Malnutrition in Patients with Parkinson's Disease: Associated Clinical Factors

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Abstract |

Objective: Parkinson's disease (PD) is a chronic, progressive disease commonly affecting the elderly. Among patients with PD (pwPD), the ones above 60 years old are considered to be high risky in terms of malnutrition. Weight loss is a common complaint in pwPD. Thus, we aimed to define the risk factors of malnutrition in geriatric pw PD.

Materials and Methods: We enrolled 66 pwPD above the age of 60 years old. Socio-demographic features were recorded, comprehensive geriatric assessments were evaluated. Malnutrition was assessed with mini-nutritional assessment questionnaire. Anthropometric measurements including body mass index, mid- upper arm circumference, calf circumference (CC) was recorded.

Results: Seven (10.6%) pwPD had malnutrition, 22 (33.3%) pwPD were under the risk of malnutrition. Univariate logistic regression analysis results revealed that low CC, presence of dyskinesia, advanced Hoehn&Yahr stage, levodopa doses of \geq 400 mg/day, difficulty in swallowing (p=0.035, p=0.041, p=0.048, p=0.027 and p=0.007, respectively) were strongly related to malnutrition among the pwPD. Difficulty in swallowing was independently related to malnutrition in pwPD [odds ratio: 7.81 (confidence interval: 2.17-28.10), p=0.002].

Conclusion: PD is the second common neurodegenerative disease in geriatric population that is likely to cause malnutrition as a result of several disabling symptoms in the progressive course of the disease such as dysphagia. To avoid or delay the poorer outcomes, clinicians should be careful to identify malnutrition with appropriate screening tools during follow-up of pwPD.

Keywords: Parkinson disease, geriatric population, malnutrition, risk factors, mini-nutritional assessment

Introduction

Parkinson's disease (PD), the second most common neurodegenerative disease among the population above 65 worldwide, is characterized by cardinal motor including bradykinesia, rigidity, rest tremor, postural instability and nonmotor symptoms (1). Due to its progressive course, not only the disabling symptoms and complications, which are more likely to be seen as the disease advances, but bradykinesia itself affecting the gastrointestinal tract as well as other motor systems, autonomic involvement also acts an important role in the occurrence of malnutrition (2). In patients with PD (pwPD) there are many factors affecting malnutrition. It is stated that non-motor and motor symptoms, diagnosis in older age, higher levodopa equivalent daily dose/body weight, depression, dementia and hallucinations are related with malnutrition among pwPD (3). Moreover, dysphagia, delayed gastric emptying, constipation, malabsorption like disturbances in gastrointestinal system and weak hand-mouth coordination may affect the dietary status (4).

Malnutrition can be described as an imbalance between nutritional intake and requirements that eventually causes changes in body weight, body composition, and physical function

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(3). As a well-known entity, the prevalence of malnutrition in the geriatric population increases with aging, and comorbid diseases, as well as the level of care (4). In accordance with aging, the reported risk of malnutrition is 3-60%, and the prevalence of malnutrition was found to be 0-24% in pwPD who are mostly geriatric (2,5). Detecting possible malnutrition early and related lifestyle characteristics is crucial. Being prone to infection, decubital ulcer and behavioural and autonomic disorders, pwPD that do not feed well, have deficits in their quality of life (6). An improved nutritional status will also improve quality of life in pwPD.

Regarding these aspects, we aimed to investigate the possible factors affecting the risk of malnutrition in older pwPD patients who are an important part of the geriatric population.

Materials and Methods

This study was cross-sectionally designed. Sixty-six pwPD above the age of 60 were enrolled in the present study. Written approvals were taken from all the patients and the relatives of the patients with dementia before enrolment. The ethical approval was taken from the Local Ethical Committee of Erciyes University (decision no: 2016/595). The exclusion criteria were active malignancy, active infectious disease, history of cerebrovascular disease, hepatic failure, and renal failure. The socio-demographic features of the participants were recorded.

The nutritional status of the pwPD was evaluated by mininutritional assessment (MNA) questionnaire, which had 18 questions with a total score of 30 (7). The normal nutrition scores were 23.5 to 30 points. Scores 17 to 23 points were considered as "risk of malnutrition", and scores below 17 points were considered "malnutrition". The study population was categorized into two groups according to MNA scores for analysis. Patients with a MNA score \leq 23.5 points defined as group with malnutrition (including both patients with malnutrition and at risk of malnutrition), and the second group with a MNA score \geq 24 points defined as normal nutritional status.

Anthropometric assessments of the groups were performed by height in centimeter (cm), weight measurements in kilogram (kg), body mass index (BMI kg/m²), mid-upper arm circumference (MUAC) in cm, calf circumference (CC) in cm and triceps skinfold thickness (TSF) in cm. Since all the participants were pwPD, the severity of PD was assessed with the Hoehn&tYahr score (8). Disease characteristic features, therapy regimens and daily levodopa equivalent doses were recorded. Associated complaints of the pwPD including dyspepsia, constipation, and weight loss in one year were also investigated by self-reported questions regarding the related symptomatology. Swallowing function was evaluated subjectively by the question "Do you have difficulty in swallowing with solid food?"

The cognitive status of pwPD was assessed with a mini-mental

status exam (MMSE). This assessment included eleven questions with a total score of 30/30 points. Since the cut-off score for cognitive impairment is 24 points, patients with the MMSE score of 24 to 30 points were considered to be normal, while mild scores between 18-23 points were considered "mild dementia", and those \leq 17 points "severe dementia" (9).

Mood assessments for depression were performed through the geriatric depression scale which has 30 items. The scores \geq 14 points were considered as depression (10).

Statistics

The Shapiro-Wilks test, histogram, and q-q plots were examined to evaluate data normality. Independent samples t-test and Mann-Whitney U test was applied for continuous variables. Pearson chi-square test or Fisher's Exact test was applied for categorical variables. Univariate and multivariate binary logistic regression models were applied to examine the risk effect of variables on malnutrition. The odds ratios (OR) were estimated with 95% confidence intervals (CI). Significant variables with potential risk factors of malnutrition on univariate analysis with p<0.1 were taken into multivariate analysis including marital status, H&Y scale, levodopa dose, DBS, difficulty in swallowing and depression. Backward stepwise selection was applied using likelihood ratio statistic at p<0.10 stringency level. Goodness of fit was assessed by the Hosmer-Lemeshow test (p=0.840). SPSS Version 22.0 (Statistical Package for Social Sciences for Windows) database was used to organize data. P-values <0.05 were accepted to remark statistical significance.

Results

Sixty-six pwPD (37 men, 29 women) above the age of 60 participated in the study. The mean age of the pwPD was 67.50 years (minimum-maximum =60-86 years). The mean BMI of the pwPD was 30.13 ± 5.04 kg/m² (minimum-maximum =22.9-42.0 kg/m²). In the study group, 37 (56.1%) pwPD had normal nutrition status, while 22 (33.3%) had malnutrition risk, and 7 (10.6%) had malnutrition. Although malnutrition was more common among widowed patients, there were no statistically significant differences in terms of socio-demographic features between the patients regarding nutrition status (p>0.05). The comparison of the socio-demographic and clinical characteristics of the pwPD according to their nutrition status is given in Table 1.

As shown in Table 1, pwPD with malnutrion had lower CC (p=0.041) and MUAC (p=0.080) than pwPD with normal nutrition. In addition in the group of pwPD with malnutrition the stage of PD was more advanced (p=0.048) and levodopa daily doses were higher (p=0.027) than the group with normal nutritional status. Dyskinesia was also more frequent in patients with malnutrition (p=0.048). Moreover, most of the pwPD in

Table 1. The comparison of the socio-demographic and clinical characteristics of the people with Parkinson's disease according to their nutrition status

	Nutritional status					
	Malnutrition Normal					
Variables	All	(MNA score ≤23.5)	(MNA score ≥24)			
	n=66 (100)	n=29 (43.9)	n=37 (56.1)	р		
Age	67.5 (63.0-72.0)	66.0 (63.0-73.0)	68.0 (63.0-71.0)	0.990		
Gender						
Men	29 (43.9)	16 (55.2)	13 (35.1)			
Women	37 (56.1)	13 (44.8)	24 (64.9)	0.136		
BMI kg/m ²	30.1+5.1	29.3+4.8	30.9+5.2	0.231		
Education						
Illiterate	17 (25.8)	9 (31.0)	8 (21.9)			
5 years	29 (43.9)	11 (37.9)	18 (48.6)	0 600		
Over 5 years	20 (30.3)	9 (31.0)	11 (29.7)	0.009		
Marital status						
Married	51 (77.3)	19 (65.5)	32 (86.5)	0.074		
Widow	15 (22.7)	10 (34.5)	5 (13.5)	0.074		
Income						
Low	49 (74.2)	20 (69.0)	29 (78.4)	0 410		
Middle/high	17(25.8)	9 (31.0)	8 (21.6)	0.110		
MUAC, cm	29.4 <u>+</u> 3.6	28.5 <u>+</u> 4.1	30.2 <u>+</u> 3.0	0.075		
CC, cm	36.1 <u>+</u> 4.4	35.6±4.4	37.9 <u>+</u> 4.1	0.035		
TSF, mm	17.2 <u>+</u> 7.3	15.8±7.1	18.4 <u>+</u> 7.4	0.158		
Dyskinesia						
Yes	17 (26.2)	11 (39.3)	6 (16.2)	0.048		
No	48 (73.8)	17 (60.7)	31 (83.8)	0.040		
H&Y scale						
Early stages	30 (45.5)	9 (31.0)	21 (56.8)	0.048		
Advanced stages	36 (54.5)	10 (69.0)	16 (43.2)	0.010		
Levodopa dose						
≥400 mg	35 (53.0)	20 (69.0)	15 (40.5)	0.027		
<400 mg	31 (47.0)	9 (31.0)	22 (59.5)			
DBS						
Yes	8 (12.1)	7 (24.1)	1 (2.7)	0.018		
No	58 (87.9)	22 (75.9)	36 (97.3)			
Dyspepsia	10 (20.0)	10 (25.7)	0 (00 5)			
res	18 (29.0)	10 (35.7)	8 (23.5)	0.400		
NO Constinution	44 (71.0)	18 (64.3)	26 (76.5)			
	41 (C2 1)	10 (CE E)	22 (FO F)			
No	41 (02.1) 25 (37.9)	19 (05.5)	22 (59.5)	0.799		
	23 (37.3)	10 (34.3)	13 (40.3)			
Difficulty in swallowing						
Yes	22 (33 3)	15(517)	7(18.9)			
No	44 (66.7)	14(48.3)	30(81.1)	0.008		
Weight loss						
Yes	25 (37.9)	16 (55.2)	9 (24.3)			
No	41 (62.1)	13 (44.8)	28 (75.7)	0.020		
Depression						
(GDS score ≥14)						
Yes	28 (42.4)	16 (55.2)	12 (32.4)	0.000		
No	38 (57.6)	13 (44.8)	25 (67.6)	0.082		
Cognitive impairment						
(IVIIVISE score <24)						
No	10 (15.2)	6 (20.7)	4 (10.8)	0.315		
	56 (84.8)	23 (79.3)	33 (89.2)	0.313		
Values are stated as n (%), mean ± SD or median (1 st -3 rd quartiles). BMI: Body mass index, CC: Calf circumference, DBS: Deep brain stimulation, H&Y:Hoehn and Yahr, MUAC: Mid-upper arm circumference, TSF: Triceps skin fold thick ness, GDS: Geriatric depression scale, SD: Standard deviation, MMSE: Mini-mental status exam						

malnutrion group were under the treatment of deep brain stimulation (DBS) (p=0.018), had difficulty in swallowing (p=0.008) and had weight loss (p=0.020) (Table 1).

The univariate logistic regression analysis of the data revealed a strong relation between the marital status, CC, dyskinesia, stage of PD, levodopa doses, DBS procedure, difficulty in swallowing, weight loss and malnutrition. Multiple regression analysis demonstrated an independent relationship between difficulty in swallowing and malnutrition (OR: 7.81, Cl: 2.17-28.10, p=0.002). The results of the univariate and multivariate logistic regression analysis determining the risk factors of malnutrition are shown in Table 2.

Discussion

PD is an important disabling neurodegenerative disease which interferes with patients' quality of life and more prone to affect the geriatric population (11,12). Not only the motor problems including rigidity, tremor, postural instability, and bradykinesia leading to dysphagia, constipation and other problems in daily activities, but also health-related problems such as mood changes, cognitive decline, and fatigue may lead to malnutrition in pwPD (4). In this study we observed that presence of dyskinesia, advanced stages of PD, higher levodopa doses, DBS procedure, difficulty in swallowing and depression were strongly related with malnutrition in pwPD. Among these variables difficulty in swallowing was independently related with malnutrition in pwPD.

Malnutrition is common in pwPD but often under-reported both by the patients and clinicians. The main reasons why pwPD are at high risk of malnutrition are, firstly, disease characteristic features, defined as the motor findings of the disease, secondly, the negative effects of the disease on nutrition in older individuals such as depression, cognitive damage, which are highly prevalent in pwPD as non-motor symptos, and thirdly, the drugs used for the treatment of PD (3). In addition studies have shown that PD patients have a lower BMI than age matched healthy controls (13). Since there is an increased risk of malnutrition reported in the literature for pwPD, it is crucial to screen nutrition in pwPD (14,15). Similar to our study, the common methods used to assess the nutritional status are anthropometric measurements including the weight and BMI, and the MNA questionnaire which is the most frequently utilized tool for nutritional status (14).

In the literature, the prevalence of malnutrition and malnutrition risk has been stated up to 24% for malnutrition, and 60% for malnutrition risk in PD (3,11). Similar to the literature, our results revealed a malnutrition rate of 43.9 % which approximately corresponds to half of our study population. Tomic et al. (3) examined 96 patients, and from among 96 patients, 55.2% were at risk of malnutrition, while 8.3% has already been malnourished. There are several determinants of malnutrition has been implicated in PD patients. It is announced that age, severity of motor symptoms, duration of the disease and intensity of stage, especially "off" states, rigidity dominant type with "off" periods, mostly affect the nutritional status (3,16). Moreover, Fávaro-Moreira et al. (17) analysed the risk factors of malnutrition among older adults above 65 years old and reported that age and PD were independent risk factors for malnutrition. In the above-mentioned study, the presence of PD in older individuals was found to be independently associated with malnutrition, reflecting that PD poses a very high risk for malnutrition.

Interestingly, pwPD are shown to be overweight in the beginning stages of the disease, but as the disease progresses and the patients end up in the advanced conversely, lower BMI and weight loss are reported to be extremely common and the latter was shown to be associated with nigrostriatal depletion, cognitive impairment, deteriorated motor functions and a poorer quality of life (18). In this study, BMI, which is one of the anthropometric determinants of malnutrition, which is frequently used in clinics, was not observed to be associated with malnutrition in PD, whereas decreased MUAC and CC were closely associated with malnutrition. In this case, although it has been stated that BMI in PD patients is lower than normal healthy controls, especially in advanced stages, we observed that BMI alone may not be sufficient in PD patients in the evaluation of malnutrition. It may be more effective to use a valid and safe screening tool such as MNA in the evaluation of malnutrition in these patients. Since one of the main manifestations of malnutrition is weight loss, the possible risk factors of weight loss in pwPD may include dysphagia which may lead to low dietary uptake, slowed gastric motility and emptying as a result of bradykinesia, and increased energy consumption as a consequence of levodopa-induced dyskinesia in some patients (19). Similar to the literature, our results disclosed that the risk of malnutrition was significantly higher in patients experiencing the advanced stage, in need of increased daily doses of levodopa, have difficulty in swallowing, levodopa-induced dyskinesia, and weight loss. However, the possible reasons for malnutrition in pwPD, apart from weight loss, and considered "not related" to weight loss actually, are hyposmia, reduced appetite, changed reward mechanism due to degeneration in the mesocorticolimbic network, and decreased levels of orexin (20,21).

In this study difficulty in swallowing was only clinical determinant that was independently related with malnutrition in PD and 33 percent of the patients had difficulty in swallowing. It has been reported that about 80% of patients with PD evolve dysphagia in the progress of the disease. Swallowing disorder complicates drug intake in pwPD, leads to malnutrition and aspiration pneumonia, and in this way, it reduces quality of life and increases mortality in pwPD. Although the fundamental

Table 2 The univariate and multivariate logistic regression analysis results, determining the risk factors of malnutrition						
	Univariate		Multivariate			
Variables	OR (95% CI)	р	OR (95% CI)	р		
Age	0.99 (0.91-1.08)	0.807	Not selected			
Gender						
Men	1	0.106	Not selected			
vvomen	2.27 (0.84-6.15)	0.000	Niet este d			
Bivil (Kg/m²)	0.940 (0.85-1.04)	0.230	Not selected			
Illiterate	1					
5 years	0.543 (0.16-1.83)	0.324	Not selected			
Over 5years	0.727 (0.19-2.67)	0.630				
Marital status						
Married	1	0.050				
Income	3.37 (1.001-11.345)	0.050				
Low	1		Not selected			
Middle/high	0.61 (0.20-1.86)	0.387				
MUAC (cm)	0.88 (0.77-1.16)	0.080	Not selected			
CC (cm)	0.88 (0.78-0.99)	0.041	Not selected			
TSF (mm)	0.95(0.89-1.02)	0.159	Not selected			
Dyskinesia						
No	1	0.041	Not selected			
	3.34 (1.05-10.64)	0.041				
Farly stages	1					
Advanced stages	2.92 (1.05-8.09)	0.040				
Levodopa dose						
<400 mg	1	0.004				
>400 mg	3.26 (1.17-9.08)	0.024				
DBS	1					
Yes	11.46 (1.32-99.46)	0.027				
Dyspepsia						
Yes	1		Not selected			
No	1.81 (0.60-5.46)	0.296				
Constipation	1		Not selected			
Yes	1.30 (0.47-3.55)	0.615	Not selected			
No						
Yes	1	0.007	1	0.002		
Weight loss	4.59 (1.55-15.76)		7.01 (2.17-20.10)			
No	1		Not selected			
Yes	3.83 (1.34-10.93)	0.012				
Depression						
No	1	0.000				
TES	2.56 (0.94-7.00)	0.066				
Cognitive impairment						
No	1		Not colected			
105	2.15 (0.55-8.49)	0.274				
OR: Odds ratio, CI: Confidence interval, p<0.005, BMI: Body mass index, CC: Calf circumference, DBS: Deep brain stimulation, H&Y: Hoehn and Yahr, MUAC: Mid-upper arm circumference,						

pathophysiology is not fully realized, dopaminergic and nondopaminergic mechanisms have been shown to play a role in the development of dysphagia in PD. Clinical assessment of dysphagia in PD patients is difficult and often yields discordant results (22). However, in this study, it was observed that PD patients who were evaluated only with a single question and described difficulty in swallowing were highly associated with risk of malnutrition.

Study Limitations

There are some limitations in this study. The most important limitation of the study is that swallowing function was not evaluated with an objective method such as functional swallowing tests, an instrumental method such as videofluoroscopic evaluation, or a PD-specific swallowing questionnaire. However, studies in the literature have shown that single screening question for dysphagia as difficulty in swallowing in is closely related to the results of evaluations made by dysphagia diagnostic tools, both in cancer patients and older people living in the community (23,24). In addition, the relationship between a one-question dysphagia screening test and difficulty in swallowing pills was investigated in Parkinson's patients, and the sensitivity of the single question in estimating dysphagia was found to be moderate, while the specificity was found to be high (25). In addition, it is a cross-sectional study with a relatively small sample size. It is important to emphasize that further investigation through large-scaled longitudinal studies is mandatory to detect early malnutrition risk in the geriatric pwPD. Since aging and neurodegenerative diseases such as PD have a strong impact on patients' nutritional status, leading to weight loss, and malnutrition via direct, and indirect mechanisms, it is important to be aware of the risk of malnutrition especially in the geriatric pwPD. To prevent the damaging effects of weight loss on motor function in pwPD, especially in the geriatric population, clinicians should be aware of the malnutrition risks such as advanced stages of the disease, increased doses of daily levodopa, dysphagia, dyskinesia. When swallowing dysfunction is detected, treatment approaches should be applied with pharmacological interventions and therapy by speech and language therapists. Regular screening of malnutrition in pwPD with a validated tool like MNA alongside anthropometric measures such as body weight and BMI in the follow-ups is another clue for early recognition of malnutrition and applying essential interventions for malnutrition to maintain a better quality of life in pwPD.

Conclusion

pwPD are at risk for malnutrition. pwPD should be regularly followed up for malnutrition by the clinicians, particularly those with high risk factors associated with disease characteristics like dysphagia. In addition, the common treatment plan of PD should include a nutritional consultation with a dietary regime.

Information: This study presented in International Congress of Parkinson's Disease and Movement Disorders in 2018, in Hong Kong, as a poster.

Ethics

Ethics Committee Approval: The ethical approval was taken from the Local Ethical Committee of Erciyes University (decision no: 2016/595).

Informed Consent: Informed consent was obtained.

Peer-review: Externally peer-reviewed.

Authorship Contributions

Surgical and Medical Practices: M.G., F.F.Ö., S.A., A.Ö., Y.D., Concept: M.G., F.F.Ö., S.A., A.Ö., Y.D., Design: M.G., F.F.Ö., S.A., A.Ö., Y.D., Data Collection or Processing: M.G., F.F.Ö., S.A., Analysis or Interpretation: M.G., F.F.Ö., S.A., A.Ö., Y.D., Literature Search: M.G., F.F.Ö., Writing: M.G., F.F.Ö.

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