

# Research Article

# Impact of Chronic Kidney Disease on Clinical Outcomes during Hospitalization and Five-Year Follow-Up after Coronary Artery Bypass Grafting

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Background. Chronic kidney disease (CKD) is often associated with multiple comorbidities including diabetes mellitus, and each has its own complications and impact after cardiac surgery including coronary revascularization. The objective of this work was to study the impact of CKD on clinical outcomes after coronary artery bypass grafting (CABG) and to compare outcomes in patients with different grades of renal functions. We retrospectively reviewed all patients who underwent CABG from January 2016 to August 2020 at our tertiary care hospital using electronic medical records. Results. The study included 410 patients with a median age of 60 years, and 28.6% of them had CKD and hospital mortality of 2.7%. About 71.4% of the patients had GFR > 60 mL/min per  $1.73 \text{ m}^2$ , 18.1% had early CKD (GFR 30-60), 2.7% had late CKD (GFR < 30), and 7.8% of them had end-stage renal disease (ESRD) requiring dialysis. The CKD group had significantly more frequent hospital mortality (p = 0.04), acute cerebrovascular stroke (p = 0.03), acute kidney injury (AKI) (p < 0.001), longer ICU stay (p = 0.002), post-ICU stay (p = 0.001), and sternotomy wound debridement (p = 0.03) compared to the non-CKD group. The frequencies of new need for dialysis were 2.4% vs. 14.9% vs. 45.5% (p < 0.001) in the patients with GFR > 60 mL/min per 1.73 m<sup>2</sup>, early CKD, and late CKD, respectively. Acute cerebral stroke (OR: 10.29, 95% CI: 1.82–58.08, and p = 0.008), new need for dialysis (OR: 25.617, 95% CI: 13.78–85.47, and p < 0.001), and emergency surgery (OR: 3.1, 95% CI: 1.82–12.37, and p = 0.036) were the independent predictors of hospital mortality after CABG. The patients with CKD had an increased risk of strokes (HR: 2.14, 95% CI: 1.20-3.81, and p = 0.01) but insignificant mortality increase (HR: 1.44, 95% CI: 0.42–4.92, and p = 0.56) during follow-up. Conclusion. The patients with CKD, especially the late grade, had worse postoperative early and late outcomes compared to non-CKD patients after CABG. Patients with dialysis-independent CKD had increased risks of needing dialysis, hospital mortality, and permanent dialysis after CABG.

#### 1. Background

Coronary artery bypass grafting (CABG) is surgical coronary revascularization in patients with advanced coronary artery disease who are unsuitable for or after unsuccessful trials of percutaneous coronary interventions (PCI) [1]. Chronic kidney disease (CKD) is often associated with multiple comorbidities including diabetes mellitus, and each has its impact after cardiac surgeries and revascularization. There is a proven association between CKD and a high prevalence of cardiovascular disorders. Moreover, patients with end-stage renal disease (ESRD) have a 10–30-fold increased mortality compared with general people [2]. Chronic renal failure was linked to cardiovascular mortality due to myocardial dysfunction, systemic hypertension, chronic anemia, dyslipidemia, hyperhomocysteinemia and hyperfibrinogenemia, nitric oxide/endothelin imbalance, oxidative stress, and chronic inflammation [2–5]. Many studies proved the association between ESRD and different worse outcomes after CABG [6–8]. We conducted this study to identify the impact of CKD on outcomes after CABG and to compare the outcomes in patients with different grades of renal functions.

#### 2. Methods

2.1. Study Design and Data Collection. We conducted this retrospective study including all adult patients who underwent CABG between 2016 and 2020 in our tertiary care hospital. We collected the variables studied from electronic hospital records. The study was approved by the Ethical Committee of King Faisal Specialist Hospital and Research Center and waived from specific consent as there were no personally identifiable data or photos. The study reference number is 2211015, and the publication number is 22350215021. The data collected included demographic and perioperative variables. The primary outcome was hospital mortality, while the secondary outcomes included cerebrovascular stroke, new need for dialysis, sternotomy wound debridement, and length of hospitalization. The studied variables included patients' characteristics, risk factors of cardiovascular and cerebrovascular diseases, prior revascularization, left ventricular ejection fraction, perioperative mechanical circulatory support, and laboratory workup including blood lactate, troponin, and glycated hemoglobin (HbA1c). The operative data collected included cardiopulmonary and aortic cross-clamping times, urgency and the approach of surgery, surgical bleeding, and need for exploration. The postoperative data collected included acute kidney injury (AKI), new need for dialysis, acute cerebrovascular stroke, mortality, length of stay, gastrointestinal bleeding, arrhythmias, sternotomy wound infection, and need for debridement. Follow-up data included mortality and acute cerebrovascular stroke.

According to the Kidney Disease: Improving Global Outcomes (KDIGO) clinical practice guidelines, CKD was defined as a glomerular filtration rate (GFR) less than 60 mL/min/1.73 m<sup>2</sup> for  $\geq$ 3 months irrespective of etiology [9, 10]. According to the degree of kidney injury, there were 5 stages of CKD. Because of imprecision to accurately estimate GFR at higher ranges, it is difficult to distinguish between stages 1 and 2. Clinically, CKD was classified according to severity into early CKD, late CKD, and ESRD [9] (Table 1). AKI was defined as an acute reduction of renal functions as monitored by urine output or GFR according to the risk, injury, failure, loss, and end-stage renal disease (RIFLE) criteria [11].

2.2. Statistical Analysis. Data were summarized using the median with an interquartile range in quantitative data and frequency (count) with relative frequency (percentage) for categorical data. The nonparametric Kruskal-Wallis and Mann-Whitney tests were used for quantitative variable comparison. The Chi-square (c2) test was performed to compare the categorical variables. Two-sided p values were considered statistically significant if < 0.05. Multivariate logistic regression was performed to get the predictors of hospital mortality and acute cerebrovascular stroke. We obtained the Kaplan-Meier survival curves using the logrank test. The Hosmer-Lemeshow test was used to detect the goodness of fit of the regression models. Variance inflation testing was performed to detect multicollinearity of the regression models. The Statistical Package for Social Sciences (SPSS) version 26 (IBM Corp., Armonk, NY, USA) was used in our study analysis.

#### 3. Results

3.1. Baseline Clinical Variables of the Studied Patients. Four hundred and ten adult patients with a median age of 60 (55–68) years were enrolled in our study and divided into 2 groups according to presence of CKD. A total of 117 (28.6%) patients had CKD, while 293 (71.4%) patients did not have CKD before CABG (Table 2).

According to the glomerular filtration rate (GFR), the studied patients were subdivided into 4 groups; 292 (71.2%) patients had GFR > 60 mL/min/1.73 m<sup>2</sup>, 74 (18.05%) patients had early CKD (GFR 30–60 mL/min/1.73 m<sup>2</sup>), 11 (2.7%) patients had late CKD (GFR less than 30 mL/min/1.73 m<sup>2</sup>), and 32 (7.8%) patients had end-stage renal disease (ESRD) requiring dialysis (Table 3).

3.2. Laboratory Variables of the Studied Patients. Preoperatively, the studied patients with CKD had significantly lower median levels of hemoglobin (p < 0.001), albumin (p < 0.001), and bilirubin (p < 0.001) but a higher median HBA1c level (p = 0.003) compared to the patients without CKD, respectively. Postoperatively, the patients with CKD had significantly higher blood lactate (p = 0.006) and troponin levels (p < 0.001) compared to the non-CKD group, respectively (Table 2).

3.3. Operative Details and Outcomes of the Studied Patients. The operative details and use of mechanical circulatory support were statistically insignificant among the studied patients with different grades of GFR. The frequencies of new need for dialysis were 2.4% vs. 14.9% vs. 45.5% (p < 0.001) in the patients with GFR > 60 mL/min per 1.73 m<sup>2</sup>, early CKD, and late CKD, respectively. The CKD group had significantly more frequent acute ischemic cerebrovascular stroke (p = 0.03), higher hospital mortality (p = 0.04), longer ICU (p = 0.002), and post-ICU stay (p = 0.001) than the non-CKD group (Tables 4 and 5).

Kaplan–Meier curves revealed that the CKD group had an increased risk of cerebrovascular strokes (HR: 2.14, 95% CI: 1.20–3.81, and p = 0.01) but insignificant mortality

Stages	Description	Classification			
		GFR $(mL/min/1.73 m^2)$	Terms		
1	Kidney damage with normal GFR	≥90	Proteinuria, albuminuria		
2	Kidney damage with a mild decrease of GFR	60-89	Proteinuria, albuminuria		
3	Moderately reduced GFR	30-59	Early renal insufficiency		
4	Severely reduced GFR	15–29	Late renal insufficiency, pre-ESRD		
5	Kidney failure	<15	ESRD		

TABLE 1: Classes of chronic kidney disease [9].

ESRD: end-stage renal disease; GFR: glomerular filtration rate.

Studied variables		All patients $(n = 410)$	Patients with CKD $(n = 117, 28.6\%)$	Patients without CKD ( <i>n</i> = 293, 71.4%)	P value
Age (years)		60 (55-68)	63 (58–71)	59 (54-66)	0.07
Gender, male (n, %)		337 (82.2)	87 (74.4)	250 (85.3)	0.09
BMI (kg/m <sup>2</sup> )		28.05 (25.8-31.8)	28.3 (25.8-32)	28 (25.9-31.5)	0.79
Smoking (n, %)		159 (38.8)	35 (29.9)	124 (42.3)	0.02
Diabetes mellitus (n, %)		338 (82.4)	111 (94.9)	227 (77.5)	< 0.001
Hyperlipidemia (n, %)		245 (59.8)	77 (65.8)	168 (57.3)	0.11
Hypertension (n, %)		351 (85.6)	110 (94)	241 (82.3)	0.002
Bronchial asthma (n, %)		44 (10.7)	25 (21.4)	19 (6.5)	< 0.001
Hypothyroidism (n, %)		42 (10.2)	11 (9.4)	31 (10.6)	0.72
Hyperthyroidism (n, %)		3 (0.7)	0 (0)	3 (1)	0.56
Preoperative AF ( <i>n</i> , %)		26 (6.3)	9 (7.7)	17 (5.8)	0.47
Autoimmune disease (n,	, %)	4 (1)	2 (1.7)	2 (0.7)	0.32
Prior stroke (n, %)		28 (6.8)	7 (6)	21 (7.2)	0.66
CAS (n, %)		37 (9)	17 (14.5)	20 (6.8)	0.014
PVD ( <i>n</i> , %)		44 (10.7)	20 (17.1)	24 (8.2)	0.009
Prior MI ( <i>n</i> , %)		257 (62.7)	87 (74.4)	170 (58)	0.002
Prior PCI (n, %)		137 (33.4)	45 (38.5)	92 (31.4)	0.17
Prior cardiac surgeries (n, %)		27 (6.6)	12 (10.3)	15 (5.1)	0.58
c .	>55%	141 (34.4)	34 (29.1)	107 (36.5)	
Duran and the INEEO/	45-55%	106 (25.9)	25 (21.4)	81 (27.6)	0.00
Preoperative LVEF%	35-45	88 (21.5)	36 (30.8)	52 (17.7)	0.08
	<35%	75 (18.3)	22 (18.8)	53 (18.1)	
Preoperative IABP (n, %	5)	15 (3.7)	2 (1.7)	13 (4.4)	0.25
Preoperative ECMO (n,	%)	2 (0.5)	0 (0)	2 (0.7)	1
Preoperative hemoglobin (g/L)		131 (115–142)	117 (101–132)	134 (121–145)	< 0.001
Preoperative platelets (10 <sup>9</sup> /L)		246 (202-308)	263 (208-316)	241 (201-305)	0.06
Preoperative serum bilirubin (µmol/L)		6.6 (4.6–11.1)	5 (3.7-8.4)	7.6 (5–12.5)	< 0.001
Preoperative INR		1 (1-1.1)	1.1 (1-1.1)	1 (1-1.1)	0.107
Preoperative fibrinogen (g/L)		3.2 (2.85-3.65)	3.26 (3.1-3.78)	3.17 (2.84-3.52)	0.051
Preoperative serum albumin (g/L)		39.8 (36.7-42.4)	37.1 (32.7-40)	41 (38-42.6)	< 0.001
HBA1c (%)		7.2 (6.4–8.2)	7.5 (6.8-8.4)	6.9 (6.3-8.1)	0.003
Peak lactate (mmol/L)		5.5 (3.8-8.9)	6.3 (3.9–10.8)	5.1 (3.7-8.4)	0.006
Peak troponin (ng/L)		634 (392–1180)	814 (490–1427)	579 (379–1090)	< 0.001

TABLE 2: Demographic and clinical variables of the patients studied.

BMI: body mass index; CAS: carotid artery stenosis; CKD: chronic kidney disease; PVD: peripheral vascular disease; MI: myocardial infarction; PCI: percutaneous coronary intervention; AF: atrial fibrillation; IABP: intra-aortic balloon pump; ECMO: extracorporeal membrane oxygenation; LVEF: left ventricle ejection fraction.

increase (HR: 1.44, 95% CI: 0.42–4.92, and p = 0.56) compared to the non-CKD group (Figures 1 and 2).

3.4. Predictors of Hospital Mortality and Cerebrovascular Stroke. Acute postoperative stroke (OR = 10.29, 95% CI: 1.823–58.08, and p = 0.008), new need for dialysis (OR = 25.617, 95% CI: 13.78–85.47, and p < 0.001), and

emergency surgery (OR = 3.1, 95% CI: 1.82–12.37, and p = 0.036) were the independent predictors of hospital mortality after CABG in our logistic regression analysis. Compared to survivors, nonsurvivors had significantly more frequent CKD, prior cardiotomies, postoperative atrial fibrillation, exploration because of mediastinal bleeding, and CABG plus valve surgeries with higher peak levels of troponin and blood lactate levels, but these variables were insignificant in the

TABLE 3: Demographic and clinical variables of the patients according to GFR.

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Studied variables		Patients with $GFR > 60$ ( $n = 292$ ,	Patients with early CKD	Patients with late CKD	Patients on dialysis $(n = 32,$	P value
Studied variable		71.4%)	(n = 74, 18.1%)	(n = 11, 2.7%)	7.8%)	1 value
Age (years)		59 (54-66)	66 (59–74)	60 (59-64)	65 (55-71)	0.057
Gender, male (n, %)		249 (85.3)	53 (71.6)	9 (81.8)	26 (81.3)	0.04
BMI (kg/m <sup>2</sup> )		28 (25.75-31.5)	28.9 (27.4-31.8)	26.6 (24.9-33.5)	27.05 (24.5-32)	0.139
Smoking (n, %)		123 (42.1)	23 (31.1)	5 (45.5)	8 (25)	0.118
Diabetes mellitus (n, %	)	227 (77.7)	72 (97.3)	11 (100)	28 (87.5)	< 0.001
Hyperlipidemia (n, %)		167 (57.2)	51 (68.9)	5 (45.5)	22 (68.8)	0.108
Hypertension (n, %)		240 (82.2)	69 (93.2)	11 (100)	31 (96.9)	0.016
Bronchial asthma (n, %	5)	19 (6.5)	17 (23)	1 (9.1)	7 (21.9)	< 0.001
Hypothyroidism (n, %)		31 (10.6)	9 (12.2)	1 (9.1)	1 (3.1)	0.55
Hyperthyroidism (n, %)		3 (1)	0 (0)	0 (0)	0 (0)	1
Preoperative AF $(n, \%)$		17 (5.8)	6 (8.1)	1 (9.1)	2 (6.3)	0.7
Autoimmune disease $(n, \%)$		2 (0.7)	2 (2.7)	0 (0)	0 (0)	0.49
Prior stroke (n, %)		20 (6.8)	5 (6.8)	2 (18.2)	1 (3.1)	0.049
CAS (n, %)		19 (6.5)	12 (16.2)	2 (18.2)	4 (12.5)	0.026
PVD (n, %)		23 (7.9)	12 (16.2)	0 (0)	9 (28.1)	0.002
Prior MI $(n, \%)$		170 (58.2)	58 (78.4)	10 (90.9)	19 (59.4)	0.004
Prior PCI (n, %)		91 (31.3)	37 (50)	1 (9.1)	8 (25)	0.003
Prior cardiac surgeries		15 (5.1)	7 (9.5)	0 (0)	5 (15.6)	0.08
Ū.	>55%	106 (36.3)	22 (29.7)	3 (27.3)	10 (31.3)	
	45-55%	81 (27.7)	17 (23)	2 (18.2)	6 (18.8)	0.22
Preoperative LVEF%	35-45	52 (17.8)	20 (27)	4 (36.4)	12 (37.5)	0.33
	<35%	53 (18.2)	15 (20.3)	3 (27.3)	4 (12.5)	
Preoperative IABP (n,	%)	13 (4.5)	2 (2.7)	0 (0)	0 (0)	0.79
Preoperative ECMO (n, %)		2 (0.7)	0 (0)	0 (0)	0 (0)	1

BMI: body mass index; CAS: carotid artery stenosis; CKD: chronic kidney disease; PVD: peripheral vascular disease; MI: myocardial infarction; PCI: percutaneous coronary intervention; AF: atrial fibrillation; IABP: intra-aortic balloon pump; ECMO: extracorporeal membrane oxygenation; LVEF: left ventricle ejection fraction.

	TABLE 4: Operative	details and	outcomes	of the	studied	patients.
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S	tudied variables	All patients	Patients with CKD	Patients without CKD	P value
TT C	Isolated CABG (n, %)	370 (90.2)	106 (90.6)	264 (90.1)	0.07
Type of surgery	CABG plus valve surgery (n, %)	40 (9.8)	11 (9.4)	29 (9.9)	0.87
Cardiopulmonary by		100 (77-126)	98 (75-128)	100 (78-125)	0.96
Aortic cross-clampin	g (minutes)	58 (43-83)	50 (42-77.5)	63 (44-84)	0.11
Annroach	Sternotomy ( <i>n</i> , %)	398 (97.1)	112 (95.7)	286 (97.6)	0.33
Approach	Minimally invasive	12 (2.9)	5 (4.3)	7 (2.4)	0.55
Urgency of surgery	Elective $(n, \%)$	397 (96.8)	114 (97.4)	283 (96.6)	0.76
Orgenicy of surgery	Emergent (n, %)	13 (3.2)	3 (2.6)	10 (3.4)	0.76
Exploration for bleed	ling ( <i>n</i> , %)	27 (6.6)	6 (5.1)	21 (7.2)	0.45
New onset AF (n, %)	)	109 (26.6)	31 (26.5)	78 (26.6)	0.9
Ventricular arrhythm	nias (n, %)	24 (5.9)	8 (6.8)	16 (5.5)	0.59
Acute kidney injury	( <i>n</i> , %)	105 (25.6)	45 (38.5)	60 (20.5)	< 0.001
New need for hemoc	lialysis (n, %)	24 (5.9)	17 (14.5)	7 (2.4)	< 0.001
Gastrointestinal bleeding (n, %)		20 (4.9)	9 (7.7)	11 (3.8)	0.09
Sternotomy wound infection ( <i>n</i> , %)		42 (10.2)	11 (9.4)	31 (10.6)	0.9
Wound debridement (n, %)		18 (4.4)	7 (5.9)	11 (3.6)	0.031
ICU days		4 (3-6)	5 (3-7)	4 (3-5)	0.002
Inotropes days		2 (1-2)	2 (1-4)	1 (1-2)	< 0.001
Ventilator days		1 (1-2)	1 (1-2)	1 (1-1)	0.019
Post-ICU ward days		5 (3-8)	6 (4–10)	4 (3-6)	0.001
Postoperative ECMC	) (n, %)	8 (2)	3 (2.6)	5 (1.7)	0.69
Postoperative IABP (n, %)		32 (7.8)	8 (6.8)	24 (8.2)	0.64
Tracheostomy (n, %)		12 (2.9)	6 (5.1)	6 (2)	0.11
Hospital mortality (n, %)		11 (2.7)	5 (4.3)	6 (2.04)	0.04
Acute ischemic stroke $(n, \%)$		30 (7.3)	12 (10.3)	18 (6.14)	0.03
Intracranial bleeding (n, %)		1 (3.2)	0 (0)	1 (0.34)	1
Stroke during follow-up ( <i>n</i> , %)		15 (3.7)	8 (6.8)	7 (2.4)	0.01
Mortality during foll	ow-up ( <i>n</i> , %)	28 (6.8)	12 (10.3)	16 (5.5)	0.5

TABLE 5: Operative details and outcomes of the studied patients according to GFR.

S	tudied variables	Patients with GFR > 60	Patients with early CKD	Patients with late CKD	Patients on dialysis	P value
Type of surgery	Isolated CABG (n, %) CABG plus valve surgery (n, %)	263 (90.1) 30 (10.3)	67 (90.5) 7 (9.5)	10 (83.3) 1 (9.1)	30 (93.8) 2 (6.3)	0.73
Cardiopulmonary by	ypass (minutes)	100 (79–125)	96 (75-128)	120.5 (87-153)	98 (73-110)	0.28
Aortic clamping (mi		63 (44.5-85)	49.5 (40-77)	62 (50-120)	48 (41-75)	0.07
A mmuo a ah	Sternotomy (n, %)	286 (97.9)	71 (95.9)	10 (90.9)	31 (96.9)	0.28
Approach	Minimally invasive	7 (2.4)	3 (4.1)	1 (9.1)	1 (3.1)	0.28
TT	Elective $(n, \%)$	282 (96.6)	71 (95.9)	11 (100)	32 (100)	0.0
Urgency of surgery	Emergent (n, %)	11 (3.8)	3 (4.1)	0 (0)	0 (0)	0.8
Exploration for blee	ding ( <i>n</i> , %)	21 (7.2)	4 (5.4)	0 (0)	2 (6.3)	0.97
New onset AF (n, %	) )	78 (26.7)	24 (32.4)	2 (18.2)	5 (15.6)	0.3
Ventricular arrhythr	nias ( <i>n</i> , %)	16 (5.5)	6 (8.1)	1 (9.1)	1 (3.1)	0.61
Acute kidney injury	( <i>n</i> , %)	59 (20.2)	36 (48.6)	10 (90.9)	0 (0)	< 0.001
New need for hemo	dialysis (n, %)	7 (2.4)	11 (14.9)	5 (45.5)	0 (0)	< 0.001
Gastrointestinal blee	ding (n, %)	11 (3.8)	8 (10.8)	1 (9.1)	0 (0)	0.044
Sternotomy wound i	infection (n, %)	38 (13)	11 (14.9)	1 (9.1)	3 (9.4)	0.9
Wound debridemen	t (n, %)	23 (7.9)	11 (14.9)	1 (9.1)	2 (6.3)	0.28
ICU days		4 (3-5)	5 (3-7)	12 (4-26)	4 (3-5)	0.002
Inotrope days		1 (1-2)	2 (1-4)	4 (2-12)	2 (1-2)	< 0.001
Ventilator days		1 (1-1)	1 (1-2)	4 (1-18)	1 (1-2)	0.001
Post-ICU ward days		4 (3-6)	6 (3-11)	7 (5–16)	5 (4-7)	0.006
Postoperative ECMC	D (n, %)	5 (1.7)	2 (2.7)	1 (9.1)	0 (0)	0.27
Postoperative IABP	( <i>n</i> , %)	24 (8.2)	6 (8.1)	1 (9.1)	1 (3.1)	0.81
Tracheostomy (n, %)	)	6 (2.1)	4 (5.4)	2 (18.2)	0 (0)	0.02
Hospital mortality (	n, %)	6 (2.1)	3 (4.1)	1 (9.1)	1 (3.1)	0.03
Hospital cerebrovaso	cular stroke (n, %)	19 (6.5)	10 (13.5)	2 (18.2)	0 (0)	0.03
Acute ischemic strol	ke ( <i>n</i> , %)	18 (6.2)	10 (13.5)	2 (18.2)	0 (0)	0.03
Acute intracerebral	bleeding (n, %)	1 (0.34)	0 (0)	0 (0)	0 (0)	1
Stroke during follow	<i>n</i> -up ( <i>n</i> , %)	7 (2.4)	3 (4.1)	1 (9.1)	4 (12.5)	0.028
Mortality during fol	low-up ( <i>n</i> , %)	16 (5.5)	10 (13.5)	0 (0)	2 (6.3)	0.104

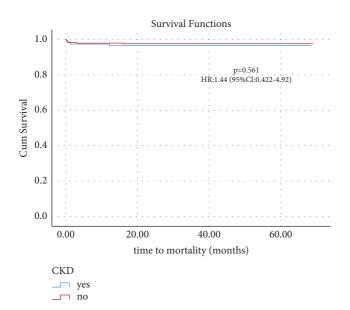


FIGURE 1: Kaplan-Meier curves of survival for 5-year follow-up.

logistic regression analysis. The regression model had a goodness of fit by the Hosmer–Lemeshow test (Pearson chi<sup>2</sup> = 57.61, p value = 1), and the mean variance inflation factor was 1.19 (Tables 6 and 7).

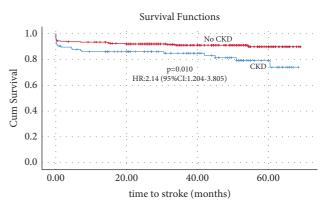


FIGURE 2: Kaplan-Meier curves of cerebrovascular stroke for 5year follow-up.

Postoperative cerebrovascular stroke was independently predicted with carotid artery stenosis (OR: 9.38, 95% CI: 3.7–23.7, and p < 0.001), atrial fibrillation (OR: 2.35, 95% CI: 1.013–5.4, and p = 0.016), CABG plus valve surgery (OR: 7.44, 95% CI: 2.75–20.15, and p < 0.001), HBA1c (OR: 1.43, 95% CI: 1.07–1.91, and p = 0.016), CKD (OR: 2.33, 95% CI: 1.06–5.17, and p = 0.036), and preoperative history of stroke (OR: 3.73, 95% CI: 1.18–11.77, and p = 0.025) in the logistic regression analysis. The regression model had a goodness of fit by the Hosmer–Lemeshow test (Pearson chi<sup>2</sup> = 38.37, p value = 0.64), and the mean variance inflation factor was 1.15 (Table 7).

TABLE 6: Univariate analysis according to hospital mortality.

Variables	Survivors ( <i>n</i> = 399, 97.3%)	Nonsurvivors ( $n = 11, 2.7\%$ )	P value
Age (years)	60 (55–68)	63 (48-80)	0.39
Diabetes mellitus (n, %)	331 (83)	7 (63.6)	0.109
Hyperlipidemia (n, %)	238 (59.6)	7 (63.6)	0.92
Hypertension (n, %)	343 (86)	8 (72.7)	0.201
Bronchial asthma (n, %)	43 (10.8)	1 (9.1)	1
CKD ( <i>n</i> , %)	112 (28.2)	5 (45.5)	0.01
PVD (n, %)	43 (10.8)	1 (9.1)	1
Preoperative AF (n, %)	24 (6)	2 (18.2)	0.15
Prior stroke (n, %)	27 (6.8)	1 (9.1)	0.54
CAS (n, %)	35 (8.8)	2 (18.2)	0.26
Prior MI (n, %)	249 (62.4)	8 (72.7)	0.75
Prior cardiac surgeries (n, %)	23 (5.8)	4 (36.4)	0.004
Isolated CABG (n, %)	365 (91.5)	5 (45.5)	(0.001
Type of surgery CABG plus valve surgery ( <i>n</i> , %)	34 (8.5)	6 (54.5)	< 0.001
Emergency surgery ( <i>n</i> , %)	10 (2.5)	3 (27.3)	0.004
Cardiopulmonary bypass (minutes)	100 (77–125)	94 (89–168)	0.33
Aortic cross-clamping (minutes)	58 (43-82)	59 (45-109)	0.36
Exploration for bleeding $(n, \%)$	24 (6)	3 (27.3)	0.03
New onset AF (n, %)	100 (25.1)	9 (81.8)	< 0.001
Acute ischemic stroke (n, %)	23 (5.8)	7 (87.5)	< 0.001
Acute kidney injury (n, %)	94 (23.6)	11 (100)	< 0.001
New need for hemodialysis (n, %)	14 (3.5)	10 (90.9)	< 0.001
Peak lactate (mmol/L)	5.4 (3.7-8.9)	15.9 (8.3-16.6)	< 0.001
Peak troponin (ng/L)	620 (391–1170)	1250 (850-2350)	0.003

TABLE 7: Logistic multivariate regression to obtain the predictors of postoperative stroke and hospital mortality.

Variables	Odds ratio	95% CI	P value
Logistic regression for hospit	tal mortal	ity	
Emergency surgery	3.1	1.82-12.37	0.036
Acute postoperative stroke	10.289	1.823-58.08	0.008
New need for dialysis	25.617	13.78-85.47	< 0.001
Prior cardiac surgery	0.369	0.016-8.46	0.53
Postoperative AF	0.403	0.017-9.382	0.57
Peak troponin level	0.9999	0.99-1.0001	0.522
Peak lactate level	0.9997	0.9992-1.00001	0.062
Logistic regression for posto	perative c	erebrovascular str	oke
Carotid artery stenosis	9.38	3.72-23.66	< 0.001
Atrial fibrillation	2.35	1.03-5.41	0.016
CABG plus valve surgery	7.44	2.75-20.15	< 0.001
HBA1c	1.43	1.071-1.91	0.016
CKD	2.33	1.06-5.17	0.036
Prior stroke	3.73	1.18-11.77	0.025

#### 4. Discussion

Our study showed that the presence of CKD before CABG was associated with increased hospital mortality, multiple morbidities, and prolonged hospitalization. The overall mortality in this cohort was 2.7%, and it ranged from 2.1% in patients with GFR >60 mL/min/1.73 m<sup>2</sup> to 9.1% in patients with late CKD. Our mortality report was similar to Cooper et al.'s study [12] that reported a hospital mortality of 2.5%, and it was 9.3% in patients with late CKD and 9% in hemodialysis dependents before CABG. Reddan et al. [13]

studied the relation between GFR and hospital mortality after revascularization and documented that each GFR decline of  $10 \text{ ml/min}/1.73 \text{ m}^2$  was associated with a 14% increased risk of mortality, but at GFR >85 mL/min/1.73 m<sup>2</sup>, the relation was attenuated. The subgroup analysis revealed a lower mortality in preoperative dialysis-dependent patients compared to the late group which may be related to the small group size. Moreover, in our cohort, 5% of the late CKD group required dialysis during hospitalization, and the new need for dialysis was a predictor of hospital mortality. Liu et al. [6] reported that preoperative hemodialysis had a three-fold increase in hospital mortality after CABG. Yamauchi et al. [7] reported significantly higher operative and 30-day mortality in hemodialysis compared to nonhemodialysis patients after CABG. Safaie et al. [8] reported a hospital mortality of 10.5% in CKD patients who underwent CABG, and there was an insignificant difference between dialysis-dependent and nondialysis-dependent patients.

Acute cerebrovascular stroke occurred in 7.3% of the patients after CABG in our study. Compared to the non-CKD group, the patients with CKD, especially those with late CKD, had an increased risk of acute stroke during hospitalization and the 5-year follow-up. Moreover, preoperative CKD was a predictor of acute postoperative stroke in the logistic regression analysis. Cooper et al. [12] reported that late CKD and dialysis dependence were associated with postoperative stroke. Liu et al. [6] reported that patients with preoperative hemodialysis dependence were 2.1 times more likely to have postoperative stroke. Preoperative CKD carried an increased risk of acute cerebrovascular stroke during the follow-up after CABG [5, 14].

Regarding other postoperative morbidities, the study results revealed a higher frequency of sternotomy wound debridement in the CKD group compared to the non-CKD group. There was a statistically insignificant difference between both groups regarding mediastinal exploration for bleeding. The CKD group had longer hospitalizations with more days of vasopressor and IABP support compared to the non-CKD group. These findings were similar to those of Cooper et al.'s report [12] which revealed prolonged hospitalization, deep sternal wound infection, and exploration for bleeding, especially in late CKD and dialysis-dependent patients. Liu et al. [6] reported that preoperative hemodialysis dependence carried a higher risk of mediastinitis but an insignificant difference for exploration because of bleeding compared to nondialysis dependence. CKD was linked to postcardiotomy 30-day and 1-year mortality [15, 16]. However, Powell et al. [17] conducted a small study and reported that dialysis dependence was associated with prolonged hospitalization without an increase in perioperative mortality and morbidities. Kan and Yang [18] reported that uremia was associated with bleeding tendency, prolonged ICU stay, and late mortality after CABG.

The patients with CKD have abnormal hemostatic profiles with abnormal risks of bleeding and thrombotic events. Ocak et al. [19] studied 10,347 patients and reported that patients with GFR  $<45 \text{ mL/min per } 1.73 \text{ m}^2$  and albuminuria had a 3.5-fold increased risk of bleeding. Platelet dysfunction was described in patients with advanced renal impairment as uremic thrombocytopathy and was related to decreased thromboxane A2 formation and von Willebrand factor (vWF) defect [20]. Renal impairment has been included in multiple bleeding risk scores such as ATRIA and HAS-BLED scores [21, 22]. The abnormal hemostatic profiles of CKD with added effects of cardiopulmonary bypass may lead to increased postoperative bleeding and thromboembolic complications. Cardiopulmonary bypass circulation leads to acute phase reaction and systemic inflammatory response with platelet activation and consumption and possible thromboembolic events [23]. The enrolled patients in this study underwent on-pump CABG, and there were insignificant differences regarding cardiopulmonary bypass and aortic clamping times in the CKD and non-CKD groups. It is still controversial to select onpump or off-pump during CABG for better postoperative outcomes. Lamy et al. [24] reported the advantages of offpump CABG regarding lower rates of AKI and respiratory and bleeding complications but similar rates of 30-day mortality, cerebrovascular stroke, myocardial infarction, and new need for dialysis compared to on-pump CABG. Li et al. [25] reported higher 30-day mortality and worse outcomes including new need for dialysis in the CKD group after off-pump CABG compared to the non-CKD group. Ueki et al. [26] reported that off-pump CABG significantly decreased the postoperative mortality in patients with CKD and decreased the need for dialysis in patients with late CKD.

In our study, deep sternal wound infection that required debridement was significantly more frequent in the CKD group compared to the non-CKD group. Our findings were consistent with Cooper et al.'s report [12]. Ishigami et al. [27] studied 9,697 patients with a 13.6-year median follow-up and reported increased hazard ratios (HRs) of infection-related hospitalizations and death with the decline of GFR. CKD is associated with a chronic low-grade inflammatory state with increased proinflammatory cytokines, resulting in increased cardiovascular risks, infections, and malignancy [28, 29].

CKD is associated with increased cardiovascular and cerebrovascular insults due to accelerated atherosclerosis, chronic inflammation, vascular calcification, electrolytes abnormalities, and anemia in addition to associated comorbidities including diabetes mellitus, dyslipidemia, and hypertension. Vascular calcification affects both the intima and media of arteries causing stiffness which increase the systolic and decrease diastolic blood pressure, resulting in decreased coronary perfusion, increased left ventricular afterload, and increased cardiovascular mortality [5, 29]. Calcification of the vascular media is the characteristic of CKD patients, described as Monckeberg's sclerosis, and results from hyperphosphatemia [30]. Impaired renal phosphate secretion results in hyperphosphatemia and increased cardiovascular mortality [5]. CKD is associated with anemia due to decreased erythropoietin secretion and results in decreased oxygen supply, increasing cardiovascular risks [31].

Another challenging point in management of patients with CKD is postoperative analgesia. Postoperative sternotomy pain requires strong analgesia to control the sympathetic stimulation and cardiovascular side effects. However, impaired pharmacokinetics of opiates in the presence of CKD can result in side effects including prolonged sedation, seizures, delirium, nausea, vomiting, and respiratory complications [32]. Appropriate postcardiac surgery analgesia is critical, especially in CKD, to achieve a balance of pain control with patient satisfaction and avoid drug accumulation with side effects [33, 34].

Finally, with the advances of medical care, the patients presented to CABG have advanced age and multiple comorbidities including CKD and dialysis dependence. Meticulous perioperative management including hemodynamics optimization, appropriate fluid and electrolyte management, and appropriate analgesia should be addressed to high-risk patients with CKD to minimize perioperative morbidities and mortality.

#### **5.** Conclusion

The patients with CKD, especially the late grade, had worse postoperative early and late outcomes compared to non-CKD patients after CABG. Patients with dialysisindependent CKD had increased risks of needing dialysis hospital mortality and permanent dialysis after CABG.

#### 6. Limitations

This work was a single-center retrospective study with a relatively small sample size. It missed perioperative blood sugar control and the amount of postoperative bleeding and blood transfusions. We used preoperative glycated hemoglobin as a marker of blood sugar control in the studied patients. We could not get data on blood glucose variability and control during hospitalization. We studied the impact of renal impairment on outcome, but the study missed some key details about dialysis modalities and efficiency and the laboratory finding of parathormone and electrolytes. The enrolled patients in this study underwent on-pump CABG as this is the standard approach in our hospital, and we do not know the validity of our result for off-pump CABG. We studied the surgical wound infection without other infections during hospitalization.

#### Abbreviations

CKD:	Chronic kidney disease
ESRD:	End-stage renal disease
AKI:	Acute kidney injury
ATRIA:	Anticoagulation and risk factors in atrial
	fibrillation
CABG:	Coronary artery bypass grafting
CPB:	Cardiopulmonary bypass
CAS:	Carotid artery stenosis
OR:	Odds ratio
HAS-	Hypertension, abnormal liver/renal profile,
BLED:	cerebrovascular stroke, bleeding history, labile
	INR, elderly, drugs/alcohol concomitantly
IABP:	Intra-aortic balloon pump
vWF:	von Willebrand factor
ECMO:	Extracorporeal membrane oxygenation.

## Data Availability

The data used to support the findings of this study are available on request from the corresponding author.

#### Disclosure

The abstract was orally presented at the Cardiovascular Research Technology Congress, CRT 2022, which was held in Washington DC, USA, and it was published in the Cardiovascular Revascularization Medicine journal. "Laimoud M, Maghirang M, Al-Mutlaq S, Althibait S, Ghamry R, Alanazi B, Alanazi M, AlHalees Z. The Impact of Chronic Kidney Disease on Outcomes after Coronary Artery Bypass Graft Surgery. Cardiovascular Revascularization Medicine. 2022 Jul 1; 40:37-8."

#### **Conflicts of Interest**

The authors declare that they have no conflicts of interest.

#### **Authors' Contributions**

ML participated in study design, data collection, statistical analysis, and drafting the manuscript. ZA and MNA participated in study design, data interpretation, and revision of the manuscript. MM, SAA, RQ, RG, BMA ZB, SMA, and MA participated in data collection and interpretation. All the authors have read and approved the final manuscript for publication.

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