1	Primary causes and direct medical cost of heart failure among adults admitted with acute
2	decompensated heart failure in a public tertiary hospital, Kenya
3	
4	Authors: Victor M. Wauye* ¹ , Chrispine O. Oduor ¹ , Felix A. Barasa ² , G. Titus Ngeno ³
5	
6 7 8 9 10	Affiliations ¹ Department of Internal Medicine, Moi University, Eldoret, Kenya ² Department of Cardiology, Moi Teaching and Referral Hospital, Eldoret, Kenya ³ Division of Cardiology, Department of Medicine, Duke University, Durham, USA
11	*Corresponding author:
12 13 14 15 16 17	Dr. Victor Mwaka Wauye, P. O. Box 85 – 50405, Butula, Kenya, Tel: +254 729-771-653 Email: <u>vmwauye@gmail.com</u>
18	
19	
20	
21	
22	
23	
24	
25	
26	
27	
28	
29	

NOTE: This preprint reports new research that has not been certified by peer review and should not be used to guide clinical practice.

Abstract 31

32 Heart failure (HF) is a major contributor of cardiovascular morbidity and mortality globally. 33 Despite its adverse impact on health outcomes in low- and middle-income countries such as 34 Kenva, data on the direct medical cost of HF hospitalization is limited.

35

36 This was a prospective study conducted at Moi Teaching and Referral Hospital. Patients with 37 HF were identified by sequential medical chart abstraction. Primary causes were extracted from 38 echocardiogram reports and adjudicated by a cardiologist. Direct medical cost of 39 hospitalization was derived using activity-based costing, micro-costing method, and payers' 40 system perspective. Drivers of overall cost were explored using linear regression models.

41

42 142 participants were consecutively recruited from September to November 2022. 51.4% were 43 females, and the overall mean age was 54 (SD 20). The leading primary cause was cor 44 pulmonale (CP), 28.9%; then dilated cardiomyopathy (DCM), 26.1%; rheumatic heart disease 45 (RHD), 19.7%; hypertensive heart disease (HHD), 16.9%; ischaemic heart disease (IHD), 46 6.3%; and pericardial disease (PD), 2.1%. Overall direct cost of HF hospitalization was Kshs. 47 11,470.94 (SD 8,289.57) per patient per day, with the mean length of hospital stay of 10.1 (SD 7.1). RHD incurred the highest costs, Kshs. 15,299.08 (SD 13,196.89) per patient per day, then 48 49 IHD, Kshs. 12,966.47 (SD 6656.49), and DCM, Kshs.12,268.08 (SD 7,816.12). Cost of 50 medications was the leading driver, $\beta = 0.56 (0.55 - 0.56)$, followed by inpatient fees, $\beta = 0.27$ 51 (0.27 - 0.28) and laboratory investigations, $\beta = 0.19 (0.18 - 0.19)$.

52

53 Cor pulmonale, CM, RHD and HHD were the major causes of HF. The overall direct medical 54 cost of hospitalization was extremely expensive compared with the average monthly household 55 income per capita in Kenya. Widespread insurance cover is therefore recommended to cushion 56 families against such catastrophic health expenditures beside public health measures aimed at 57 addressing primary causes of HF.

58

59 Keywords: Primary causes, heart failure, direct medical cost, hospitalization, Moi Teaching 60 and Referral Hospital

- 61 62 63 64 65 66 67
- 68

3

69 **1 Introduction**

70 Cardiovascular diseases (CVDs) are the most common cause of global morbidity and mortality, 71 accounting for more than 17.9 million deaths annually in the recent years. More than 75% of 72 these deaths occur in the middle and low income countries (1). Heart failure (HF), is one of the 73 most common primary CVD diagnosis among hospitalized medical patients in Sub-Saharan Africa (SSA), Kenya included (2,3). The main causes of HF in SSA have been traditionally 74 75 described as hypertensive heart disease (HHD), dilated cardiomyopathy (DCM) and rheumatic 76 heart disease (RHD) and less commonly, other causes of HF such as ischemic and congenital 77 cardiomyopathies (4,5).

Treatment of HF confers significant financial burden both to the healthcare system as well the affected families, being largely driven by direct medical costs of hospitalization (6,7). Direct medical costs accounted for about 60% of the total costs in a review by Cook et al. (6). There is however limited data published on the cost of treatment of HF in Africa. Ogah et al. reported a total cost of USD 2,128 per patient per year in Nigeria, using a broader societal costing approach (8).

To the best of our knowledge, there is no published study on medical cost of hospitalization of 84 85 HF in Kenya despite the efforts towards the achievement of universal health coverage. 86 According to the 2023 Kenya Demographic Health Survey, healthcare expenditure in Kenya is 87 mainly paid for out of pocket, with only 26.5% of the Kenyans having any form of health 88 insurance (9). Of these, only 24% are covered by the government sponsored national health 89 insurance fund (NHIF), while the rest are covered by private insurance. At tertiary government 90 health facilities, about 35% of the hospitalized patients have NHIF cover (10). Furthermore, it 91 is notable that about 83% of workforce in Kenya is comprised of informal sector (11). A study 92 assessing the uptake of NHIF among informal workers in Western Kenya showed that only

93 12% had health insurance coverage, with the majority experiencing catastrophic health
94 expenditure (12), thus implying that patients with HF would be at a high risk of financial
95 catastrophe.

In our study, we sought to further develop the understanding of the cost of HF treatment in a
low-middle-income economy, by describing the primary causes of HF at Moi Teaching and
Referral Hospital (MTRH), Kenya, and determining the direct medical cost of hospitalization
among patients hospitalized with acute decompensated HF (ADHF).

100 2 Methodology

101 2.1 Study design and population

102 This was a prospective study conducted at the Moi Teaching and Referral Hospital (MTRH). 103 Kenya, over a period of three months from 01 September to 30 November 2022. MTRH is a national referral hospital located in Western Kenva, with a catchment area of about 24 million 104 105 people. It has two wings: public and private wing, with the former being where the general 106 public seek treatment. All those who were hospitalized to the public wing medical wards and 107 cardiac care unit (CCU) with the diagnosis of ADHF based on the Modified Framingham 108 clinical criteria for HF and were aged 18 years and above were included in the study. Those 109 who were less than 18 years old, readmitted during the study period and discharged to or from 110 the MTRH private wing were excluded from the study.

111 2.2 Case definition for the primary causes of HF

112 Comprehensive clinical data including clinical symptoms and signs, past medical history, and 113 results of laboratory as well as imaging investigations were gathered and collated with the two-114 dimensional doppler echocardiography (ECHO) and 12-lead electrocardiography (ECG)

5

115	findings. Both electrocardiograms and echocardiograms were performed by well-trained
116	ECHO/ECG technicians at MTRH ECG/ECHO centre, according to the American Society of
117	Cardiology guidelines (13). The findings were summarized by the principal investigator, and
118	further confirmed by the consultant cardiologist. The most likely primary cause of HF was then
119	established using a predetermined case definition criteria (Table 1), which was guided by the
120	European Society of Cardiology guidelines (14). A similar criteria was previously applied in
121	the Heart of Soweto study in South Africa (2,15), and the Sub-Saharan Africa Survey of Heart
122	Failure which recruited participants from 9 African countries, Kenya included (4). The criteria

123 was however tweaked to fit our set up with limited diagnostic resources.

Cause of HF	Case Definition
Hypertensive Heart Disease	Documented history or new diagnosis of HTN plus ECHO report of LVH or ECG features of LVH: LAD or Positive Cornell Voltage ECG criteria for LVH (Sum of R wave in aVL and S wave in V3 > to 2.8mV in males and 2.0mV in females) or Sokolow-Lyon criteria: sum of S in V1 or V2 and R in V5 or V6 >35mm.
Rheumatic Heart Disease	Pathological MR with at least 2 of: AMVL thickening, chordal thickening, restricted leaflet motion or excessive leaflet tip motion during systole or
	MS mean gradient ≥4mmHg or
	Pathological AR with at least 2 of: AV focal thickening, coaptation defect, restricted leaflet motion or prolapse.
Ischemic Heart Disease	Regional wall abnormality plus/minus pathological Q waves defined as any Q waves $>0.02s$ in V2-V3 or QS complex in V2-V3, or Q waves $\ge 0.03s$ and $\ge 1mm$ deep or QS complex in other leads in at least 2 contiguous leads.
Dilated Cardiomyopathy	Dilated ventricular chambers with reduced LVEF or dilated ventricular chambers with ECG features of LVH in the absence of HHD and IHD.
Cor Pulmonale	ECHO: Pulmonary HTN of >35mmHg (in the absence of valvular heart disease as RHD and LVEF \leq 40%), TAPSE <1.7cm, right ventricular enlargement, and dilated non-collapsing IVC, and/or
	ECG: Right Axis Deviation, R/S amplitude ratio>1 in V1, R/S amplitude ratio<1 in V6, $S_1Q_3T_3$, or Right Bundle Branch Block and/or
	Documented chronic lung disease.
Pericardial Disease	Pericardial echo free fluid plus/minus any features of cardiac tamponade and indices of cardiac contusion in the absence of any identifiable causes above.

124

Table 1:Case definition for the primary causes of heart failure

125 Abbreviations: AR, Aortic valve; AMVL, anterior mitral valve leaflet; AV, aortic valve; ECG, electrocardiogram; ECHO,

126 echocardiogram; HHD, hypertensive heart disease; HTN, hypertension; IHD, ischaemic heart disease; IVC, inferior vena cava;

- 127 LAD, left axis deviation; LVEF, left ventricular ejection fraction; LVH, left ventricular hypertrophy; MR, mitral regurgitation;
- 128 TAPSE, transannular plane systolic excursion.

6

Direct medical costing approaches 129 2.3

130 Costing was done using payers' perspective(16), with a prevalence and prospective methods 131 (17). Detailed data regarding direct medical costs were obtained from the MTRH finance 132 department in printed forms. The data included a breakdown of all the costs that each patient 133 incurred during hospitalization, from the date of admission to the point of discharge (the date 134 on which the attending physician decided that the patient be discharged, or the patient died). 135 Micro-costing technique was used to detail the cost components, which included costs related 136 to inpatient charges (nutrition, admission, physiotherapy, bed and nursing charges), 137 medications, laboratory investigations, imaging, oxygen therapy, and other utilities 138 (consumables such as syringes, gauzes, needles, nasal prongs among others) (18,19). Further, 139 direct costs included both out of pocket (OOP) and national health insurance (NHIF) payments. 140 Notably, being a public health facility with human resource remuneration largely covered by the government, charges related to remuneration were not included. Moreover, indirect costs 141 such as transport and productivity losses were not included. 142

143

Data collection 2.4

144 Data were collected using interviewer administered questionnaires, and included information 145 on socio-demographic characteristics, past medical history, New York Heart Association 146 assessment (NYHA), vital signs such as blood pressure and heart rate, ancillary laboratory 147 investigations such as haemoglobin, electrolytes, urea and creatinine, lipid profile, thyroid function tests, liver function tests and international normalizing ratio, chest Xray, select ECG 148 149 and ECHO findings. Comorbidities were physician-diagnosed and documented in the files, 150 largely based on the patients' past medical history and laboratory investigations.

151

Statistical analysis 2.5 153

154 Data were analysed using RStudio statistical software version R.4.2.2. Continuous variables 155 were reported using mean and median with respective standard deviations (SD) and 156 interquartile range (IQR). Primary causes of HF were reported using proportions and 157 percentages, with respective 95% confidence intervals (CI). Costs were reported using mean 158 (SD) as cost (in Kenya shillings) per patient per day; calculated by dividing the total cost during 159 hospitalization by the mean length of hospital stay (LOS) by the total number of participants, 160 N, and compared between the CCU and the general ward. Linear regression model with 161 standardized β -coefficients was used to determine how each cost component drove the overall 162 cost. Due to inability to delineate direct medical cost of comorbidities, the mean costs were all-163 inclusive. However, the effect of each comorbidity on the overall cost was explored using generalized linear regression model with gamma distribution and log-link function. 164 165 Regressions involved both bivariate and multivariate analyses. Comparisons of mean costs 166 between general ward and CCU was done using Mann-Whitney and Kruskal-Wallis non-167 parametric tests of association. Further, sub-analysis of the overall costs was done by NYHA, 168 grade of ejection fraction, and in-hospital outcome (discharged alive or dead). Costs were further converted to American dollars (USD) per the Central Bank of Kenya foreign exchange 169 170 rates during the study period, for which 1USD = Kshs 122.6129(20).

Ethics statement 171 2.6

A formal written and signed informed consent was obtained from the participants before 172 173 enrolment into the study. The informed consent described the details of the study, detailing 174 both the benefits and potential harms. Participants' data were decoded anonymously for confidentiality. Further, the study was approved by the Moi University Institutional Research 175 176 Ethics Committee (approval number 003738 and reference number IREC/2020/187).

8

177 **3. Results**

- 178 A total of 142 participants were included in the study, after 178 cases were screened for
- 179 eligibility (Fig 1). The mean age was 54 (SD 17) years, with the median age of 60 (IQR 37.3,
- 180 70). 51.4% were females. Further, 73.9% had unskilled labour, and only 52.8% had NHIF
- 181 cover. The mean length of hospital stay was 10.1 (SD 7.1) days. Table 2 shows further socio-
- 182 demographic, laboratory, comorbidities, medications and ECG/ECHO findings.
- 183

[Fig 1 Recruitment schema of the participants]

Baseline characteristics		(n, %)
Gender	Male	69 (48.59)
	Female	73 (51.41)
Age (years) (mean (SD) M	Iedian (IQR) ¹	54.0 (20.0) 60 (37.3, 70)
Marital Status	Married	92 (64.8)
	Single	25 (17.6)
	Divorced/Separated	7 (4.93)
	Widowed	18 (12.7)
Level of Education	Primary	85 (59.9)
	Secondary	36 (25.4)
	Tertiary	21 (14.8)
Occupation	Unskilled	105 (73.9)
	Students	11 (7.7)
	Skilled	15 (10.6)
	Professional	8 (5.6)
	Retired	3 (2.1)
Mode of Payment	NHIF	75 (52.8)
	OOP	67 (47.2)
LOS (mean (SD) median (10.1 (7.1) 8 (5, 12.8)
Ward	General ward	73 (51.4)
	CCU	69 (48.6)
Discharge Status	Alive	115 (81.0)
	Dead	27 (19.0)
HF History	De novo HF	68 (47.9)
	ADCHF	74 (52.11)
	HIV	8 (5.6)
	HTN	45 (31.7)
	DM	17 (12.0)
	Anaemia	37 (26.1)
	Chronic Lung Disease	31 (21.8)
Comorbidities	Liver Disease	47 (33.1)
	Renal Insufficiency	97 (68.3)
	Thyroid Dysfunction	26 (18.3)
	Dyslipidaemia	87 (61.3)
	Cancers	6 (4.2)
	II	2 (1.4)
NYHA Stage	III	87 (61.2)
	IV	53 (37.3)
	HR (bpm) ¹	105 (29) 102 (85, 121)
	Sinus	110 (77.5)
	Atrial fibrillation	32 (22.5)
ECG/ECHO summary	Reduced	70 (49.3)
	Mildly reduced	18 (12.7)
	Preserved	54 (38.0)
	RVSP ¹	60 (20) 56 (45, 75)
	TAPSE ¹	1.5 (0.5) 1.3 (1, 1.9)
	Diuretics	131 (92.3)
	MRAs	83 (58.5)
	ACEIs/ARBs	74 (52.1)
	SGLT2 Inhibitors	35 (24.7)
Medications	Vasopressors	32 (22.5)
	B -Blockers	30 (21.1)
	Digoxin	29 (20.4)
	Amiodarone	27 (19.0)

Table 2: Baseline characteristics of the participants 209

> ¹Mean (SD) |Median (IQR). Abbreviations: ACEIs, angiotensin converting enzyme inhibitors; ADCHF, acute decompensation of chronic heart failure; ARBs, angiotensin receptor blockers; CCB, calcium channel blockers; CCU, Cardiac care unit; DM, diabetes mellitus; HIV, human immunodeficiency virus; HR, Heart rate; MRAs, mineralocorticoids; NHIF, National health insurance fund; OOP, Out of pocket payment; RVSP, right ventricular systolic pressure; SGLT2, sodium glucose transporter-2; TAPSE, Transannular plane systolic excursion.

10

Primary causes of HF 3.1 210

- 211 The most common primary cause of HF was cor pulmonale (CP), accounting for 28.9% (95%
- CI 21.1% 37.9%) of the cases, followed by dilated cardiomyopathy (CM), 26.1% (95% CI 212
- 213 18.3% - 35.0%), rheumatic heart disease (RHD), 19.7% (95% CI 12% - 28.7%) and
- 214 hypertensive heart disease (HHD), 16.9% (9.2% - 25.9%) (Table 3).

Primary Cause of Heart Failure	n (%)	95% CI
Cor Pulmonale	41 (28.9)	21.1 - 37.9
Dilated cardiomyopathy	37 (26.1)	18.3 - 35.0
Rheumatic Heart Disease	28 (19.7)	12.0 - 28.7
Hypertensive Heart Disease	24 (16.9)	9.2 - 25.9
Ischaemic Heart Disease	9 (6.3)	0 - 15.3
Pericardial Disease	3 (2.1)	0 - 11.1
Total, N	142	

215 Table 3: Primary causes of heart failure

Direct medical cost of HF hospitalization 216 3.2

217 The mean direct medical cost of HF hospitalization was Kshs. 11,470.94 (SD 8,289.57) per 218 patient per day, with the cost in the CCU being as twice as the cost in the general ward (table 219 4). Of the cost components, cost of medications was the leading driver of the overall cost, with 220 standardized β -coefficient of 0.56 (95% CI 0.55 – 0.56), followed by the cost of inpatient fees and laboratory investigations respectively (Fig 2). Stratified by the primary cause of HF, RHD 221 222 incurred the highest mean direct medical cost, Kshs. 15,299.08 (SD 13,196.89) per patient per 223 day (Fig 3). Overall, direct medical cost of HF hospitalization was increasingly higher among 224 those with NYHA stages III and IV, HFrEF and HFmrEF, comorbid renal and liver diseases, 225 and those who died during hospitalization (S Table 1, S Table 2, S Table 3 and S Table 4, and 226 S Fig 1).

227

228

11

Item Line	Amount (Kshs) (Mean, SD)			
	Overall	General ward	CCU	P-value
Inpatient Fees	18091.72 (20861.32)	7919.78 (4856.96)	28853.34 (25471.93)	< 0.0001
Lab Investigations	22733.39 (14135.5)	20953.01 (11167.76)	24619.99 (16590.92)	0.5124
Imaging	5139.44 (4928.71)	6434.25 (5814.52)	3769.57 (3296.85)	0.0004
Medications	36694.92 (42403.40)	18833.61 (19719.78)	55591.67 (51105.17)	< 0.0001
Oxygen	10044.44 (9743.60)	7960.53 (6530.91)	12373.53 (12076.73)	0.167
Other Utilities	11427.48 (12925.69)	7505.31 (4583.15)	15577.02 (17035.38)	0.0011
Cost/patient (Kshs)	99065.65 (76339.71)	65775.94 (36922.10)	134285.20 (90510.95)	< 0.0001
Total cost/patient (USD)	807.44 (622.21)	536.11 (300.94)	1094.50 (737.71)	
LOS (days)	10.1 (7.1)	10.0 (6.6)	10.3 (7.7)	
Cost/patient/day (Kshs)	11470.94 (8289.57)	7687.57 (3783.92)	15473.63 (9782.79)	
Cost/patient/day (USD)	93.49 (67.56)	62.66 (30.84)	126.12 (79.74)	
able 4: Direct medical cost of he	eart failure hospitalization			
	1			

231 232

230

233

234

[Fig 3. Stratified direct medical costs by the primary cause of HF]

[Fig 2. Drivers of the overall direct medical cost of HF hospitalization]

235

4. Discussion 236

237 This study found that CP was the most common primary cause of HF among patients 238 hospitalized with ADHF in MTRH, accounting for 28.9%. Other leading causes of HF were 239 HHD, DCM and RHD, consistent with several previously published studies in Sub-Saharan Africa (SSA) (4,15,21–23). Our findings highlight the importance of CP, as well as the role of 240 chronic lung diseases (CLDs), alongside other drivers of HF morbidity in SSA (24-26). 241 242 Notably, 46.3% of those who had CP in this study had chronic obstructive lung disease, 24.4% 243 had indeterminate pulmonary hypertension, 17.1% had post-tuberculosis lung fibrosis and 244 bronchiectasis, 7.3% had lung masses and 4.9% had pulmonary embolism. Although we did 245 not collect data on indoor air pollution, it is possible that indoor air pollution contributed to the high prevalence of chronic lung diseases hence CP. Recent studies conducted in Western 246 247 Kenva reported significantly high levels of indoor air pollution due to biomass fuel use (27,28). 248 and this was strongly associated with different cardiac abnormalities, differentially affecting women and those with low socioeconomic status (29,30). It is notable that more women than 249

12

250 men had HF in this study; 51.4% vs 48.6%, had CP; 56.1% vs 43.9%, and 73.9% overall had
251 unskilled labour, reflecting low socio-economic status.

252 This study found that the mean direct medical cost of HF hospitalization at MTRH was Kshs. 253 11,470.94 (SD 8,289.57) {USD 93.49 (67.56)} per patient per day. Kwok et al., in the US 254 reported a cost of USD 11,845 (SD 22,710)(31), while Zaour et al., in Canada reported USD 10,123 per patient per hospitalization (32). Other studies from observational cohorts in SSA 255 256 have estimated the of societal cost impact of HF hospitalization to be USD 2,128 per patient 257 per year (8). Comparatively, these costs are much higher than what we found, highlighting the 258 importance of healthcare service delivery context in cost modelling. It is possible that higher 259 healthcare resource utilization in the developed countries, alongside different costs of health system inputs, variable health insurance coverage, and heterogeneity in the study designs 260 261 resulted in the differences observed (33–36). In addition, our cost estimates represent costs in 262 a government-subsidized health care system wherein, indirect costs as well as the human 263 resource costs are not charged directly to patients.

The 2022 Economic Survey by the Kenya National Bureau of Statistics showed that the gross 264 265 national income per capita in 2021 was Kshs. 20,122.23 (USD 164.01) per month(37). We 266 were not able to get data on the average monthly income of the participants in this study, but it 267 is notable that 83.5% of the participants did not complete high school, and 73.9% were unskilled labourers. Therefore, our finding of Kshs 11,470.94 (USD 93.49) indirect costs per 268 269 patient per day would be extremely high against the Kenvan gross national income per capita. 270 posing a high risk of financial catastrophe to the affected family members and public healthcare 271 system. Although higher proportion of patients (52.8%) had national health insurance cover 272 compared to the national average (26.5%), the higher coverage rate likely represents increased 273 awareness of the importance of insurance cover and utilization among patients with chronic 274 disease seeking tertiary facility care. This notwithstanding, there remains a need to intensify

13

advocacy for and implementation of widespread national health insurance coverage among patients with HF towards the achievement of universal health coverage (UHC) and sustainable development goal (SDG) 3.8)(38,39). Further, there is need to intensify primary management and prevention strategies among patients with HF to reduce chances of hospitalization hence the high direct medical costs of hospitalization.

To the best of our knowledge, this was the first study in the region to stratify direct medical 280 281 cost of HF hospitalization by aetiology. This study found that treatment of HF caused by RHD 282 was the most expensive, costing Kshs 15,299.08 per patient per day. Sub-analysis showed that 283 of the patients with RHD, 67.9% were admitted in the cardiac intensive care unit (CCU). RHD 284 associated HF accounted for 27.5% of the total CCU admissions. Further, 28.6% of those admitted with RHD associated HF died, accounting for 29.6% of all those who died (S Table 285 286 5). Notably, direct medical cost of those who died was twice the cost of those who were 287 discharged alive. Finally, RHD accounted for 37.8% of all the patients with atrial fibrillation 288 in this study. It is possible that additional treatment modalities such as cardioversion and 289 anticoagulation among patients with RHD and atrial fibrillation (40) increased the cost of 290 treatment. It is further arguable that most patients with RHD were critically ill at presentation, 291 were admitted to the CCU for intensive care, and a significant number died, driving the cost 292 upwards. Further studies are needed to characterize costs attributable to RHD, as well as 293 community-based prevention measures noting that RHD is a preventable disease.

The cost of medications was the leading driver of the overall cost as has been evidenced in prior literature (41,42). The overall cost also expectedly increased with advanced NYHA (S Table 1), EF (S Table 2), comorbidities particularly (S Table S, and S Fig 1), and discharge status; overall cost of those who died was as twice as that of those who were discharged alive (S Table 3). Additionally, overall treatment cost among those who were hospitalised in CCU was twice as high as treatment cost in the general wards (Table 4). This would be because of

14

increased resource utilisation following intensive care for critically ill patients with advancedstages of HF (43–49).

302 This study however had the following limitations. First, participants were recruited from a 303 public healthcare facility, hence the findings regarding cost may not be applicable to private 304 healthcare facilities. Second, the utility of the findings on cost in designing healthcare budgets 305 should be taken with caution because this study was conducted over a short period of time, and 306 it may be difficult to predict how the cost may change over a longer time horizon. However, 307 the costs reported per patient per day are intuitive and reflects daily consumption of healthcare resources during hospitalization with ADHF. Finally, payer's perspective used for costing in 308 309 this study could be narrow and may not reflect the total cost of HF treatment during 310 hospitalization.

311 **5. Conclusion**

312 In conclusion, cor pulmonale was the most common primary cause of HF, followed by CM, 313 RHD and HHD. The mean direct medical cost of hospitalization was Kshs. 11,470.94 per 314 patient per day. Weighted against Kenyan monthly gross national income per capita, this was 315 extremely expensive. Stratified by primary cause of HF, RHD incurred the highest direct 316 medical costs. Therefore, policies and community-oriented interventions should be put in place 317 for early recognition and treatment of CP, among other primary causes of HF in view of the 318 ongoing epidemiological transition in SSA since these causes are completely preventable. 319 Further studies are also recommended exploring causal relationship between wider causes of 320 HF in SSA, particularly with the current public health challenges of global concern such as 321 climate change and air pollution. Additionally, widespread and improved health insurance 322 coverage should be implemented to protect the affected families and the healthcare system 323 from catastrophic spending resulting from high direct medical cost of HF hospitalization amid

the efforts towards the achievement of universal health coverage in Kenya. Finally, more

324

- 325 extensive cost of illness studies such as societal perspective are recommended, detailing lost 326 productivity among patients hospitalized with HF. Acknowledgements 327 328 The authors thank all the staff and study participants for their co-operation during the study. 329 References 330 1. 331 World Health Organization. Cardiovascular diseases (CVDs) [Internet]. Cardiovascular 332 Diseases (CVDS). 2017. https://www.who.int/news-room/fact-sheets/detail/cardio-333 vascular-diseases-(cvds) 334 2. Sliwa K, Wilkinson D, Hansen C, Ntyintyane L, Tibazarwa K, Becker A, et al. Spectrum 335 of heart disease and risk factors in a black urban population in South Africa (the Heart of Soweto Study): a cohort study. Lancet. 2008;371(9616):915-22. 336 Kimani K, Namukwaya E, Grant L, Murray SA. What is known about heart failure in 337 3. 338 sub-Saharan Africa: A scoping review of the English literature. Vol. 7, BMJ Supportive 339 and Palliative Care. BMJ Publishing Group; 2017. p. 122-7. 340 4. Damasceno A, Mayosi BM, Sani M, Ogah OS, Mondo C, Ojji D, et al. The Causes, Treatment, and Outcome of Acute Heart Failure in 1006 Africans From 9 Countries. 341 342 Arch Intern Med. 2012 Oct 8;172(18):1386. 5. Bloomfield GS, Barasa FA, Doll JA, Velazquez EJ. Heart Failure in Sub-Saharan Africa. 343 Curr Cardiol Rev. 2013 May 17;9(2):157-73. 344 Cook C, Cole G, Asaria P, Jabbour R, Francis DP. The annual global economic burden 345 6.
- 346
 of heart failure. Int J Cardiol. 2013 Feb 15;171(3):1–9.

 247
 7
 6
- 347 7. Stevens B, Pezzullo L, Verdian L, Tomlinson J, George A, Bacal F. The economic
 348 burden of heart conditions in Brazil. Arq Bras Cardiol. 2018;111(1):29–36.
- Ogah OS, Stewart S, Onwujekwe OE, Falase AO, Adebayo SO, Olunuga T, et al.
 Economic burden of heart failure: Investigating outpatient and inpatient costs in
 Abeokuta, Southwest Nigeria. PLoS One. 2014 Nov 21;9(11):1–17.

16

352 9. KNBS and ICF. Kenya Demographic and Health Survey 2022.Key Indicators Report..

- Nairobi, Kenya, and Rockville, Maryland, USA; 2023. https://dhsprogram.com/pubs/pdf/PR143/PR143.pdf
- Binanay CA, Akwanalo CO, Aruasa W, Barasa FA, Corey GR, Crowe S, et al. Building
 Sustainable Capacity for Cardiovascular Care at a Public Hospital in Western Kenya. J
 Am Coll Cardiol. 2015;66(22):2550–60.
- 11. Chuma J, Okungu V. Viewing the Kenyan health system through an equity lens:
 Implications for universal coverage. Int J Equity Health. 2011 May 26;10(1):1–14.
- Maritim B, Koon AD, Kimaina A, Lagat C, Riungu E, Laktabai J, et al. "It is like an
 umbrella covering you, yet it does not protect you from the rain": a mixed methods study
 of insurance affordability, coverage, and financial protection in rural western Kenya. Int
 J Equity Health. 2023 Dec 1;22(1):1–13.
- Rudski LG, Lai WW, Afilalo J, Hua L, Handschumacher MD, Chandrasekaran K, et al.
 Guidelines for the Echocardiographic Assessment of the Right Heart in Adults: A Report
 from the American Society of Echocardiography: Endorsed by the European
 Association of Echocardiography, a registered branch of the European Society of
 Cardiology, and the Canadian Society of Echocardiography. J Am Soc Echocardiogr.
 2010 Jul 1;23(7):685–713.
- McDonagh TA, Metra M, Adamo M, Gardner RS, Baumbach A, Böhm M, et al. 2021
 ESC Guidelines for the diagnosis and treatment of acute and chronic heart failure. Eur
 Heart J. 2021;42(36):3599–726.
- 373 15. Stewart S, Wilkinson D, Hansen C, Vaghela V, Mvungi R, McMurray J, et al.
 374 Predominance of Heart Failure in the Heart of Soweto Study Cohort. Circulation. 2008
 375 Dec 2;118(23):2360–7.
- 376 16. Choi H-J, Lee E-W. Methodology of Estimating Socioeconomic Burden of Disease
 377 Using National Health Insurance (NHI) Data. In: Reddy S, Tavares AI, editors. Rijeka:
 378 IntechOpen; 2019. p. Ch. 8.
- 379 17. Costa N, Derumeaux H, Rapp T, Garnault V, Ferlicoq L, Gillette S, et al.
 380 Methodological considerations in cost of illness studies on Alzheimer disease. Health
 381 Econ Rev. 2012 Sep 11;2(1):1–12.

17

Turner HC, Archer RA, Downey LE, Isaranuwatchai W, Chalkidou K, Jit M, et al. An
Introduction to the Main Types of Economic Evaluations Used for Informing Priority
Setting and Resource Allocation in Healthcare: Key Features, Uses, and Limitations.
Front Public Heal. 2021 Aug 25;9:722927.

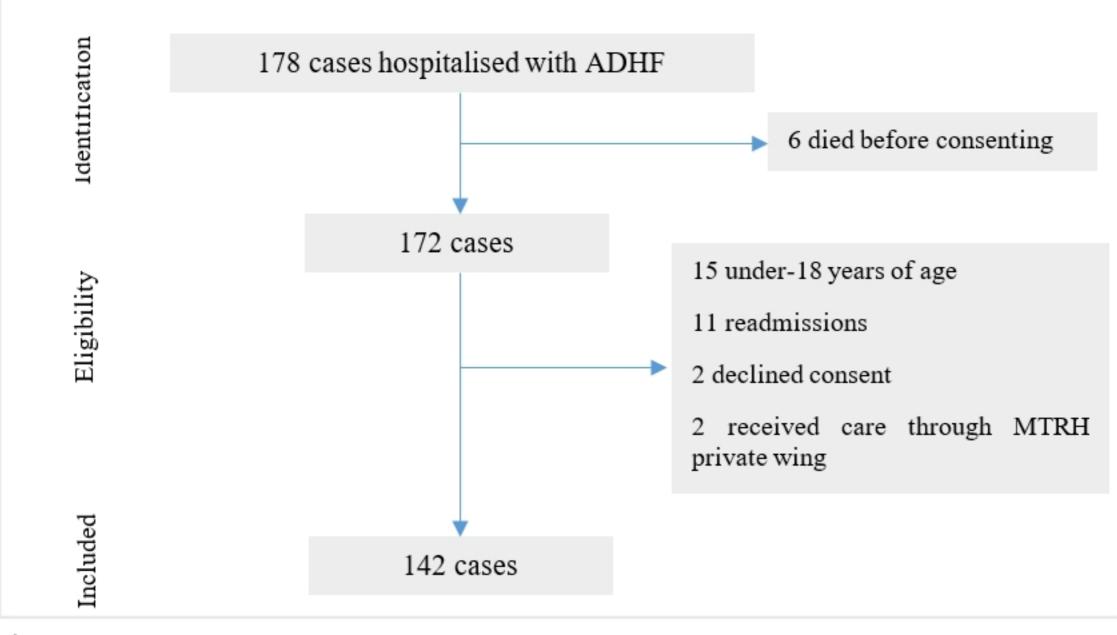
- Jo C. Cost-of-illness studies: concepts, scopes, and methods. Clin Mol Hepatol.
 2014;20(4):327–37.
- 20. Central Bank of Kenya. Foreign Exchange Rates | CBK. 2022. https://www.centralbank.go.ke/rates/forex-exchange-rates/
- 390 21. Karaye KM, Dokainish H, ElSayed A, Mondo C, Damasceno A, Sliwa K, et al. Clinical
 391 Profiles and Outcomes of Heart Failure in Five African Countries: Results from INTER392 CHF Study. Glob Heart. 2021 Jul 30;16(1):50.
- 22. Eberly LA, Rusingiza E, Park PH, Ngoga G, Dusabeyezu S, Mutabazi F, et al.
 Understanding the Etiology of Heart Failure Among the Rural Poor in Sub-Saharan
 Africa: A 10-Year Experience From District Hospitals in Rwanda. J Card Fail.
 2018;24(12):849–53.
- Makubi A, Hage C, Sartipy U, Lwakatare J, Janabi M, Kisenge P, et al. Heart failure in
 Tanzania and Sweden: Comparative characterization and prognosis in the Tanzania
 Heart Failure (TaHeF) study and the Swedish Heart Failure Registry (SwedeHF). Int J
 Cardiol. 2016 Oct 10;220:750.
- 401 24. Onyango EM, Onyango BM. The Rise of Noncommunicable Diseases in Kenya: An
 402 Examination of the Time Trends and Contribution of the Changes in Diet and Physical
 403 Inactivity. J Epidemiol Glob Health. 2018;8(1–2):1.
- 40425.Xie M, Liu X, Cao X, Guo M, Li X. Trends in prevalence and incidence of chronic405respiratory diseases from 1990 to 2017. Respir Res. 2020 Feb 11;21(1):1–13.
- 406 26. Binegdie AB, Meme H, El Sony A, Haile T, Osman R, Miheso B, et al. Chronic
 407 respiratory disease in adult outpatients in three African countries: a cross-sectional
 408 study. Int J Tuberc Lung Dis. 2022 Jan 1;26(1):18.
- 27. Dida GO, Lutta PO, Abuom PO, Mestrovic T, Anyona DN. Factors predisposing women
 and children to indoor air pollution in rural villages, Western Kenya. Arch Public Heal.
 2022 Dec 1;80(1):1–13.

412 413	28.	Musyoka D, Muindi K. A Descriptive Assessment of Household Air Pollution in Rural Kitchens in Kenya. Atmos 2022, Vol 13, Page 2115. 2022 Dec 16;13(12):2115.
414 415 416	29.	Lagat DK, Delong AK, Wellenius GA, Carter EJ, Bloomfield GS, Velazquez EJ, et al. Factors associated with isolated right heart failure in women a pilot study from western kenya. Glob Heart. 2014;9(2):249–54.
417 418 419	30.	Agarwal A, Kirwa K, Eliot MN, Alenezi F, Menya D, Mitter SS, et al. Household air pollution is associated with altered cardiac function among women in Kenya. Am J Respir Crit Care Med. 2018 Apr 1;197(7):958–61.
420 421 422	31.	Kwok CS, Abramov D, Parwani P, Ghosh RK, Kittleson M, Ahmad FZ, et al. Cost of inpatient heart failure care and 30-day readmissions in the United States. Int J Cardiol. 2021;329:115–22.
423 424	32.	Zaour N, Barbeau M, Liu N, Borrelli R, Fischer A. The cost of hospitalization and length of stay for chronic heart failure cases in Canada. Can J Cardiol. 2015 Oct 1;31(10):S273.
425 426	33.	Garcia-Escribano M, Mogues T, Juarros P. Patterns and Drivers of Health Spending Efficiency. IMF Work Pap. 2022;2022(048):1.
427 428 429	34.	Urbich M, Globe G, Pantiri K, Heisen M, Bennison C, Wirtz HS, et al. A Systematic Review of Medical Costs Associated with Heart Failure in the USA (2014–2020). PharmacoEconomics 2020 3811. 2020 Aug 19;38(11):1219–36.
430 431 432	35.	Osenenko KM, Kuti E, Deighton AM, Pimple P, Szabo SM. Burden of hospitalization for heart failure in the United States: a systematic literature review. J Manag Care Spec Pharm. 2022;28(2):157–67.
433 434	36.	Tarricone R. Cost-of-illness analysis. What room in health economics? Health Policy (New York). 2006;77(1):51–63.
435	37.	Kenya National Bureau of Statistics. Economic Survey 2022. 2022.
436 437 438	38.	WHO. Targets of Sustainable Development Goal 3. 2023. https://www.who.int/europe/- about-us/our-work/sustainable-development-goals/targets-of-sustainable-development- goal-3
439 440	39.	WHO. Universal health coverage (UHC). 2022. https://www.who.int/news-room/fact-sheets/detail/universal-health-coverage-(uhc)

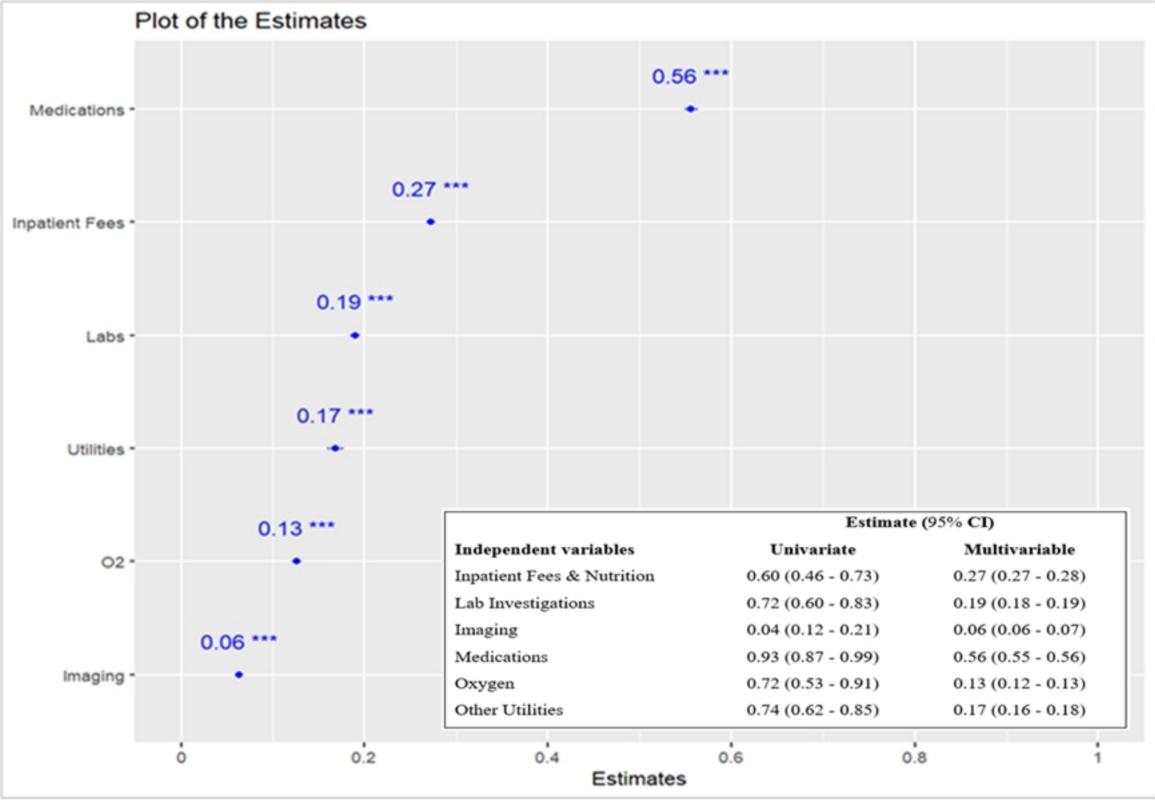
441 442	40.	Katritsis DG, Gersh BJ, John Camm A. Anticoagulation in Atrial Fibrillation – Current Concepts. Arrhythmia Electrophysiol Rev. 2015 Sep 1;4(2):100.
443 444	41.	Wang S, Petzold M, Cao J, Zhang Y, Wang W. Direct Medical Costs of Hospitalizations for Cardiovascular Diseases in Shanghai, China. Med (United States). 2015;94(20):1–8.
445 446 447	42.	Ong SC, Low JZ, Yew WY, Yen CH, Abdul Kader MASK, Liew HB, et al. Cost analysis of chronic heart failure management in Malaysia: A multi-centred retrospective study. Front Cardiovasc Med. 2022 Nov 2;9.
448 449 450	43.	Alghamdi A, Algarni E, Balkhi B, Altowaijri A, Alhossan A. Healthcare Expenditures Associated with Heart Failure in Saudi Arabia: A Cost of Illness Study. Healthcare. 2021 Aug 1;9(8).
451 452 453	44.	Delgado JF, Oliva J, Llano M, Pascual-Figal D, Grillo JJ, Comín-Colet J, et al. Health care and nonhealth care costs in the treatment of patients with symptomatic chronic heart failure in Spain. Rev Esp Cardiol (Engl Ed). 2014 Aug;67(8):643–50.
454 455 456	45.	Givertz MM, Yang M, Hess GP, Zhao B, Rai A, Butler J. Resource utilization and costs among patients with heart failure with reduced ejection fraction following a worsening heart failure event. ESC Hear Fail. 2021 Jun 1;8(3):1915–23.
457 458 459	46.	Stafylas P, Farmakis D, Kourlaba G, Giamouzis G, Tsarouhas K, Maniadakis N, et al. The heart failure pandemic: The clinical and economic burden in Greece. Int J Cardiol. 2017 Jan 15;227:923–9.
460 461 462 463	47.	Rihova B, Paremca J, Jarkovsky J, Miklik R, Sulcova A, Littnerova S, et al. Cardiology department hospitalization costs in patients with acute heart failure vary according to the etiology of the acute heart failure: Data from the AHEAD Core registry 2005–2009. Cor Vasa. 2013 Feb 1;55(1):e7–14.
464 465 466	48.	Chamberlain AM, Sauver JLS, Gerber Y, Manemann SM, Boyd CM, Dunlay SM, et al. Multimorbidity in heart failure: A community perspective. Am J Med [Internet]. 2015;128(1):38–45.
467 468 469	49.	Mehrotra S, Sharma TM, Bahl A. Impact of Comorbidities in Heart Failure – Prevalence, Effect on Functional Status, and Outcome in Indian Population: A Single-Center Experience. J Clin Prev Cardiol. 2019;(8):166–72.
470		

Supporting information 471

- 472 S Table 1. Direct medical cost by NYHA
- 473 S Table 2. Direct medical cost by EF
- 474 S Table 3. Direct medical costs by outcome status
- 475 S Table 4. Effect of comorbidities on the overall cost
- 476 S Table 5. Baseline Characteristics per primary cause of HF
- 477 S Fig 1. Coefficient plot of the effect of the comorbidities on the overall cost

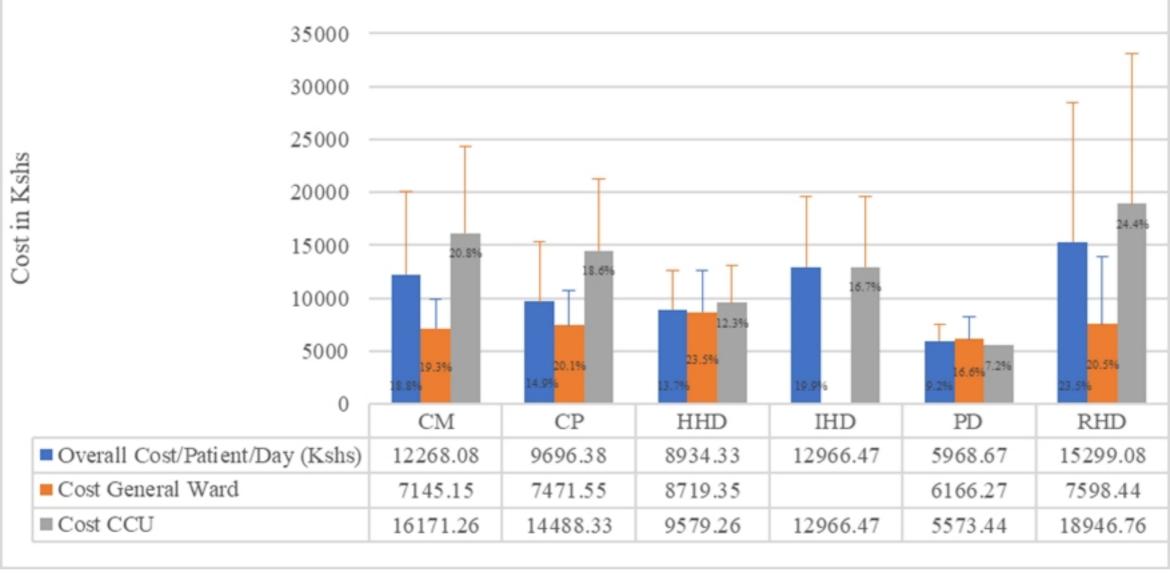


Figure



Figure

Stratified Direct Medical Cost of HF Hospitalization



Figure