# Experience in the use of a multimodal staging system for Parkinson's disease in clinical practice

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# ABSTRACT

**Objective.** The aim of the study was to evaluate the information content of the multimodal staging system for Parkinson's disease in clinical practice.

**Material and methods.** The study was conducted on the basis of the Regional Clinical Hospital (Odesa). 364 patients with Parkinson's disease were examined. Clinical manifestations were assessed according to the recommendations of Levin O.A. et al. (2019).

**Outcomes.** Average age of the patients was  $64.6\pm0.5$  years, the sample was dominated by men – the gender ratio was 1/1.14. Left-sided lesions were noted in 126 (34.6%) patients, right-sided – in 127 (34.9%), bilateral – in 111 (30.5%).

The akinetic-rigid form was observed in 92 (25.3%) cases, trembling – in 27 (7.4%) cases, mixed rigid-trembling in 157 (43.1%) cases, tremulous-rigid in 92 (7.4%) cases).

Stage 1 according to Hoehn-Yahr was determined in 33 (9.1%), stage 1.5 - in 58 (15.9%), stage 2 - in 104 (28.6%), stage 2.5 - in 36 (9.9%), stage 3 - in 102 (28.0%), stage 3.5 - in 6 (1.6%), stage 4 - in 25 (6.9%) cases. No cases of stage 5 were reported.

65.4% of patients had cognitive impairments. 121 (33.2%) patients had pain syndrome of varying severity. Vegetative manifestations of varying severity were observed in 201 (55.2%) patients, affective manifestations – in 170 (46.7%) patients.

**Conclusion.** The MOSCOVA scale is advisable to detail the criteria for staging within motor and non-motor manifestations of PD.

Keywords: Pakinson disease, diagnosis, non-motor manifestations

## INTRODUCTION

Parkinson's disease (PD) is a progressive neurodegenerative disease accompanied by motor disorders (tremor, hypokinesia, rigidity), as well as a number of non-motor disorders (cognitive, affective, vegetative etc.) [1,2]. Such non-motor disorders how depression and dementia affect the quality of life of patients and their relatives and make the greatest contribution to the development of social maladaptation [1,3]. In this regard, the search for effective ways to display the clinical manifestations of PD does not stop.

The modified Hoehn and Yahr scale is the most commonly used classification of PD stages [4]. It re-

lies mainly on the prevalence of movement disorders, their lateralization, as well as the severity of postural instability and other disorders that limit the patient's mobility. Until recently, there was no effective system for presenting a wide range of both motor and non-motor disorders in PD [5]. Recently, Russian specialists have proposed a fundamentally new concept of staging PD, based on the consideration of both motor and non-motor (sensory, autonomic, psychiatric etc.) manifestations of PD, as well as motor or non-motor fluctuations and dyskinesias that occur during long-term therapy with levodopa [6].

The authors propose to divide the clinical symptoms in PD into six categories:

- Motoric
- Obstacles or Complications of treatment (motor fluctuations and dyskinesias associated with drug therapy)
- Sensory (including pain, chronic fatigue, disturbed sleep and wakefulness)
- Cognitive
- Vegetative
- Affective

The latter category includes, along with affective disorders proper, other neuropsychiatric disorders, primarily psychotic ones.

To facilitate the mnemonization of this approach, the authors proposed to combine the first letters of the names of the above categories into the abbreviation MOCKBA (Moscow in Russian) [6,7]. Taking into account the fact that the Cyrillic alphabet is used as the official alphabet only in 15 countries of the world, we propose an abbreviation for the Latin alphabet - MOSCOVA (Motoric, Obstacles, Sensory, Cognitive, Vegetative, Affective)<sup>1</sup>. Another feature of this scale is the ability to quantify each domain using a rank scale from 0 to 5 for motor impairments and from 0 to 4 for non-motor disorders. At the beginning of 2022, there was no information on the validation of the scale in large-scale clinical trials, but the proposed approach, in our opinion, deserves attention.

The aim of the study was to evaluate the information content of the multimodal staging system for Parkinson's disease in clinical practice.

### MATERIAL AND METHODS

The study was conducted on the basis of the Regional Clinical Hospital (Odessa). 364 patients with Parkinson's disease included in the regional register of extrapyramidal pathology were examined. The scope of the examination complied with the requirements of the Guidelines for the Diagnosis and Treatment of Parkinson's Disease, approved by the Scientific Council of the State Institution "Institute of Gerontology" and MDS recommendations [8].

# There were used Beck questionnaire, The King's Parkinson's Disease Pain

Questionnaire (KPPQ), MMSE, UPDRS. The symptoms of the disease were assessed taking into account its clinical picture for the previous month.

Motor disorders were assessed in accordance with the Hoehn-Yahr scale, non-motor disorders according to the recommendations of Levin O.A. et al. (2019) [7] (Table 1). The staging of development of sensory disorders (including pain syndromes, akathisia, anosmia, visual impairment) was assessed together with chronic fatigue and sleep and wakefulness disorders.

Additionally, the psychometric parameters (reliability, validity, sensitivity) of the proposed scale were evaluated [9]. Internal consistency and reproducibility were evaluated as reliability parameters. Internal consistency was studied using an analogue of the Cronbach's alpha coefficient for a dichotomous scale, the Kuder–Richardson test. The reproducibility of the scale, i.e., its resistance to measurement errors over time, was determined by the test-retest method. Statistical analysis of the obtained results was performed using the TIBCO Statistica 13.5 software (USA) [10].

### OUTCOMES

The average age of the patients was  $64.6\pm0.5$  years, the sample was dominated by men - the gender ratio was 1/1.14. At the onset of the disease, left-sided lesions were noted in 126 out of 364 patients, that is, 34.6%, right-sided - in 127 (34.9%), in the remaining 111 (30.5%) - bilateral lesions.

Mixed forms prevailed in the structure of the disease. The akinetic-rigid form was observed in 92 (25.3%) cases, trembling - in 27 (7.4%) cases, mixed rigid-trembling in 157 (43.1%) cases, tremulous-rigid in 92 (7.4%) cases).

The examined patients were distributed according to severity as follows: stage 1 according to Hoehn-Yahr was determined in 33 (9.1%), stage 1.5 - in 58 (15.9%), stage 2 - in 104 (28.6%), stage 2.5 - in 36 (9.9%), stage 3 - in 102 (28.0%), stage 3.5 - in 6 (1.6%), stage 4 - in 25 (6.9%) %). No cases of stage 5 were reported.

Fluctuations associated with taking levodopa were noted only in 87 (23.9%), in most patients they were mild and did not significantly limit activity.

A significant number of patients (238 or 65.4%) had cognitive impairments, the average score on the MMSE scale was  $25.3 \pm 0.3$ . A more detailed analysis of the distribution of patients according to the age of onset of the disease indicates that pre-dementia changes were in 82 (15.6%) patients. Accordingly, mild dementia was diagnosed in 15 (2.8%) patients, moderate dementia - in 18 (3.4%).

When analyzing the prevalence of pain syndrome in patients with CP, who were included in the regional registry, it was found that 121 (33.2%) patients had pain syndrome of varying severity. The most common were night pains (21.5%) and muscular-skeletal pains, both isolated (23.1%) and in various combinations. In general, pain with fluctuations of various localization occurred in 27 patients (5.1%) of the total number of persons in the registry (n =

 $<sup>^1\,\</sup>rm MOSCOVA$  – name of Moscow city in Romanian and in Interlingua, in Milano (Italy) – the name of historical district and metro station

Domain	Scores	Description							
Motoric	MO	No motor disorders (H-Y 0)							
meterie	M1	Unilateral involvement only usually with minimal or no functional disability (H-Y 1)							
	M2	Bilateral or midline involvement without impairment of balance (H-Y 2)							
	M3	Bilateral disease: mild to moderate disability with impaired postural reflexes: physically independent (H-Y 3)							
	M4	Severely disabling disease: still able to walk or stand unassisted (H-Y 4)							
	M5	Confinement to bed or wheelchair unless aided (H-Y 5)							
Obstacles	00	No complications							
	01	Mild fluctuations or dyskinesias limit certain activities							
	02	Moderate fluctuations or dyskinesias limit many activities							
	03	Manifested functional disorders caused by fluctuations or dyskinesias are so pronounced that the patient usually							
		does not carry out many activities or his interaction with others is significantly limited							
	04	Severe functional disorders caused by fluctuations or dyskinesias are so pronounced that the patient usually does							
		not carry out most activities or his interaction with others is severely limited							
Sensory	S0	No sensory disorders							
	S1	Occasional mild disorders limit certain types of activity							
	S2	Disorders are frequent, moderate, limit many types of activity							
	S3	Permanent, pronounced disorders limit the patient's activity to such an extent that they usually do not allow him							
		to carry out certain activities or limit his interaction with others							
	S4	Disorders (including fatigue, sleep and wakefulness), persistent, severe, and usually exclude the possibil							
	-	most activities and severely limit interaction with others							
Cognitive	CO0	No cognitive disorders							
0	CO1	Mild disorders limit certain types of activity							
	CO2	Disorders are frequent, moderate, limit many types of activity (moderate cognitive impairment)							
	CO3	Manifested disorders that exclude the possibility of certain activities and limit interaction with others (mild							
		dementia)							
	CO4	Severe cognitive impairment, usually excluding the possibility of most activities and significantly limiting the							
		patient's interaction with others (dementia)							
Vegetative	V0	No disorders							
	V1	Occasional mild disorders limit certain types of activity							
	V2	Disorders are frequent, moderate, limit many types of activity							
	V3	Constant manifested vegetative disturbances precluding the implementation of certain activities and restricting							
		interaction with others							
	V4	Permanent severe vegetative disturbances, usually hindering the implementation of most activities and							
		significantly limiting interaction with others							
Affective	A0	No disorders							
	A1	Mild anxiety or anxiety-depressive disorders, periodic anhedonia, limiting the implementation of certain activities							
	A2	Persistent moderately expressed anxiety or anxiety-depressive disorders, extracampine phenomena that limit							
		the implementation of many activities							
	A3	Expressed anxiety, depression, apathy, episodic hallucinations, paranoid ideas that are persistent in nature, which							
	_	usually exclude the possibility of certain activities and limit interaction with others							
	A4	severe persistent affective and neuropsychiatric disorders (depression or apathy, hallucinosis, paranoid syndrome.							
		delirium), which usually exclude the possibility of most activities and severely limit interaction with others							
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## TABLE 1. Evaluation of the clinical manifestations of PD by the MOSCOVA scale

527). Nocturnal pain occurred in 94 (17.8%) cases, orofacial pain in 29 (5.5%) cases. Pain associated with edema and signs of inflammation were in 58 (11.0%), radicular pain - in 37 (7.0%).

Some patients had other sensory disturbances (akathisia, anosmia, etc.), chronic fatigue, sleep and wakefulness disorders.

Vegetative manifestations of varying severity were observed in 201 (55.2%) patients, affective manifestations - in 170 (46.7%) patients (Table 2). When calculating the criterion values, it was found that the global Cronbach's alpha for the proposed method for assessing the severity of the disease was 0.86, while for assessing motor function (M) the values of Cronbach's alpha were 0.77, for complications (O) - 0.68, for sensory disorders - 0.65, for cognitive impairments - 0.69, for autonomic disorders - 0.65, for affective manifestations - 0.66. The reliability coefficient of Krueder-Richardson was rKR=0.78 for the studied data array.

Subscale	0		1		2		3		4		5	
	abs.	%	abs.	%	abs.	%	abs.	%	abs.	%	abs.	%
М	-	-	33	9,1	162	44,5	138	37,9	31	8,5	-	-
0	277	76,1	56	15,4	28	7,7	3	0,8	-	-		
S	243	66,8	55	15,1	22	6,0	21	5,8	3	0,8		
со	126	34,6	134	36,8	93	25,5	11	3,0	-	-		
V	163	44,8	109	29,9	69	18,9	22	6,0	-	-		
A	194	53,3	148	40,7	13	3,6	9	2,5	-	-		

TABLE 2. Distribution of patients according to the criteria of the multimodal classification system (MOSCOVA)

### CONCLUSIONS

The described approach to the complex determination of the stage of PD with the assessment of certain categories of symptoms (motor, cognitive, affective, sensory etc.) was validated on a sample of 364 patients with Parkinson's disease. The results of the

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#### REFERENCES

- Cabreira V, Massano J. Doença de Parkinson: Revisão Clínica e Atualização [Parkinson's Disease: Clinical Review and Update]. Acta Med Port. 2019 Oct 1;32(10):661-70
- Armstrong MJ, Okun MS. Diagnosis and Treatment of Parkinson Disease: A Review. JAMA. 2020 Feb 11;323(6):548-60.
- Sauerbier A, Jenner P, Todorova A, Chaudhuri KR. Non motor subtypes and Parkinson's disease. *Parkinsonism Relat Disord*. 2016 Jan;22(1):S41-6.
- Opara J, Małecki A, Małecka E, Socha T. Motor assessment in Parkinson's disease. Ann Agric Environ Med. 2017 Sep 21;24(3):411-415.
- Krüger R, Klucken J, Weiss D, Tönges L, Kolber P, Unterecker S et al. Classification of advanced stages of Parkinson's disease: translation into stratified treatments. *J Neural Transm* (Vienna). 2017 Aug;124(8):1015-27.

validation indicate that this scale can be used in clinical practice to assess the dynamics of the state of patients with PD and plan their treatment. It is advisable to detail the criteria for staging within each category of PD symptoms.

- [Parkinson's disease and movement disorders] / ed. S.N. Illarioshkina, O.S. Levina M, 2017 381 p. [Rus]
- Levin O, Vasenina E, Chimagomedova A, Skripkina N. [MOCKBA: new staging system for Parkinson's disease] [Rus]. Retrieved from https:// www.xn--80aocaipeaifmp.xn--p1ai/moskva-novaja-sistemastadirovanija-bolezni-parkinsona/. Accesed 12.08.2022
- MDS guidelines. Retrieved from: https://www.movementdisorders. org/MDS-Files1/Resources/PDFs/ TreatmentsforMotorSymptomsofPD-2018.pdf on 12.08.2022
- Godinho C, Domingos J, Cunha G, Santos AT, Fernandes RM, Abreu D et al. A systematic review of the characteristics and validity of monitoring technologies to assess Parkinson's disease. J Neuroeng Rehabil. 2016 Mar 12:13:24.
- Manual of Statistica software. Retrieved from: https:// statisticasoftware.wordpress.com/tag/manuals/ on 12.08.2022