



Therapeutic Hypothermia induced Hypokalemia and Hyperglycemia in Cardiac Surgery

Authors

**Tuhinshra Medda¹, Kamal Das², Pritam Nandy³, Nripendra Kr.Tiwari⁴,
Sudipta Patra⁵, Subrata Dey⁶**

^{1,3,4}BSc Perfusion Technology, Dept of Cardiothoracic & Vascular Surgery, R G Kar Medical College, Kolkata

²MSc in Perfusion Sciences, Chief Perfusionist, Department of Cardiothoracic & Vascular Surgery, R G Kar Medical College, Kolkata

⁵MSc in Perfusion Sciences, Chief Perfusionist, Department of Cardiothoracic & Vascular Surgery, R G Kar Medical College, Kolkata

⁶H.O.D Department of Cardiothoracic & Vascular Surgery, R G Kar Medical College, Kolkata

Abstract

Hyperglycemia is common in critically ill patients and has been associated with poor outcomes. The effect of hypothermia, whether induced or spontaneous, on insulin resistance and hyperglycemia is not well understood and sometimes overlooked. In cardiac surgery hypothermia impairs blood glucose homeostasis and insulin sensitivity⁽¹⁾. However, the impact of therapeutic hypothermia on blood glucose levels and insulin requirements is unknown⁽¹⁾.

Keywords: *Therapeutic Hypothermia, Hyperglycemia.*

Introduction

Hyperglycemia has been associated with poor neurological outcomes for patients treated with therapeutic hypothermia, but it is unclear whether hyperglycemia directly contributes to deleterious outcomes or is simply a marker for overall poor health⁽²⁾. Hypothermia indirectly increases the glucose levels in the blood stream, resulting in hyperglycemia. As hypothermia is initiated, sympathetic activity is increased resulting in elevated levels of catecholamines and free fatty acids, results in decreased insulin secretion and increased tissue resistance to insulin.⁽³⁾ The increase in catecholamine release is compounded by impaired peripheral glucose uptake at the tissue

level because of hypothermia⁽³⁾. The decrease of is also caused by cooling of the islets of Langerhans, responsible for insulin secretion⁽⁴⁾. A third effect of hypothermia is hypokalemia⁽⁵⁾. Hypokalemia is associated with insulin resistance and hyperglycemia⁽⁶⁾. Therefore, three factors play a role in increasing glucose levels in hypothermic conditions: increased catecholamine levels, hypothermic effect on the pancreas, and hypothermic induction of hypokalemia⁽³⁾.

In cardiac surgery hypothermia is an important method to protect organs during low flow. But hypothermia induced hypoglycaemia is also a challenge to surgical outcome. In surgical patients, perioperative hyperglycemia increases risk of

postoperative mortality, and cardiovascular, respiratory, neurologic, and infectious morbidity ⁽¹⁾.

Method

In this study 50 consecutive adult patients were taken in a randomised nonblinded manner elective open heart surgery like ASD, VSD, MVR, DVR, CABG ON PUMP under general anaesthesia, moderate hypothermia (28-32 °C) and alpha stat pH management.

Preoperative exclusion criteria were

- 1) Less than 18 years,
- 2) Diabetic patients,
- 3) Organ dysfunction.

During surgery we divided total CPB Bypass period in to three phases- Cooling Phase, Plateau Phase and Rewarming Phase. We monitored nasopharyngeal temperature. In cooling phase we monitored the blood glucose level and serum potassium every after 2°C decrease of body temperature. During Plateau phase the temperature was remain same and we check glucose every after 20 minutes. In rewarming phase we check blood glucose level and serum potassium every after 2°C decrease of body temperature.

We found a higher blood glucose level and hypokalemia with decreasing temperature. We added insulin as our institutional protocol. During bypass we added insulin as bolus dose according to our institutional protocol (Table-1).

Table-1. Protocol for bolus insulin dose.

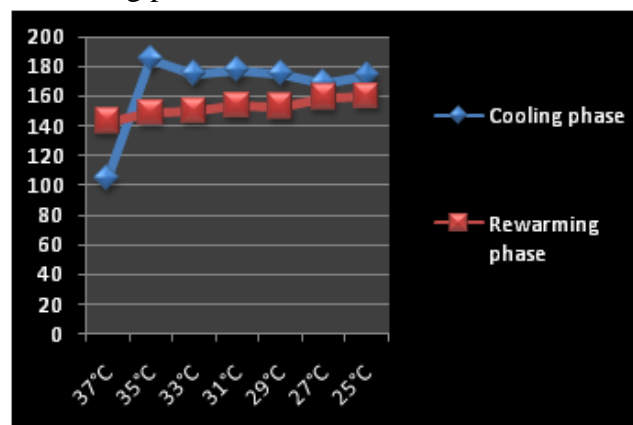
Blood glucose level (mg/dl)	Insulin Bolus Units (IU)
170-180	5
170-190	8
190-200	10
>200	15

The Society of Thoracic Surgeons recommends insulin for cardiac surgical patients when glucose concentrations are persistently >180 mg/dL ⁽⁷⁾

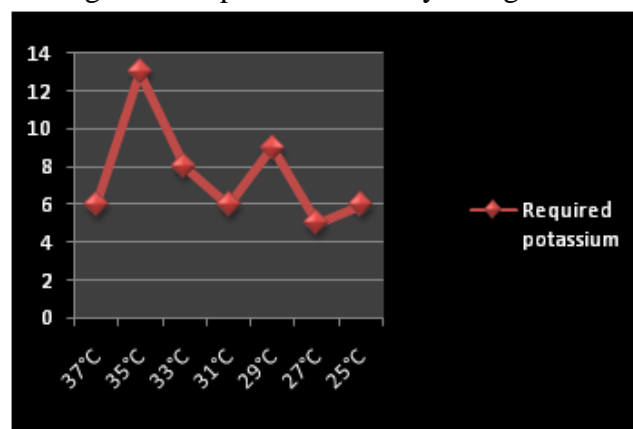
Result

In our study we found that blood glucose level was increasing with decreasing temperature. Before anaesthesia an average blood glucose level of 50 patients was 90-100mg/dl. After anaesthesia

it was 100-110 mg/ dl. Just 10 minutes after the starting of CPB bypass we found a raised blood glucose level (average-180-190 mg/dl) and hypokalemia. We added insulin as our protocol and hypokalemia is also corrected. But after starting the cooling phase the blood glucose level was increased with decreasing body temperature. Then insulin was added but blood glucose level was remaining same. Sometimes we found a raised blood glucose level. We found that hyperglycemia and hypokalemia both increased simultaneously in cooling phase. But in rewarming phase hyperglycemia and also the requirement of potassium both were gradually decreased with increasing temperature. We had no need to correct potassium except 8 cases during rewarming phase.



Plot-1 is showing the changes of blood glucose level with respect to the change of temperature in both cooling phase and rewarming phase. Temperature is plotted horizontally in °C and blood glucose is plotted vertically in mg/dl unit.



Plot 2 is showing the requirement of potassium to correct hypokalemia during cooling phase.

Temperature is plotted horizontally and requirement of potassium in meq is plotted vertically.

In both plot we found a large difference between 37°C-35°C. This is because at that time CPB was initiated and a dilution problem was occurred.

Discussion

Hyperglycemia accompanies initiation of CPB regardless of the anaesthetic technique^(8,9,10).

Concentrations of glucose, insulin, and glucagon are higher during normothermic than hypothermic CPB^(11,12).

Markedly high serum catecholamine concentrations, which occur during CPB, inhibit the pancreatic β -cell the insulin secretory response to hyperglycemia CPB⁽¹³⁾, during moderate hypothermia, which results blood glucose concentrations raise steadily at initiation of CPB. In our study we found it from Plot-1. Despite marked hyperglycemia, insulin concentrations decline from their control values during hypothermic bypass^(14,15,16).

Counter-regulatory hormones also decline from pre-bypass concentrations during hypothermic bypass⁽¹⁷⁾.

Another effect of hypothermia is hypokalemia⁽¹⁸⁾. Hypokalemia is associated with insulin resistance and hyperglycemia⁽¹⁹⁾. With rewarming, insulin concentrations rise spontaneously to appropriate high levels; nonetheless, blood glucose remains elevated⁽²⁰⁾.

Conclusion

In cardiac surgery therapeutic hypothermia is most important and also the Hypothermia induced hyperglycemia and hypokalemia is most common. Hyperglycemia and hypokalemia both increase with decreasing temperature and both occurs simultaneously. These two factors are most important for outcome of cardiac surgery.

Reference

1. Cueni-Villoz N, Devigili A, Delodder F, Cianferoni S, Feihl F, Rossetti AO, et al.

Increased blood glucose variability during therapeutic hypothermia and outcome after cardiac arrest. *Critical care medicine*. 2011;39(10):2225-31.

2. Nielsen NM, Sunde K, Hovdenes J, et al. The Hypothermia Network. Adverse events and their relation to mortality in out-of-hospital cardiac arrest patients treated with therapeutic hypothermia. *Crit Care Med* 2011;39:57–64.
3. Hyperglycemia as an Effect of Cardiopulmonary Bypass: Intra-operative Glucose Management Samira Najmaii, MS; Daniel Redford, et al. *JECT*. 2006;38:168–173
4. Mallet ML. Pathophysiology of accidental hypothermia. *QJM*. 2002; 95:775–85.
5. Zydlewski AW, Hasbargen JA. Hypothermia-induced hypokalemia. *Mil Med*. 1998;163:719–21.
6. Plavinik FL, Rodrigues CI, Zanella MT, et al. Hypokalemia, glucose intolerance, and hyperinsulinemia during diuretic therapy. *Hypertension*. 1992;19:II26–9.
7. Lazar HL, McDonnell M, Chipkin SR, et al.: The Society of Thoracic Surgeons practice guideline series: Blood glucose management during adult cardiac surgery. *Ann Thorac Surg* 2009; 87: 663-9
8. Malatinsky J, Vigas M, Jezova D, et al. The effects of open heart surgery on growth hormone, cortisol and insulin levels in man. *Hormone levels during open heart surgery*. *Resuscitation* 1984;11(1-2):57-68.
9. Allison SP, Prowse K, Chamberlain MJ. Failure of insulin response to glucose load during operation and after myocardial infarction. *Lancet* 1967;1(7488):478-481.
10. Abe T. Influence of cardiac surgery using cardio-pulmonary bypass on metabolic regulation. *Jpn Circ J* 1974;38(1):13-21.
11. Kuntschen FR, Galletti PM, Hahn C. Glucose-insulin interactions during cardiopulmonary bypass. *Hypothermia*

- versus normothermia. *J Thorac Cardiovasc Surg* 1986;91:451-459.
12. Lehot JJ, Piriz H, Villard J, et al. Glucose homeostasis: comparison between hypothermic and normothermic cardiopulmonary bypass. *Chest* 1992;102:106-111
 13. Porte D Jr, Graber AL, Kuzuya T, et al. The effect of epinephrine on immunoreactive insulin levels in man. *J Clin Invest* 1966;45(2):228-236.
 14. Nagaoka H, Innami R, Watanabe M, et al. Preservation of pancreatic beta cell function with pulsatile cardiopulmonary bypass. *Ann Thorac Surg* 1989;48:798-802
 15. Rogers AT, Zaloga GP, Prough DS, et al. Hyperglycemia during cardiac surgery: central vs peripheral mechanisms [abstract]. *Anesth Analg* 1990;70:S328
 16. Kuntschen FR, Galletti PM, Hahn C. Glucose-insulin interactions during cardiopulmonary bypass. Hypothermia versus normothermia. *J Thorac Cardiovasc Surg* 1986;91:451-459.
 17. Ridley PD, Ratcliffe JM, Alberti KGMM, et al. The metabolic consequences of a “washed” cardiopulmonary bypass pump-priming fluid in children undergoing cardiac operations. *J Thorac Cardiovasc Surg* 1990;100:528-537.
 18. Zydlewski AW, Hasbargen JA. Hypothermia-induced hypokalemia. *Mil Med.* 1998;163:719–21.
 19. Plavinik FL, Rodrigues CI, Zanella MT, et al. Hypokalemia, glucose intolerance, and hyperinsulinemia during diuretic therapy. *Hypertension.* 1992;19:II26–9. 172 S. *NAJMAII ET AL*
 20. Rogers AT, Zaloga GP, Prough DS, et al. Hyperglycemia during cardiac surgery: central vs peripheral mechanisms [abstract]. *Anesth Analg* 1990;70:S328.