

A Rare Case of Ethambutol Induced Optic Neuritis in a Peripheral Health Institute of Northern India

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ABSTRACT

Ethambutol (EMB) is one of the first-line drugs in the treatment of tuberculosis. EMB induces ocular toxicity which is a rare side-effect that is either dose-related or duration-related effect and is often reversible on therapy discontinuation. The toxicity has been reported at 4 to 12 months of the initiation of the therapy. We reported a rare case of EMB-induced optic neuritis after 1 month of initiation of the therapy. After the withdrawal of the drug, there was complete recovery of the patient. This case description would improve the understanding of the clinical presentation as well as the treatment of ethambutol optic neuritis, especially in recourse-limited settings. It would also enrich the knowledge regarding the prevention strategies and timely recognition of the side effect.

Keywords: Ethambutol, Optic neuritis, Isoniazid

INTRODUCTION

Ethambutol (EMB) hydrochloride is one of the first-line drugs which is used to treat tuberculosis. Though infrequent, ocular toxicity in the form of optic neuritis (most commonly retro bulbar neuritis) is well recognized and a serious complication that has been documented since the drug was first used in the 1960s¹. EMB induced ocular toxicity which is often dependent on the dose and also on the duration of the therapy; however the toxicity is reversible when the therapy is discontinued². Optic neuritis is the most important side-effect observed after the administration of Ethambutol. The optic neuritis is generally

reversible and is specific to the dose and duration of treatment². Prompt recognition of the side effect can help in preventing this sight-threatening side effect.

CASE DESCRIPTION

A 65-year female weighing 60 Kg visiting our OPD in November 2019 and was diagnosed with Pulmonary tuberculosis, based on the sputum microscopy. After performing the necessary routine workup, she was started on the Anti-tubercular treatment consisting of the Isoniazid, Rifampicin, pyrazinamide, and Ethambutol (HRZE). The dose of ethambutol administered to the patient was 15mg/kg^{3,4}. Before the treatment eye examination was conducted which showed a corrected visual acuity of 6/6 in both eyes. However, after one month the patient reported blurring of vision. Ophthalmologic examination revealed best corrected visual acuity of 6/12 in the right eye and 6/18 in the left. After 2 months, an ophthalmologic examination revealed that best corrected visual acuity was 6/60 in both eyes. Confrontation tests revealed bilateral central scotomas. Colour vision was tested using Ishihara charting, which was impaired in both eyes. On distant direct ophthalmoscopy, the optic disc of both the eyes was normal. Considering the possibility of EMB induced optic neuritis. EMB therapy was promptly stopped; however, the patient was continued on isoniazid, rifampicin, and pyrazinamide. There was a progressive but slow

improvement in her vision. Within 3 months, the best corrected visual acuity was 6/12 in the right eye and 6/9 in the left eye. The patient is on regular follow-up and is improving.

DISCUSSION

Ethambutol (EMB) is the most common drug used in the treatment of tuberculosis. The drug is regarded as the least toxic of the first-line anti-tuberculosis drugs, it also has a very low incidence of ocular toxicity.^{5, 6} The ocular toxicity induced from ethambutol is not common at standard doses used in the anti-tubercular regimen. Optic neuritis is the most important and potentially the most serious side effect of EMB. Retro bulbar neuritis is the most common form, there is involvement of either the axial fibres or less commonly paraxial fibers.^{6, 7} Clinically, and EMB optic neuropathy presents with the bilateral involvement of both the eyes as was seen in our case. However, the onset may be unilateral, but gradually, both eyes are involved. Symptoms of ethambutol toxicity have been reported between four and twelve months after starting EMB and are dose dependent, but rarely have been reported within few days of the start of therapy.⁸ Our patient was receiving standard 15mg/kg of the dose, despite that the symptoms appeared within one month of the treatment. On examination central scotoma is common with the loss of colour vision.⁹ Various risk factors have been identified which can predispose to the ocular toxicity of EMB, these include increasing age, prolonged duration of EMB, a higher dose, poor renal function, hypertension, diabetes, and concurrent optic neuritis, related to alcohol and tobacco use.^{10, 11, 12} The only treatment which has proven beneficial in EMB induced optic neuropathy is the discontinuation of the drug. Most patients will recover within weeks to months after stopping the drug,¹³ However, there are reports that vision may continue to decline or fail to recover even after the drug is stopped.¹⁴ The other drug, isoniazid which is

frequently administered with EMB can also cause ocular toxicity and this often leads to confusion regarding the aetiology of ocular toxicity in patients receiving EMB.¹⁵ If visual impairment persists even after stopping of EMB, the stoppage of INH can be considered. Routine visual acuity testing is recommended before initiation of the EMB by various international societies.¹⁶ ¹⁷Pre-treatment renal function should be measured, any history of eye diseases, pre-treatment record of visual acuity, routine visual acuity tests, and colour vision tests should be done in the patients on regular basis to avoid complications related to EMB toxicity. Physicians should frequently ask patients about changes in vision and make sure all patients understand that EMB should be immediately stopped and take medical advice immediately.

CONCLUSION

EMB, though being a safe anti-tubercular drug has the potential to cause serious ocular side effects in susceptible patients. Though the side-effect is known to be either dose or duration-related, sometimes it may occur even at lower doses and at shorter duration of treatments in susceptible individuals, leading to permanent vision loss. Even though, International guidelines on prevention and early detection of EMB induced ocular toxicity have been published, the physicians in recourse limited settings should be aware of the side effects and the management of this potential sight-threatening side effect. There is also a need to educate the patients about side-effects and check regular vision tests for early toxicity detection. If there is no vision improvement even after withdrawal of EMB, isoniazid toxicity should also be considered and the drug should also be stopped.

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