## **ORIGINAL ARTICLE**

# Impact of Elevated Glycosylated Haemoglobin on Outcomes Following CABG in Patients with Diabetes Mellitus

FAISAL ALI<sup>1</sup>, SHAHBAZ AHMAD KHILJI<sup>2</sup>, SHUJA TAHIR<sup>3</sup>

<sup>1</sup>Consultant Cardiologist, <sup>2</sup>Associate Professor of Cardiac Surgery, <sup>3</sup>Resident, Faisalabad Institute of Cardiology, Faisalabad Correspondence to: Dr. Shahbaz Ahmad Khilji, E-mail: drshahbazkhilji@gmail.com, Cell 0321-9490717

#### ABSTRACT

**Objective:** To determine the impact of HbA1c on clinical outcome after coronary artery bypass graft surgery in patients with diabetes mellitus.

Study Design: Randomized control trial.

**Place and Duration of study:** Department of Cardiac Surgery, Faisalabad Institute of cardiology, Faisalabad from 1<sup>st</sup> June 2019 to 30<sup>th</sup> June 2020.

**Methodology:** Five hundred patients who underwent isolated coronary artery bypass graft surgery were included and stratified into two groups according to HbA1c levels. Control group have patients with HbA1c levels less than 7% while Study group have patients with HbA1c levels greater than 7%. Postoperative atrial fibrillation (AF), myocardial infarction (MI), hospital stay, time on ventilator, stroke, wound infection, in-hospital and 30 day mortality were recorded and compared.

**Results:** There are statistically significant results in terms of hospital stay, time on ventilator and wound infection (p-value 0.001).

**Conclusion:** HbA1c is good predictor of in hospital morbidity, so it is necessary to have strict glycaemic control to prevent postoperative complications.

**Keywords:** Glycosylated haemoglobin, Coronary artery bypass grafting, Diabetes, Myocardial infarction, Wound infection, Stroke, Mortality

## INTRODUCTION

Diabetes mellitus is a metabolic syndrome and is known to be a major independent risk factor after optimizing other factors such as age, smoking, hypertension and hypercholesterolemia for development and progression of coronary artery disease and surgical complications.<sup>1,2</sup> Recently more than 55% of the patients with diabetes mellitus have coronary artery disease which was 19% in the past.<sup>3</sup> Of all these diabetic patients 20% of the patient underwent coronary artery bypass graft surgery.<sup>4,5</sup> Glycosylated haemoglobin (HbA1C) is found to be major predictor for assessment of glycaemic control of diabetic patients as it measures the mean glucose levels over previous 3 to 4 months<sup>6,7</sup> and there is about 4 time increase in risk of mortality in patients with HbA1C levels greater than 8.6%.<sup>8,9</sup>

There is strong association of diabetes mellitus and postoperative morbidity/mortality affecting short term and long term survival after coronary artery bypass graft surgery.<sup>11-12</sup> Moreover, patients with diabetes mellitus having coronary artery bypass graft surgery are more prone to develop surgical site infections and new onset of atrial fibrillation (AF).<sup>13,14</sup> Current guidelines of the American Diabetes Association suggests to have HbA1C levels of at least less than 7%.<sup>15</sup> The aim of this study is to assess the postoperative adverse outcome of raised HbA1C levels in diabetic patients after coronary artery bypass graft surgery.

#### MATERIALS AND METHODS

This randomized control trial was conducted at Department of Cardiac Surgery, Faisalabad Institute of Cardiology Faisalabad from 1<sup>st</sup> June 2019 to 30<sup>th</sup> June 2020. A total of 500 patients were enrolled who were scheduled for coronary artery bypass graft surgery. Preoperative investigation was carried out in the cardiac surgery ward. Two groups were formed depending upon HbA1c levels. 250 patients having HbA1c levels greater than 7% were included in study group while 250 patients with HbA1c levels of less than 7% were in control group. Postoperative atrial fibrillation (AF), myocardial infarction (MI), stroke, time on ventilator, surgical site infection, hospital stay, in hospital and 30 day mortality were recorded. Patients of both genders and age (25-70years), isolated coronary artery disease and left main disease were included. Patients with known atrial fibrillation (AF), moribund preoperative states, valvular heart diseases, previous cardiac surgery, preoperative history of stroke and preoperative multi-organ disease were excluded.

Surgery was performed using standard techniques of median sternotomy, grafts harvesting, heparinization and standard cannulation to establish standard cardiopulmonary bypass (CPB), cold blood cardioplegia, grafts anastomosis, coronary endarterectomy in diffusely diseased vessels, CPB weaning off, decannulation, drains and pacing wires and chest closure. Left internal mammary artery (LIMA) was used as the conduit to graft LAD while a saphenous vein graft was used for all other coronary vessels. Heparin 100 mg/kg was administered to keep the activated clotting time (ACT) between 200 and 400 sec. Patients were shifted to ICU and their post-operative outcomes were recorded. The data was entered and analyzed through SPSS-26.

#### RESULTS

Mean age in control group was  $55.19\pm6.273$  while in study group  $53.44\pm6.254$ . Male ratio in control group was 200 (80%) in study group there were 196 (78.4%). Patients with new onset atrial fibrillation (AF) postoperatively in control group was 25 (10%) while in study group 40 (16%) with p value (0.062). There was no statistically significance difference in incidence of mortality (p-value 0.020), postoperative myocardial infarction (p value 0.237) and stroke (p-value 0.036). Contrarily there is significantly higher incidence of hospital stay, ventilation time and wound infection (p-value 0.001) in study group as compared to control group (Table 1).

Table 1. Morbidity	/ and mortality i	in control and study group	
	y and montailly i	in control and study group	

Variable	Control group HBA1C<7	Study group HBA1C >7	P value		
Age	55.19±6.273 (35–69)	53.44±6.254 (35-69)	0.002		
Gender					
Male	200 (80%)	196 (78.4%)	0.741		
Female	50 (20%)	54 (21.6%)			
Atrial Fibrillation					
Negative	225 (90%)	210 (84%)	0.062		
Positive	25 (10%)	40 (16%)			
Myocardial Infarction					
Negative	229 (91.6%)	220 (88%)	0.237		
Positive	21 (8.4%)	30 (12%)			
Hospital stay	5.98±0.703	6.97±1.097	0.001		
(days)	(5–7)	(5–8)			
Ventilation time	8.35±1.506	13.25±4.078	0.001		
(hours)	(4–12)	(6–24)			
Stroke					
Negative	248 (99.2%)	240 (96%)	0.036		
Positive	2 (0.8%)	10 (4%)			
Wound Infection					
Negative	225 (90%)	175 (70%)	0.001		
Positive	25 (10%)	75 (30%)			
Mortality					
Negative	244 (97.6%)	232 (92.8%)	0.020		
Positive	6 (2.4%)	18 (7.2%)			

#### DISCUSSION

For over a decade studies have been suggested that patients with diabetes mellitus have increased incidence of developing postoperative morbidity, mortality and recurrence of angina.<sup>16</sup> However, there is increased risk of mortality (16%) in patients who are discovered raised glucose levels after admission in hospital.<sup>17,18</sup> HbA1c independently affects the outcome of patients after CABG. Carson and Colleagues<sup>19</sup> examined large group of patients in which 41663 patients were in diabetic group having 23% to 37% rise in in-hospital morbidity/mortality as compared to 105123 non-diabetic patients. Alserius et al<sup>20</sup> demonstrated increased surgical site infections and decreased 3 year survival in patients while Halkos et al<sup>21</sup> and Narayan et al<sup>3</sup> suggests significant rise in deep sternal wound infections having HbA1c levels greater than 6%. However in terms of hospital mortality and post op myocardial infarction we found no statistical significant results and is consistent with results by Faritous et al<sup>22</sup> and Goksedef et al<sup>23</sup> while Kuhl et al<sup>24</sup> found significant long term results in term of mortality after coronary artery bypass graft surgery. Umpierrez et al<sup>25</sup> showed no significant results in terms of postoperative AF and neurological complication in contrast to hospital stay, time on ventilator and respiratory complication. However, we still agree with the most authors that strict preoperative glycaemic control is mandatory to prevent adverse surgical outcomes and to improve quality of life after coronary artery bypass graft surgery.

#### CONCLUSION

HbA1c levels of <7% is associated with decreased postoperative morbidity or mortality and surgical complications.

#### REFERENCES

- Rollins KE, Varadhan KK, Dhatariya K, Lobo DN. Systematic review of the impact of HbA1c on outcomes following surgery in patients with diabetes mellitus. Clin Nutr 2016;35(2):308–16.
- 2. Forouhi NG, Wareham NJ. Epidemiology of diabetes. Medicine (Abingdon) 2014; 42(12):698–702.
- 3. Narayan P, Kshirsagar SN, Mandal CK, Ghorai PA, Rao YM, Das D, et al. Preoperative Glycosylated Hemoglobin: A Risk Factor for Patients Undergoing Coronary Artery Bypass. Ann Thorac Surg 2017;104(2):606–12.
- Ramadan M, Abdelgawad A, Elshemy A, Sarawy E, Emad A, Mazen M, et al. Impact of elevated glycosylated hemoglobin on hospital outcome and 1 year survival of primary isolated coronary artery bypass grafting patients. Egypt Hear J 2018;70(2):113–8.
- Greco G, Ferket BS, D'Alessandro DA, Shi W, Horvath KA, Rosen A, et al. Diabetes and the Association of Postoperative Hyperglycemia With Clinical and Economic Outcomes in Cardiac Surgery. Diabetes Care 2016;39(3):408–17.
- Saudek CD, Derr RL, Kalyani RR. Assessing Glycemia in Diabetes Using Self-monitoring Blood Glucose and Hemoglobin A1c. JAMA 2006 12;295(14):1688–97.
- Reddy P, Duggar B, Butterworth J. Blood glucose management in the patient undergoing cardiac surgery: A review. World J Cardiol 2014;6(11):1209–17.
- Tennyson C, Lee R, Attia R. Is there a role for HbA1c in predicting mortality and morbidity outcomes after coronary artery bypass graft surgery? Interact Cardiovasc Thorac Surg 2013;17(6):1000–8.
- 9. Girish G, Agarwal S, Satsangi DK, Tempe D, Dutta N, Pratap H. Glycemic control in cardiac surgery: rationale and current evidence. Ann Card Anaesth 2014;17(3):222–8.
- Holzmann MJ, Rathsman B, Eliasson B, Kuhl J, Svensson A-M, Nyström T, et al. Long-term prognosis in patients with type 1 and 2 diabetes mellitus after coronary artery bypass grafting. J Am Coll Cardiol 2015;65(16):1644–52.
- Gallagher S, Kapur A, Lovell MJ, Jones DA, Kirkwood A, Hassan S, et al. Impact of diabetes mellitus and renal insufficiency on 5-year mortality following coronary artery bypass graft surgery: a cohort study of 4869 UK patients. Eur J cardio-thoracic Surg Off J Eur Assoc Cardio-thoracic Surg 2014;45(6):1075–81.
- Koechlin L, Schurr U, Gahl B, Grapow M, Reuthebuch O, Berdajs DA. Glycosylated hemoglobin and adverse events following elective coronary bypass revascularization. Curr Res Cardiol 2018;05(02):1–10.
- Liu Y, Han J, Liu T, Yang Z, Jiang H, Wang H. The Effects of Diabetes Mellitus in Patients Undergoing Off-Pump Coronary Artery Bypass Grafting. Biomed Res Int 2016;2016.
- Ogawa S, Okawa Y, Sawada K, Goto Y, Yamamoto M, Koyama Y, et al. Continuous postoperative insulin infusion reduces deep sternal wound infection in patients with diabetes undergoing coronary artery bypass grafting using bilateral internal mammary artery grafts: a propensitymatched analysis. Eur J cardio-thoracic Surg Off J Eur Assoc Cardio-thoracic Surg 2016;49(2):420–6.
- 15. Navaratnarajah M, Rea R, Evans R, Gibson F, Antoniades C, Keiralla A, et al. Effect of glycaemic control on

complications following cardiac surgery: literature review. J Cardiothorac Surg 2018;13(1):10.

- Tsai LL, Jensen HA, Thourani VH. Intensive Glycemic Control in Cardiac Surgery. Curr Diab Rep 2016;16(4):25.
- Stoodley L, Wung S-F. Hyperglycemia after cardiac surgery: improving a quality measure. AACN Adv Crit Care 2014;25(3):221–7.
- Zheng J, Cheng J, Wang T, Zhang Q, Xiao X. Does HbA1c Level Have Clinical Implications in Diabetic Patients Undergoing Coronary Artery Bypass Grafting? A Systematic Review and Meta-Analysis. Int J Endocrinol 2017;2017.
- Carson JL, Scholz PM, Chen AY, Peterson ED, Gold J, Schneider SH. Diabetes mellitus increases short-term mortality and morbidity in patients undergoing coronary artery bypass graft surgery. J Am Coll Cardiol 2002; 40(3):418–23.
- Alserius T, Anderson RE, Hammar N, Nordqvist T, Ivert T. Elevated glycosylated haemoglobin (HbA1c) is a risk marker in coronary artery bypass surgery. Scand Cardiovasc J 2008;42(6):392–8.
- 21. Halkos ME, Puskas JD, Lattouf OM, Kilgo P, Kerendi F, Song HK, et al. Elevated preoperative hemoglobin A1c level

is predictive of adverse events after coronary artery bypass surgery. J Thorac Cardiovasc Surg 2008; 136(3):631–40.

- 22. Faritous Z, Ardeshiri M, Yazdanian F, Jalali A, Totonchi Z, Azarfarin R. Hyperglycemia or High Hemoglobin A1C: Which One is More Associated with Morbidity and Mortality after Coronary Artery Bypass Graft Surgery? Ann Thorac Cardiovasc Surg 2014;20(3):223–8.
- Göksedef D, Ömeroğlu SN, Yalvaç ESD, Bitargil M, Ipek G. Is elevated HbA1c a risk factor for infection after coronary artery bypass grafting surgery? Turkish J Thorac Cardiovasc Surg 2010;18(4):252–8.
- Kuhl J, Sartipy U, Eliasson B, Nyström T, Holzmann MJ. Relationship between preoperative hemoglobin A1c levels and long-term mortality after coronary artery bypass grafting in patients with type 2 diabetes mellitus. Int J Cardiol 2016;202:291–6.
- Umpierrez G, Cardona S, Pasquel F, Jacobs S, Peng L, Unigwe M, et al. Randomized Controlled Trial of Intensive Versus Conservative Glucose Control in Patients Undergoing Coronary Artery Bypass Graft Surgery: GLUCO-CABG Trial. Diabetes Care 2015;38(9):1665–72