# Peri-operative dexamethasone therapy and post-operative psychosis in patients undergoing major oral and maxillofacial surgery

#### Chethan Manohara Koteswara, Pritish Patnaik<sup>1</sup>

Department of Anaesthesiology, AJ Institute of Medical Sciences, <sup>1</sup>Department of Oral and Maxillofacial Surgery, AJ Institute of Dental Sciences, Mangalore, Karnataka, India

# <u>Abstract</u>

A broad array of behavioral symptoms, including psychosis, can transpire post-operatively following a variety of surgeries. It is difficult to diagnose the exact cause of post-operative psychosis. We report three cases, which developed psychosis post-operatively after undergoing major oral and maxillofacial surgeries. All the three patients were administered dexamethasone peri-operatively. Dexamethasone is used to prevent or reduce post-operative edema. The exact dose of dexamethasone, which can cause psychosis, is unknown. It is important to raise awareness about this potential complication so that measures for management can be put in place in anticipation of such an event.

Key words: Dexamethasone, oral and maxillofacial surgery, post-operative psychosis

## Introduction

Glucocorticosteroids have been used in an attempt to minimize or prevent post-operative edema after oral and maxillofacial surgery. Steroid use is associated with more complications especially in long-term administration. Dexamethasone seems to be the most suitable because it has the highest anti-inflammatory activity, no mineralocorticoid activity and a longest available half-life of 36-54 h. Psychosis is a generic psychiatric term for a mental state often described as involving a "loss of contact with reality." People experiencing psychosis may report hallucinations or delusional beliefs and they may exhibit personality changes and disorganized thinking. We report three maxillofacial cases who experienced post-operative psychosis after peri-operative dexamethasone therapy, as a result of which, the patients had a prolonged stay in the surgical intensive care unit.

Address for correspondence: Dr. Chethan Manohara Koteswara, Department of Anaesthesiology, AJ Institute of Medical Sciences, Mangalore, Karnataka, India.

E-mail: major.chethan@gmail.com

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# **Case Reports**

#### Case 1

An 18-year-old, female patient, presented for surgery for mandibular prognathism. She was taken up for orthognathic surgery. Pre-operatively, as per protocol, she received intravenous (IV) dexamethasone 16 mg on the day of surgery at 6 am. Patient was pre-medicated with IV fentanyl 75 µg. General anesthesia was induced with propofol and intubated with vecuronium. Anesthesia was maintained with N<sub>2</sub>O-O<sub>2</sub>-sevoflurane. Surgery lasted for 4 h. After extubation, she was transferred to the post-operative care unit (PACU) awake and calm. On arriving at PACU, she was given another dose of dexamethasone, 8 mg. After 2 h, she became agitated, restless and aggressive. IV midazolam, propofol and IV fentanyl were administered for sedation, but the patient remained agitated and confused. Over the next 3 h, IV fentanyl 100 µg, propofol 40 mg, midazolam 4 mg was administered after which the patient was relatively calm. Then, she became restless and delirious. Psychiatry consultation was sought. She was diagnosed with an acute psychotic episode possibly induced by steroids. IV midazolam 2 mg and intramuscular (IM) haloperidol 4 mg were given. An IV dexmedetomidine infusion at 0.4 microgm/kg/min tapered to 0.2 microgm/kg/min, provided sedation for 6 h, after which it was stopped. As the patient continued to be calm over the next 12 h, she was transferred to the ward. She continued to be stable and was discharged on the post-operative day 5.

#### Case 2

A 58-year-old, female patient, presented for surgery for wide local excision of squamous cell carcinoma of maxilla and closure with split skin graft. Pre-operatively, as per protocol, she received IV dexamethasone 16 mg on the day of surgery at 6 am. Patient was pre-medicated with IV fentanyl 100 µg. General anesthesia was induced with propofol and intubated with vecuronium. Anesthesia was maintained with N<sub>2</sub>O-O<sub>2</sub>-sevoflurane. Surgery lasted for 2 h. After extubation, she was awake and calm. On arriving at PACU, Another dose of 8 mg of dexamethasone was administered. After 6 h, she became restless and delirious. Psychiatry consultation was sought immediately. She too was diagnosed with an acute psychotic episode possibly induced by steroids. IV midazolam 2 mg and IM haloperidol 5 mg were administered. An IV dexmedetomedine infusion providing sedation was administered for 3 h at 0.4 µg/kg/h. She remained calm over the next 4 h after which she again became agitated. Midazolam 2 mg and haloperidol 2.5 mg were repeated. The dexmedetomidine infusion was restarted for next 6 h, initially at 0.4 µg/kg/h tapered to 0.2 µg/kg/h and then stopped. As the patient remained calm over the next 12 h, she was transferred to the ward. She continued to be psychologically stable and was discharged on the post-operative day 6.

#### Case 3

An 80-year-old, female patient, weighing 35 kg presented with biopsy proven squamous cell carcinoma of the left side of the mandible. She had no other comorbidities. She was taken up for wide excision with segmental mandibulectomy, modified radical neck dissection and reconstruction with pectoralis major myocutaneous flap. Pre-operatively, as per protocol, she received IV dexamethasone 8 mg on the day of surgery at 6 am. Patient was pre-medicated with fentanyl 50 µg. General anesthesia was induced with propofol and intubated with vecuronium. Anesthesia was maintained with N<sub>2</sub>O-O<sub>2</sub>-sevoflurane. Surgery lasted for 4 h. She was electively placed on endotracheal tube (ETT) with T-piece in the post-operative period in view of extensive surgery. In the

PACU, another dose of 8 mg was administered. Fourteen hours after surgery, she became restless and she pulled out her ETT and Ryle's tube. She started bleeding from her suture site. She was wheeled into the operation theatre immediately for secondary suturing. After the surgery, she was placed on dexmedetomidine infusion for 12 h, initially at 0.3  $\mu$ /kg/h and then tapered to 0.1 $\mu$ /kg/h (the infusion was stopped intermittently whenever the patient developed bradycardia or hypotension and whenever the patient remained calm). IV midazolam 1 mg and IM haloperidol 2.5 mg were continued 8 hourly for 2 days, after which the rest of post-operative period was uneventful. She was extubated 16 h after the surgery and then shifted back to the ward. She continued to be psychologically stable and was discharged on the post-operative day 8.

#### **Discussion**

Glucocorticosteroids are used extensively for minimizing and preventing post-operative oedema after oral and maxillofacial surgery.[1] Dexamethasone seems to be the most suitable because it has the highest anti-inflammatory activity, no mineralocorticoid activity and a longest available half-life of 36 to 54 h.[2] Dexamethasone has also been in use in the peri-operative period for post-operative nausea and vomiting<sup>[3,4]</sup> pain management<sup>[5-7]</sup> and in neurosurgery. [8] Even though, efficacious for preventing and minimizing edema in oral and maxillofacial surgeries, there have been reports attributing post-operative psychosis to peri-operative dexamethasone [Table 1].[9,10] In fact in one of the reports a single dose has been implicated for the development of post-operative psychosis. [9,10] While the incidence of post-operative delirium is 36.8%,[11] acute steroid related psychiatric disorder frequency is 5%. [12] In view of such a high incidence, it may be imperative that, with every case where peri-operative dexamethasone is administered, a post-operative psychotic reaction should be anticipated. In patients who are being post-operatively ventilated, it may be necessary to use restraints to prevent self-extubation and a pre-operative consent for the same would be required. Haloperidol prophylaxis may be used for these cases. The episodes of psychosis can prolong

Table 1: Patient profile in previously published case reports								
Authors	Age (years)	Gender	Surgery	Co-existing systemic disorders	Total dose of dexamethasone	Day of presentation of symptoms	Treatment given	
Silva and Tolstunov 1995 <sup>[13]</sup>	45	Male	Left hemimaxillectomy	Ischemic heart disease, deep vein thrombosis, major depression	60 mg	2 <sup>nd</sup> post-operative day	Haloperidol, perphenazine	
Ferris and Eisele 2003 <sup>[12]</sup>	62	Male	Parotidectomy	Nil	24 mg	2 <sup>nd</sup> post-operative day	Sertraline, risperidone, lorazepam	
Artukoglu and Asenjo 2007 <sup>[10]</sup>	33	Female	Biopsy of humerus	Nil	8 mg	Immediate post- operative period	Chlorpromazine, haloperidol, propofol infusion	
Bansal et al. 2009 <sup>[9]</sup>	19	Female	Enucleation of periapical cyst	Nil	8 mg	3 <sup>rd</sup> post-operative day	Lorazepam, risperidone	

the hospital stay<sup>[13]</sup> or may result in readmission<sup>[12]</sup> and increase the total cost for the patient. Early recognition and management with sedatives/anti-psychotic drugs is crucial. Once there is a psychotic event, the stay of the patient is prolonged in the intensive care units thereby increasing the costs. The exact dose of dexamethasone that can cause a psychotic reaction is unknown. Since it has been reported that a single dose can cause post-operative psychosis [Table 1] all patients receiving dexamethasone have to be monitored carefully post-operatively. Special care has to be taken in patients who have a previous history of psychiatric disorders.<sup>[13]</sup>

## **Conclusion**

Awareness and the ability to diagnose post-operative psychosis following use of peri-operative dexamethasone are crucial to provide better care and effective treatment.

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