## Paliperidone-Induced Dystonic Dysphagia

To the Editor: Acute dystonic reactions are well recognized, but isolated dystonia of the pharyngeal muscle causing dysphagia is a rare occurrence with neuroleptic drugs. Extended-release paliperidone is a newly introduced atypical antipsychotic medication formulated using the principal active metabolite of risperidone, 9-hydroxyrisperidone. In contrast to risperidone, it provides consistent and smooth plasma drug concentrations over 24 hours.<sup>1</sup> We report a young man who developed acute-onset dysphagia after taking paliperidone.

## Case Report

Mr. S, an 18-year-old male student, presented with a history of being suspicious, irritable, withdrawn, muttering to self, and having grandiose ideas and disturbed biological functions for a duration of 1 year. He was diagnosed with schizophrenia (DSM-IV-TR criteria). He was treated with divalproex sodium, 750 mg/day, olanzapine, 30 mg/day, and clozapine, 100 mg/day, with marked recovery in psychotic symptoms. He remained stabilized for 3 months while continuing the same treatment. Subsequently, he stopped taking medicine and had a relapse within 2 weeks. This time he was treated with divalproex sodium, 500 mg/day, and paliperidone, 3 mg/day. At day 3, the dosage of paliperidone was increased to 6 mg/day. After taking only two doses of paliperidone at 6 mg/day and within 12 hours of his last dose of paliperidone, he started complaining of a choking sensation in his

throat if he tried to ingest any solid or semisolid food and made gurgling sounds. He had little difficulty taking liquids, but solid food would stick in his throat. He presented with severe anxiety and was calling for help. By the time he was brought to the treating psychiatrist, within 4 hours, he also had difficulty making sounds of any kind.

Neurological examination did not show any evidence of organic illness, and no abnormal movements were noticed on other parts of the body. He was given intramuscular promethazine, 50 mg, and within 30 minutes he showed remarkable recovery in swallowing. Otolaryngological and gastroenterological evaluation did not find any organic causes and confirmed a case of drug-induced dysphagia. His organic workup for dystonia, including EEG and brain MRI, was negative. Paliperidone was replaced by clozapine, 200 mg/day, with no further recurrence of dysphagia.

## Discussion

The association between paliperidone and dystonic dysphagia is irrefutable in this report; the complaints developed shortly after the introduction of paliperidone and resolved with an anticholinergic drug. Dysphagia may be a manifestation of extrapyramidal side effect when it is associated with neuroleptic therapy.<sup>2</sup> Literature shows that patients with dystonic dysphagia frequently have oromandibular dystonia or neck dystonia,3 but in our case it was an isolated phenomenon. The mechanism of action of paliperidone is unclear. Paliperidone may cause dystonia due to its dopamine D<sub>2</sub>-serotonin 5HT2A antagonist property. Further, it can block alpha 2 adrenoceptors which can increase noradrenergic activity by inhibiting the autoreceptors.<sup>4</sup> Animal study has suggested that enhanced activity of central noradrenergic nuclei is one of the pathophysiologies of dystonia.<sup>5</sup>

Complaints of dysphagia in patients with schizophrenia are often overlooked, and this report shows that early diagnosis of neuroleptic-induced dysphagia can avoid potential complications and prevent the patient from undergoing extensive diagnostic testing and various drug trials.

DATTATREYA NAMDEORAO
MENDHEKAR, D.P.M.
Neuropsychiatry and Headache Clinic, New Delhi, India
ASHISH AGARWAL, M.D.
Department of Psychiatry, GB
Pant Hospital, New Delhi

## References

- 1. Spina E, Cavallaro R: The pharmacology and safety of paliperidone extended-release in the treatment of schizophrenia. Expert Opin Drug Saf 2007; 6:651–662
- Nair S, Saeed O, Shahab H, et al: Sudden dysphagia with uvular enlargement following the initiation of risperidone which responded to benztropine: was this an extrapyramidal side effect? Gen Hosp Psychiatry 2001; 23:231–232
- 3. Tolosa E, Mart M: Adult-onset idiopathic torsion dystonia, in Movement Disorders: Neurologic Principles and Practice. Edited by Watts RL, Koller WC. New York, McGraw-Hill, Health Professions Division, 1997, pp 511–531
- 4. Frankhuijzen AL, Wardeh G, Hogenboom F, et al: Alpha 2-adrenoceptor mediated inhibition of the release of radiolabelled 5-hydroxytryptamine and noradrenaline from slices of the dorsal region of the rat brain. Naunyn Schmiedebergs Arch Pharmacol 1988; 337:255–260
- 5. Loscher W, Annies R, Richter A, et al: Marked regional disturbances in brain metabolism of monoaminergic neurotransmitters in the genetically dystonic hamster. Brain Res 1994; 658:199–208