

Propofol Induced Acute Pancreatitis-A Rare Adverse Effect For A Common Drug

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Abstract

Acute pancreatitis caused by a widely and commonly used anaesthetic drug, Propofol, is an uncommon event rarely reported in Malaysia's literature. Drugs induced acute pancreatitis is also a rare phenomenon in Malaysia, where the two common causes of acute pancreatitis here are gallstones and alcohol consumptions. The mechanism of the incidence of Propofol induced acute pancreatitis is still unknown and remain unclear. While this condition is a diagnosis of exclusion, management's principle remains the same as acute pancreatitis caused by other risk factors. We present a case of Propofol induced acute pancreatitis in an elderly lady after a gynaecological surgery to highlight the rarity of the case and stress the possibility of acute pancreatitis after Propofol administration in a patient without other risk factors.

Keywords: Propofol, acute pancreatitis,

Introduction

Acute pancreatitis is a sudden inflammation of the pancreas that can manifest clinically in various spectrum of clinical situations. The two most common causes of acute pancreatitis are gallstones and alcohol consumption. Rarely, acute pancreatitis is drug-induced where the exact incident rate and the mechanism of action of drug-induced acute pancreatitis are still unknown and remains unclear (1). Drugs commonly listed as the causative agent of acute pancreatitis include angiotensin-converting enzyme inhibitors, statins, steroids, oral contraceptives, hormone replacement therapy, diuretics, and highly active antiretroviral therapy, valproic acid and hypoglycaemic agents (2).

Propofol is a short-acting anaesthetic agent mostly used as a parenteral anaesthetic agent worldwide (3). Propofol is a lipid-based hypnotic agent widely used during major surgeries and short procedures such as endoscopy and sedation of patients undergoing mechanical ventilation. While Propofol usage is common, the incidence of acute pancreatitis induced by Propofol is still rare, and only a few cases were reported in this region. We are reporting a case of an elderly lady who was diagnosed to have acute pancreatitis one day after undergoing general anaesthesia with Propofol administration

during gynaecological surgery. She was not documented to have any evidence of other risk factors to develop acute pancreatitis, including consuming any other drugs.

Case Report

A 67yearsold lady presented to the Emergency Department complaining of severe abdominal pain, nausea, vomiting and fever. A day before this episode, she had undergone colposcopy and cervical biopsy under general anaesthesia to investigate the cause of her post-menopausal bleeding. During the daycare procedure, she was given 200mg of IV Propofol during induction by the anaesthetist. As this short procedure lasted about 15 minutes, she was not given any infusion of Propofol to maintain the general anaesthesia. She was also given 70% nitrous oxide and oxygen throughout the short duration of general anaesthesia, and intravenous Paracetamol was given for analgesia. She was then monitored post-surgically and was discharged well on the same day. She was discharged only with oral Paracetamol and given follow up appointment. She started to develop symptoms of acute abdomen the next morning and was rushed to hospital by the afternoon when her symptoms worsen.

During her post-operative presentation to the hospital, the patient was alert, conscious but appeared dehydrated. She was, however, hemodynamically stable. On palpation, the epigastric region was tender. However, no evidence of peritonitis found on her. Bowel sound was normal on auscultation. Examination by the gynaecologist who performed the first surgery revealed no significant complications from the surgery.

Blood investigations revealed a raise in total white cell count ($17 \times 10^9/L$). Serum amylase was significantly high documented at 3397u/L. All other blood investigations, including renal profile, liver function test and arterial blood gas (ABG), were within normal parameters. A erect chest X-ray was performed on the patient, and there was no gas under the diaphragm to indicate perforated viscus. Contrast-enhanced CT abdomen and pelvis of the patient indicated hypodensities in the peripancreatic areas of the right and left pararenal spaces (Figures 1, 2 and 3), suggesting acute pancreatitis with minimal ascites around the liver.

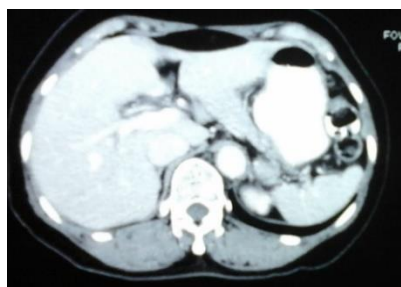


Figure 1



Figure 2

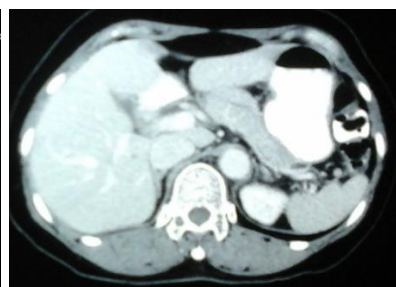


Figure 3

Figures 1,2 and 3: Showing oedematous pancreatic parenchyma and prominent vessel suggesting acute pancreatitis in the Contrast Enhance CT of Abdomen

The patient was then admitted, managed, and monitored in-patient with intravenous fluids, opioids analgesia, a proton-pump inhibitor, and allowed orally within a day of admission. The patient recovered well with conservative management and was discharged well three days later.

Discussion

Acute pancreatitis after propofol administration is a rare adverse event of this anaesthetic drug (4). Propofol-induced acute pancreatitis has been reported at 0.5% - 2 % of all cases of acute pancreatitis (5). A short-acting Propofol is mostly used as an anaesthetic agent in the United States and worldwide (3). Propofol was approved by the US Food and Drug Administration (FDA) almost three decades ago. Propofol has been extensively used during General Anaesthesia and induction of sedation for patients who require mechanical ventilation.

Furthermore, Propofol also used during short procedures such as endoscopy, transoesophageal echocardiogram and abscess drainage (6). In some rare cases, the use of Propofol has been associated with idiopathic pancreatitis (7), and when using for a long period and in high doses, it can cause "Propofol infusion syndrome" (3). Propofol is insoluble in water. Hence when given, it is usually given in a lipid emulsion. The typical dose for Propofol is 1.5 to 2.5 mg/kg, followed by small boluses of low doses of infusion as maintenance of anaesthesia (3). Propofol is classified as a Class II drug in the Badalov classification (8) of drugs that induce acute pancreatitis (1). In a literature review by Scholten, it is reported that there was no definite causal relationship between Propofol and the incidence of acute pancreatitis (4), which make the mechanism of Propofol inducing acute pancreatitis remains unclear (1). A few theories were reported on the possible mechanism of propofol-induced acute pancreatitis, including hypertriglyceridemia, hypersensitivity, or direct pancreatic toxicity of the drug (6).

Csomor, 2017, reported a case of acute pancreatitis after an elective thyroidectomy where Propofol was used as an anaesthetic agent. All other common causes were ruled out, and they concluded that their patient had developed acute necrotising pancreatitis confirmed through a contrast-enhanced CT scan (5). The incident was similar to our patient diagnosed and confirmed to have acute pancreatitis using a contrasted CT scan. The finding shows the importance of a contrast-enhanced CT scan of the abdomen in confirming the incidence of acute pancreatitis presenting with acute abdomen in the patient with doubtful presentation or without any risk factors.

In another report by Muniraj, the patient developed acute pancreatitis after administering propofol infusion in an intensive care setting and was found to have underlying hypertriglyceridemia (6). This report was different from our patient, who did not have any underlying illnesses before the surgery. She was not a critically ill patient, and she was not given any Propofol infusion but rather just a bolus. This finding highlights that Propofol can cause acute pancreatitis in isolation without other risk factors such as hypertriglyceridemia, and Propofol can cause acute pancreatitis after both infusion and bolus administration.

Gottschling reported that they encountered a similar situation of acute pancreatitis hours after their patient was administered with propofol bolus before a magnetic resonance imaging procedure (9). This is comparable to our patient who developed acute pancreatitis within a short duration of Propofol administration. This information leads us back to the question of the mechanism of Propofol induced pancreatitis and whether the direct toxicity of the drug to the pancreatic parenchyma could be the most likely reason as acute pancreatitis in these cases occurs almost immediately after Propofol administration.

Conclusion

Propofol-induced acute pancreatitis is rare even though Propofol is one of the most used anaesthetic drugs daily for various reasons worldwide. Clinicians should always keep in mind that patient who received Propofol administration may be at risk of developing acute pancreatitis especially when other causes of the acute abdomen have been ruled out.

References

1. Haffar S, Kaur RJ, Garg SK, et al. Acute pancreatitis associated with intravenous administration of Propofol: evaluation of causality in a systematic review of the literature. *Gastroenterol Rep (Oxf)*. 2019;7(1):13-23.
2. Kaurich T. Drug-induced acute pancreatitis. *Proc (Bayl Univ Med Cent)*. 2008;21(1):77-81.
3. Propofol. In: *LiverTox: Clinical and Research Information on Drug-Induced Liver Injury* [Internet]. Bethesda (MD): National Institute of Diabetes and Digestive and Kidney Diseases; 2012–. PMID: 31643176
4. Bethesda (MD): National Institute of Diabetes and Digestive and Kidney Diseases; 2012.

5. Scholten JG, Buijs E. Acute pancreatitis na het toedienen van propofol [Acute pancreatitis after propofol administration]. Ned Tijdschr Geneesk. 2014;158: A7115.
6. Csomor J, Murínová I, Broulíková K, et al. Propofol-induced acute pancreatitis. J Clin Pharm Ther. 2017;42(4):495-498.
7. Asghar MU, Cheema HA, Tanveer K, Leinwand J. Propofol Infusion and Acute Pancreatitis. Am J Ther. 2019;10.1097
8. Muniraj T, Aslanian HR. Hypertriglyceridemia independent propofol-induced pancreatitis. JOP. 2012;13(4):451-453.
9. Badalov N, Baradaran R, Iswara K, Li J, Steinberg W, Tenner S. Drug-induced acute pancreatitis: an evidence-based review. Clin Gastroenterol Hepatol. 2007;5(6):648-644.
10. Gottschling S, Larsen R, Meyer S, Graf N, Reinhard H. Acute pancreatitis induced by short-term propofol administration. Paediatr Anaesth. 2005;15(11):1006-1008.