



Comment on “The Effect of Nitroglycerine Infusion on Postoperative Pain in Lower Limb Surgery: A Clinical Double-Blind Study”

Mohammad Sistanizad^{1,2}, Ali Saffaei³ and Seyedpouzhia Shojaei^{4,*}

¹Prevention of Cardiovascular Disease Research Center, Shahid Beheshti University of Medical Sciences, Tehran, Iran

²Department of Clinical Pharmacy, School of Pharmacy, Shahid Beheshti University of Medical Sciences, Tehran, Iran

³Student Research Committee, Department of Clinical Pharmacy, School of Pharmacy, Shahid Beheshti University of Medical Sciences, Tehran, Iran

⁴Department of Critical Care Medicine, Emam Hossein Medical and Educational Center, Shahid Beheshti University of Medical Sciences, Tehran, Iran

*Corresponding author: Department of Critical Care Medicine, Emam Hossein Medical and Educational Center, Shahid Beheshti University of Medical Sciences, Tehran, Iran. Tel: +98-2173432309, Email: poujsh@gmail.com

Received 2019 October 02; Accepted 2019 December 03.

Keywords: Nitroglycerine, Pain, Surgery, Fentanyl

Dear Editor,

We had this chance to read the valuable article entitled “The Effect of Nitroglycerine Infusion on Postoperative Pain in Lower Limb Surgery: A Clinical Double-Blind Study” published by Rahimzadeh et al. (1). This is a well-designed study that emphasized a novel strategy regarding pain management in postoperative settings. The authors prepared three stock formulations of fentanyl and nitroglycerin and subsequently administered them via infusion pumps. These three formulations were as follows:

A) Fentanyl 10 mc/kg + 10 mL distilled water + 100 mL normal saline;

B) Fentanyl 10 mc/kg + nitroglycerin 500 mc diluted in 10 mL distilled water and then diluted with 100 mL normal saline;

C) Fentanyl 10 mc/kg + nitroglycerin 1000 mc diluted in 10 mL distilled water and then diluted with 100 mL normal saline.

They started infusion right after the operation and followed patients for 48 hours after surgery. The authors found that nitroglycerine, as an adjuvant agent, could be added to intravenous fentanyl in patients undergoing surgery for better pain control. However, the authors did not consider the compatibility and stability of the prepared mixtures. The compatibility and stability of intervening agents are important, especially in intensive care units where patients must receive several intravenous medicines. This can be easily prevented through checking by online databases such as the handbook on injectable drugs or Lexi intravenous compatibility database (2). Ac-

ording to these databases, the mixture of fentanyl at a concentration of 50 µg/mL and nitroglycerin at a concentration of 0.2 mg/mL is visually compatible for up to 24 hours at 24°C. Also, the mixture of fentanyl at a concentration of 0.05 mg/mL and nitroglycerin at a concentration of 0.4 mg/mL is visually compatible for up to 4 hours at 27°C. All mixtures also should be diluted with dexterosus water 5% (3). On the other hand, nitroglycerin is a sensitive medicine in terms of stability and physical condition can easily affect its potency (4). Hence, in conclusion, it was better to evaluate the compatibility and stability of prepared stock mixtures after 24 hours. Alternatively, they could prepare fresh solutions instead of preparing stock solutions. Checking the compatibility and stability with online databases is also recommended before all these strategies.

Footnotes

Authors' Contribution: Mohammad Sistanizad, Seyedpouzhia Shojaei, and Ali Saffaei designed the study and they prepared the first draft of the manuscript. All authors approved the final version of the manuscript.

Conflict of Interests: No conflict of interest is reported.

Funding/Support: No funding source is reported.

References

1. Rahimzadeh P, Faiz SHR, Imani F, Soltani A, Derakhshan P. The effect of nitroglycerine infusion on postoperative pain in lower limb surgery: A clinical double-blind study. *Anesth Pain Med.* 2019;9(4).

- e93848. doi: [10.5812/aapm.93848](https://doi.org/10.5812/aapm.93848). [PubMed: [31754611](https://pubmed.ncbi.nlm.nih.gov/31754611/)]. [PubMed Central: [PMC6825369](https://pubmed.ncbi.nlm.nih.gov/PMC6825369/)].
- Smith WD, Karpinski JP, Timpe EM, Hatton RC. Evaluation of seven i.v. drug compatibility references by using requests from a drug information center. *Am J Health Syst Pharm*. 2009;**66**(15):1369-75. doi: [10.2146/ajhp080373](https://doi.org/10.2146/ajhp080373). [PubMed: [19635773](https://pubmed.ncbi.nlm.nih.gov/19635773/)].
 - McEvoy GK; Pharmacists ASHP's. *Handbook on injectable drugs*. USA: American Society of Health-System Pharmacists; 2016.
 - Scheife AH, Grisafe JA, Shargel L. Stability of intravenous nitroglycerin solutions. *J Pharm Sci*. 1982;**71**(1):55-9. doi: [10.1002/jps.2600710114](https://doi.org/10.1002/jps.2600710114). [PubMed: [6799641](https://pubmed.ncbi.nlm.nih.gov/6799641/)].