Original Article

A Study on Non-motor Manifestations in Young Onset Parkinsons Disease (YOPD) in Eastern Indian Population

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Background : Non-motor Symptoms (NMS) are extremely common in Young Onset Parkinson Disease (YOPD) with high incidence of neurobehavioral and autonomic dysfunction which should be given greater emphasis as it will affect the Quality of Life.

Materials and Methods : This is a cross-sectional study in which all consecutive patients of age less than 50 years satisfying the UPKRDS Diagnostic criteria for Idiopathic Parkinson's Disease (IPD) attending the Neurology Outpatient services at a super speciality clinic during study period July 1, 2021 to April 30, 2022 were included in the sample. All secondary causes of parkinsonism like drug induced, multi-infarct state and normal pressure hydrocephalus were excluded. All patients were examined, and demographic data and non-motor symptoms were documented using the Non-motor Scale (NMSS) of International Parkinson's and movement disorder society and modified Hoehn & Yahr staging used for staging.

Results : A total 32 patients were diagnosed with YOPD during the study period. Out of which 19 (59.37%) were Males and 13 (40.32 %) were Females. Majority of the patients were in the age group of 40-50 years (84%) and onset of illness in 1 month to 5 years (78.17%) with 37% each of patients were in the H & Y Stages I-1.5 and 1.5-2. Common Non-motor Symptoms (NMS) observed were anxiety (71.87%), memory loss (59.37%) and depression (56.25%). Most of the patients had one or more autonomic symptoms and sleep disturbances.

Conclusion : Non-motor Symptoms were present in all patients (100%) with anxiety, memory loss, depression and constipation are being the commonest of the NMS.

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Key words : Non-motor Manifesations, Young Onset Parkinsons Disease (YOPD).

Parkinson's Disease (PD) is a neurodegenerative disorder characterized by bradykinesia, rest tremor, rigidity and postural instability along with variety of Non-motor Symptoms (NMS)¹. It is usually considered as age related disease with mean age in early to mid 60s, but it can occur in early life also². Based on age of onset, PD can be divided into early and late [Late Onset PD (LOPD)]. Early Onset PD (EOPD) further subdivided into Juvenile Onset PD (JOPD) (before 21 years) and Young Onset PD (YOPD) (between 21-40 years of age) while onset after 60 years defines LOPD³⁻⁵. Due to lack of consensus, the maximal age for YOPD has varied from 40 to 55^{3,6,7} and minimal age for LOPD has varied from 50 to 70^{3,8,9}.

The prevalence of PD in the Western World has been reported to range from 130 to 200 per 100,000 in community-based studies but reported as high as Editor's Comment :

Non-motor manifestations are very commonly seen in YOPD. Depression, anxiety and constipation are the commonest to be seen in this study.

2000/100,000 in individuals over 80 years of age¹⁰⁻¹³ and YOPD represents 5-7% of this. As per WHO (14 June, 2022) prevalence of Parkinson disease has doubled in the past 25 years with global estimates in 2019 showing over 8.5 million individuals living with PD, 5.8 million disability-adjusted life years and 329,000 deaths.

YOPD and LOPD are similar in both clinical and pathological features, except for higher rate of treatment-related dyskinesias¹⁴, slow onset of progression and impact on Quality of Life (QOL) especially due to NMS¹⁴⁻¹⁸. YOPD patients facing occupational and life style challenges than LOPD not only due to motor symptoms but also due to NMS like depression, sexual dysfunction, marital conflict, loss of occupation and future uncertainty¹⁹⁻²¹. It also leads to behavioural disorder like addiction, impulse control disorders like gambling, compulsive shopping and sexual addiction.

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MATERIALS AND METHODS

This is a cross sectional study in which all Consecutive patients less than fifty years satisfying the UPKRDS Diagnostic criteria for Idiopathic Parkinson's Disease (IPD) attending the Neurology Outpatient services at a Super Speciality Neurology Clinic, Durgapur during study period July 1, 2021 to April 30, 2022 were included in the sample. All patients were examined and demographic data and Non-motor Symptoms were documented using the Non-motor Scale (NMSS) of International Parkinson's and movement disorder society and modified Hoehn & Yahr staging used for staging.Local Ethical committee clearance was taken by Asansol Durgapur Ethics Committee before starting the study, Letter no -(15/ 21). Exclusion criteria- All secondary causes of parkinsonism like drug related, multi-infarct state and normal pressure hydrocephalus were excluded.

RESULTS

A total 32 patients were diagnosed with YOPD and satisfied the inclusion criteria of study. Out of which 19 (59.37%) were Males and 13 (40.32%) were Females. In 27 patients were within 40-50 years (84%) and remaining 5 patients were within 30-39 years (16%). The Hoehn & Yahr (H& Y) stages were 1-1.5 (37%), 2(37%) and least common H & Y stage was 3-5 (25%) (Table 1). Onset of illness was most commonly between 1 month - 5 years (78.17%) followed by 6 years -10 years (21.87%) and 11 years - 15 years (6.25%) in descending order (Fig 1).

Non-motor Symptoms (Table 2) seen in decending order were Memory loss (59.37%), constipation (46.87%), Sexual dysfunction (40.62%), Anxiety



Fig 1 — Onset of illness

Table 2 — NMS in descending order of prevalence in studypopulation	
Non-motor Symptoms Percentage of	Study Population (%)
Non-motor Symptoms Percentage of 3 ANXIETY MEMORY LOSS DEPRESSION CONSTIPATION NOCTURIA INSOMNIA SEXUAL DISORDER DAY TIME SLEEPINESS EXCESSIVE SWEATING FROZEN SHOLDER URINARY FREQUENCY LOSS OF TASTE & SMELL PANIC ATTACK URINARY URGENCY RESTLESS LEG SYNDROME FAINTING DIZZNESS DIZZNESS	Study Population (%) 71.87 59.37 56.25 46.87 43.75 43.75 40.62 31.25 31.25 31.25 31.25 31.25 28.12 28.12 28.12 28.12 18.75 15.62 12.05 12.05 9.37
REM SLEEP BEHAVIOURAL DISORDER HYPER SALIVATION	9.37 3.12

(71.87%), Depression (56.25%), Nocturia (43.75%), Urinary Frequency (28.12%), Insomnia (43.75%), Urinary Urgency (15.62%), Day Time Sleepiness (13.25%), Loss of Taste and Smell (28.12%), Excess Sweating (31.25%), Frozen Shoulder (31.25%), Restless Leg Syndrome (12.05%), Dizziness (9.37%), Panic Attack (18.75%), Fainting (15.05%), RBD (9.37%) and Hypersalivation (3.12%).

DISCUSSION

YOPD is almost similar to LOPD in clinical picture except in slow disease progression²², less falls, freezing³ and increased treatment related motor complications¹⁴. In this study, it is only focussed on clinical profile of Non-motor Symptoms of YOPD regarding their relation to stage and duration of disease. Out of 32 patients included in this study majority are coming under the age group of 40-50 years (84%) followed by 30 - 39 years. Onset of illness in majority of patients are 1 month to 5 years (78.17%) followed by 6 years -10 years (21.87 %) and 11 years-15 years (6.25 %) with equal distribution in both 1-1.5 and 2 H&Y staging (37% each) and remaining 26% patients are in 3 -5 stage. NMS included in this study consist of autonomic dysfunction, sleep disorders, psychological and behavioural disorders, sensory and cognitive symptoms and all patients were shown one or more NMS.

The most common NMS is Anxiety (71.87%) followed by Memory loss (59.37%), Depression (56.25%) and Constipation (46.87%) compared to a study conducted in Indian populations which shows Depression (45.6%), Anxiety (45.4%) and Apathy

(30.5%)²³. A similar study in USA shows similar rate of Depression (48.3%)²⁴. YOPD usually defines between 21-40 years but there is lack of consensus regarding the upper limit which varies from 40 to 55 years in different studies. We considered the upper limit of age as 50 years. Cognitive decline is less in YOPD compared to LOPD^{25,26}, even though it is more dependent on age. In this study Memory loss was the 3rd most common NMS which is probably due to the increased age of patient (84% is between 40-50 years). Genetic study was not included in this study even though genetic predisposition is well recognised with age of onset-younger the age higher the genetic association²⁷. Family history is reported in 20% of YOPD patients compared to 6.9% of LOPD patients, and the age-specific risk of PD is 7.8-fold higher in the relatives of patients with YOPD compared to 2.9-fold among the relatives of patients with LOPD^{7,28}. Many genes are considered causing PD which mainly include SNCA, LRRK2, GBA. Duplication of PARK 1 gene is associated with NMS like severe psychiatric features. In 1/3rd of the study population frozen shoulder was seen and panic attack was documented in 18.75% of the study population. In several studies restless leg syndrome had a higher rate^{5,9,18,29-33} compared to 12.05% in our study. Other common manifestations were constipation, urinary urgency, frequency, sexual dysfunction, hyper salivation, excessive sweating, fainting and loss of smell and taste. Sleep disturbances like REM Behavioural Disorders, excessive day time sleeping and insomnia were seen commonly in the study population. NMS in YOPD has to be diagnosed early and treated because it affects the peak and productive years of life which leads to loss of employment, social and family conflicts, loss of selfesteems and various addictive behavioural problems. There is a diagnostic challenge in YOPD mainly due to rarity and age of onset because PD is mainly considered as a disease of elderly and this leads to multiple neurologist visit, multiple investigations and delay in diagnosis and treatment^{19,6}.

Limitations: Small sample size, cross sectional study, genetic study is not done in the patients

CONCLUSION

Non-motor Symptoms were seen in all the patients with Young Onset Parkinsons Disease (YOPD) in our study. Anxiety, Memory Loss, Depression and Constipation are the commonest non motor manifestations.

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Conflict of interest : There is no conflict of interest.

REFERENCES

- 1 Obeso JA Past, present, and future of Parkinson's disease: A special essay on the 200th Anniversary of the Shaking Palsy. *Movement Disorders* 2017; **32(9):** 1264-310.
- 2 Samii A, Nutt JG, Ransom BR Parkinson's disease. Lancet 2004; 363: 1783-93.
- 3 Schrag A, Schott JM Epidemiological, clinical, and genetic characteristics of early-onset parkinsonism. *Lancet Neurol* 2006; 5: 355-63.
- 4 Morales-Briceno H, Mohammad SS, Post B, Fois AF, Dale RC, Tchan M, Fung VSC — Clinical and neuroimag- ing phenotypes of genetic parkinsonism from infancy to adolescence. *Brain* 2020; **143**: 751-70.
- 5 Thenganatt MA, Jankovic J Parkinson disease subtypes, *JAMA Neurol* 2014; **71:** 499-504.
- 6 Rana AQ, Siddiqui I, Yousuf M Challenges in diagnosis of young onset Parkinson's disease, *J Neurol Sci* 2012; **323**: 113-6.
- 7 Mehanna R, Moore S, Hou JG, Sarwar AI, Lai EC Comparing clinical features of young onset, middle onset and late onset Parkinson's disease. *Park Relat Disord* 2014; 20: 530-4.
- 8 Wickremaratchi MM, Perera D, O'Loghlen C Prevalence and age of onset of Parkinson's disease in Cardiff: a community based cross sectional study and meta-analysis. *J Neurol Neurosurg Psychiatry* 2009; **80:** 805-7.
- 9 Wickremaratchi MM, Ben-Shlomo Y, Morris HR The effect of onset age on the clinical features of Parkinson's disease. *Eur J Neurol* 2009; **164:** 450-6.
- 10 Schrag A, Ben-Shlomo Y, Quinn N Cross sectional prevalence survey of idio- pathic Parkinson's disease and parkinsonism in London. *BMJ* 2000; **321:** 21-2.
- 11 Dorsey ER, Bloem BR The Parkinson pandemic-A call to action. JAMA Neurol 2018; 75: 9-10.
- 12 Pringsheim T, Jette N, Frolkis A, Steeves TD The prevalence of Parkinson's (disease: a systematic review and metaanalysis. *Mov Disord* 2014; 29: 1583-90.
- 13 Wanneveich M, Moisan F, Jacqmin-Gadda H, Elbaz A, Joly P — Projections of (prevalence, lifetime risk, and life expectancy of Parkinson's disease (2010-2030) in France. *Mov Disord* 2018; **33**: 1449-55.
- 14 Schrag A, Hovris A, Morley D, Quinn N, Jahanshahi M Young- versus older-onset Parkinson's disease: impact of disease and psychosocial consequences. *Mov Disord* 2003; 18: 1250-6.
- 15 Kupsch A, Oertel WB Selegiline, pregnancy, and Parkinson's disease. *Mov Disord* 1998; **13:** 175-6.
- 16 Schrag A, Banks P Time of loss of employment in Parkinson's disease. *Mov Disord* 2006; 21: 183943.
- 17 Murphy R, Tubridy N, Kevelighan H, O'Riordan S Parkinson's disease, How is employment affected? *Ir J Med Sc* 2013; 182: 4159.
- 18 Knipe MD, Wickremaratchi MM, Wyatt-Haines E, Morris HR, Ben-Shlomo Y — Quality of life in young- compared with late-onset Parkinson's disease. *Mov Disord* 2011; 26: 2011-8.

- 19 Schrag D, Morley N, Quinn M Jahanshahi, Impact of Parkinson's disease on patients' adolescent and adult children, *Park Relat Disord* 2004; **10**: 391-7.
- 20 Carter, Julie H Living with a person who has Parkinson's disease: the spouse's perspective by stage of disease." Movement disorders: official journal of the Movement Disorder Society 1988; 13(1): 20-8.
- 21 Ravenek M, Rudman DL, Jenkins ME, Spaulding S Understanding uncertainty in young-onset Parkinson disease. *Chronic Illness* 2017; **13:** 288-98.
- 22 Alves G, Wentzel-Larsen T, Aarsland D, Larsen JP Progression of motor impair- ment and disability in Parkinson disease: a population-based study. *Neurology* 2005; 65: 1436-41.
- 23 Kukkle P, Goyal V, Geetha T, Mridula K, Kumar H, Borgohain R, et al — Clinical Study of 668 Indian Subjects with Juvenile, Young, and Early Onset Parkinson's Disease. Canadian Journal of Neurological Sciences / Journal Canadien Des Sciences Neurologiques 2022; 49(1): 93-101. doi:10.1017/ cjn.2021.40
- 24 Mehanna R, Moore S, Hou JG, Sarwar AI, Lai EC Comparing clinical features of young onset, middle onset and late onset Parkinson's disease. *Parkinsonism Relat Disord* 2014; 20: 530-4.
- 25 Schrag A, Ben-ShlomoY, Brown R, Marsden CD, Quinn N Young-onsetParkinson's disease revisited—clinical features, natural history, and mortality. *Mov Disord* 1998; **13**: 885-94.

- 26 Chaudhary S, Joshi D, Pathak A, Mishra VN, Chaurasia RN, Gupta G — Comparison of cognitive profile in young- and late-onset Parkinson's disease pa- tients *Ann Indian Acad Neurol* 2018; 21: 130-2.
- 27 Deng H, Wang P, Jankovic J The genetics of Parkinson disease. Ageing Res Rev 2018; 42: 72-85.
- 28 Payami H, Zareparsi S, James D, Nutt J Familial aggregation of Parkinson disease: a comparative study of early-onset and late-onset disease. Arch Neurol 2002; 59: 848-50.
- 29 Calne SM, Lidstone SC, Kumar A Psychosocial issues in young-onset Parkinson's disease: current research and challenges. *Park Relat Disord* 2008; **142:** 143-50.
- 30 Spica V, Pekmezovic T, Svetel M, Kostic VS Prevalence of non-motor symptoms in young-onset versus late-onset Parkinson's disease. J Neurol 2013; 260: 131-7.
- 31 Cilia ER Cereda, Klersy C, Canesi M, Zecchinelli AL, Mariani CB, Tesei S, et al — Parkinson's disease beyond 20 years, J Neurol Neurosurg Psychiatry 2015; 86: 849-55.
- 32 Kostic KS, Filipovic SR, Lecic D, Momcilovi D, Sokic D, Sternic N Effect of age at onset on frequency of depression in Parkinson's disease. *J Neurol Neurosurg Psychiatry* 1994; 57: 1265-7.
- 33 Dissanayaka NN, Sellbach A, Matheson S, O'Sullivan JD, Silburn PA, Byrne GJ, et al — Anxiety disorders in Parkinson's disease: pre- valence and risk factors. *Mov Disord* 2010; 25: 838-45.

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