



Cefixime induced Erythema Multiforme (Case Study)

Dr Kasi Jagadeesh M

Assistant Professor, Department of Pharmacy Practice, Vikas Institute of Pharmaceutical Sciences, Nidigatla Road, Rajahmundry, Andhra Pradesh-533289.

Dr Achuta Lakshmi

Assistant Professor, Department of Pharmacy Practice, Vikas Institute of Pharmaceutical Sciences, Nidigatla Road, Rajahmundry, Andhra Pradesh-533289.

K Prathyusha

Student, Doctor of Pharmacy, Vikas Institute of Pharmaceutical Sciences, Nidigatla Road, Rajahmundry, Andhra Pradesh-533289.

K Dhanavardhan

Student, Doctor of Pharmacy, Vikas Institute of Pharmaceutical Sciences, Nidigatla Road, Rajahmundry, Andhra Pradesh-533289.

KEYWORDS :

INTRODUCTION

Erythema multiforme (EMF) is an acute skin and mucous membranes disease. It is characterized by symmetrical distribution of lesions on the skin, primarily on the extremities, and has a tendency of recurrences. Erythema multiforme is likely to be uncommon in childhood, and very less paediatric series of EMF^{1,2}. In EMF, pathologic changes during initial skin lesions consist of mononuclear cells at the superficial dermal blood vessels, epidermal damage, and keratinocyte necrosis. These are more characteristic of EMF leading to multilocular intra-epidermal blisters³. Depending on the absence or presence of mucous membrane erosions, causes may be classified as major EMF or minor EMF⁴. In Erythema multiforme, Eosinophils produce basic protein, eosinophil cationic protein, neurotoxin derived by eosinophils, and eosinophil peroxidase that could directly exploit involved tissue or augment the inflammatory cascade into the inflammatory loci by initiating other effector lymphocytes^{5,6,7}. Thus, eosinophil predominantly plays a vital role in drug eruptions pathogenesis. Yawalkar N et al, data indicated that eosinophil counts positively correlated with poor liver function, extended hospitalization, prolonged corticosteroid use in patients with Erythema multiforme-type drug eruption.

Eosinophilia occurs in 4 stages:

- (1) Eosinophil differentiation and proliferation in bone marrow
- (2) Rolling, adhesion, and migration
- (3) Migration through chemo-attraction and
- (4) Activation and consecutive cell death.

During the initial stage of drug eruption, circulating eosinophils initiated to cause cutaneous lesions or consume in larger numbers than bone marrow, immediate replenishment might contribute to low range of circulating eosinophils⁸. Circulating eosinophils of high levels in patients with Erythema multiforme-type drug eruption may be due to IL-5, IL-3, and GM-CSF over production in lesions and peripheral blood during drug eruption⁹. Longitudinal analyses signified increased eosinophil counts decreased notably in Erythema multiforme-type drug eruption after corticosteroids drug employment⁹. Cytokines high levels could stimulate differentiation and proliferation of eosinophils in bone marrow and augment eosinophil employ to peripheral blood and lesions^{10,11}.

Cefixime is an antibiotic belonging to third generation cephalosporins, they exert effect by adhering to penicillin-binding proteins (PBP) and by inhibiting peptidoglycan synthesis, thus making damage to cell wall of bacteria. Cefixime attains serum peak levels approximately 4 hours by oral dosing and half-life is of 3 to 4 hours and not dose dependent. It is excreted by renal mechanism and also biliary mech-

anism. About 50% of absorbed dose excreted unchanged in urine within 24 hours. There is no evidence for Cefixime metabolism *in vivo*^{12,13}. Even though dermatological reactions due to the Cefixime are rare, there are reports for pruritus, rashes, urticaria, and drug fever as hypersensitivity reactions. In fewer patients severe reactions are Stevens-Johnson syndrome, Erythema multiforme, and toxic epidermal necrolysis have also been reported¹⁴.

CASE REPORT

A twenty five old female patient visited to our dermatology department with the complaints of on and off fever, gangrene on the right foot, swelling of feet and hands with pain in joints. A diagnosis of foot ulcer had been made in government hospital and prescribed with tablet Cefixime (oral 200mg BID). After taking Cefixime 200mg orally, the very next day erythematous rashes were seen on hands and legs. The rashes were progressive day by day. After 15 days the patient got admitted to dermatology ward presenting Erythema multiforme lesions with palpable purpura along with feet and hand swelling. Based on the history and physical examination findings, a diagnosis of Cefixime induced EMF was made in our hospital & Treated with Dexamethasone 2cc IV OD was given along with tablet ascorbic acid, topical local application of liquid paraffin is advised. Patient recovered after 5 weeks with progressive response to the treatment.

DISCUSSION & CONCLUSION

EMF is an acute mucocutaneous, which is an inflammatory and hypersensitivity reaction characterized by symmetrical distribution of lesions as skin eruption, erythematous oedematous bullous lesions on skin or mucous membranes¹². EMF major is likely severe, typically involving 2 or more mucous membranes with more variable skin involvement. This feature is used to distinguish it from Stevens-Johnson syndrome, where there is extensive skin involvement, significant morbidity, and a mortality rate of 5% to 15%.

J Yang et al, in a study did horizontal comparison with 3 groups has ratified that eosinophil count significantly are higher in Erythema multiforme-type eruption patients. Yawalkar N et al., study data indicated the eosinophil counts positively and likely correlated with poor liver function, extended hospitalization, prolonged corticosteroid use in patients with Erythema multiforme-type drug eruption⁹. Farthing P et al, observed EM more in males, similar incidence of drug-related EMF is found in both males and females¹³. Cefixime is a third-generation oral cephalosporin which is commonly utilized in the recent world for bacterial infections. It is an antibiotic that exerts effect by adhering to penicillin-binding proteins (PBP) and by inhibiting peptidoglycan synthesis, thus making damage to cell wall of bacteria. Gupta's

et al also reported dermatological fixed drug eruptions in an outpatient due to the use of Cefixime. Even though reactions by Cefixime are not likely, there are findings of pruritus, rash, urticaria, and drug sensitive fever as hypersensitivity reactions in not more but less than 2% of patients. Severe reactions such as erythematous multiforme, Stevens - Johnson syndrome, and toxic epidermal necrolysis have been also reported by cefixime.

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