Research Report

Screening for Cognitive Impairment in Movement Disorders: Comparison of the Montreal Cognitive Assessment and Quick Mild Cognitive Impairment Screen in Parkinson's Disease and Lewy Body Dementia

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Abstract.

Background: The Montreal Cognitive Assessment (MoCA) is recommended by the Movement Disorder Society for cognitive testing in movement disorders including Parkinson's disease (PD) and lewy body dementia. Few studies have compared cognitive screening instruments in these diseases, which overlap clinically.

Objective: To compare the MoCA and Quick Mild Cognitive Impairment (Qmci) screen in this population.

Methods: Patients attending memory and movement disorder clinics associated with a university hospital had the MoCA and Qmci screen performed and diagnostic accuracy compared with the area under the receiver operating characteristic curve (AUC). Duration and severity of movement disorders was assessed using the Unified PD Rating Scale (UPDRS).

Results: In total, 133 assessments were available, median age 74 ± 5 . Median education was 11 ± 4 years and 65% were male. Median total UPDRS score was 37 ± 26 . Median Qmci screen was 51 ± 27 , median MoCA was 19 ± 10 . There were statistically significant differences in test scores between those with subjective symptoms but normal cognition, mild cognitive impairment (MCI) and dementia (p < 0.001). The Qmci screen had significantly greater accuracy differentiating normal cognition from MCI versus the MoCA (AUC 0.90 versus 0.72, p = 0.01). Both instruments had similar accuracy in identifying cognitive impairment and separating MCI from dementia. The median administration time for the Qmci screen and MoCA were 5.19 and 9.24 minutes (p < 0.001), respectively.

Conclusions: Both the MoCA and Qmci screen have good to excellent accuracy in a population with movement disorders experiencing cognitive symptoms. The Qmci screen was significantly more accurate for those with early symptoms and had a shorter administration time.

Keywords: Alzheimer's disease, cognitive assessment screening instrument, lewy body disease, Montreal Cognitive Assessment, movement disorders, Parkinson's disease

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INTRODUCTION

The prevalence of movement disorders is increasing in tandem with population aging [1]. Approximately 28% of adults aged over 50 years can be classified as having a movement disorder, most commonly a tremor [2]. One of the most prevalent movement disorders worldwide is Parkinson's disease (PD); the number of persons with PD is expected to double by 2030 [3] with an increasing incidence and burden associated with this condition in most regions and countries globally [4]. As cognitive impairment (CI) is common in PD, particularly as the disease advances, this will result in an overall increase in the number of patients reporting cognitive symptoms associated with PD. Many of these will be older patients. Prevalence rates of PD-related CI approach 80% [5], while the rate of progression to Parkinson's disease dementia (PDD) is approximately 10% per year [6]. Mean duration from onset of PD symptoms to dementia is estimated at 10 years [7]. Many patients with PD have CI from the time of diagnosis (approximately 15-20%), though the prevalence in prodromal disease is less clear [8]. Lewy body dementia (LBD) is one of the most common dementia subtypes and is also associated with the development of a movement disorder [9]. Similar to PD, LBD is a dementia associated with parkinsonism and Lewy body formation and it likewise increases in prevalence with age, representing approximately 5% of dementia cases in older populations [9, 10]. As with CI in PD, it is likely under-reported [10].

Mild cognitive impairment (MCI) affects at least one-third (27%-42.5%) of people with PD [11-13] and is associated with a greater chance of developing dementia, especially if present when patients are first diagnosed [14]. Single domain, non-amnestic is the most common subtype [12]. The diagnosis of PD-MCI is challenging with uncertainty surrounding the duration of follow-up and number of repeated cognitive tests required to make the diagnosis [15]. Reflecting this, some guidelines including the Movement Disorder Society Task Force Guidelines [11], suggest that up to a minimum of 10 repeat assessments are required to diagnose and subtype PD-MCI [15]. PD-MCI features exaggerated attention and executive function deficits [16]. LBD is also associated with an MCI syndrome referred to as LBD-MCI, which, similar to PD-MCI, is characterized by executive, visuospatial, and attentional deficits [17, 18]. CI in persons with movement disorders impacts upon life

expectancy [19], quality of life [20], healthcare costs [21] and activities of daily living (ADLs), even among those without dementia, i.e., at prodromal stages [20].

Early diagnosis of CI is important to facilitate prompt treatment, identify reversible or compounding factors and plan for the future [22]. Monitoring change over time is important in this context and relies on the use of accurate short cognitive screening instruments (CSIs) [23]. Repeated neuropsychological testing [14] is recommended to increase prognostic accuracy, particularly conversion from MCI syndromes to clinical dementia. The progression from PD-MCI and LBD-MCI to dementia is particularly challenging to diagnose. An ideal short CSI in this setting would be reliable, brief, sensitive to early change, have normative data available, cover core cognitive domains relevant to these conditions (i.e., attention-working memory, memory, executive functioning, visuospatial skills, and language) and be largely immune to the effects of motor limitations [23, 24]. Such CSIs would also ideally correlate well with functional measures and emerging biomarkers [23].

At present, the Movement Disorder Society recommends the use of the Montreal Cognitive Assessment (MoCA) [25] for cognitive testing in movement disorders including PD and LBD [26-28]. The reason why the MoCA is well suited for use in this population is because it lacks ceiling effects and is weighted towards domains such as executive and visuospatial functioning and less towards orientation and language, which are relatively well preserved in PD [29]. Other widely-used CSIs may be less suitable. For example, the Mini-Mental State Examination (MMSE), one of the most established and widelyused CSIs, is still commonly used for these patients [28, 30]. However, the MMSE, because of its low ceiling effects and selected cognitive domains, may miss early cognitive deficits in these conditions [29] and is recommended for use only with this caveat [28]. This may result in failed opportunities to initiate early appropriate treatment and discontinue inappropriate medications like anti-cholinergics (medications that bind to muscarinic receptors and block acetylcholine neurotransmission), which is particularly important in diseases associated with a cholinergic deficit such as Alzheimer's disease, PDD and LBD [31]. Despite the increasing use of the MoCA, questions remain about its accuracy; studies suggest that up to onequarter of patients with PD with normal MoCA scores report functional cognitive difficulties, while threequarters with low MoCA scores report none [32].

On this basis, other tests with a greater emphasis on visio-cognitive impairment, may be preferable [33].

Despite the advantages of the MoCA and its extensive validation [34], some challenges remain with its use in persons with movement disorders and other populations such as in primary care and community-based memory clinics (where access to neuropsychology may be limited) including elements of redundancy, a relatively long administration time for a CSI (10-12 min), its known floor effects, its low specificity and a high false positive rate in older adults, particularly at its recommended cutoff score (> 26) [35–39]. Given this, we sought to investigate if the Quick Mild Cognitive Impairment (Qmci) screen [40], which has yet to be validated in patients with movement disorders and cognitive symptoms, is accurate in detecting cognitive impairment in this population and how it compares to the MoCA.

MATERIALS AND METHODS

Patients

This is a secondary analysis of a cross-sectional study that was conducted parallel to a larger study evaluating the Qmci screen in an Irish clinic population. The methods have been reported elsewhere [38] but in summary, a consecutive sample of patients with cognitive symptoms attending either a university hospital movement disorder clinic or a geriatric medicine memory clinic in Cork City, Ireland between January 2013 to December 2014, were included. Patients were classified as having normal cognition (subjective symptoms only), MCI or dementia by a consultant physician, specialized in the diagnosis and management of cognitive disorders. A diagnosis of dementia (PDD or LBD) was based on Diagnostic and Statistical Manual of Mental Disorders, (5th edition) criteria [41]. A diagnosis of PD-MCI was made using level I diagnostic criteria according to the Movement Disorder Society Task Force Guidelines [12], defined as recent, subjective but corroborated memory loss without obvious loss of social or occupational function, with evidence of objective deficits on a global cognitive scale: the Unified Parkinson's Disease Rating Scale (UPDRS) part I, in those with established PD. LBD-MCI was diagnosed using the third report of the LBD Consortium [42]. Patients were excluded if they were aged < 40 years of age, if they had other

movement disorders such as essential tremor (n = 3), if they were unable to communicate verbally in English, if they had depression (as defined by a Geriatric Depression Scale-short [GDS-SF] form with a cut-off ≥ 7 points to increase specificity [43]), or if a reliable collateral was not available but required. The GDS-SF, scored out of 15 points, is validated in older adults with PD [44] and has been used in LBD [45], again taking a higher cut-off score than in Alzheimer's disease [46].

Data collection

Patients underwent comprehensive clinical assessment and were screened for cognitive impairment. This assessment included a full history, physical examination, laboratory testing and neuroimaging. A short neuropsychological battery including the Standardized MMSE [47] and two informantrated assessments, the AD8 questionnaire [48] and IQCODE Short Form [49] were conducted by a consultant geriatrician, blind to the results of the CSIs, to inform the clinical diagnosis. The MoCA and Qmci screen were scored in random order, by a trained rater, prior to and independent of the clinical assessment. The presence and severity of any extrapyramidal symptoms were graded using the UPDRS performed by an Advanced Nurse Practitioner in movement disorders (MJF). The study adhered to the tenets of the Declaration of Helsinki and ethical approval was obtained from the Clinical Research Ethics Committee of the Cork Teaching Hospitals (reference number: ECM 4 (aa) 03/04/12) and patients provided informed consent. Assent was obtained from individuals who were felt to lack capacity.

Measures

Montreal cognitive assessment

The MoCA has seven subtests covering five cognitive domains; specifically, it includes visuospatial, attention, processing speed, language, memory, and cognitive control scored out of 30 points with lower points indicating cognitive impairment. A cut-off of < 26/30 is suggested for use in routine practice [25], although lower cut-offs have been suggested. Lower scores imply more impaired cognition.

Quick mild cognitive impairment screen

The Qmci screen is a brief and accurate CSI for MCI and has been studied in persons attending mem-

Comparison of characteristi	es of patients with noi	imai cogintion, innu c	ogintive iniparment (MCI) and dementia	
Variable	Total $(Q3-Q1 = \pm IQR)$	Normal ($n = 34, 25\%$) (Q3-Q1 = ±IQR)	MCI ($n = 42, 32\%$) (Q3-Q1 = ±IQR)	Dementia ($n = 57, 43\%$) (Q3-Q1 = ±IQR)	P = x
Age (Median & IQR)	74 (79–72 = ± 5)	$73(74-72 = \pm 2)$	75 (79–70 = \pm 9)	76 (81–72 = \pm 9)	0.02*
Education (Median & IQR)	$11(14-10 = \pm 4)$	$14(19-14 = \pm 5)$	$13(14 - 10 = \pm 4)$	$11(12-10 = \pm 2)$	0.28
Years since Diagnosis (Median & IQR)	$7(10-3 = \pm 7)$	$3(7-2=\pm 5)$	$7.5(11-5=\pm 6)$	$6(11-3=\pm 8)$	0.10
Gender (% male)	65%	65%	71%	61%	0.58
UPDRS – Total (Median & IQR)	$37(50-24 = \pm 26)$	$36(50-22 = \pm 28)$	$37 (45 - 24 = \pm 21)$	$38(54-32 = \pm 22)$	0.51
UPDRS-Cog (Median & IQR)	$3(5-2=\pm 3)$	1.5(4.5 - 1 = 3.5)	$3(5-2=\pm 3)$	$4(5-3=\pm 2)$	0.06
SMMSE (Median & IQR)	$26(28-22 = \pm 6)$	28 (30–27 = \pm 3)	27 (29–26 = \pm 3)	$22(25-19 = \pm 6)$	< 0.001*
MoCA (Median & IQR)	$19(23 - 13 = \pm 10)$	$25(26-23 = \pm 3)$	$22(24-18 = \pm 6)$	$13(16-10 = \pm 6)$	< 0.001*
Qmci screen (Median & IQR)	51 (63–36 = \pm 27)	$70(73-65 = \pm 8)$	55 (59–53 = ± 6)	$36(44-29 = \pm 15)$	< 0.001*

Table 1 Comparison of characteristics of patients with normal cognition, mild cognitive impairment (MCI) and dementia

IQR, Interquartile range. *Statistically significant. UPDRS, Unified Parkinson's Disease Rating Scale; SMMSE, Standardized Mini-Mental State Examination; *Qmci* screen, Quick Mild Cognitive Impairment Screen; MoCA, Montreal Cognitive Assessment.

ory clinics but not in those with PD or LBD [38]. The Qmci screen has six subtests covering five cognitive domains; specifically, it includes five orientation items, five registration items, a clock drawing test, a delayed recall question, verbal fluency (e.g., animals named in 1 min) and a test of logical memory (immediate verbal recall of a short story) [50]. It takes 3–5 min to complete [50]. The optimal Qmci screen cut-off score for cognitive impairment (MCI or dementia) is < 62/100 [38]. Again, lower scores suggest greater cognitive impairment.

Unified Parkinson's disease rating scale

The severity of PD was determined using the UPDRS [51]. The UPDRS total score consists of the sum of parts I (mentation, behavior, and mood), II (ADLs) and III (motor examination), with scores ranging from 0 (not affected) to 176 (most severely affected). A score of one or more on item one (intellectual impairment) of part I was taken as supportive of CI (PD-MCI or PDD).

Analysis

Data were analyzed using R version 4.2.2 (2022-10-31) -"Innocent and Trusting" (R Core Team, 2022). All significance tests were two sided, and a *p*-value of < 0.05 was considered statistically significant. Most data were non-normally distributed and analyzed with non-parametric tests. Spearman's rho (*r*) measured rank correlation. The Chi Squared test assessed differences between the distributions of categorical variables. The Mann-Whitney U test and the Kruskal-Wallis test was used to examine differences between non-parametric continuous variables. Diagnostic accuracy was determined from analysis of the Area Under the Curve (AUC) generated by receiver operating characteristics (ROC) curves. These were compared using the compared using the DeLong method [52] Sensitivity and specificity were determined and optimal cut-offs calculated based on Youden's Index [53].

RESULTS

In all, 133 patient assessments were available and included. In addition to those excluded with other movement disorders, one duplicate was removed. The characteristics of patients according to their diagnostic category are presented in Table 1. The median age of the sample was 74 years, interquartile range $(IQR) \pm 5$. Most, 65%, were male. The median number of years in education was 11 IQR \pm 4 years. The majority of patients were diagnosed with parkinsonism with 109(82%) having idiopathic PD and 11(8%)having vascular-type parkinsonism. The remaining 10% (*n* = 13) had a diagnosis of LBD. Most patients had either dementia (43%) or MCI (32%). The median time between diagnosis and the assessment was 7 IQR \pm 7 years and the median UPDRS (total score) was 37 IOR \pm 26. While patients with normal cognition were statistically younger (p = 0.02), there was no clinically meaningful difference in age according to cognitive status. There was no statistically significant difference in sex, years of education or time since diagnosis between the diagnostic categories. The median SMMSE score was 26/30 $(IQR \pm 6)$ for the total sample.

Table 2 Sensitivity, Specificity, and diagnostic accuracy for each cognitive screening instrument based upon area under the receiver operating characteristic c for the ontimal cut-off scores for coentive inmairment	ble	pecificity, and diagnostic accuracy for each cognitive screening instrument based upon area under the r	optimal cut-off scores for cognitive impairm
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3	Diagnostic		MoCA				Qmci screen	creen		Comparison
	category	Youden's Index (Optimal cut-off)	Sensitivity	Specificity	AUC (95% CIs)	Youden's Index (Optimal cut-off)	Sensitivity	Sensitivity Specificity	AUC (95% CIs)	AUC comparison $p = x$
o M	MCI from Normal Cognition	0.45 (<23/30)	0.66	0.79	0.72 (0.67-0.88)	0.75 (<60/100)	0.84	0.91	0.90 (0.80-0.99)	<i>p</i> =0.014
P P	MCI and Dementia	0.58 (<19/30)	0.87	0.71	0.92 (0.81-0.95)	0.71 (<49/100)	0.87	0.84	0.89 (0.82-0.99)	<i>p</i> =0.397
a IL C N	Normal from Cognitive Impairment (MCI and Dementia)	0.67 (<22/30)	0.76	16.0	0.89 (0.84-0.95)	0.85 (<60/100)	0.94	16.0	0.94 (0.87-0.99)	<i>p</i> =0.143
d L	Dementia	0.7 (< 19/30)	0.87	0.83	0.95 (0.88-0.97)	0.78 (<53/100)	0.93	0.85	0.93 (0.88-0.99)	p = 0.315

A strong, positive correlation was seen between both CSIs, r=0.87, p<0.001. The median MoCA score for all patients was 19/30 (IQR ± 10) with statistically significant differences between the diagnostic groupings (p<0.001). The median Qmci screen score was 51/100 (IQR ± 17) and again there was a statistically significant gradient in median values between persons with normal cognition, MCI and dementia (p<0.001). The median administration time for the Qmci screen was 5.19 IQR ± 1.4 minutes compared to 9.24 IQR ± 1.3 minutes for the MoCA, a statistically significant difference, p<0.001.

Table 2 shows the optimal cut-off scores with sensitivity and specificity values according to Youden's Index for both CSIs for different comparisons. The diagnostic accuracy based on AUC are also provided and presented visually as ROC curves in Fig. 1. The Qmci screen had excellent accuracy, which was statistically significantly greater for separating patients with normal from those with MCI compared to the MoCA (AUC 0.90 versus 0.72, p = 0.01). The Qmci screen at a cut-off of < 60/100 had a sensitivity of 84% and specificity of 91%, while the MoCA's optimal cut-off for MCI < 23/30 with lower sensitivity (66%) and specificity (79%) in this sample. Both instruments had similar accuracy in identifying cognitive impairment (either MCI or dementia) from normal and for differentiating MCI from dementia.

DISCUSSION

This study compares the diagnostic accuracy of two short CSIs, the MoCA and the Qmci screen, in their ability to differentiate those with subjective cognitive symptoms but normal cognition from those with CI due to movement disorders (Parkinsonism and LBD). The results show that the Qmci screen had statistically significantly greater accuracy in separating normal cognition from MCI than the MoCA but equivalence for other comparisons. This is reflected by a greater and better balance between sensitivity and specificity. The Qmci screen also had a significantly shorter administration time, approximately half that of the MoCA, another key feature of an ideal CSI for this population [23].

This middle-aged and older sample was predominantly comprised of patients with idiopathic PD and despite a moderate median UPDRS (total) scores, a large proportion (75%) of this sample had CI. Given that the median time since diagnosis in this study to the date of the assessment was seven years

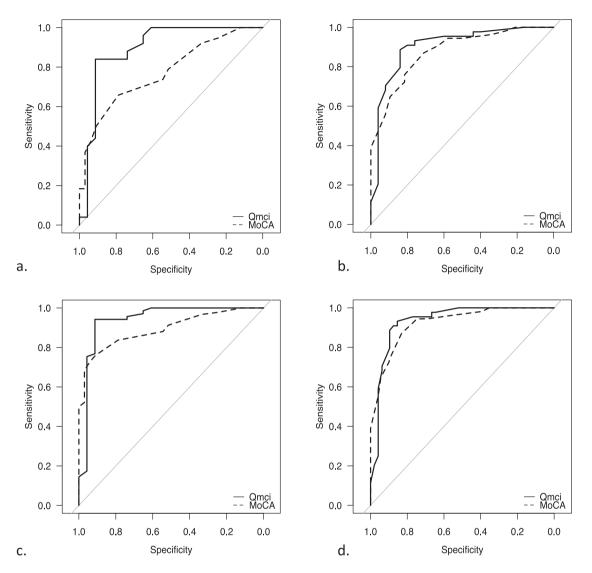


Fig. 1. Receiver Operating Characteristic curves demonstrating the accuracy of the Quick Mild Cognitive Impairment (Qmci) screen and Montreal Cognitive Assessment (MoCA) in differentiating patients with movement disorders experiencing cognitive symptoms (Parkinson's disease and Lewy Body Dementia) in separating (a) mild cognitive impairment (MCI) from normal cognition, (b) MCI and dementia, (c) normal from cognitive impairment (MCI and dementia), (d) Dementia from everything else.

and this is likely an under-estimation, this likely explains the high prevalence of CI found in this sample. Evidence suggests that the mean time from PD onset to PDD is approximately 10 years [54]. While this study included middle-aged adults > 40 years, the median age of the sample was much older (median age 74 years \pm 5), likely reflecting the service (geriatrician-led clinics). Most patients, 65%, were male, consistent with the higher prevalence of movement disorders and in particular PD, among male patients [55]. The optimal cut-off for the Qmci screen for differentiating CI (MCI and dementia) and MCI from normal cognition at < 60/100 in this sample is similar to that of the established cut-off found in an Irish sample attending a memory clinic (mainly with Alzheimer's disease related cognitive decline) [38]. It is also similar to that found in a large pooled analysis of patients in Canada [56]. It also highlights that the traditional MoCA cut-off of < 26/30 [25] is unsuitable for classifying older patients in this population. Instead, a lower cut-off of < 23/30, produced a good sensitivity of 76% and excellent specificity of 91% for CI (MCI and dementia). This lower value has been found in other studies [57] including other language versions of the MoCA and populations with lower levels of educational attainment [58].

Limitations

This study has a number of limitations. As this study was a secondary analysis of an existing database (gathered approximately ten years ago), rather than a primary data collection and thus lacks a priori power calculation, limits the available sample. This small sample size hence likely under-powers the study, limiting the inferences that can be drawn. The timing of data collection, however, is unlikely to have affected the results. This study mainly included those with idiopathic PD, who represented the majority of the attendees. This reduces the generalizability of the study for other movement disorders especially LBD. Further, the population was largely comprised of older adults, reducing the generalizability of the findings, particularly to patients attending neurology clinics. This said, movement disorders are predominantly conditions of older people [5, 6]. In addition, the study was conducted at a single center, with a homogenous population, further reducing generalizability. Another limitation is the potential for misclassification. The diagnosis of PD-MCI remains contentious with much variation in how ADLs are evaluated across studies, a challenge that is compounded by difficulty in distinguishing cognitive and motor effects of PD [59]. In this study the diagnosis was made independently and clinically by physician experts in cognition and movement disorders in conjunction with the cognitive component of the UPDRS and an established battery of cognitive assessments. While this approach is not a gold standard, it is widely used and accepted in clinical research. Furthermore, the diagnosis of cognitive impairment was sometimes based on a single cognitive assessment, which may have reduced accuracy. That said, it is argued that where clinical findings are robust, more detailed neuropsychological assessment may not be required [15]. Nevertheless, there is potential for misclassification bias. As is common to such studies, the prevalence of cognitive impairment was high in this sample (75%), potentially resulting in in spectrum bias.

Further research comparing the Qmci screen and MoCA in a larger and adequately powered sample and to other cognitive scales, as recommended by the Movement Disorder Society [28] such as the Mattis Dementia Rating Scale Second Edition [60] and the Parkinson's Disease-Cognitive Rating Scale [61], is now required.

In conclusion, the Qmci screen is a valid measure of CI in a population with movement disorders and compares favorably with the widely-used MoCA. Indeed, it has a significantly shorter administration time and greater accuracy in separating patients with PD and LBD related MCI from those reporting subjective cognitive symptoms but with normal cognition. Given the recognized challenges of using the MoCA in this sample, in particular its floor and ceiling effects in some populations and especially those with lower levels of education [62], the Qmci screen may be the better CSI, albeit more research is required to investigate this further.

AUTHOR CONTRIBUTIONS

Rónán O'Caoimh (Conceptualization; Data curation; Formal analysis; Writing – original draft); Mary J. Foley (Investigation); Suzanne Timmons (Supervision); D. William Molloy (Supervision).

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CONFLICT OF INTEREST

Dr. Rónán O'Caoimh and Prof. D. William Molloy are co-copyright holders of the Quick Mild Cognitive Impairment Screen. The authors report no other conflict of interest.

DATA AVAILABILITY

Available on request.

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