Check for updates

OPEN ACCESS

EDITED BY John Yfantopoulos, National and Kapodistrian University of Athens, Greece

REVIEWED BY Abdene Weya Kaso, Dilla University, Ethiopia Malaisamy Muniyandi, National Institute of Research in Tuberculosis (ICMR), India

*CORRESPONDENCE Girma Tekle Gebremariam girma.tekle@aau.edu.et

[†]These authors share first authorship

SPECIALTY SECTION This article was submitted to Health Economics, a section of the journal Frontiers in Public Health

RECEIVED 18 June 2022 ACCEPTED 13 September 2022 PUBLISHED 10 October 2022

CITATION

Tito K, Gebremariam GT, Beyene K, Sander B and Gebretekle GB (2022) Health-related quality of life and treatment satisfaction of patients with cardiovascular disease in Ethiopia. *Front. Public Health* 10:972378. doi: 10.3389/fpubh.2022.972378

COPYRIGHT

© 2022 Tito, Gebremariam, Beyene, Sander and Gebretekle. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.

Health-related quality of life and treatment satisfaction of patients with cardiovascular disease in Ethiopia

Kebron Tito^{1†}, Girma Tekle Gebremariam ^{1*†}, Kebede Beyene ², Beate Sander ^{3,4,5,6} and Gebremedhin Beedemariam Gebretekle ^{3,4}

¹School of Pharmacy, Addis Ababa University, Addis Ababa, Ethiopia, ²Department of Pharmaceutical and Administrative Sciences, University of Health Sciences and Pharmacy in St. Louis, St. Louis, MO, United States, ³Institute of Health Policy, Management, and Evaluation, University of Toronto, Toronto, ON, Canada, ⁴Toronto Health Economics and Technology Assessment (THETA) Collaborative, University Health Network, Toronto, ON, Canada, ⁵Institute for Clinical Evaluative Sciences (ICES), Toronto, ON, Canada, ⁶Public Health Ontario, Toronto, ON, Canada

Purpose: Cardiovascular disease is the most prevalent health problem associated with poorer health-related quality of life (HRQoL). We aimed to assess HRQoL and treatment satisfaction of cardiovascular disease patients in Ethiopia.

Methods: A cross-sectional survey was conducted among adults attending the outpatient cardiac clinic at Tikur Anbessa Specialized Hospital from July to September 2021. Patients were recruited consecutively during follow-up visits. Treatment Satisfaction Questionnaire for Medication and European Quality of life questionnaires were used to evaluate treatment satisfaction and HRQoL, respectively. Kruskal-Wallis and Mann-Whitney *U*-tests were used to compare utility weights between patient subgroups. Utility values were computed using disutility weights of the Ethiopian general population derived using a hybrid regression model. Tobit regression modeling was used to explore factors associated with poor HRQoL. Statistical significance was determined at p < 0.05.

Results: A total of 357 patients participated in the study with a mean age of 49.3 ± 17.8 years. The most frequently reported health problems were pain/discomfort (75.4%), followed by mobility (73.4%). The median (interquartile range) European Quality questionnaires five dimensions with five levels utility (EQ-5D-5L) and European Quality of life Visual Analog Scale scores were 0.84 (0.55–0.92) and 70.0 (50.0–85.0), respectively. The highest and lowest mean (standard deviation) treatment satisfaction scores were for the convenience and safety satisfaction dimensions: 87.7 (17.9) and 53.1 (33.5), respectively. Unemployment, older age, previous hospital admission, non-adherence to lifestyle modification, and presence of three or more cardiovascular disease factors were significantly negatively associated with HRQoL.

Conclusions: Overall, the study found that cardiovascular disease had a profound negative effect on HRQoL and patient treatment satisfaction. We suggest that interventions to enhance HRQoL and treatment

satisfactions should focus on modifiable associated factors including lifestyle changes and controlling disease progression.

KEYWORDS

cardiovascular disease, EQ-5D-5L, Ethiopia, HRQoL, treatment satisfaction

Introduction

Cardiovascular disease (CVD) is a group of disorders affecting the heart and/or blood vessels (1, 2). It is the leading cause of disability and premature death globally and has significant health and economic burden on patients, families, and healthcare systems (1). According to 2022 American Heart Association report, an estimated 19 million CVD-related deaths occurred globally, with three-quarters of these deaths occurring in low- and middle-income countries (LMICs) (3). In Ethiopia, the reported prevalence of CVD ranges from 1 to 24% with a pooled prevalence of 5%, and significantly contributes to increasing healthcare costs (4, 5). CVD and its treatments pose significant burden on patients' health-related quality of life (HRQoL) and can affect their ability to function (6, 7).

HRQoL is a patient-reported outcome measure that evaluates how a patient's health status is affected by disease, complications, and therapy (8, 9). It can provide information about a person's overall health state that is linked with physical, social, and mental health as well as their respective impact on health status of the patient that is affected by individual's beliefs, perceptions, experiences, and expectations (10, 11). Thus, HRQoL is an important metric to assess the impact of health interventions and is a useful input for valuing health outcomes in economic evaluations. Likewise, treatment satisfaction is a patient-reported outcome measure that considers the patient's perspective of medical treatment and healthcare services (12) and can be used to predict adherence and persistence of medication use over time (13, 14).

To improve patient outcomes, patient-reported problems must be addressed alongside cardiovascular pharmacotherapy. Previous studies showed that a more active involvement of patient's in disease management can substantially enhance HRQoL (15, 16). In LMICs, lower socioeconomic status, low access and high cost of medications could negatively affect HRQoL and treatment satisfaction. On the other hand, adequate physical activity and younger age were associated with better HRQoL (17). Similarly, patients with controlled blood pressure have better overall satisfaction with their medications and HRQoL. Patients who had no adverse events and experienced less anxiety and depression had significantly higher overall treatment satisfaction and better HRQoL (18–21).

Several generic and disease-specific tools have been developed for measuring HRQoL of patients with CVD (22-25). The European Quality of life five-dimension-5 level scale questionnaire (EQ-5D-5L) is a widely used generic, preferencebased multi-attribute utility instrument to quantify the global burden of disease in clinical practice and aids health technology assessment in many jurisdictions (26, 27). The tool measures the health state of the patients by generating a single summary utility value, which reflects how good or bad a health state is based on the preferences of the general population of a country (28). The utility weight generated is an important measure to quantify the impact of disease, measure potential benefits and harms of different interventions; and inform resource allocation (29). However, data on HRQoL and treatment satisfaction of patients with CVD in Ethiopia remains limited. This study aimed to assess HRQoL and its associated factors in a sample of patients with CVD at a tertiary care teaching hospital in Ethiopia. The findings could aid in the development of tailored interventions that improve patient health outcomes.

Methods

Study design and setting

An interview-based cross-sectional survey was conducted from July to September 2021 among patients attending an outpatient cardiac clinic at Tikur Anbessa Specialized Hospital in Addis Ababa, Ethiopia. The hospital is the largest tertiary care teaching hospital in the country with over 800 beds and serves over half a million patients per year. Of these, approximately 5,500 patients attend the cardiac clinic annually.

Patient recruitment and data collection procedure

All patients visiting the cardiac clinic comprised the source population while those who fulfilled the eligibility criteria during the data collection period comprised the study population. The sample size was estimated based on a single population proportion formula (30), considering a *Z*-value of 1.96 with

Abbreviations: CVD, Cardiovascular disease; EQ-5D-5L, EuroQoL life questionnaries five dimensions with five levels; EQ-VAS, EuroQoL Visual Analog scale; HRQoL, Health-Related Quality of Life; IQR, interquartile range; LMICs, Low-middle and low-income countries; MGL, Morisky Green-Levine questionnaire; SD, Standard deviation; TSQM, Treatment Satisfaction Questionnaire for Medication.

a 95% level of confidence and a 5% margin of error. The prevalence (P) for sample size estimation was based on a previous study conducted in Ethiopia using the World Health Organization Quality of Life questionnaire (WHOQoL-BREF) instrument (15), in which 45.6% of patients with heart failure rated their overall HRQoL as poor. We applied the finite population correction factor formula (31) for the source population of 5,500 CVD patients, and recruited a total of 360 eligible patients using a consecutive sampling method. Patients were eligible if they were 18 years of age or older and diagnosed with CVD. We excluded patients under the age of 18 years, unwilling to participate, and those who had severe cognitive/mental problems as confirmed by the treating physician.

Three clinical pharmacists conducted face-to-face interviews with the patients during their follow-up clinic visit. All data collectors were trained to ensure uniformity and reduce interobserver bias in data collection. The purpose and procedure of the study were explained to all study participants before data collection. Written informed consent was obtained from all study participants, and personal identifiers were not collected. During the data collection process; data collectors clarified queries raised by patients. Information on sociodemographic characteristics such as sex, age, marital status, occupation, level of education, alcohol habit, smoking status, and lifestylerelated factors were collected through face-to-face interviews. Chart review was undertaken to gather information on clinical characteristics of the patients including duration of CVD, complications, non-cardiac comorbidities, and the number of medications taken. We collected patients' treatment satisfaction and HRQoL using the Treatment Satisfaction Questionnaire for Medication (TSQM) and EQ-5D-5L instruments, respectively. Patients' information regarding medication adherence was collected using Morisky Green-Levine (MGL) questionnaire.

Instruments

EQ-5D-5L

EQ-5D-5L is a generic instrument that consists of five dimensions with five levels, of which four are physical dimensions and one is psychological dimension (28, 32). Validated Amharic version of the EQ-5D-5L was used for the interview. The instrument has two parts: the EQ-5D-5L descriptive system short questionnaires and European Quality Visual Analog Scale (EQ-VAS). In the first part of the instrument, patients select the statement most reflective of their health state in the descriptive system that has five dimensions (mobility, self-care, usual activities, pain/discomfort, and anxiety/depression). Under each dimension, there are five levels (no problem, slight problem, moderate problem, severe problem, and extreme problem) which represent the severity of problems. We asked participants to choose one level that reflects their health state on the interview date for each of the five dimensions. The EQ-VAS is used for subjective assessment of one's current health state from the patient's perspective. Using this scale, each interviewed patient self-rated his/her health status on a vertical scale that ranges from zero (the worst health one can imagine) to 100 (the best health one can imagine).

TSQM

TSQM is a 14-item instrument used to evaluate treatment satisfaction with medication over the past 2–3 weeks, or since the patient's last medication use (33). The TSQM has been translated to Amharic language, the national language used in Ethiopia, using forward and backward translation technique. It has four dimensions: effectiveness (three items), safety (five items), convenience (three items), and global satisfaction (three items). Item scores are summed to give four dimension scores, which are in turn transformed to a scale of zero (extremely dissatisfied) to 100 (extremely satisfied) (34). The TSMQ have also been utilized in previous studies in Ethiopia to assess satisfaction with treatment.

MGL

We used the four-item Morisky Green-Levine (MGL) questionnaire to assess medication adherence. The tool results in a score ranging from 0 to 4. A dichotomous definition of adherence based on MGL used with "0" indicating perfect adherence and "1" indicating some level of non-adherence (35).

Statistical analysis

Descriptive statistics such as mean and percentage are used to summarize patients' demographic and clinical characteristics. Differences in percentages of reported problems with patients' characteristics were tested using the χ^2 -test. The level of severity for each dimension of the EQ-5D-5L descriptive system was reported using 5 levels to determine the proportion of health problems reported in different subgroups (1 = no)problem to 5 = extreme problem). The EQ-5D-5L utility score were computed using disutility coefficients (decrement in utility) obtained from the Ethiopian general population using a hybrid regression model (36). For each of the five health dimension [mobility (mo), self-care (sc), usual activities (ua), pain/discomfort (pd), and anxiety/depression (ad)]; we set four incremental dummies (a total of 20 parameters) by considering "1 = no problem" as a reference and representing the utility decrement when moving from lower level to the next higher level (e.g., moving from sc1 to sc2 resulted in a decrease in utility of 0.0235125). The utility value for CVD patients was computed using Equation (1). As the EQ-5D-5L utility and

EQ-VAS scores were non-normally distributed (Shapiro-Wilk *test*, p < 0.05), we present median [interquartile range (IQR)] scores. We applied Kruskal-Wallis and Mann-Whitney U tests to determine the differences in the EQ-5D-5L utility scores and EQ-VAS among subgroups of patients. A multivariate Tobit regression model was used to examine the predictors of HRQoL. Some covariates, i.e., smoking status and alcohol consumption habits, were not entered into regression models due to a smaller number of responses. We censored the utility score at one and the EQ-VAS score at 100. In addition, Pearson's correlation coefficient was used to assess correlation between EQ-VAS scores, EQ-5D-5L utility index, and TSQM dimension scores. To distinguish higher treatment satisfaction from lower satisfaction in each dimension; we dichotomized the mean TSQM score into >75 (higher treatment satisfaction) and <75 (lower treatment satisfaction) (37). Statistical analyses were performed using STATA Version 14. All statistical tests were performed using a level of significance of *p*-value < 0.05.

 $\begin{array}{l} \textbf{Utility value} = 1-(mo2 * 0.0337341 + mo3 * 0.0644715 + \\ mo4 * 0.2276493 + mo5 * 0.3598963) + (sc2 * 0.0235125 + \\ sc3 * 0.0394815 + sc4 * 0.1419238 + sc5 * 0.2223553) + (ua2 \\ * 0.0323013 + ua3 * 0.0482993 + ua4 * 0.1573934 + ua5 * \\ 0.2721253) + (pd2 * 0.0360808 + pd3 * 0.0515949 + pd4 * \\ 0.2703189 + pd5 * 0.4063984) + (ad2 * 0.0258862 + ad3 * \\ 0.0848133 + ad4 * 0.2987388 + ad5 * 0.4577938) (1) \end{array}$

Results

Sociodemographic characteristics

A total of 360 patients were approached to take part in the study of which 357 (99.2%) consented and completed the study questionnaires. The majority of the study participants were females (65.8%), married (57.4%), unemployed (79.6%), and 18–60 years old (69.7%; mean age = 49.3 \pm 17.8 years). One in five participants were illiterate, approximately three-fourths of participants were living in Addis Ababa and most participants (60.2%) had health insurance coverage. Detailed sociodemographic patient characteristics are presented in Table 1.

Clinical characteristics of patients with CVD

The majority (72.8%) of participants had CVD for 5 years or more with a mean duration of 9.57 ± 8.59 years. Three hundred (84%) patients were adherent to lifestyle modifications; most (60.3%) used dietary lifestyle modification (Table 2). Nearly all patients (98.3%) were non-smokers, and 332 (93%) had no alcohol consumption habits within 1 month prior to the interview date. One in five (20.4%) patients had a history of TABLE 1 Sociodemographic characteristics of patients with cardiovascular disease.

Variables	N (%)
Age category (in years)	
18-60	249 (69.7)
>60	108 (30.3)
Sex	
Male	122 (34.2)
Female	235 (65.8)
Marital status	
Single	84 (23.5)
Married	205 (57.4)
Divorced	26 (7.3)
Widow	42 (11.8)
Educational status	
Unable to read and write	72 (20.2)
Primary education	105 (29.4)
Secondary education	113 (31.7)
Post-secondary education	67 (18.8)
Employment status	
Employed	73 (20.4)
Unemployed	284 (79.6)
Residence	
Addis Ababa	263 (73.7)
Out of Addis Ababa	94 (26.3)
Health insurance usage	
Yes	215 (60.2)
No	142 (39.8)

hospital admission in the past year, and the majority (87.7%) of patients had more than two CVDs. Almost half of the patients (45.1%) had non-cardiac comorbidities. Most patients (65.3%) were taking more than five medications, and 204 (57.1%) were adherent to their medications. Unavailability of medications (37.5%) and forgetfulness (29%) were the main reasons to medication non-adherence.

Distribution of EQ-5D-5L descriptive dimensions

Patients' self-reported health status for the five dimensions of EQ-5D-5L is presented in Figure 1. The most frequent health problems reported were "pain/discomfort" (75.4%, all levels) followed by "mobility" (73.4%, all levels), while the least health problem was "self-care" (23%, all levels). Thirtyseven (10.4%) participants reported "no problem" or a "perfect health state" (11,111) in the EQ-5D-5L descriptive dimension, whereas only 30 (8.4%) reported the "best health state" (100) in the EQ-VAS. The frequencies of any problem reported TABLE 2 Clinical characteristics of patients with cardiovascular disease.

Duration of CVD (in years)	
<5 years	97 (27.2)
5–10 years	125 (35.0)
>10 years	135 (37.8)
Use of lifestyle modifications	
Yes	300 (84.0)
No	57 (16.0)
Type of lifestyle modifications used	
Diet	181 (60.3)
Exercise	17 (5.7)
Both	102 (34.0)
Smoking habit	
Yes	5 (1.4)
No	351 (98.3)
Alcohol use	
Yes	25 (7.0)
No	332 (93.0)
Previous hospital admission	
Yes	73 (20.4)
No	284 (79.6)
Non-cardiac comorbidities	
Yes	161 (45.1)
No	196 (54.9)
Number of CVD	
<3 conditions	313 (87.7)
\geq 3 conditions	44 (12.3)
Number of medications	
<5	124 (34.7)
≥5	233 (65.3)
Adherent to medications	
Yes	204 (57.1)
No	153 (42.9)
Reasons for non-adherence	
Forgetfulness	49 (32.1)
Unavailability	57 (37.3)
Fear of side effects	7 (4.5)
Others*	40 (26.1)

 * Lack of transportation, felt well, no belief in effectiveness of medications, got worsen, not afford the cost.

CVD, Cardiovascular disease.

with all dimensions were significantly different across patient characteristics such as sex, educational status, age, noncardiac comorbidities, types of lifestyle modifications, previous hospitalization history, medication adherence status as well as the different dimensions of treatment satisfaction. Patients with higher treatment satisfaction (\geq 75) had lower problem levels in EQ-5D-5L dimensions (mobility, activities, self-care, pain/discomfort, and anxiety/depression) Supplementary File 1.

EQ-5D-5L index value and EQ-VAS score

The median (IQR) EQ-5D-5L index and EQ-VAS scores were 0.84 (0.55-0.92) and 70 (50-85), respectively. Whereas, the mean (SD) EQ-5D-5L index and EQ-VAS scores were 0.69 (0.34) and 68.7 (21.6), respectively. The median EQ-5D-5L index and EQ-VAS scores were significantly higher in patients with higher educational status compared to illiterate (p < 0.05). There were significant differences in both EQ-5D-5L utility value and EQ-VAS scores among patients with previous hospital admission history compared to those without [(0.67 vs. 0.86 and (65 vs. 75)], respectively. Significant positive statistical correlations were observed with both median scores of EQ-5D-5L utility and EQ-VAS, and adherence to different types of lifestyle modifications. Conversely, no significant correlation was observed between EQ-5D-5L utility value and EQ-VAS scores, and medication adherence status, number of medications used, non-cardiac comorbidities, and number of cardiac conditions. A significant difference in EQ-5D-5L utility and EQ-VAS scores were found between all satisfaction dimensions score. Patients with higher treatment satisfaction (>75) had higher EQ-5D-5L utility index and EQ-VAS scores (Table 3).

Treatment satisfaction and its correlation with HRQoL

The mean (SD) treatment satisfaction scores for convenience and overall satisfaction were 87.7 (17.9) and 78.9 (27.7), respectively. On the other hand, the effectiveness and safety satisfaction dimensions (experience of patient encountered side effects) measured over the previous 2–3 weeks were found to be 73.8 (36.7) and 53.1 (33.5), respectively. There were significant modest positive correlations between effectiveness, convenience, and overall treatment satisfaction dimensions with EQ-5D-5L and EQ-VAS scores. The Pearson's correlation coefficient between mean scores of effectiveness and the EQ-5D-5L score was 0.32 while the correlation between the convenience dimension showed a negative correlation with the EQ-5D-5L index (correlation coefficient = -0.37) and EQ-VAS score (correlation coefficient = -0.34) (Table 4).

Factors associated with health-related quality of life

The multivariable Tobit regression model (Table 5) showed that older age ($\beta = -0.033$, 95%CI = -0.151, 0.005),



being unemployed ($\beta = -0.087$, 95%CI = -0.174, -0.001), previous hospital admission history ($\beta = -0.165$, 95%CI = -0.249, -0.082), and presence of three or more cardiovascular diseases ($\beta = -0.103$, 95%CI = -0.205, -0.001) were significantly negatively associated with the EQ-5D-5L utility index. Conversely, overall treatment satisfaction dimension (B = 0.003, 95%CI = 0.001, 0.004) and using lifestyle modification $(\beta = 0.164; 95\%$ CI = 0.255, 0.072) were significantly positively associated with EQ-5D-5L utility. Similarly, overall treatment satisfaction dimension ($\beta = 0.002, 95\%$ CI = 0.001, 0.003) and lifestyle modifications ($\beta = 0.086$, 95%CI = 0.147, 0.026) were positively associated with EQ-VAS score while previous hospitalization history was negatively associated with EQ-VAS score. Marital status, sex, education status, insurance usage, non-cardiac comorbidities, years since CVD diagnosis, and polypharmacy were not significantly associated with either the EQ-5D-5L index or EQ-VAS scores.

Discussion

The symptoms and complications of CVD as well as its treatment have a significant impact on patients' HRQoL (6, 7, 19, 37, 38). As a result, HRQoL measures are now widely used to assess disease burden and treatment outcomes in both clinical practice and research. Thus, this study aimed to assess the HRQoL and treatment satisfaction of patients with CVD; compare EQ-5D-5L index and EQ-VAS scores across different levels of patient characteristics, and identify factors associated with HRQoL. Consistent with the previous literature (18, 19, 39, 40), our findings demonstrated that CVD diagnosis was significantly associated with impaired HRQoL. The mean EQ-5D-5L utility value in this study was 0.69, which is lower

than the mean utility value of 0.92 for the Ethiopian general population (36). The most frequently reported health problems were pain/discomfort followed by mobility. Further, our study identified factors associated with HRQoL that could be targeted by interventions to improve patients' HRQoL and reduce burden of the disease.

A perfect health state (11,111) was reported only in 37 (10.4%) patients in the descriptive dimension of EQ-5D-5L which is much lower than reported in a Chinese study (55.6%) (39). Similar to previous studies (18, 37, 40), we found that pain/discomfort was the most affected dimension (75.4%). This indicates that CVD could have a detrimental impact on the physical component of patient's HRQoL; particularly due to frequent use of injectable medications, cardiovascular complications such as stroke, and general disease progression (19, 37). The mobility dimension was the second most frequently reported health problem, which could be due to the inclusion of patients with stroke and heart failure in this study. These conditions may cause significant difficulty in climbing stairs and walking. The EQ-5D-5L usual activity dimension was the third most frequently reported problem and this might be due to cardiovascular complications that might have contributed to a decline in daily routine activities of patients. Overall, our findings highlight the importance of paying attention to patients' physical functioning by fostering comprehensive lifestyle changes with physical activity, controlling CVD complications, and non-cardiac comorbidities. Additionally, HRQoL should be part of routine patient assessment and be an integral part of CVD management beyond pharmacotherapy.

The mean EQ-5D-5L utility value of our finding was 0.69, which is lower than that reported by studies from China (0.85), European countries (0.78), Vietnam (0.82), and Hong Kong (0.889) (18, 41-43). The mean EQ-VAS score in our study

TABLE 3 Median (IQR) differences of EQ-5D-5L utility and EQ-VAS scores with patient characteristics.

Variables	EQ-5D-5L index	Mean rank	<i>p</i> -value	EQ-VAS median	Mean rank	<i>p</i> -value
	median score (IQR)			score (IQR)		
Sex						
Male	0.70 (0.57-0.94)	190.6	0.127	75 (50.0-85.0)	176.9	0.784
Female	0.82 (0.52-0.91)	173.0		70 (50.0-85.0)	180.1	
Age category						
<60	0.84 (0.58–0.92)	181.1	0.552	70 (50.0-85.0)	180.0	0.500
≥60	0.82 (0.50-0.92)	174.1		75 (50.0-85.0)	176.9	
Marital status						
Single	0.82 (0.36-0.90)	164.3	0.195	70 (50.0-88.8)	171.8	0.240
Divorced	0.88 (0.61-0.92)	191.1		66 (48.8-81.3)	161.9	
Married	0.86 (0.58–0.93)	187.2		75 (60.0-85.0)	188.1	
Widow	0.77 (0.45-0.92)	160.8		65 (50.0-80.0)	159.8	
Employment status						
Employed	0.86 (0.64–0.93)	194.7	0.114	75 (60.0–90.0)	198.8	0.068
Unemployed	0.82 (0.51-0.92)	166.7		70 (50.0-85.0)	173.9	
Education status						
Unable to read and write	0.72 (0.49-0.89)	167.5	0.001	85 (47.5–77.5)	178.5	0.009
Primary education	0.74 (0.42–0.90)	179.2		70 (50.0-80.0)	177.4	
Secondary education	0.74 (0.42-0.90)	178.3		75 (60.0-85.5)	175.8	
Higher education	0.88 (0.59-0.94)	192.3		78 (63.8–86.3)	187.4	
Residence						
Addis Ababa	0.86 (0.58-0.92)	187.6	0.008	75 (55.0-85.0)	184.9	0.086
Other	0.69 (0.44–0.90)	155.0		70 (50.0-80.0)	163.4	
Health insurance usage						
Yes	0.82 (0.51-0.92)	170.0	0.042	70 (50.0-85.0)	171.5	0.088
No	0.86 (0.58–0.94)	192.7		75 (58.8–85.0)	190.4	
Lifestyle modification						
Yes	0.85 (0.59–0.92)	185.6	0.006	75 (55.0-85.0)	185.7	0.005
No	0.72 (0.25-0.90)	144.3		60 (50.0-75.0)	144.0	
Lifestyle modification type						
Diet	0.72 (0.47–0.90)	126.5	0.000	70 (50.0-80.0)	130.6	0.000
Exercise	0.87 (0.83–0.93)	178.5		80 (67.5–90.0)	180.1	
Both	0.90 (0.80-0.96)	188.5		80 (70.0-90.0)	180.8	
Smoking habit						
Yes	0.85 (0.59–0.92)	225.8	0.300	75 (55.0-85.0)	140.7	0.406
No	0.72 (0.25–0.89)	177.8		60 (50.0-75.0)	179.0	
Alcohol use						
Yes	0.90 (0.72–1.00)	219.5	0.042	75 (62.5–90.0)	196.7	0.371
No	0.84 (0.54–0.92)	176.0		70 (50.0-85.0)	177.7	
History of hospital admission						
Yes	0.67 (0.34–0.88)	138.1	0.000	65 (45.0-80.0)	150.3	0.008
No	0.86 (0.59–0.93)	189.5		75 (56.3–85.0)	186.4	
Time since CVD diagnosis						
<5 years	0.84 (0.39-0.92)	174.9	0.586	70 (50.0-86.0)	176.6	0.944
5–10 years	0.82 (0.53-0.92)	174.4		75 (50.5–80.0)	181.3	
>10 years	0.86 (0.60-0.92)	186.2		75 (50.0-85.0)	178.6	

(Continued)

TABLE 3 Continued

Variables	EQ-5D-5L index median score (IQR)	Mean rank	<i>p</i> -value	EQ-VAS median score (IQR)	Mean rank	<i>p</i> -value
Number of cardiac conditions						
<3 condition	0.85 (0.59-0.92)	178.6	0.160	70 (50.0-85.0)	177.0	0.37
\geq 3 conditions	0.87 (0.58-0.92)	183.6		78 (53.7–90.0)	199.4	
Non-cardiac comorbidities						
Yes	0.85 (0.57-0.92)	180.7	0.730	70 (50.0-85.0)	178.2	0.87
No	0.84 (0.54-0.92)	176.9		75 (50.0-82.5)	180.0	
Number of medications						
<5	0.85 (0.58-0.92)	183.3	0.131	70 (50.0-80.0)	183.4	0.63
≥5	0.74 (0.58-0.93)	163.0		70 (50.0-85.0)	173.9	
Adherence to medications						
Yes	0.84 (0.56-0.92)	178.2	0.960	75 (55.0-85.0)	181.5	0.456
No	0.84 (0.51-0.92)	177.7		70 (50.0-80.0)	173.3	
Effectiveness dimension						
<75	0.67 (0.32-0.90)	143.4	0.001	60 (50.0-80.0)	140.0	0.001
≥75	0.87 (0.66-0.92)	195.5		75 (60.0-90.0)	197.1	
Safety dimension						
<75	0.87 (0.61-0.92)	53.7	0.005	70 (55.0-85.0)	53.4	0.008
≥75	0.49 (0.48-0.87)	36.5		53 (44.0-72.5)	37.1	
Convenience dimension						
<75	0.65 (0.29-0.89)	138.8	0.001	60 (50.0-80.0)	141.8	0.001
≥75	0.86 (0.60-0.92)	188.0		75 (60.0-85.0)	187.3	
Overall satisfaction dimension						
<75	0.68 (0.47-0.90)	137.0	0.001	70 (50.0-80.0)	137.6	0.001
≥75	0.88 (0.97-0.94)	194.0		75 (60.0-90.0)	193.8	

CVD, Cardiovascular disease; EQ-5D-5L, EuroQoL life questionnaires five dimensions with five levels; EQ-VAS, EuroQoL Visual Analog scale; IQR, Interquartile range. Bold text indicates a significant statistical association.

TABLE 4 Pearson correlation coefficient between treatment satisfaction and HRQoL.

Satisfaction dimension	Number of items	Mean (SD)	EQ-5D-5L score	EQ-VAS score
Effectiveness	3	73.8 (36.8)	0.32	0.28
Convenience	3	87.7 (17.9)	0.26	0.22
Safety	5	53.1 (33.5)	-0.37	-0.34
Overall satisfaction	3	78.9 (27.7)	0.26	0.27

EQ-5D-5L, EuroQoL life questionnaires five dimensions with five levels; EQ-VAS, EuroQoL Visual Analog scale; HRQoL, Health-related quality of life.

was 68.7; this is comparable with Henan et al.'s finding among Chinese patients, which was 69.44 (40). We infer from these results that CVD is associated with impaired HRQoL. These differences in utility values could be due to variations in patients' profiles, sociocultural beliefs, and types of CVD across study settings as well as access to medical care and differences in EQ-5D-5L tariffs utilized.

Similar to previous studies (26, 38, 44), we found statistically significant differences in EQ-5D-5L index and EQ-VAS scores among participants with different levels of education, lifestyle modification use, social habits, previous hospital admission, and TSQM dimensions score. Our findings showed that higher treatment satisfaction (TSQM overall score \geq 75) was associated with improved HRQoL, which is consistent with previous findings (44). Our findings confirmed that improving treatment satisfaction and improving patients' safety by reducing medication side effects could improve health outcomes. Our findings may help in designing intervention strategies for CVD patients through identifying modifiable factors associated with poorer HRQoL. The modest correlations between TQSM, EQ-5D-5L and EQ-VAS scores could be due to different constructs of the scales. TSQM focuses more on patient judgment about treatment-related experiences and healthcare services, while the EQ-5D-5L dimensions

TABLE 5 Predictors of HRQoL in patients with cardiovascular disease.

Variables	EQ-5D-5L index score		EQ-VAS score	EQ-VAS score	
	β-Coeff. (95 % CI)	<i>p</i> -value	β-Coeff. (95 % CI)	<i>p</i> -value	
Female	0.011 (-0.063, 0.085)	0.778	-0.027 (-0.077, 3.33)	0.756	
Level of education (ref = illiterate)					
Primary	-0.085 (-0.182, 0.011)	0.072	-0.011 (-0.075, 0.054)	0.746	
Secondary and higher	-0.080(-0.167, 0.007)	0.082	0.012 (-0.070, 0.046)	0.686	
Marital status (ref = unmarried)					
Married	-0.048 (-0.117, 0.022)	0.179	-0.034 (-0.081, 0.012)	0.148	
Age (Ref = <60 years)					
\geq 60 years	-0.033 (-0.151,0.005)	0.027*	-0.036 (-0.085, 0.013)	0.144	
Alcohol use $(ref = no)$					
Yes	-0.105 (-0.239, 0.028)	0.120	-0.059 (-0.149, 0.721)	0.191	
Lifestyle modification (ref $=$ no)					
Yes	0.164 (0.255, 0.072)	0.001*	0.086 (0.147, 0.026)	0.006*	
Insurance usage (ref = no)					
Yes	0.048 (-0.023, 0.118)	0.183	0.033 (-0.013, 0.080)	0.161	
Employment status (ref = employed)					
Non-employed	-0.087 (-0.174, -0.001)	0.049*	-0.055 (-0.113, 0.001)	0.056	
History of previous admission (ref = no)					
Yes	-0.165 (-0.249, -0.082)	0.001*	-0.081 (-0.137, -0.025)	0.005*	
Years since CVD diagnosis	-0.003 (-0.001, -0.007)	0.191	-0.001 (-0.002, 0.003)	0.663	
Non-cardiac comorbidity (ref = no)					
Yes	-0.037 (-0.110, 0.037)	0.322	-0.012 (-0.059, -0.035)	0.614	
Number of CVD (ref = <3 conditions)					
\geq 3 conditions	-0.103 (-0.205, -0.001)	0.049*	-0.025 (-0.093, 0.044)	0.478	
Number of medication (ref = <5)					
>5 medications	-0.010 (-0.083, 0.064)	0.791	-0.028 (-0.077, 0.021)	0.269	
Overall satisfaction	0.003 (0.001, 0.004)	0.001*	0.002 (0.001, 0.003)	0.001*	

*p-value < 0.05.

CI, Confidence interval; CVD, Cardiovascular disease; EQ-5D-5L, EuroQoL life questionnaires five dimensions with five levels; EQ-VAS, EuroQoL Visual Analog scale; Ref, Reference. Bold text indicates a significant statistical association.

focus more on physical functioning for a health state (12, 44).

The present study identified unemployment, older age, previous hospital admission, non-adherence to lifestyle changes, presence of three or more CVD, and lower overall satisfaction as significant negative predictors of HRQoL. The decline of HRQoL with older individuals could be attributed to increased deterioration of physical functioning, which might increase CVD progression and non-cardiac comorbidities leading to reduced overall HRQoL. Likewise, lower HRQoL among previously hospitalized patients could be explained by hospital admission being mainly due to progression and worsening of the disease conditions (45). Consistent with previous research (37, 38, 41), our findings illustrate that the use of lifestyle changes is positively associated with better HRQoL, and those patients who used physical activity had significantly better HRQoL than patients who did not. Our study also demonstrated that the presence of more than three CVD was an independent predictor of impaired HRQoL. Thus, our findings indicate the importance of regular physical activity as well as reducing the progression of disease in the patients (37). Further, our finding showed that higher overall treatment satisfaction was significantly associated with better HRQoL, demonstrating the importance to focus on increasing patients' treatment satisfaction by ameliorating medication side effects. Therefore, healthcare providers should evaluate patient satisfaction with treatment during routine follow-up.

There are some limitations to this study. First, since our study was a cross-sectional survey, it is not possible to demonstrate causal relationships between associated factors and HRQoL. Second, patients were recruited from one tertiary hospital, and thus, our conclusions cannot be generalized to CVD patients in Ethiopia. Despite these limitations, this was the first study assessing the association between HRQoL and treatment satisfaction in patients with CVD in Ethiopia. Thus, our findings have broad implications for enhancing HRQoL of patients with CVD by identifying modifiable factors for poorer HRQoL that could inform context-specific interventions in resource limited settings. In addition, the utility weight generated is important measure that helps to conduct economic evaluation in the future.

Conclusions

Our study found that CVD diagnosis had a profound negative effect on HRQoL and patient treatment satisfaction. Most patients reported problems in the pain/discomfort and mobility physical dimensions of the EQ-5D-5L. The median EQ-5D-5L utility and EQ-VAS scores were found to be 0.84 and 70, respectively. Older age, previous hospital admission, non-adherence to lifestyle modifications, and presence of three or more CVD factors were significantly negatively associated with HRQoL. Therefore, future intervention efforts aimed at improving HRQoL and treatment satisfaction in CVD patients should be designed with a focus on modifiable factors such as controlling progression of CVD and promoting lifestyle change. Further, the utility values derived in our study could support economic evaluations to identify cost-effective interventions that can improve health outcomes for patients with CVD in the future.

Data availability statement

The original contributions presented in the study are included in the article/Supplementary material, further inquiries can be directed to the corresponding authors.

Ethics statement

The study received ethics approval from the Ethics Review Board of the School of Pharmacy, Addis Ababa University

References

(Protocol#:ERB/SOP/296/13/2021). The patients/participants provided their written informed consent to participate in this study.

Author contributions

KT, GTG, KB, and GBG contributed to the study concept and design. KT and GTG took the lead in data collection and analysis, in results interpretation, and drafting the first manuscript. KB, BS, and GBG contributed towards data analysis, result interpretation, and revising the manuscript for publication. All authors read and approved the final manuscript.

Funding

This research was supported, in part, by Canadian Research Chair in Economics of Infectious Diseases at the University Health Network in Toronto, Ontario held by BS (CRC-950-232429).

Acknowledgments

The authors would like to extend their sincere thanks to all research participants as well as pharmacists, nurses, and physicians working in the study hospitals.

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

^{1.} Roh GA, Mensah GA, Johnson CO, Addolorato G, Ammirati E, Baddour LM, et al. Global burden of cardiovascular diseases and risk factors, 1990-2019: update from the GBD 2019 study. *J Am Coll Cardiol.* (2020) 76:2982-3021. doi: 10.1016/j.jacc.2020.11.010

^{2.} Virani SS, Alonso A, Aparicio HJ, Benjamin EJ, Bittencourt MS, Callaway CW, et al. Heart disease and stroke statistics-2021 update a report from the American Heart Association. *Circulation*. (2021) 143:254–743. doi: 10.1161/CIR.000000000000950

^{3.} Tsao CW, Aday AW, Almarzooq ZI, Alonso A, Beaton AZ, Bittencourt MS, et al. Heart disease and stroke statistics--2 022 update: a report from the American Heart Association. *Circulation.* (2022) 145:153-639. doi: 10.1161/CIR.000000000001052

^{4.} Walelgne W, Yadeta D, Feleke Y, Kebede T. *Guidelines on Clinical and Programmatic Management of Major Non Communicable Diseases.* Addis Ababa: Federal Democratic Republic of Ethiopia Ministry of Health (2016).

5. Angaw DA, Ali R, Tadele A, Shumet S. The prevalence of cardiovascular disease in Ethiopia: a systematic review and meta-analysis of institutional and community-based studies. *BMC Cardiovasc Disord.* (2021) 21:37. doi: 10.1186/s12872-020-01828-z

6. Bahall M, Legall G, Khan K. Quality of life among patients with cardiac disease: the impact of comorbid depression. *Health Qual Life Outcomes.* (2020) 18:1–10. doi: 10.1186/s12955-020-01433-w

7. Ko HY, Lee JK, Shin JY, Jo E. Health-related quality of life and cardiovascular disease risk in Korean adults. *Korean J Fam Med.* (2015) 36:349. doi: 10.4082/kjfm.2015.36.6.349

8. Singh R, Dixit S. Health-related quality of life and health management. J Health Manag. (2010) 12:153–72. doi: 10.1177/097206341001200204

9. Harper A, Power M, Orley J, Herrman H, Schofield H, Murphy B, et al. Development of the world health organization WHOQOL-BREF quality of life assessment. *Psychol Med.* (1998) 28:551–8. doi: 10.1017/S0033291798006667

10. Guyatt GH, Feeny DH, Patrick DL. Measuring healthrelated quality of life. *Ann Intern Med.* (1993) 118:622– 9. doi: 10.7326/0003-4819-118-8-199304150-00009

11. Zyoud SH, Al-Jabi SW, Sweileh WM, Arandi DA, Dabeek SA, Esawi HH, et al. Relationship of treatment satisfaction to health-related quality of life among Palestinian patients with type 2 diabetes mellitus: findings from a cross-sectional study. *J Clin Transl Endocrinol.* (2015) 2:66–71. doi: 10.1016/j.jcte.2015.03.002

12. Xue C, Ye J, Dong Y, Xu C. Home medication cabinets and medication taking behavior of the staffs in a University in China. *IOP Conf Ser Mater Sci Eng.* (2018) 301:8. doi: 10.1088/1757-899X/301/1/012107

13. Liberato ACS, Rodrigues RCM, São-João TM, Alexandre NMC, Gallani MCBJ. Satisfaction with medication in coronary disease treatment: psychometrics of the treatment satisfaction questionnaire for medication. *Rev Lat Am Enfermagem.* (2016) 24:e2705. doi: 10.1590/1518-8345.0745.2705

14. Alam MM, Sikdar P, Kumar A, Mittal A. Assessing adherence and patient satisfaction with medication: validation of TSQM in emerging markets. *Int J Pharm Healthc Mark.* (2018) 12:409–32. doi: 10.1108/IJPHM-10-2016-0053

15. Tarekegn GE, Gezie LD, Birhan TY, Ewnetu F. Health-related quality of life among heart failure patients attending an outpatient clinic in the University of Gondar Comprehensive Specialized Hospital Northwest, Ethiopia, 2020: using structural equation modeling approach. *Patient Relat Outcome Meas.* (2021) 12:279–90. doi: 10.2147/PROM.S322421

16. Johansson I, Joseph P, Balasubramanian K, McMurray JJV, Lund LH, Ezekowitz JA, et al. Health-related quality of life and mortality in heart failure: the global congestive heart failure study of 23 000 patients from 40 countries. *Circulation*. (2021) 143:2129–42. doi: 10.1161/CIRCULATIONAHA.120.050850

17. Chen K, Chiou CF, Plauschinat CA, Frech F, Harper A, Dubois R. Patient satisfaction with antihypertensive therapy. *J Hum Hypertens.* (2005) 19:793–9. doi: 10.1038/sj.jhh.1001899

18. Liang Z, Zhang T, Lin T, Liu L, Wang B, Fu AZ, et al. Healthrelated quality of life among rural men and women with hypertension: assessment by the EQ-5D-5L in Jiangsu, China. *Qual Life Res.* (2019) 28:2069– 80. doi: 10.1007/s11136-019-02139-3

19. Petek D, Petek-Ster M, Tusek-Bunc K. Health behavior and health-related quality of life in patients with a high risk of cardiovascular disease. *Zdr Varst.* (2018) 57:39–46. doi: 10.2478/sjph-2018-0006

20. Pinheiro LC, Reshetnyak E, Sterling MR, Richman JS, Kern LM, Safford MM. Using health-related quality of life to predict cardiovascular disease events. *Qual Life Res.* (2019) 28:1465–75. doi: 10.1007/s11136-019-02103-1

21. Kazukauskiene N, Burkauskas J, Macijauskiene J, Mickuviene N, Brozaitiene J. Exploring potential biomarkers associated with health-related quality of life in patients with coronary artery disease and heart failure. *Eur J Cardiovasc Nurs.* (2018) 17:645–51. doi: 10.1177/1474515118772828

22. EuroQol Group. EuroQol-a new facility for the measurement of health-related quality of life. *Health Policy*. (1990) 16:199–208. doi: 10.1016/0168-8510(90)90421-9

23. Juniper EF, Sonia Buist A, Cox FM, Ferrie PJ, King DR. Validation of a standardized version ofthe asthma quality of life questionnaire. *Chest.* (1999) 115:1265–70. doi: 10.1378/chest.115.5.1265

24. Power MJ, Green AM. Development of the WHOQOL disabilities module. *Qual Life Res.* (2010) 19:571–84. doi: 10.1007/s11136-010-9616-6

25. Lins L, Carvalho FM. SF-36 total score as a single measure of health-related quality of life: scoping review. *SAGE Open Med.* (2016) 4:725. doi: 10.1177/2050312116671725

26. Nguyen LH, Tran BX, Hoang Le QN, Tran TT, Latkin CA. Quality of life profile of general Vietnamese population using EQ-5D-5L. *Health Qual Life Outcomes.* (2017) 15:199. doi: 10.1186/s12955-017-0771-0

27. Pennington B, Hernandez-Alava M, Pudney S, Wailoo A. The impact of moving from EQ-5D-3L to-5L in NICE technology appraisals. *Pharmacoeconomics.* (2019) 37:75-84. doi: 10.1007/s40273-018-0701-y

28. EuroQol Research Foundation. *EQ-5D-5L User Guide* v3.0. (2019). p. 169–232. Available online at: https://euroqol.org/publications/user-guides (accessed May 20, 2022).

29. Montgomery AA, Fahey T, Ben-Shlomo Y, Harding J. The influence of absolute cardiovascular risk, patient utilities, and costs on the decision to treat hypertension: a Markov decision analysis. *J Hypertens.* (2003) 21:1753–9. doi: 10.1097/00004872-200309000-00026

30. Arifin WN. Introduction to sample size calculation. *Educ Med J.* (2013) 5:e89–96. doi: 10.5959/eimj.v5i2.130

31. Kasiulevičius V, Šapoka V, Filipavičiute R. Sample size calculation in epidemiological studies. *Gerontologija*. (2006) 7:225–31.

32. Power MJ, Green AM. Development of the WHOQOL disabilities module. *Qual Life Res.* (2010) 19:571–84.

33. Atkinson MJ, Sinha A, Hass SL, Colman SS, Kumar RN, Brod M, et al. Validation of a general measure of treatment satisfaction, the Treatment Satisfaction Questionnaire for Medication (TSQM), using a national panel study of chronic disease. *Health Qual Life Outcomes.* (2004) 2:12. doi: 10.1186/1477-7525-2-12

34. Ting J, Liu Y, Petrillo J, Giannattasio G, Sabatella G. Treatment satisfaction with disease modifying therapies in multiple sclerosis: a systematic review of studies using the treatment satisfaction questionnaire for medication (Tsqm). *Value Heal.* (2015) 18:760–1. doi: 10.1016/j.jval.2015.09.2484

35. Morisky DE, Green LW, Levine DM. Concurrent and predictive validity of a self-reported measure of medication adherence. *Med Care.* (1986) 24:67–74. doi: 10.1097/00005650-198601000-00007

36. Welie AG, Gebretekle GB, Stolk E, Mukuria C, Krahn MD, Enquoselassie F, et al. Valuing health state: an EQ-5D-5L value set for Ethiopians. *Value Heal Reg Issues.* (2020) 22:7–14. doi: 10.1016/j.vhri.2019.08.475

37. Schriefer D, Haase R, Kullmann JS, Ziemssen T. Health-related quality of life and the relationship to treatment satisfaction in patients with multiple sclerosis: insights from a large observational study. *Patient Prefer Adherence*. (2020) 14:869–80. doi: 10.2147/PPA.S248272

38. Arija V, Villalobos F, Pedret R, Vinuesa A, Jovani D, Pascual G, et al. Physical activity, cardiovascular health, quality of life and blood pressure control in hypertensive subjects: randomized clinical trial. *Health Qual Life Outcomes*. (2018) 16:184. doi: 10.1186/s12955-018-1008-6

39. Wong ELY, Xu RH, Cheung AWL. Health-related quality of life among patients with hypertension: population-based survey using EQ-5D-5L in Hong Kong SAR, China. *BMJ Open.* (2019) 9:e032544. doi: 10.1136/bmjopen-2019-032544

40. Mei YX, Wu H, Zhang HY, Hou J, Zhang ZX, Liao W, et al. Health-related quality of life and its related factors in coronary heart disease patients: results from the Henan Rural Cohort study. *Sci Rep.* (2021) 11:1–10. doi: 10.1038/s41598-021-84554-6

41. Ludt S, Wensing M, Szecsenyi J, van Lieshout J, Rochon J, Freund T, et al. Predictors of health-related quality of life in patients at risk for cardiovascular disease in European primary care. *PLoS ONE.* (2011) 6:e0029334. doi: 10.1371/journal.pone.0029334

42. Tran BX, Moir MP, Thai TPT, Nguyen LH, Ha GH, Nguyen THT, et al. Socioeconomic inequalities in health-related quality of life among patients with cardiovascular diseases in Vietnam. *Biomed Res Int.* (2018) 2018:2643814. doi: 10.1155/2018/2643814

43. Xu RH, Cheung AWL, Wong ELY. The relationship between shared decision making and health-related quality of life among patients in Hong Kong SAR, China. *Int J Qual Heal Care.* (2017) 29:534–40. doi: 10.1093/intqhc/m zx067

44. Al-Jabi SW, Zyoud SH, Sweileh WM, Wildali AH, Saleem HM, Aysa HA, et al. Relationship of treatment satisfaction to health-related quality of life: findings from a cross-sectional survey among hypertensive patients in Palestine. *Heal Expect.* (2015) 18:3336–48. doi: 10.1111/hex.12324

45. Pocock S, Brieger DB, Owen R, Chen J, Cohen MG, Goodman S, et al. Healthrelated quality of life 1-3 years post-myocardial infarction: its impact on prognosis. *Open Hear.* (2021) 8:1–11. doi: 10.1136/openhrt-2020-001499