

Exploration of Risk Factors for Mild and Major Neurocognitive Disorders in A Sample of Elderly Population

Azza Saber Abdelaziz*, Haydy Hassan Sayed, Omneya Ibrahim, Aya Elhusseiny, Ashraf El Tantawy

Department of Psychiatry and Neurology, Faculty of Medicine, Suez Canal University, Egypt

*Corresponding author: Azza Saber Abdelaziz, Mobile: (+20) 01095008139, Email: azzasaber@med.suez.edu.eg

ABSTRACT

Background: Early identification and management of modifiable risk factors for neurocognitive disorders is becoming more important to slow progression of the disease, which would be very beneficial for both the patient and the caregiver.

Aim: Assessing risk factors for mild and major neurocognitive disorders among a sample of elderly population in Suez Canal Area.

Patients and Methods: This cross-sectional comparative analytical study was conducted on a sample of 156 elderly people ≥ 60 years old in Suez Canal Area over the period from March 2022 to February 2023. Study tools included a semi-structured clinical interview to assess sociodemographic, medical and lifestyle risk factors, DSM-5 criteria to diagnose mild and major neurocognitive disorders, The Montreal Cognitive Assessment scale to assess cognitive function, and The Activities of Daily Living Questionnaire to assess functional impairment and dependency.

Results: Mild and major neurocognitive disorders have multiple sociodemographic, medical and lifestyle risk factors, including: aging, lower education, female gender, non-married status, unemployment and physical work, lower income, less physical, cognitive and social activities, increased number of chronic diseases and family history of cognitive impairment.

Conclusion: Multiple modifiable risk factors for mild and major neurocognitive disorders could be identified, and their management may contribute to lowering burden of neurocognitive disorders.

Keywords: Risk factors, Neurocognitive disorder, Cognitive impairment, Elderly.

INTRODUCTION

Mild neurocognitive disorder (mild NCD) and major neurocognitive disorder (major NCD) are characterized by cognitive decline that develops slowly in elderly population. The difference between mild and major neurocognitive disorders is mainly based on the severity of symptoms. Major NCD is characterized by significant cognitive decline and the development of dependence in daily life activities, while mild NCD is characterized by mild cognitive decline not affecting independence in daily activities. To be diagnosed, delirium and other mental disorders should be ruled out ^[1].

Previous studies have identified risk factors for neurocognitive disorders, including sociodemographic factors (e.g., sex, age, years of education and living conditions), health factors (e.g., hearing loss, cardiovascular diseases, hypertension, diabetes) and lifestyle factors (e.g., smoking and lack of physical activity) ^[2].

Identifying modifiable risk factors is essential element of preventive measures for mild and major neurocognitive disorders and is necessary for designing and implementing public health programs and healthcare policies that target high-risk groups ^[3].

This study aimed to investigate risk factors for neurocognitive disorders among a sample of elderly population in Suez Canal Area.

PATIENTS AND METHODS

Participants

This cross-sectional study was performed on 156 participants aged ≥ 60 years old in Suez Canal Area, Egypt, over 12 months from March 2022 to February 2023. Study participants were recruited from two settings: 78 subjects from geriatric homes and 78 subjects from primary health care centers by convenience sampling.

Eligibility criteria

The following subjects were excluded from participation in the study: people with mental retardation or a severe uncontrolled medical condition, people with a mental disorder or drug intake (e.g.: anticholinergic drugs, antipsychotics, etc.) that may interfere with cognitive testing, as well as people with severe visual, hearing, speech, or functional impairment that would limit their participation in the study.

Procedure

The researchers conducted a detailed semi-structured interview with each participant to collect the following data:

- 1) Sociodemographic data, occupational data regarding current work status and occupational style (physical work vs. brainwork) and income.
- 2) Lifestyle factors such as smoking, substance use, participation in cognitive, physical and social

activities, and family history of cognitive impairment.

- 3) Chronic health conditions, medication history and adherence to treatment.

We diagnosed subjects having neurocognitive disorders by the DSM-5 criteria [4] and the Montreal Cognitive Assessment Scale (MoCA), and classified study subjects according to cognitive status into three groups: normal cognition, mild neurocognitive disorder, and major neurocognitive disorder. We explored association of cognitive function with variable risk factors.

Tools:

1. The Montreal Cognitive Assessment Scale (MoCA):

It is a 30-point test assessing several cognitive domains including attention/working memory, short-term memory, visuospatial abilities, language, executive functioning, and orientation. One additional point is added to illiterate participants. The Arabic version was translated and validated by **Rahman and El Gaafary** [5].

2. The Activities of Daily Living Questionnaire (ADLQ):

It assesses the degree of dependency in performing basic activities of daily living including six functional domains: bathing, dressing, going to the toilet, transferring, continence, and feeding.

Each domain had a score of 0 (complete dependence), 0.5 (partial independence), or 1 (complete independence). A score of 6 indicated complete independence and 0 complete dependence. The used Arabic version was translated and validated by **Nasser and Doumit** [6].

Ethical Approval

Participants signed informed written consent after acknowledgement about the study, and the research proposal was approved by the Ethical Committee of Suez Canal University, Faculty of Medicine (IRB: 4778; 3 January 2022). This study has been carried out following The Helsinki Declaration.

Statistical analysis

All statistical analyses were performed using the *SPSS* (Statistical Package for The Social Sciences) version 26. Continuous data were presented as mean, standard deviation (SD), median, and range and were compared by independent t-test for normally distributed data and by Mann-Whitney U test and Kruskal Wallis test for non-normally distributed data. Categorical data were presented as frequency and percentage and were compared by Chi-Square test or Fisher's exact test. Spearman correlation test was used for correlations and multiple logistic regression analysis for predicting factors was also used. The $p < 0.05$ value indicated the level of significance.

RESULTS

Mean of age was significantly higher in major NCD group compared to normal and mild NCD groups, and most males had normal cognition compared to females. The majority of normal cognition group were married (51.4%) and highly educated compared to mild and major NCD groups with highly significant difference. Most of major NCD group lived in geriatric homes (85.7%) while 62.8% of normal cognition group lived with their families. Most of unemployed participants had mild NCD, and subjects who were still working had less neurocognitive disorders compared to retired and unemployed subjects. The majority of subjects with brainwork and enough income with saving had normal cognition (Table 1).

Table 1: Association between sociodemographic variables and cognitive function

Variables	Normal cognition (n=70)	Mild NCD (n=58)	Major NCD (n=28)	p value
Gender, n (%)				
Male	50 (71.4)	29 (50)	15 (53.6)	0.035^{*a}
Female	20 (28.6)	29 (50)	13 (46.4)	
Age, n (%)				
Mean± SD	67.0± 4.8	68.1± 6.2	73.6± 6.7	<0.001^{**b}
60-69	46 (65.7)	28 (48.3)	8 (28.6)	
70-79	22 (31.4)	27 (46.6)	14 (50)	
≥ 80	2 (2.9)	3 (5.2)	6 (21.4)	
Marital status, n (%)				
Single	0	2 (3.4)	2 (7.1)	<0.001^{**c}
Married	36 (51.4)	18 (31)	2 (7.1)	
Widowed	18 (25.7)	24 (41.4)	18 (64.3)	
Divorced	16 (22.9)	14 (24.1)	6 (21.4)	
Educational level, n (%)				
Illiterate	0	6 (10.3)	14 (50)	<0.001^{**a}
Primary school	8 (11.4)	16 (27.6)	0	
Preparatory school	8 (11.4)	22 (37.9)	10 (35.7)	
Secondary school	26 (37.1)	10 (17.2)	4 (14.3)	
University graduate	28 (40)	4 (6.9)	0	
Number of children				
Mean± SD	3.1± 1.4	3± 1.5	3± 1.6	0.684 ^b
Living condition, n (%)				
Geriatric homes	26 (37.1)	28 (48.3)	24 (85.7)	<0.001^{**c}
With a family member	16 (22.8)	14 (24.1)	2 (7.1)	
Nuclear family	14 (20)	6 (10.3)	2 (7.1)	
Extended family	14 (20)	10 (17.2)	0	
Work status, n (%)				
Retired	50 (80)	30 (51.7)	16 (57.1)	<0.001^{**c}
Unemployed	2 (2.9)	16 (27.6)	10 (35.7)	
Still working	12 (17.1)	12 (20.7)	2 (7.1)	
Occupational style, n (%)				
None	2 (2.9)	16 (27.6)	10 (35.7)	<0.001^{**a}
Brainwork	34 (48.6)	2 (3.4)	0	
Physical work	34 (48.6)	40 (69)	18 (64.3)	
Income, n (%)				
Not enough	18 (25.7)	20 (34.5)	18 (64.3)	0.001^{*a}
Enough	32 (45.7)	32 (55.2)	8 (28.6)	
Enough with saving	20 (28.6)	6 (10.3)	2 (7.1)	

a: Chi-square test, b: Mann-Whitney test, c: Fisher Exact test, *: significant, **: highly significant

Mild NCD and major NCD were significantly associated with pulmonary disease and heart disease. There was a highly significant association with bone and joint disorders, hyperlipidemia, and with adherence to treatment (Table 2).

Table 2: Association between chronic medical diseases and cognitive function

Variables	Normal Cognition (n=70)	Mild NCD (n=58)	Major NCD (n=28)	p value
Diabetes mellitus, n (%)				
No	52 (74.3)	38 (65.5)	14 (50)	0.068 ^a
Yes	18 (25.7)	20 (34.5)	14 (50)	
Duration of diabetes mellitus, n (%)				
< 10 years	2 (11.1)	4 (20)	0	0.215 ^b
≥ 10 years	16 (88.9)	16 (80)	14 (100)	
Treatment of diabetes mellitus, n (%)				
Insulin	12 (66.7)	12 (60)	4 (28.6)	0.078 ^a
Oral hypoglycemic	6 (33.3)	10 (50)	10 (71.4)	0.102 ^a
Hypertension, n (%)				
No	40 (57.1)	24 (41.4)	14 (50)	0.207 ^a
Yes	30 (42.9)	34 (58.6)	14 (50)	
Cerebrovascular (Stroke or TIA), n (%)				
No	70 (100)	56 (96.6)	26 (92.6)	0.063 ^b
Yes	0	2 (3.4)	2 (7.1)	
Heart disease, n (%)				
No	60 (85.7)	40 (69)	18 (64.3)	0.027**^a
Yes	10 (14.3)	18 (31)	10 (35.7)	
Pulmonary disease, n (%)				
No	70 (100)	52 (89.7)	24 (85.7)	0.003**^b
Yes	0	6 (10.3)	4 (14.3)	
Bone and joint disorders, n (%)				
No	48 (68.6)	42 (72.4)	8 (28.6)	<0.001***^a
Yes	22 (31.4)	16 (27.6)	20 (71.4)	
Thyroid disease, n (%)				
No	68 (97.1)	56 (96.6)	28 (100)	>0.624 ^b
Yes	2 (2.9)	2 (3.4)	0	
Cancer, n (%)				
No	64 (91.4)	58 (100)	26 (92.6)	0.079 ^b
Yes	6 (8.6)	0	2 (7.1)	
Hyperlipidemia, n (%)				
No	50 (71.4)	26 (44.8)	10 (35.7)	<0.001***^a
Yes	20 (28.6)	32 (55.2)	18 (64.3)	
Obesity, n (%)				
Normal	34 (48.6)	28 (48.3)	12 (42.9)	0.322 ^b
Overweight	4 (5.7)	0	2 (7.1)	
Obese	32 (45.7)	30 (51.7)	14 (50)	
Number of physical illnesses, n (%)				
Mean± SD	1.6± 1.5	2.3± 1.4	3± 1.8	0.074 ^c
Median (Range)	1 (0-5)	2 (0-5)	3 (0-7)	
Adherence to treatment, n (%)				
No	18 (34.6)	8 (16)	16 (61.5)	<0.001***^a
Yes	34 (65.4)	42 (84)	10 (38.5)	

a: Chi square test, b: Fisher Exact test; c: Kruskal Wallis test*: significant,**: highly significant
TIA: Transient ischemic attack

No significant association was found between cognitive function and smoking or history of substance use (Table 3).

Table 3: Association of smoking and substance use with cognitive function

Variables	Normal cognition (n=70)	Mild NCD (n=58)	Major NCD (n=28)	p value
Smoking				
Never smoker	15 (21.4)	15 (25.9)	8 (28.6)	0.535 ^a
Ex-smoker	15 (21.4)	9 (15.5)	2 (7.1)	
Current smoker	40 (57.1)	34 (58.6)	18 (64.3)	
History of Substance use				
No	66 (94.3)	56 (96.6)	27 (96.4)	0.878 ^a
Yes	4 (5.7)	2 (3.4)	1 (3.6)	

a: Fisher Exact test

Mild NCD patients had significantly lower physical, cognitive, and social activities than elderly with normal cognition and significantly higher than major NCD. Family history of cognitive impairment was significantly higher among mild NCD group. Major NCD group had significantly lower ADLQ score (Table 4).

Table 4: Association of lifestyle risk factors, family history of cognitive impairment and ADL score with cognitive function

Variables	Normal cognition (n=70)	Mild NCD (n=58)	Major NCD (n=28)	p value
Physical activity (hours/week), n (%)				
<1	8 (11.4)	12 (20.7)	16 (57.1)	<0.001 ^{**a}
1-2	32 (45.7)	30 (51.7)	10 (35.7)	
> 2	30 (42.9)	16 (27.6)	2 (7.1)	
Cognitive activity, n (%)				
No	44 (62.9)	56 (96.6)	28 (100)	<0.001 ^{**a}
Yes	26 (37.1)	2 (3.4)	0	
Social activity (times/week), n (%)				
0-3	10 (14.3)	16 (27.6)	21 (75)	<0.001 ^{**b}
4-6	36 (51.4)	30 (51.7)	7 (25)	
7-9	24 (34.3)	12 (20.7)	0	
Family history of cognitive impairment, n (%)				
No	66 (94.3)	46 (79.3)	26 (92.9)	0.029 ^{*b}
Yes	4 (5.7)	12 (20.7)	2 (7.1)	
ADLQ score				
Mean± SD	6± 0	6± 0	4.2± 0.9	<0.001 ^{**b}

a: Chi-square test, b: Fisher Exact test, *: significant, **: highly significant. ADLQ: Activities of Daily Living Questionnaire.

MoCA total score showed significant positive correlations with educational level, income, physical activity duration and ADLQ score. It also showed significant negative correlations with age, number of physical illnesses, family history of cognitive impairment, female gender, single marital status, and retired work status (Table 5).

Table 5: Correlation between MoCA total score and other study variables

Variables	MoCA Total Score	
	r	p value
Gender (female)	-0.174	0.030*
Age	-0.334	0.000 ^{**}
Marital status (single)	-0.219	0.006*
Education (university)	0.700	0.000 ^{**}
Work status (retired)	-0.206	0.010*
Income	0.359	0.000 ^{**}
Number of physical illnesses	-0.321	0.000 ^{**}
Physical activity duration (hour/week)	0.388	0.000 ^{**}
Family history of cognitive impairment	-0.373	0.006*
ADLQ score	0.468	<0.001 ^{**}

*: significant, **: highly significant, MOCA: Montreal Cognitive Assessment.

Family history of cognitive impairment, physical work, bone and joint disorders, and age were significant positive predictors for mild NCD. Meanwhile, adherence to treatment, education, increased cognitive activity and physical activity duration were significant negative predictors for mild NCD (Table 6).

Table 6: Multiple logistic regression analysis for predictors of mild neurocognitive disorder

Predictor	B	95% CI	Exp(B)	p value
Age	2.040	0.22-0.970	1.041	0.016*
Education (preparatory)	-4.667	6.297-1795.774	106.337	0.001*
Occupation style (physical work)	3.457	0.003-0.292	0.032	0.002*
Bone and joint disorders	2.636	2.399-81.288	13.964	0.003*
Adherence to treatment	-4.262	0.002-0.122	0.014	0.000**
Cognitive activity	-3.878	4.155-562.565	48.346	0.002*
Family history of cognitive impairment	3.943	0.002-0.236	0.019	0.002*
Physical activity duration	-2.771	0.031-0.950	3.276	0.011*

*: significant,**: highly significant.

Number of chronic diseases, family history of cognitive impairment and age were significant positive predictors for major NCD, while physical activity duration, preparatory education and income were significant negative predictors for major NCD (Table 7).

Table 7: Multiple logistic regression analysis for predictors of major neurocognitive disorder

Predictor	B	95% CI	Exp(B)	p value
Age	2.200	0.014-0.871	0.111	0.037*
Education (preparatory)	-1.524	0.038-1.165	3.716	0.021*
Income	-4.554	0.000-0.611	0.011	0.028*
Number of chronic diseases	3.219	2.249-277.940	25.000	0.009*
Physical activity duration	-3.672	0.15-0.412	9.872	0.002*
Family history of cognitive impairment	1.691	0.073-0.708	1.997	0.011*

*: significant.

DISCUSSION

In this study, we aimed to assess risk factors associated with mild and major neurocognitive disorders, in order to provide interventions for modifiable risk factors in elderly population.

Age correlated negatively with MoCA total score, and it was a predictor for both mild and major neurocognitive disorders. This could be explained by aging-related factors such as nerve cell degeneration, higher risk of cardiovascular disease, and changes in nucleic acid and cell structure. This finding agrees with multiple previous studies [7-9].

Education had a positive correlation with MoCA total score, and low education was a predictor for both mild and major neurocognitive disorders. These results are consistent with previous research [7,9,10]. Higher education levels may lead to increased cognitive reserve of the brain that protects against age-related cognitive impairment, and can also be linked to better socioeconomic status and following a healthy lifestyle.

Our study showed a significant association between female gender and neurocognitive disorders. This may be attributed to hormonal differences and decreased estrogen levels in females after menopause [11].

However, this association is controversial, as it was supported by some studies [9,10] while other studies showed no gender difference [7].

Regarding marital status, neurocognitive disorders had significant association with being widowed, while majority of normal subjects were married. This is in agreement with other studies [9,12]. This may be attributed to lack of emotional support and social interaction that can provide daily cognitive stimulation. However, other studies found no association [8].

Our study found no association between number of children and cognitive function, although an Egyptian study found a significant relationship between contact frequency with children and positive cognitive function [13]. We did not assess contact with offspring, which may have had a confounding effect.

Association of neurocognitive disorders with being retired or unemployed is consistent with other studies [8,14]; as cognitive impairment would make it difficult to work, or employment may act the same as education in improving cognitive reserve of the brain.

Neurocognitive disorders were more common among those who had physical work compared to

brainwork, and physical work was a positive predictor for mild NCD. Our results are in line with a literature review of Egyptian studies by **Sabry et al.** [8]. Physical work may provide less cognitive stimulation compared to brainwork, and may be associated with other factors like low education and lower income.

Participants with higher income had better cognitive function, and it was a negative predictor for major NCD. This is consistent with other studies [15,16], and may be linked to the level of education and access to care and treatment.

Our results showed significant association with heart disease, in line with previous research [17] interpreting this association by heart disease resulting in decreased cerebral blood flow and being a risk factor for cerebral emboli.

Pulmonary disease was also associated with cognitive impairment, which is supported by other studies [13,18] explaining this by hypoxic damage to the brain resulting from poor pulmonary function, or the presence of shared risk factors.

There was a significant association of neurocognitive disorders with bone and joint disorders, which were predictors of mild NCD. This finding is consistent with other studies [19,20] attributing it to shared risk factors such as aging, systemic inflammation, vitamin D deficiency, and physical inactivity.

We found no significant association between neurocognitive disorders and history of cancer, although previous research linked cognitive impairment to cancer and its treatments [21]. This finding may be due to the low presentation of cancer cases in our sample (5.1% only).

The current study showed highly significant association between hyperlipidemia and cognitive impairment, which is in line with previous research [12,22].

There was no significant association between obesity and neurocognitive disorders, similar to another Egyptian study by **Abdulsalam et al.** [23]. The relationship between obesity and cognitive function is conflicting as some researchers found that obesity in mid-life is a predictor of cognitive impairment at old age while it is protective at late life [24,25]. Our relatively small sample may have led to the inability to observe any association.

We found a highly significant association between adherence to treatment and normal cognitive function, and it was a negative predictor for mild NCD. This association is established in previous studies [26,27]. The relationship is reciprocal as non-adherence to treatment causes increased severity and complication of diseases, while impaired memory may decrease adherence to treatment.

No association was found between neurocognitive disorders and diabetes mellitus or hypertension, in agreement with **Khater and Abouelezz** [28], but previous

studies have shown a significant relationship [18,29], while **Amer et al.** [30] found a significant association with hypertension but not with diabetes mellitus.

We found no significant association between neurocognitive disorders and history of cerebrovascular disease or thyroid disease, which was inconsistent with the literature that shows an increased risk [18,31,32]. This may be due to the very low presentation of those subjects in our sample (2.6% each).

In the current study, neurocognitive disorders were associated with number of chronic diseases, which were predictor for major NCD. This result is in agreement with other studies showing that multimorbidity was significantly associated with cognitive impairment, even more than the severity of each disease separately [9,16].

There was no significant association between smoking and neurocognitive disorders, which agrees with results of an Egyptian literature review by **Sabry et al.** [8], although cognitive impairment was linked to smoking in some other studies [10,33].

There was no significant association between history of substance use and cognitive impairment in our study; although one study [34] documented this association in old age people. This result may be due to the small number of substance users in our sample (4.5%). Substance use is often denied due to stigma in the community [9].

Physical activity had a highly significant association with normal cognition and was a negative predictor for both mild NCD and major NCD. These results are consistent with previous studies [33,35] proposing that physical activity significantly improves the management of cardiovascular risk factors, and may increase the production of neurotransmitters, and support neurogenesis and synaptic plasticity.

There was a highly significant association between cognitive activity and normal cognition, and it was a negative predictor for mild NCD. This is in line with other studies [33,36]. Activities requiring mental effort such as reading, crosswords or Sudoku puzzles may stimulate more brain wiring and neuroplasticity.

There was a highly significant association between social activity and normal cognition. This result is in line with previous research [36]. Social interaction on a regular basis stimulates the mind and can enhance cognitive reserve.

Family history of cognitive impairment was found to have a significant association with neurocognitive disorders, being a positive predictor for both mild and major neurocognitive disorders. This is in agreement with previous studies [9,12,29], and emphasizes the hereditary component of neurocognitive disorders.

In the current study, the score of Activities of Daily Living Questionnaire (ADLQ) was associated with major NCD and correlated positively with MoCA total

score. This result is in line with previous research [37]. Difficulties in performing the activities of daily living may be due to the poor physical function and muscle strength coexisting with cognitive impairment in elderly.

CONCLUSION

Mild NCD and major NCD have multiple risk factors, from which the most important are higher age, lower education, female gender, non-married status, unemployment and physical work, lower income, less physical, cognitive and social activities, increased number of chronic diseases and family history of cognitive impairment. Controlling modifiable risk factors can contribute to lowering burden of neurocognitive disorders.

DECLARATIONS

Conflicting interests:

The authors declare no potential conflict of interest.

Funding:

The authors received no financial support.

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