Case Report

Predicting Prognosis of Psychosis in Huntington’s Disease: Case Report and Review of Literature

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Huntington’s disease (HD) is a rare autosomal dominant disorder that results from progressive neurodegeneration at basal ganglia due to the formation of abundant abnormal protein “Huntingtin.” Expansion of the trinucleotide “CAG” repeat sequence (>35 repeats) on the short arm of chromosome 4, gives rise to the formation of protein “Huntingtin.” The onset of HD usually occurs in the fourth to fifth decade of life and choreiform movement is the prominent manifestation, however as the disease advances cognitive impairment and psychiatric manifestations become evident. Nearly, one-third to three-fourth of patients with HD has psychiatric manifestations. Mood disorders, aggression, anxiety are the most common psychiatric manifestations and psychosis being relatively rare. Psychiatric disorders often occur before the onset of motor symptoms of HD. Psychotic symptoms often appear in the course of HD, after the motor symptoms develop and hence, in the early days, it was thought to be the forerunner of the HD. Schizophrenia-like psychotic symptoms were also known to precede the onset of abnormal body movements of HD. Psychotic symptoms may mask the motor symptoms or may delay the diagnosis of HD.

Case Report

A 50-year-old, nondiabetic and nonhypertensive male was hospitalized in our psychiatric unit for his aggressive behavior, refusal for food and disturbed sleep. He also had delusions of control (people are controlling him by putting a computer chip in his brain) and persecution (neighbors were plotting plans to kill him by poisoning his food). Due to this, he was not going out of his house, keeping himself locked inside. These symptoms had started since 5 years but worsened over past 1 year. In the past 5 years, once he had consulted a psychiatrist for illness and received treatment (risperidone 6 mg/day and lorazepam 2 mg/day) for 2 months with little improvement in symptoms. Nonadherence to treatment subsequently further worsened his symptoms. Family members of the patient reported about abnormal body movements involving his extremities, trunk, and neck as well as his orofacial musculature since past 8 years (i.e., 3 years before the onset of psychiatric symptoms). These movements were involuntary, bizzare and were of facial grimacing, hand twisting, tongue darting, and pelvic thrusting type in nature which was causing significant impairment in talking, eating, drinking, holding objects in hand as well as walking. His history was insignificant. Family history of the patient revealed similar movement disorder in his elder brother in his fourth decade of life, who subsequently developed psychotic features and absconded from home and was missing till date. Patient’s father had died early at the age of 35 years.
in an accident, and by that time he did not have any movement disorder or any psychiatric illness. There was no history of substance abuse. No cognitive decline was reported and observed on evaluation.

General physical examination was normal. Neurological examination revealed choreiform movements involving his extremities as well as trunk. Other systemic examinations were within normal limits. His hematological investigations were within normal limits. Neuroimaging of the brain (computed tomography scan and magnetic resonance imaging) revealed diffuse cortical atrophy [Figure 1]. Genetic evaluation revealed 43 repeats of CAG in one allele of Huntingtin gene and 17 CAG repeats in the other allele. He was diagnosed with HD with organic delusional (schizophrenia-like) disorder (F06.2 as per International Classification of Diseases-10 diagnostic criteria). He was prescribed olanzapine (up to 20 mg/day) for schizophrenia. After opinion of neurologist, sodium valproate (600 mg/day) was started for the management of abnormal movements. Over a period of 4 weeks, the severity of psychotic symptoms and abnormal movements were reduced.

**DISCUSSION**

Psychiatric manifestations are common in HD and can occur anytime in the course of illness (before as well as after the onset of motor symptoms). The psychiatric symptoms evolve in the course of HD. Evidences suggest worsening of irritability, apathy, cognitive impairment as well as motor symptoms worsen with disease progression.

Progressive neurodegeneration occurs in HD as well as in schizophrenia causing loss of brain volume. The presence of both the entities together is likely to hasten this process further. Hence, coexistence of both the entities may be considered as a poor prognostic factor for schizophrenia. In HD, there occurs loss of volume of gray matter, white matter, and expansion of the cerebrospinal fluid (CSF) spaces. The presence of neurological signs and symptoms are also considered as poor prognostic indicator of schizophrenia. Our patient had neurological signs and symptoms (movement disorder, ataxia, speech abnormality) along with radiological structural brain changes. The degree of mutation of CAG triplet has a correlation with the expansion of the CSF spaces, hence can be considered as a prognostic indicator of schizophrenia also. However, another gene KCNN3 located in chromosome 1q21.3 also produces polymorphic CAG repeat sequences, is associated with better cognitive performance in schizophrenia. Similarly, another earlier research concludes that degree of expansion of CAG repeats encoded by KCNN3 gene had a positive correlation with various symptoms of schizophrenia such as lack of energy, paranoia, and negative symptoms.

Our patient had HD for past 8 years and psychosis for 5 years. Here, psychosis followed the motor symptoms of HD. The patient had significant cerebral atrophy, leading to dilatation of the ventricular system and cortical thinning [Figure 1]. In our patient, it can be a poor prognostic factor. CAG repeat sequence in one allele of Huntingtin gene was 43, which can be again a poor prognostic factor and have a positive correlation with the CSF space expansion (ventricular dilatation). The progressive neurodegeneration due to HD as well as schizophrenia can be double jeopardy in this case. Hence, the presence of HD may be considered as a poor prognostic indicator of psychosis.

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**Conflicts of interest**

There are no conflicts of interest.

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