

ACE- inhibitor induced angioedema requiring an emergency surgical airway

Bhargav Patel MD, Hemant Raval MD, Gaurav Patel MD
Ahmed Al-Chalabi MD, Robert Fleishman MD

ABSTRACT

Angiotensin converting enzyme (ACE) inhibitors cause approximately 30% of the cases of angioedema which present to the emergency department; 11% of these patients require ICU admission. We report two patients who required emergency surgical airways secondary to angioedema related to ACE-inhibitors. Clinicians need to remember that these situations can be extremely dangerous and plan airway management carefully in these patients.

Key words: angioedema, tracheostomy, angiotensin converting enzyme inhibitors

INTRODUCTION

Angioedema is a well-known adverse effect of angiotensin converting enzyme (ACE)-inhibitor therapy that has been seen in up to 1% of recipients.^{1,2} Out of all angioedema-related emergency department (ED) visits each year, approximately 30% of cases are attributed to ACE-inhibitors.³ Most cases are usually reversible with discontinuation of the drug; treatment in the intensive care unit is required in 11% of patients.³ We present a case with severe upper airway obstruction requiring cricothyroidotomy followed by tracheostomy to manage the airway and a second case requiring emergency tracheostomy.

CASE PRESENTATION 1

A 59-year-old African-American woman who had started lisinopril (10mg daily) 10 days prior presented with facial swelling and increasing difficulty in breathing for one day. Her other medications included aspirin (81mg for years), amlodipine, glyburide, insulin, and rosuvastatin. She had no known drug allergies. She spoke in short sentences with a muffled voice associated with drooling and had trouble swallowing. She had swollen lips, tongue, soft palate, and hard palate. She had no urticaria. The ED physicians were unable to visualize the uvula. She was diagnosed with ACE-inhibitor induced angioedema. Despite diphenhydramine, epinephrine, and methylprednisolone, continued respiratory distress led to a nasal intubation attempt by the anesthesiology service. When intubation failed, an emergent cricothyroidotomy was performed. Tracheostomy was completed after she stabilized. She was monitored in the SICU for five days; the angioedema resolved and the

Corresponding author: Bhargav Patel MD
Contact Information: drbhargavpatel@gmail.com
DOI: 10.12746/swrccc2015.0309.120

tracheostomy tube was removed.

CASE PRESENTATION 2

A 74-year-old African-American woman on fosinopril (20mg daily) presented to the ED with swelling of the tongue and difficulty in breathing. She was unable to speak full sentences or clear her secretions. Her other medications included cefdinir, hydrochlorothiazide, simvastatin, and folic acid with no known drug allergies. She was diagnosed with ACE-inhibitor induced angioedema unresponsive to diphenhydramine and methylprednisolone. After failed attempts by the anesthesiology service to obtain orotracheal or nasotracheal airway, a tracheostomy was performed with a 6 Fr cuffed tracheostomy tube. Eventually her angioedema resolved, and the patient was transferred to the regular inpatient service for further management and disposition.

DISCUSSION

ACE-inhibitor induced angioedema is a potentially life threatening emergency. The benefit of medical management with epinephrine, antihistamine medications, and steroids is uncertain as the pathophysiological pathways involve increased levels of bradykinin.⁴ Recently, results of a trial evaluating treatment with a kallikrein inhibitor (ecallantide) were disappointing.⁵ A bradykinin B2 receptor antagonist has been reported effective in several case reports, but more randomized studies are needed.^{6,7} The need for a surgical airway in ACE-inhibitor induced angioedema is estimated to be 1%.³ Frequent assessment and repeated monitoring of the airway are essential since intubation and mechanical ventilation may be required in severe cases. An increased need for intubation/tracheostomy has been reported in hospitalized patients with angioedema due to hypertension drugs.⁸ Occasionally, the usual preference for endotracheal intubation is replaced with tracheostomy for better control of the airway.⁹ While Mallampati tests have limited accuracy for predicting difficult intubation,¹⁰ some retrospective studies have reported the

value of fiberoptic laryngoscopy to predict early airway intervention based on the patient's age and various sites involved in upper respiratory tract and oropharynx.^{11,12} Patients with difficult airways may be good candidates for more invasive airway management instead of attempting intubation. Our cases illustrate the need for an emergent cricothyroidotomy and/or tracheostomy rather than repeated efforts with difficult intubations for ACE-inhibitor induced angioedema with respiratory failure and hemodynamic instability. More studies are needed to determine the best candidates for early surgical intervention when a difficult intubation seems likely.

Author Affiliation: Bhargav Patel is a resident in internal medicine at Jamaica Hospital Medical Center in New York. Hemantkumar Raval is a fellow in critical care medicine at Mount Sinai Hospital in New York. Gaurav Patel is a fellow in pulmonary medicine at Texas Tech University Health Sciences Center in Lubbock TX. Ahmed Al-Chalabi is a resident at Jamaica Medical Center in New York. Robert Fleishman is a resident at Hahnemann University Hospital in Philadelphia, PA.

Received: 10/20/2014

Accepted: 12/19/2014

Reviewers: James Tarbox MD

Published electronically: 01/15/2015

Conflict of Interest Disclosures: none

REFERENCES

1. Makani H, Messerli FH, Romero J, *et al.* Meta-analysis of randomized trials of angioedema as an adverse event of renin-angiotensin system inhibitors. *Am J Cardiol* 2012; 110(3):383-91.
2. Vleeming W, van Amsterda JG, Stricker BH, de Wildt DJ. ACE inhibitor-induced angioedema. incidence, prevention and management. *Drug Saf* 1998; 18(3):171-88.
3. Banerji A, Clark S, Blanda M, *et al.* Multicenter study of patients with angiotensin-converting enzyme inhibitor-induced angioedema who present to the emergency department. *Ann Allergy*

Asthma Immunol 2008; 100(4): 327-32.

4. Nussberger J, Cugno M, Amstutz C, *et al.* Plasma bradykinin in angio-oedema. *Lancet* 1998; 351(9117):1693-7.

5. Lewis LM, Graffeo C, Crosley P, *et al.* Ecallantide for the acute treatment of angiotensin-converting enzyme inhibitor-induced angioedema: a multicenter, randomized, controlled trial. *Ann Emerg Med* 2014.

6. Bas M, Greve J, Stelter K, *et al.* Therapeutic efficacy of icatibant in angioedema induced by angiotensin-converting enzyme inhibitors: a case series. *Ann Emerg Med* 2010; 56(3):278-82.

7. Del Corso I, Puxeddu I, Sardano E, *et al.* Treatment of idiopathic nonhistaminergic angioedema with bradykinin B2 receptor antagonist icatibant. *Ann Allergy Asthma Immunol* 2012; 108(6):460-1.

8. Lin R Y, Levine RJ, Lin H. Adverse drug effects and angioedema hospitalizations in the United States from 2000 to 2009. *Allergy Asthma Proc* 2013; 34(1):65-71.

9. Pruet CW, Kornblut AD, Brickman C, *et al.* Management of the airway in patients with angioedema. *Laryngoscope* 1983; 93(6):749-55.

10. Lee A, Fan LT, Gin T, *et al.* A systematic review (meta-analysis) of the accuracy of the Mallampati tests to predict the difficult airway. *Anesth Analg* 2006; 102(6):1867-78.

11. McCormick M, Folbe AJ, Lin HS, *et al.* Site involvement as a predictor of airway intervention in angioedema. *Laryngoscope* 2011; 121(2):262-6.

12. Zirkle M, Bhattacharyya N. Predictors of airway intervention in angioedema of the head and neck. *Otolaryngol Head Neck Surg* 2000; 123(3): 240-5.