$  $2-4$  In addition to physical post-stroke deficits,<sup>[5](#page-7-0)</sup> approximately 30–50% of stroke survivors are affected by each of depression, obstructive sleep apnea (OSA) and cognitive impairment (DOC).<sup>[6](#page-7-0)–[9](#page-7-0)</sup> These DOC comorbidities are all associated with poorer functional outcomes $^{10}$  $^{10}$  $^{10}$  and an increased risk of mortality.<sup>[11](#page-7-0)</sup>

The DOC screen was developed as a feasible and valid tool to screen and stratify stroke patients into high, intermediate and low risk groups for DOC comorbidities to facilitate detection and management in high-volume stroke clinic settings.<sup>[12](#page-7-0)</sup> The screen is efficient, yet designed to maintain the construct validity of a delayed recall task. Eighty-nine percent of patients in stroke prevention clinics are able to complete the tool in  $<6$  minutes (mean = 4.2) minutes,  $SD = 1.5$ .<sup>[12](#page-7-0)</sup> In validation studies, the cognitive component of the DOC score is helpful to quickly stratify people into "cognitively normal," "cognitively impaired" and "need more assessment" groups, compared to more detailed cognitive testing.<sup>12</sup> Although the DOC completion time was originally collected as a way to assess feasibility, practitioners can record this measure when administering the DOC screen in clinical settings. Several studies have reported the average time taken to complete other well-known cognitive screens as feasibility demonstrations, including the Montreal Cognitive Assessment (MoCA; means ranging from 9.5 minutes to 11 minutes)<sup>13,[14](#page-7-0)</sup> and the Mini-Mental State Examination (MMSE; means ranging from 8 minutes to 13.4 minutes).<sup>[14,15](#page-7-0)</sup> However, few studies have assessed the utility of using a cognitive screen's completion time as a metric to evaluate underlying cognitive abilities, such as executive functioning.

Executive dysfunction and delays in speed of processing are the most commonly reported cognitive impairments after stroke. The DOC screen specifically examines mood symptoms, cognitive (executive, memory and abstraction) dysfunction and OSA/fatigue – all of which could be associated with cognitive or psychomotor slowing.<sup>16</sup>

#### Aim

Screen completion time is an immediately available metric, requiring no additional effort from either patients or clinicians, which might reflect executive function. The objective of this study was to determine whether completion time for the DOC screen is a reliable reflection of cognitive dysfunction and whether a single completion time cut-point could indicate cognitive impairment.

#### Methods

All patients were recruited from the DOC feasibility and validity study[.12](#page-7-0) This study included English-speaking (or English-fluent) patients newly referred to stroke prevention clinics between April 23, 2012, and April 30, 2014 ( $n = 1504$ ), who could complete the screen independently (with the administrator, but without third-party support). We excluded patients with severe aphasia, severe motor dysfunction (unable to hold a pen and draw a clock) and patients who were not fluent in English. Each eligible participant was administered the DOC screen (Figure 1) as a brief screen of DOC. All DOC screens were timed from the beginning of the memory registration (first task) until the end of the five-word free recall (final task). Chart abstractions by trained research members captured demographic and clinical data on all participants from patient charts using previously published and validated methods.[17,18](#page-7-0)

To reduce sampling bias, all consecutive patients from stroke prevention clinics who completed the DOC screen were asked to complete more detailed neuropsychological assessments, including a cognitive battery and formal mood assessments as outlined in the DOC feasibility study.<sup>[12](#page-7-0)</sup> All patients who completed the detailed assessments provided written informed consent. Only the site PI could access the information that could identify individual participants, all the other authors were given anonymized study IDs that were created upon the completion of the informed consent process. A complete list of all mood and cognitive assessments completed as part of the DOC study is reported elsewhere.<sup>[12](#page-7-0)</sup> In this analysis, cognition was assessed using the 30-minute neuropsychological battery recommended by the NINDS-CSN.[19](#page-7-0) This cognitive battery consists of the Controlled Oral Word Association Test (phonemic fluency), Animal Naming task (semantic fluency), California Verbal Learning Test (CVLT), Digit Symbol Coding and Trail Making Tests (TMT-A and TMT-B). All scores were normalized (z-score or scaled score) for age using age-matched norms from each respective test manual. CVLT and Animal Naming were also education-standardized.<sup>[20,21](#page-7-0)</sup> The study was approved by the Sunnybrook Research Ethics Board (approval number SUN-2312).

#### Statistical analysis

Statistical analysis was performed using IBM SPSS Statistics for Windows version 24. Descriptive statistics, including means and standard deviations, were calculated for age, screen completion time and number of years of education.

### Multivariable linear regression analyses of the relationship between time-to-completion and NINDS-CSC standardized scores

To assess whether screen time reflects cognitive function, independent linear regression models were used to examine the association between DOC completion time and the scaled or z-scores of all neuropsychological subtests. Data from all participants were used in the regression models. A sensitivity analysis was performed using a complete case approach to assess whether missing variables affected the models. All models controlled for age, education, modified Rankin Score (mRS) and sex. Due to the established relationship between the DOC cognitive sub-scores and detailed cognitive assessments,<sup>[12](#page-7-0)</sup> we also controlled for the DOC-Cognition score in all models. To adjust for multiple (7) linear regressions, Bonferroni correction  $(0.05/7 = 0.0071)$  was used to define significance at  $p < 0.007$  for all analyses.



Figure 1. The depression, obstructive sleep apnea and cognitive impairment (DOC) screen (freely available for download at [www.docscreen.ca\)](https://www.docscreen.ca). (continued on next page)

ROC and logistic regression analyses to identify cutoffs associated with high likelihood of cognitive impairments

To identify whether a single cut-point (in seconds) for screen time could be found with high specificity and likelihood ratios for cognitive impairment, receiver operating characteristic (ROC) curves were used. ROC analyses were run for each neuropsychological assessment significantly associated with the DOC screen completion time. A logistic regression with screen time completion



Modified/Combined from: <sup>1</sup>PHQ-2: Hajek VE et al. Brief assessment of cognitive impairment in patients with stroke. Arch Phys Med Rehabil. 1989 Feb; 70(2):114-7. 'STOP: Chung F et al. STOP questionnaire: A tool to screen patients for obstructive sleep apnea. Anesthesiology. 2008 May, 108(5):812-21. 74. 'MoCA: Nasreddine 25 et al. The montreal cognitive assessment, MoCA: A brief screening tool for mild cognitive impairment. J Am Geriatr Soc. 2005 Apr;53(4):695-9. MoCA copyright of the adapted version: Z. Nasreddine MD. Adapted by Swartz et al., 2013.

Figure 1. (Continued).

(as a continuous variable) and the cognitive impairment classification on the NINDS-CSN assessments were applied to the ROC curves. The classification of cognitive impairment of NINDS-CSN was defined as scores >2.0 standard deviations from expected norms, on two or more cognitive tasks. This required participants to have completed all tests in the detailed cognitive battery; thus, a complete case approach was used for all ROC analyses. First, a single, specific cut-point (time in seconds) was

<span id="page-4-0"></span>Table 1. Demographics for participants completing detailed cognitive and neuropsychological assessments ( $n = 437$ )

<b>Variables</b>	Mean (SD)
Age (years)	62.7 (15.6)
Education (years)	15.6(3.9)
DOC screen completion time (s)	227.8 (76.6)
Language	n(%)
English	363(83.1)
English Second Language	74 (16.9)
Sex (female)	51.3%
Most responsible diagnosis	
Undetermined diagnosis	4(0.9)
Abnormal CT/MRI scan	21(4.8)
Asymptomatic carotid artery disease	4(0.9)
Definite ischemic stroke	121 (27.7)
Definite TIA	54 (12.4)
Hemorrhage ICH	17 (3.9)
Hemorrhage IVH	1(0.2)
Hemorrhage SAH	4(0.9)
Hemorrhage SDH	1(0.2)
Other non-vascular	96 (22)
Other vascular	14(3.2)
Possible/query ischemic	13(3.0)
Possible/query TIA	84 (19.2)
Sinovenous thrombosis	3(0.7)
Modified Ranking Scale (mRS)	
0	230 (52.6)
1	113 (25.9)
$\overline{2}$	69(15.8)
3	19(4.3)
4	2(0.5)
Missing	4(0.9)

<sup>†</sup> TIA = transient ischemic attack; ICH = intracerebral hemorrhage; IVH = intraventricular hemorrhage; SDH = subarachnoid hemorrhage; SDH = subdural hemorrhage;  $hemorrhage;$   $SAH = subarachnoid$  hemorrhage; DOC = depression, obstructive sleep apnea and cognitive impairment.

defined based on the ROC curve output for patients with an overall classification of impaired on the NINDS-CSN battery. The cutpoint was pre-specified to have 95% specificity for cognitive impairment. This cut-point was then applied to ROC curves from each individual assessment and evaluated using likelihood ratios.

#### Results

A total 437 patients completed the cognitive and mood gold standard assessments within a maximum of 3 months of screening, with an average time interval of 3 days<sup>[12](#page-7-0)</sup> (Supplemental Table [1\)](https://doi.org/10.1017/cjn.2024.303). Of these, 213 (48.7%) participants were male, with a mean ( $\pm$  standard deviation) age of 62.7  $\pm$  15.6 years and a mean years of education of  $15.6 \pm 3.9$  years (Table 1). Additionally, 387 patients were able to complete all assessments in the battery; 13.7 % of these were classified as impaired based on the NINDS-CSN



Table 2. Linear regression results showing the effect of the DOC screen

completion time on individual neuropsychological assessments

DOC = depression, obstructive sleep apnea and cognitive impairment.

\*All models controlled for by age, sex, years of education, DOC-cognition score and modified Rankin Scale (mRS).

<sup>†</sup>Significant results bolded and set at  $p < 0.007$ .

 $\dagger$  TMT = Trail Making Test; CVLT = California Verbal Learning Test.

classification. The DOC screen completion mean was  $3.8 \pm 1.3$ minutes (range: 1.9–9.6 minutes). Among the patients, 134 (31%) had an ischemic stroke, 138 (32%) had a probable/possible TIA and the remainder (37%) were diagnosed with other conditions (Table 1). Non-stroke/transient ischemic attack (TIA) diagnoses included patients referred with possible stroke symptoms, but whose further investigations revealed alternative diagnoses, as well as patients without specific stroke/TIA symptoms referred for either vascular risk reduction or assessment of incidental abnormal imaging findings.

We performed linear regressions with DOC screen completion time (in seconds) as a predictor for each neuropsychological assessment score (Table 2). In all models, we controlled for age, sex, years of education, screening score of cognitive function (DOCcognition score) and overall function (mRS). All regression models for screen completion time were significant  $(p < 0.001)$ (Supplemental Table [2](https://doi.org/10.1017/cjn.2024.303)). Additionally, model summaries showed that screen completion time was a significant predictor ( $p < 0.005$ ) of verbal fluency semantic score (95% confidence interval (CI) of beta-coefficient from linear regression: −0.006 to −0.001), verbal fluency phonemic score (95% CI: −0.018 to −0.006), digit symbol coding (95% CI: −0.016 to −0.004) and the Trail Making Tests (TMT-A 95% CI: −0.017 to −0.005; TMT-B 95% CI: −0.016 to −0.004). In all cases, these were negative correlations (i.e., longer





 $\dagger$  Trails = Trail Making Test; DOC = depression, obstructive sleep apnea and cognitive impairment.





Figure 2. Receiver operating characteristic (ROC) curve, model for overall cognitive impairment with a cutoff set at 95% specificity.

completion times correlated with poorer cognitive scores). DOC screen completion time was not a significant predictor of memory performance on the CVLT Short Delay Free Recall ( $p = 0.713, 95\%$ CI:  $-0.003-0.002$ ) or the CVLT Long Delay Free Recall ( $p = 0.790$ , 95% CI: −0.002–0.003). Results did not differ in the sensitivity models with complete case data (see Table [2](#page-4-0) compared to Supplemental Table [3](https://doi.org/10.1017/cjn.2024.303) with complete case data). Neither DOC mood and apnea screening scores nor SCID-D or polysomnogram scores were associated with DOC screen completion time in any multivariable regression.

Using the single cutoff point approach on the overall cognitive impairment ROC curve (Figure 2, Table 3A), the point with 95% specificity for cognitive impairment was 332.5 s. When this time was applied to ROC models for each individual cognitive task (Table 3B), the same cut-point had high specificity on all executive and speed of processing tasks. The area under the curve was greater than 0.7 for all executive and speed of processing tasks. Likelihood ratios for predicting abnormal results on executive and speed of processing tasks ranged from four to six – that is, people taking more than 332.5 s to complete the DOC screen were 4–6 times more likely to have severe cognitive impairment on executive and speed of processing tasks than those with faster completion times (see Table 3). Scatterplots demonstrating the predicted probability of impairment on each domain by completion time, derived from the logistic regression analysis can be found in the supplemental material.

## **Discussion**

Several studies $^{22}$  have shown that post-stroke cognitive impairments can be separated into independent cognitive factors including language, memory and executive function, with deficits in executive functioning and speed of processing being the most common. $23$ Screening tests for executive function and speed of processing are limited and rarely used in routine clinical care. These results demonstrate that DOC screen completion time is an independent

predictor of executive function (semantic fluency,  $^{24}$  $^{24}$  $^{24}$  TMT-B<sup>25</sup>), speed of processing (digit symbol coding, $^{26}$  TMT-A and TMT-B<sup>28</sup>) and verbal fluency<sup>28</sup> after stroke, even after controlling for age, sex, education, DOC-cognition score and stroke severity. Completion time did not predict CVLT scores, a verbal test primarily affecting verbal memory (learning/registration and recall).<sup>[29](#page-8-0)</sup> Verbal fluency, while reflecting language function, is also reflective of executive function[.30](#page-8-0) Moreover, we have demonstrated that a 332.5 s (roughly 5.5 minutes) cutoff has 95% specificity and high likelihood ratios for predicting both overall cognitive and executive function impairment. This can be used as a quick and easily obtainable measure to identify people at risk for impairment on executive and speed of processing tasks. Certainly, other timed tasks, whether pen-and-paper (like Trails) or digital (e.g., Creyos), can be used to assess executive and speed of processing deficits in detail; however, detailed cognitive batteries are too onerous for routine clinical use. Simply timing the DOC screen as it is administered provides additional information, beyond the actual DOC cognitive screening score, which can flag people at high risk of having multi-domain cognitive impairment and executive/speed of processing dysfunction.

A few notable neuropsychological measures have used completion time to assess specific cognitive functions. For instance, Trail Making Tests (TMTs) are a set of widely accepted timed neuropsychological measures that provide insight into executive abilities.<sup>[27](#page-8-0)</sup> Processing speed is highly associated with performance on TMT Part B (a task reflecting attention and executive functions such as set-shifting) and with performance on TMT Part A (which is more closely related to motor speed and attention).[25,31,32](#page-8-0) Similarly, Woods et al. discovered that a patient's question completion time on self-paced questionnaires could be used as a measure of executive functioning.<sup>33</sup> Question completion time measures processing and decision-making speeds, providing insight into motivation, effort and cognitive ability that is not measured by existing tests.<sup>[33](#page-8-0)</sup> These studies support the notion that timed measures may be useful as a measure of executive dysfunction in addition to their use as screening instruments. The findings presented in our study correspond well to those reported by Woods et al. Their analyses showed that complex tasks, akin to our DOC-cognitive tasks, were strongly related to executive function and processing speed. Their neuropsychological tests (including TMT-B and Digit Span) also correlated significantly with self-paced question completion time. Their research process was similar to ours, wherein completion time was compared to existing screens to validate completion time as a metric; both studies suggest that completion time of self-paced complex assessments may be valid markers of executive function.

Few studies use the completion time of a neuropsychological screening tool as a cognitive marker. Most timed tasks examine processing speed directly (e.g., Trails, symbol-digit modalities test<sup>34</sup>) and have been studied in clinical settings, for example, for HIV-induced cognitive dysfunction<sup>[35](#page-8-0),[36](#page-8-0)</sup> and in multiple sclerosis.[37,38](#page-8-0) However, these types of tasks are more detailed and timeconsuming, and while they can be performed in a clinic in isolation, they are more often done as part of larger batteries. In contrast, screening tasks like the MoCA or MMSE are not routinely timed when applied in clinical settings. By simply timing the DOC screen, in addition to the information generated by the screen on mood, apnea and cognitive function, the time taken to complete the entire screen is itself an indirect measure that can highlight people at risk for cognitive impairment, especially executive, speed of processing and attentional issues. Moreover, executive function deficits are not often assessed in stroke patients;

these deficits are subtle, challenging to test for and often go unrecognized.<sup>23</sup> The NINDS-CSC battery is recommended as a research battery, but it requires a trained administrator and at least 30 minutes per person plus scoring. This is not feasible for routine clinic use. The DOC screen, in contrast, takes less than 5 minutes, can be performed by clinical staff (students, administrative assistants, nurses and physicians) and can help to highlight people at risk for impairments in mood, apnea and cognition.

The interpretation of our findings is limited by our sample population. Compared to the total number of patients who were asked to volunteer from the stroke prevention clinic ( $n = 1504$ ), consenting participants ( $n = 437$ ) tended to be slightly younger and with slightly milder neuropsychological deficits (healthy participant bias).<sup>[12](#page-7-0)</sup> However, our sample also included a wide range of patients across the full spectrum of severity. As expected from stroke/TIA clinic samples, 62% had a diagnosis of stroke and/or TIA, and the rest had alternative diagnoses common in stroke prevention clinics (mimics, multiple vascular risk factors, abnormal imaging). This heterogeneity reflects the pragmatic nature of the screening and its broad generalizability to the population of patients referred to stroke prevention clinics. TIA patients are well recognized to share similar long-term risk profiles<sup>[39](#page-8-0)</sup> and are also at risk for cognitive impairment,<sup>40</sup> compared to those with imaging-confirmed strokes. While the strongest associations to DOC completion were with tests of executive function, processing speed and verbal fluency, other domains that were less well represented in the NINDS-CSC battery could also impact screen completion time. For example, visuospatial function was not specifically assessed in the NINDS-CSC battery, and while language function could also affect completion time, there was no relationship with score on the California Verbal Learning Task (a verbal memory task). Since many tasks have more than one cognitive construct underlying them (e.g., phonemic and semantic fluency tasks each require language, attention and executive functions), DOC screen time cannot be considered a reflection of only one underlying domain. However, the tasks associated with DOC screen time all share underlying cognitive constructs of attention, executive dysfunction and/or speed of processing. The relationship between DOC completion time and gold standard testing was found across a range of severity from normal function to severely impaired. It should also be noted that there is not a single perfect cutoff score for DOC completion time that indicates executive dysfunction. To facilitate clinical utility, and because this is intended as a screen in high-volume clinics, we chose to explore a cutoff with high specificity so clinicians could be confident there was a high likelihood of true cognitive impairment beyond this time; however, this cutoff will have a low sensitivity and will miss some people with cognitive impairments. Previous work has already established that the DOC-cognition score can also be a sensitive screen, effectively ruling out cognitive impairment in people who score highly.<sup>12</sup> Finally, it is important to note that although screen completion time may be a useful tool to identify people at risk for executive dysfunction, it is still not equivalent to a detailed neuropsychological assessment.

### Conclusion

Clinical cognitive screening tools have not commonly used completion time as a metric. We aimed to determine whether the DOC screen completion time could provide clinically relevant information on patients' cognitive function. DOC screen completion time reflects executive function, speed of processing and verbal fluency. When administering the DOC screen, completion time requires no additional time or patient burden to collect. This convenience is vital in busy

<span id="page-7-0"></span>stroke prevention clinic settings, where there is minimal time for detailed cognitive assessments. Exploring whether screen time can act as a predictor of future outcomes would provide further support for the utility of this measure in clinical settings.

Supplementary material. The supplementary material for this article can be found at <https://doi.org/10.1017/cjn.2024.303>.

Data availability statement. DOC screening for mood, cognition and apnea was performed in stroke prevention clinics under waiver of consent. Patients provided written consent to undergo detailed cognitive testing and to relate their screening results to the detailed neuropsychological testing. However, the public release of data was not part of the patients' consent.

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Author contributions. Sajeevan Sujanthan and Alisia Southwell contributed equally to this work.

SS – Formal analysis, writing – original draft, writing – reviewing and editing.

- AS Data collection, formal analysis, writing original draft.
- TA Formal analysis, writing original draft.
- EX Formal analysis, writing original draft.
- AK Formal analysis, writing original draft.
- XL Formal analysis, writing original draft.
- KLL Resources, supervision, writing review and editing.
- NH Resources, supervision, writing review and editing.
- BJM Resources, supervision, writing review and editing.
- KET Resources, supervision, writing review and editing.
- MLC Data collection, writing review and editing.
- MNS Data collection, writing review and editing.
- KL Data collection, writing review and editing.
- DS Resources, supervision, writing review and editing.

RHS – Funding, conceptualization, resources, supervision, writing – original draft, writing – reviewing and editing.

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Competing interests. RHS reports ownership shares in FollowMD Inc., a virtual vascular risk reduction clinic. None of the other authors have any conflicts of interest to disclose.

Data access. Data is not available to share publicly, as patients did not consent to public data release. Clinical Trials Registration Identifier: NCT02363114. Clinical Trials URL: <https://clinicaltrials.gov/ct2/show/NCT02363114>.

#### References

- 1. Feigin VL, Nichols E, Alam T, et al. Global, regional, and national burden of neurological disorders, 1990-2016: a systematic analysis for the global burden of disease study 2016. Lancet Neurol. 2019;18:459–80. DOI: [10.1016/](https://doi.org/10.1016/S1474-4422(18)30499-X) [S1474-4422\(18\)30499-X.](https://doi.org/10.1016/S1474-4422(18)30499-X)
- 2. Lakshminarayan K, Berger AK, Fuller CC, et al. Trends in 10-year survival of patients with stroke hospitalized Between 1980 and 2000. Stroke. 2014;45:2575–81. DOI: [10.1161/STROKEAHA.114.005512.](https://doi.org/10.1161/STROKEAHA.114.005512)
- 3. Rodríguez-Castro E, López-Dequit I, Santamaría-Cadavid M, et al. Trends in stroke outcomes in the last ten years in a European tertiary hospital. BMC Neurol. 2018;18:164. DOI: [10.1186/s12883-018-1164-7.](https://doi.org/10.1186/s12883-018-1164-7)
- 4. Waziry R, Heshmatollah A, Bos D, et al. Time trends in survival following first hemorrhagic or ischemic stroke Between 1991 and 2015 in the rotterdam study. Stroke. 2020;51:824–9. DOI: [10.1161/STROKEAHA.119.](https://doi.org/10.1161/STROKEAHA.119.027198) [027198.](https://doi.org/10.1161/STROKEAHA.119.027198)
- 5. Verstraeten S, Mark R, Sitskoorn M. Motor and cognitive impairment after stroke: a common bond or a simultaneous deficit? Stroke Res Ther. 2016;1:539–547.
- 6. Herrmann N, Seitz D, Fischer H, et al. Detection and treatment of post stroke depression: results from the registry of the Canadian stroke network. Int J Geriatr Psychiatry. 2011;26:1195–1200. DOI: [10.1002/gps.2663.](https://doi.org/10.1002/gps.2663)
- 7. Johnson KG, Johnson DC. Frequency of sleep apnea in stroke and TIA patients: a meta-analysis. J Clin Sleep Med. 2010;6:131–7.
- 8. Patel MD, Coshall C, Rudd AG, Wolfe CDA. Cognitive impairment after stroke: clinical determinants and its associations with long-term stroke outcomes. J Am Geriatr Soc. 2002;50:700–6. DOI: [10.1046/j.1532-5415.](https://doi.org/10.1046/j.1532-5415.2002.50165.x) [2002.50165.x](https://doi.org/10.1046/j.1532-5415.2002.50165.x).
- 9. Kapoor A, Lanctôt KL, Bayley M, et al. "Good outcome" isn't good enough. Stroke. 2017;48:1688–90. DOI: [10.1161/strokeaha.117.016728](https://doi.org/10.1161/strokeaha.117.016728).
- 10. Kapoor A, Lanctot KL, Bayley M, Herrmann N, Murray BJ, Swartz RH. Screening for post-stroke depression and cognitive impairment at baseline predicts long-term patient-centered outcomes after stroke. J Geriatr Psychiatry Neurol. 2019;32:40–8. DOI: [10.1177/0891988718819859](https://doi.org/10.1177/0891988718819859).
- 11. Swartz RH, Bayley M, Lanctôt KL, et al. Post-stroke depression, obstructive sleep apnea, and cognitive impairment: rationale for, and barriers to, routine screening. Int J Stroke. 2016;11:509–18. DOI: [10.1177/1747493016641968.](https://doi.org/10.1177/1747493016641968)
- 12. Swartz RH, Cayley ML, Lanctôt KL, et al. The "DOC" screen: feasible and valid screening for depression, obstructive sleep apnea (OSA) and cognitive impairment in stroke prevention clinics. PLOS ONE. 2017;12:e0174451. DOI: [10.1371/journal.pone.0174451](https://doi.org/10.1371/journal.pone.0174451).
- 13. Lees RA, Hendry BA, Broomfield K, et al. Cognitive assessment in stroke: feasibility and test properties using differing approaches to scoring of incomplete items. Int J Geriatr Psychiatry. 2017;32:1072–8. DOI: [10.1002/](https://doi.org/10.1002/gps.4568) [gps.4568](https://doi.org/10.1002/gps.4568).
- 14. Barnay JL, Wauquiez G, Bonnin-Koang HY, et al. Feasibility of the cognitive assessment scale for stroke patients (CASP) vs. MMSE and MoCA in aphasic left hemispheric stroke patients. Ann Phys Rehabil Med. 2014;57:422–35. DOI: [10.1016/J.REHAB.2014.05.010.](https://doi.org/10.1016/J.REHAB.2014.05.010)
- 15. Molloy DW, Standish TI. A guide to the standardized Mini-Mental State Examination. Int Psychogeriatr. 1997;9(Suppl 1):87–94. DOI: [10.1017/](https://doi.org/10.1017/s1041610297004754) [s1041610297004754.](https://doi.org/10.1017/s1041610297004754)
- 16. Schrijvers D, Hulstijn W, Sabbe BGC. Psychomotor symptoms in depression: a diagnostic, pathophysiological and therapeutic tool. J Affect Disord. 2008;109:1–20. DOI: [10.1016/j.jad.2007.10.019](https://doi.org/10.1016/j.jad.2007.10.019).
- 17. Hall R, Khan F, O'Callaghan C, et al. Ontario Stroke Evaluation Report 2013: Spotlight on Secondary Stroke Prevention and Care. Toronto, ON: The Institute for Clinical Evaluative Sciences (ICES); 2013.
- 18. Kapral MK, Fang J, Hill MD, et al. Sex differences in stroke care and outcomes. Stroke. 2005;36:809–14. DOI: [10.1161/01.STR.0000157662.](https://doi.org/10.1161/01.STR.0000157662.09551.e5) [09551.e5](https://doi.org/10.1161/01.STR.0000157662.09551.e5).
- 19. Hachinski V, Iadecola C, Petersen RC, et al. National Institute of Neurological Disorders and Stroke-Canadian Stroke Network Vascular Cognitive Impairment Harmonization Standards. Stroke. 2006;37:2220–41. DOI: [10.1161/01.STR.0000237236.88823.47](https://doi.org/10.1161/01.STR.0000237236.88823.47).
- 20. Tombaugh T. Normative data stratified by age and education for two measures of verbal fluency FAS and animal naming. Arch Clin Neuropsych. 1999;14:167–77. DOI: [10.1016/S0887-6177\(97\)00095-4.](https://doi.org/10.1016/S0887-6177(97)00095-4)
- 21. Delis DC, Kramer JH, Kaplan E, Ober BA. California verbal learning test second edition (CVLT-II). San Antonio, TX: The Psychological Corporation; 2000.
- 22. Swartz RH, Stuss DT, Gao F, Black SE. Independent cognitive effects of atrophy and diffuse subcortical and thalamico-cortical cerebrovascular disease in dementia. Stroke. 2008;39:822–30. DOI: [10.1161/STROKEAHA.](https://doi.org/10.1161/STROKEAHA.107.491936) [107.491936.](https://doi.org/10.1161/STROKEAHA.107.491936)
- 23. Zinn S, Bosworth HB, Hoenig HM, Swartzwelder HS. Executive function deficits in acute stroke. Arch Phys Med Rehabil. 2007;88:173–80. DOI: [10.1016/J.APMR.2006.11.015](https://doi.org/10.1016/J.APMR.2006.11.015).
- <span id="page-8-0"></span>24. Maseda A, Lodeiro-Fernández L, Lorenzo-López L, Núñez-Naveira L, Balo A, Millán-Calenti JC. Verbal fluency, naming and verbal comprehension: three aspects of language as predictors of cognitive impairment. Aging Ment Health. 2014;18:1037–45. DOI: [10.1080/13607863.2014.908457](https://doi.org/10.1080/13607863.2014.908457).
- 25. Arbuthnott K, Frank J. Trail making test, part B as a measure of executive control: validation using a set-switching paradigm. J Clin Exp Neuropsychol. 2000;22:518–28. DOI: [10.1076/1380-3395\(200008\)22:4;1-0;FT518.](https://doi.org/10.1076/1380-3395(200008)22:4;1-0;FT518)
- 26. Deloire MS, Bonnet MC, Salort E, et al. How to detect cognitive dysfunction at early stages of multiple sclerosis? Mult Scler J. 2006;12:445–52. DOI: [10.1191/1352458506ms1289oa](https://doi.org/10.1191/1352458506ms1289oa).
- 27. Muir RT, Lam B, Honjo K, et al. Trail making test elucidates neural substrates of specific poststroke executive dysfunctions. Stroke. 2015;46:2755–61. DOI: [10.1161/STROKEAHA.115.009936](https://doi.org/10.1161/STROKEAHA.115.009936).
- 28. Whiteside DM, Kealey T, Semla M, et al. Verbal fluency: language or executive function measure? Appl Neuropsychol Adult. 2016;23:29–34. DOI: [10.1080/23279095.2015.1004574.](https://doi.org/10.1080/23279095.2015.1004574)
- 29. Jacobs M, Donders J. Criterion validity of the California verbal learning testsecond edition (CVLT-II) after traumatic brain injury. Arch Clin Neuropsych. 2007;22:143–9. DOI: [10.1016/j.acn.2006.12.002.](https://doi.org/10.1016/j.acn.2006.12.002)
- 30. Aita SL, Beach JD, Taylor SE, Borgogna NC, Harrell MN, Hill BD. Executive, language, or both? An examination of the construct validity of verbal fluency measures. Appl Neuropsychol Adult. 2019;26:441–51. DOI: [10.1080/23279095.2018.1439830.](https://doi.org/10.1080/23279095.2018.1439830)
- 31. MacPherson SE, Cox SR, Dickie DA, et al. Processing speed and the relationship between trail making test-B performance, cortical thinning and white matter microstructure in older adults. Cortex. 2017;95:92–103. DOI: [10.1016/j.cortex.2017.07.021.](https://doi.org/10.1016/j.cortex.2017.07.021)
- 32. Fishman KN, Ashbaugh AR, Swartz RH. Goal setting improves cognitive performance in a randomized trial of chronic stroke survivors. Stroke. 2021;52:458–70. DOI: [10.1161/STROKEAHA.120.032131.](https://doi.org/10.1161/STROKEAHA.120.032131)
- 33. Woods DL, William Yund E, Wyma JM, Ruff R, Herron TJ. Measuring executive function in control subjects and TBI patients with question completion time (QCT). Front Hum Neurosci. 2015;9:288. DOI: [10.3389/](https://doi.org/10.3389/FNHUM.2015.00288) [FNHUM.2015.00288.](https://doi.org/10.3389/FNHUM.2015.00288)
- 34. Zaidi KB, Rich JB, Sunderland KM, et al. Methods for improving screening for vascular cognitive impairment using the Montreal cognitive assessment. Can J Neurol Sci. 2020;47:756–63. DOI: [10.1017/CJN.2020.121.](https://doi.org/10.1017/CJN.2020.121)
- 35. Davis HF, Skolasky RL, Selnes OA, Burgess DM, McArthur JC. Assessing HIV-associated dementia: modified HIV dementia scale versus the grooved pegboard. Aids Read. 2002;12:29–31 38.
- 36. Van Harten B, Courant MNJ, Scheltens P, Weinstein HC. Validation of the HIV dementia scale in an elderly cohort of patients with subcortical cognitive impairment caused by subcortical ischaemic vascular disease or a normal pressure hydrocephalus. Dement Geriatr Cogn Disord. 2004;18: 109–14. DOI: [10.1159/000077818](https://doi.org/10.1159/000077818).
- 37. Ezegbe C, Zarghami A, van der Mei I, Alty J, Honan C, Taylor B. Instruments measuring change in cognitive function in multiple sclerosis: a systematic review. Brain Behav. 2023;13:e3009. DOI: [10.1002/BRB3.](https://doi.org/10.1002/BRB3.3009) [3009.](https://doi.org/10.1002/BRB3.3009)
- 38. Wishart M, Everest MR, Morrow SA, Rose J, Shen L, Feinstein A. Establishing the consistency of a voice recognition symbol digit modalities test analogue. Mult Scler. 2023;29:1676–9. DOI: [10.1177/13524585](https://doi.org/10.1177/13524585231199321) [231199321](https://doi.org/10.1177/13524585231199321).
- 39. Edwards JD, Kapral MK, Fang J, Swartz RH. Long-term morbidity and mortality in patients without early complications after stroke or transient ischemic attack. Can Med Assoc J. 2017;189:E954–E961. DOI: [10.1503/cmaj.](https://doi.org/10.1503/cmaj.161142) [161142.](https://doi.org/10.1503/cmaj.161142)
- 40. Pendlebury ST, Wadling S, Silver LE, Mehta Z, Rothwell PM. Transient cognitive impairment in TIA and minor stroke. Stroke. 2011;42:3116–21. DOI: [10.1161/STROKEAHA.111.621490.](https://doi.org/10.1161/STROKEAHA.111.621490)