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## Health-related Quality of Life, Functional Status and Cardiac Event-Free Survival in Patients with Heart Failure

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### Abstract

**BACKGROUND**—Health-related quality of life (HRQOL), functional status, and cardiac event-free survival are outcomes used to assess the effectiveness of interventions in patients with heart failure (HF). However, the nature of the relationships among HRQOL, functional status and cardiac event-free survival remains unclear.

**OBJECTIVE**—The purpose of this study was to examine the nature of the relationships among HRQOL, functional status, and cardiac event-free survival in patients with HF.

**METHODS**—This was a prospective, observational study of 313 patients with HF that was a secondary analysis from a registry. At baseline, patient demographic and clinical data were collected. HRQOL was assessed using the Minnesota Living with Heart Failure Questionnaire (MLHFQ) and functional status was measured using the Duke Activity Status Index (DASI). Cardiac event-free survival data were obtained by patient interview, hospital database and death certificate review. Multiple linear and Cox regressions were used to explore the relationships among HRQOL, functional status and cardiac event-free survival while adjusting for demographic and clinical factors.

**RESULTS**—Participants (n = 313) were male (69%), Caucasians (79%), and aged 62 ± 11 years. Mean left ventricular ejection fraction of (LVEF) was 35 ± 14%. Mean HRQOL score of 32.3 ± 20.6 indicated poor HRQOL. Mean DASI score of 16.2 ± 12.9 indicated poor functional status. Cardiac event-free survival was significantly worse in patients who had worse HRQOL or poorer functional status. Patients who had better functional status had better HRQOL ( $p < .001$ ). HRQOL was not a significant predictor of cardiac event-free survival after entering functional status in the model ( $p = .54$ ) demonstrating that it was a mediator of the relationship between HRQOL and outcome.

**CONCLUSION**—Functional status was a mediator between HRQOL and cardiac event-free survival. These data suggest intervention studies to improve functional status are needed.

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**Conflict of Interest**  
None.

## Keywords

Health-related quality of life; Functional Status; Event-free Survival; Outcomes; Heart Failure

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## Introduction

Heart failure (HF) is a chronic and prevalent syndrome with a poor prognosis,<sup>1</sup> and a multitude of symptoms that reduce activity level and result in psychological distress. Health-related quality of life (HRQOL) is a multidimensional construct referring to how a health condition affects total well-being, including physical, functional, emotional, and social dimensions;<sup>2</sup> adults with HF have poorer HRQOL compared to those without HF.<sup>3–5</sup> Health-related quality of life is a predictor of hospitalization and death in patients with HF.<sup>6–14</sup> It is a subjective, patient-centered outcome that is recognized by researchers and clinicians as an important outcome for patients with HF.<sup>15</sup> However, little is known about the potential mechanisms linking HRQOL with hospitalization and death.

Poor functional status is also associated with poorer HF outcomes (i.e., hospitalizations and death).<sup>16, 17</sup> Functional status, the ability to perform activities of daily living, predicted survival in patients with HF, and was an indicator for selection of patients with HF for heart transplantation.<sup>18–20</sup> Because of the association of HRQOL, functional status, and hospitalization/death, these variables have become important endpoints of HF care.<sup>21–25</sup> Numerous investigators have tested interventions to improve HRQOL, functional status, and survival. A number of multidisciplinary disease management, and exercise intervention studies in patients with HF have led to improvement in HRQOL,<sup>21–24</sup> functional status,<sup>22–24</sup> hospitalization, and death.<sup>21, 25, 26</sup> Moreover, many demographic (age, gender, ethnicity)<sup>8, 17, 27–31</sup> and clinical factors (ejection fraction, comorbidities, beta-blocker, ACE inhibitor)<sup>5, 8, 9, 13, 16, 17, 27, 28, 30–33</sup> have been reported to influence HRQOL, functional status, hospitalization or death. However, the relationships among HRQOL, functional status, and hospitalization and death are not clear. Therefore, the purpose of this study was to examine the nature of the relationships among HRQOL, functional status, and cardiac event-free survival in patients with HF by determining whether functional status was a mediator between HRQOL and cardiac event-free survival in this patient population with and without adjustment of some important covariates.

## Methods

### Design and Sample

This was a prospective, observational study conducted as a secondary analysis<sup>34</sup> that included a sample of 313 patients enrolled from outpatient setting of multiple large community hospitals and academic medical centers in the United States. Patients were eligible for inclusion if they had a confirmed diagnosis of chronic HF, preserved or non-preserved systolic function, had been optimized on HF medications and on stable doses for three months, and were English-speaking. They were excluded for a myocardial infarction within 3 months or unstable angina, obvious cognitive impairment (i.e., not able to give informed consent or participate in an interview), discharge to a skilled nursing facility, or

were diagnosed with severe psychiatric impairment other than depression or anxiety. For the purposes of this study, patients were only included if they had complete data on HRQOL, functional status and cardiac events.

## Measurement

**Cardiac event-free survival**—Cardiac event-free survival was the composite end-point of time to the first occurrence of one of the following cardiac events from the enrollment date: cardiac emergency department (ED) visits, cardiac hospitalizations, or cardiac mortality. This composite end-point is commonly used in such research.<sup>9, 21, 25, 28, 35, 36</sup> During data collection, the date and reasons for hospitalization and death were noted. Data were obtained by trained research assistants with expertise in cardiovascular nursing by patient/family interview, hospital database review and review of death certificates and records. All outcome assessment was blinded to patient HRQOL and functional status.

**Health-related quality of life (HRQOL)**—The Minnesota Living with Heart Failure questionnaire (MLHFQ) is a measure of HRQOL that is used to assess the patient's perceptions of the influence of HF on physical and emotional aspects of life.<sup>2, 37</sup> The 21 items are summed and ranged from 0–105 with higher scores indicating worse HRQOL. This instrument has been widely used to measure quality of life in this population.<sup>38–40</sup> Researchers have demonstrated evidence for validity and reliability.<sup>2, 37</sup> The Cronbach's alpha in our study was 0.93, adding support for reliability.

**Functional Status**—Functional status was assessed using the Duke Activity Status Index (DASI).<sup>41</sup> The DASI has been used in a variety of cardiac disease populations, including HF.<sup>29, 42, 43</sup> The DASI consists of 12 items and each item has four response options ranging from 1 = “can perform activity with difficulty” to 4 = “cannot perform activity at all”. Each item is weighted based on the metabolic equivalent associated with the activity represented by that item. For example, the weight for walking indoors is 1.75, while that for running is 8. Only items that are rated 1 by the respondent receive a score. Items that are rated 2, 3, or 4, indicating that the activity can only be performed with difficulty or cannot be done at all, are scored as zero. The total score is calculated by adding the weighted score for each item. The total score can range from 0 to 58.2, with higher scores indicating better functional status. The activities in the DASI included personal care, ambulation, household tasks, sexual function and recreational activities which represent major aspects of physical function.<sup>41</sup> The reliability and validity of the DASI have been demonstrated previously.<sup>31, 41, 44, 45</sup> Cronbach's  $\alpha$  in the current study was 0.84.

**Demographic and clinical characteristics**—To describe the sample and obtain data about potential confounding variables, the following information was collected by patient interview and chart review: age, gender, ethnicity, highest education level attained, financial status, and living alone or with someone else. Financial status was measured by one question about whether patients “had more than enough, enough, or did not have enough to make ends meet”. The following clinical characteristics were collected by chart review: left ventricular ejection fraction (LVEF) within the past 3 months, and medications (e.g., taking ACE inhibitor, or  $\beta$ -blocker). Data about comorbidities were collected by chart review and

patient interview using the Charlson Comorbidity Index (CCI).<sup>46, 47</sup> The CCI is weighted for severity of comorbidity and is computed as a total score. A higher CCI score implies higher comorbidity burden.<sup>46, 47</sup>

## Procedure

Institutional Review Board approval was obtained for each site and patients gave written informed consent. The review board at the primary author's institution approved all secondary data analyses of this investigation as an exempt protocol. Patients were enrolled and recruited in cardiology clinics after referral from clinicians. These patients completed baseline assessment and were followed monthly by telephone to collect data about cardiac events and confirmed by hospital data base review and review of death certificates and records. Patients were followed for a median of 360 days to determine cardiac event-free survival.

## Data analysis

SPSS version 22.0 (Chicago, IL) was used for data analysis; a *p* value of less than 0.05 was considered significant. Patients were categorized as better or worse HRQOL based on the median score of the MLHFQ in this sample (i.e., 30) and divided into better or poorer functional status groups based on the median score of the DASI in this sample (i.e., 10.7). The median score was used in this study because there are no standard cutpoints, and the median is the most commonly used cutpoint in the literature.<sup>8, 12</sup> Differences in demographic and clinical variables between the groups formed by median MLHFQ and DASI scores were assessed with independent t-tests or chi-square tests of association based on the level of measurement.

Logistic/linear regressions, t-tests, Pearson correlation, Kaplan-Meier plots with log-rank test, and Cox regressions were used to explore the relationships among HRQOL, functional status, and cardiac event-free survival. In the linear regression models, we performed multicollinearity tests to evaluate this assumption of regression. There were no issues with multicollinearity as all variance inflation factors were < 8. The log-rank test was used to compare the time to cardiac event-free survival between patients with better and worse HRQOL, and with better and worse functional status. Kaplan-Meier plots were used to graphically depict group differences in cardiac event-free survival. Univariate and adjusted Cox proportional hazards regression modeling was used to assess the time to cardiac event between groups. Variables showing marginal association with cardiac events in univariate analyses with alpha set at < 0.10, as were those with prior evidence of association with one or more of the independent variables or dependent variable (e.g., age, gender, ethnicity, education level, financial status, and living status, comorbidity, LVEF, taking ACE inhibitor, and  $\beta$ -blocker)<sup>5, 8, 9, 13, 16, 17, 27-33</sup> were forced into the Cox regression analysis. We also conducted the analyses with HRQOL and functional status as continuous variables.

To further explore the relationships among HRQOL, functional status, and cardiac event-free survival, we conducted mediation analyses to test whether functional status was a mediator of the relationship between HRQOL and cardiac event-free survival using a series of regression models and Cox-survival analyses. The test for mediation followed the steps

outlined by Baron and Kenny<sup>48</sup> and Bennett.<sup>49</sup> Four regression models were performed to test for the mediator effect. The first model tested whether HRQOL (the independent variable) was a predictor of functional status (mediator). The second model tested whether HRQOL was a predictor of cardiac event-free survival (outcome variable). The third model tested whether functional status was a predictor of event-free survival. In the fourth model, both HRQOL and functional status (independent and mediator variables) were entered simultaneously as predictors of cardiac event-free survival (outcome variable). The following conditions must be met if a mediator effect is present: 1) the results of the first, second, and the third models should be significant, and 2) the significance level of the coefficient associated with the independent variable in the fourth model is less significant (partial mediator) or non-significant (full mediator) compared to the third model (Figure 1).<sup>48–50</sup>

## Results

### Sample Characteristics

Three hundred and thirteen patients who had complete data on HRQOL, functional status, and cardiac events were included in this study. There were no differences in characteristics between the 313 patients who had complete data and those who did not have complete data. The mean age of patients in the sample was  $62 \pm 11$  years, and about one third of patients were female. The majority of the patients were Caucasian (79%) and living with someone (69%). Full sample characteristics and comparison of better and worse HRQOL, and better and worse functional status groups are presented in Table 1.

The mean HRQOL score as measured by MLHFQ was  $32.3 \pm 20.6$  (median: 30). Of the total sample, 159 out of the 313 patients were classified as having poorer HRQOL. Patients with worse HRQOL were younger, had less education ( $p < .001$ ), and lower functional status score ( $p < .001$ ) than those with better HRQOL.

The mean functional status score as measured by the DASI was  $16.2 \pm 12.9$  (median: 10.7). We had a greater percentage of female patients in the poorer functional status group than in the better functional status group. The proportion of female patients with poorer functional status was significantly more than predicted and higher than the group with better functional status (higher functional status 39%, lower functional status 25%,  $p = .013$ ). In addition, patients in the poorer functional status group had one year less education (poorer  $13 \pm 3$  years, better  $14 \pm 3$  years,  $p = 0.001$ ), greater comorbidity burden (poorer  $3.6 \pm 2$ , better  $2.9 \pm 2$ ,  $p = 0.002$ ) and worse HRQOL scores (poorer  $44.2 \pm 19.1$ , better  $23.7 \pm 17.1$ ,  $p < 0.001$ ) compared with those in the better functional status group.

### Association of health-related quality of life with cardiac event-free survival

Kaplan-Meier plots with log-rank tests (Figure 2) demonstrated that cardiac event-free survival was significantly worse in patients who had worse HRQOL ( $p = .003$ ). In simple Cox regression modeling, HRQOL predicted cardiac event-free survival (hazard ratio [HR] = 2.01,  $p = .003$ ). After controlling for age, gender, ethnicity, education level, financial status, living status, comorbidity, LVEF, ACE inhibitor use, and  $\beta$ -blocker use, patients who had

worse HRQOL had 2.32 times the risk of experiencing a cardiac event compared to patients who had better HRQOL. Likewise, when HRQOL was analyzed as a continuous variable, HRQOL predicted cardiac event-free survival before and after adjusting for covariates (HR = 1.015 and 1.016, respectively,  $p = .008$ ) (Table 2). For every one-point increase in MLHFQ score, the risk of a cardiac event during follow-up increased 1.5–1.6%.

### Association of functional status with cardiac event-free survival

Kaplan-Meier plots with log-rank tests (Figure 3) demonstrated that cardiac event-free survival was significantly worse in patients who had poorer functional status ( $p = .02$ ). In simple Cox regression modeling, functional status predicted cardiac event-free survival (HR=1.69,  $p = .022$ ). After controlling for age, gender, ethnicity, education level, financial status, living status, comorbidity, LVEF, ACE inhibitor use, and  $\beta$ -blocker use, patients who had poorer functional status had 1.63 times the risk of experiencing a cardiac event compared to patients with better functional status. When functional status was analyzed as a continuous variable, functional status predicted cardiac event-free survival before and after adjusting for covariates (HR = .956,  $p = .001$ ) (Table 3). For every one-point increase in DASI score, the risk of cardiac event during follow-up decreased 4.4%.

### HRQOL, functional status, cardiac event-free survival—Mediation analysis

HRQOL was moderately correlated with functional status ( $r = -.55$ ,  $p < .001$ ). When patients had better functional status (higher scores on DASI), they also had better HRQOL (lower scores on MLHFQ). In a series of regression models and Cox-survival analyses, functional status mediated the relationship between HRQOL and cardiac event-free survival based on the following sequence of regression analyses. First, in Path A (Figure 1), HRQOL independently predicted functional status ( $p < .001$ ). Second, in Path B, functional status predicted cardiac event-free survival ( $p = .022$ ). Third, in Path C, HRQOL was an independent predictor of cardiac event-free survival ( $p = .003$ ). In the final Path D, HRQOL ( $p = .54$ ) was no longer a significant predictor of cardiac event-free survival when functional status ( $p = .004$ ) was entered into the model, indicating functional status mediated the relationship between HRQOL and cardiac event-free survival. When adding age, gender, ethnicity, education level, financial status, living status, comorbidity, LVEF, ACE inhibitor and  $\beta$ -blocker use to the model, functional status ( $p = .007$ ) still mediated the relationship between HRQOL and cardiac event-free survival.

### Additional Analysis

To examine the possibility that HRQOL was a mediator, we treated functional status as an independent variable. Functional status independently predicted HRQOL scores ( $p < .001$ ) and cardiac event-free survival ( $p = .022$ ). However, HRQOL scores did not predict cardiac event-free survival in combination with functional status ( $p > 0.05$ ). Therefore, HRQOL did not mediate the relationship between functional status and cardiac event-free survival, providing further support for the hypothesis that it is functional status that mediates the relationship between HRQOL and cardiac event-free survival.

## Discussion

Heart failure is characterized by worse HRQOL and functional status, and also by poor prognosis, including frequent hospitalizations, and high mortality. In this study, we examined the role of functional status in the link between HRQOL and cardiac events in patients with HF. We found that functional status was a mediator of the association of HRQOL with hospitalization and death. Patients with HF and better HRQOL have enhanced cardiac event-free survival because of superior functional status.

In the HF literature, HRQOL is a predictor of hospitalization and death in patients with HF,<sup>6-14, 37</sup> and many investigators have found that worse physical HRQOL is an indication of worsening prognosis.<sup>7, 12</sup> For example, in the EPICAL study,<sup>7</sup> a worse score in the physical, but not the mental, component of the Duke Health Profile (a HRQOL questionnaire) was associated with a greater risk of HF hospitalization or death. Similarly, Rodriguez-Artalejo and colleagues<sup>12</sup> also reported a relationship between mortality and the physical score on the MLHFQ and Short Form-36 (another generic HRQOL questionnaire), but no association with the emotional score. Our findings not only support the hypothesis, as we found that functional status, as measured by the DASI, predicted cardiac event-free survival, but also demonstrated that functional status mediated the relationship between HRQOL and cardiac event-free survival.

Patient-reported HRQOL has been demonstrated to be an independent predictor of hospitalization and mortality in patients with HF.<sup>6-14</sup> Rector and colleagues<sup>2</sup> reported that many patients with HF would accept some risk of medication-induced death in exchange for improved health-related quality of life, demonstrating the importance of HRQOL in this population. Our findings are consistent with prior studies.<sup>6-14</sup> Patients who had worse self-reported HRQOL had more than twice the risk of experiencing a cardiac event compared to patients with better HRQOL before and after controlling for potential confounders. Lupon and colleagues<sup>8</sup> assessed HRQOL at baseline, 1, 3, and 5 years and found that baseline and follow-up HRQOL assessments were independently associated with death.<sup>8</sup> Patients who died in the one-year period after any HRQOL assessment reported significantly worse HRQOL compared with those who survived.<sup>8</sup> Moser and colleagues<sup>37</sup> found that HRQOL remained impaired, but improved substantially within 1 month of hospital discharge for most patients. However, there was a 3.3 times increased risk of rehospitalization or death in those without improved 1-month HRQOL. Results from these studies and our study confirm the importance of assessing patient-reported HRQOL.

In addition to HRQOL, functional status was associated with hospitalization and mortality.<sup>16, 17</sup> In the current study, patients with poorer functional status had about twice the risk of experiencing a cardiac event, compared to patients with better functional status before and after controlling for potential confounders. Parissis and colleagues<sup>16</sup> found that functional status, as measured by the DASI, was independently associated with cardiac events after adjusting for demographic and clinical variables.<sup>16</sup> Furthermore, Koch and colleagues reported a “dose-response” relationship between baseline and follow-up functional status and risk of long-term survival (median follow-up was 8.6 years).<sup>17</sup> Our results, combined with findings from these investigators, support the need for regular

assessment of patient-reported HRQOL and functional status to identify patients who are at risk for reduced event-free survival.

A number of investigators previously provided evidence that interventions like exercise training may improve HRQOL, and also reduce cardiac events including death in patients with HF.<sup>21–24, 51, 52</sup> Our findings suggested that improving functional status may attenuate the association between poor HRQOL, and cardiac hospitalizations and mortality. Nishi and colleagues<sup>53</sup> used exercise training in patients with HF and found that functional status was improved by  $16 \pm 15\%$  in those in the exercise training group, while functional status remained unchanged in the control group ( $p < .001$ ). Gary and colleagues<sup>22</sup> tested the effects of a home-based exercise program, and found that the exercise group had improved HRQOL. In a subsequent aerobic and resistance exercise intervention study, Gary and colleagues<sup>23, 24</sup> found that exercise participants had significant improvement in physical function, muscle strength, and HRQOL compared with those in the attention control group. Similarly, Servantes and colleagues<sup>54</sup> found that a home-based exercise training intervention improved both HRQOL (as assessed by MLHFQ) and functional status (as measured by exercise testing, peak oxygen consumption) in patients with HF. In the HF-ACTION trial, the largest and longest exercise intervention trial in patients with HF published to date, the investigators found that exercise training improved functional status<sup>51</sup>, HRQOL<sup>52</sup>, and was also associated with reduction in hospitalizations and death, after adjusting for highly prognostic baseline characteristics like depressive symptoms.<sup>51</sup> The findings and conclusions from numerous intervention trials support the importance of exercise for HRQOL, functional status, and reductions in hospitalizations and prolonged survival. Our findings further define the relationship between these variables.

## Limitations

Our study has several limitations. First, the DASI is a self-report of functional status and not an objective measure. Use of an objective measure may increase accuracy of assessment. Our data, which demonstrated a strong relationship between functional status and outcomes, suggested that functional status was accurately reflected by the self-report measure in this study. Moreover, the DASI has been validated against the reference standard of peak oxygen uptake in younger patients,<sup>41</sup> patients with HF,<sup>45</sup> and patients with chronic obstructive pulmonary disease.<sup>44</sup>

Second, we measured HRQOL and functional status in a cross-sectional fashion, which limited our ability to determine a causal relationship. However, in our additional analyses, there was no evidence to support the mediating role of HRQOL on the relationship between functional status and cardiac event-free survival. Longitudinal studies are necessary to determine the causal relationship between HRQOL and functional status.

Third, HRQOL was measured by self-reported method and we only included English-speaking patients in this study. However, HRQOL is subjective and there is no other valid way to assess HRQOL than by self-report. The MLHFQ was specifically developed to measure HRQOL in patients with HF; the instrument has demonstrated good reliability and validity, and is widely used in studies with patients with HF.<sup>38–40</sup>



Fourth, the mean age of our sample was younger than patients in general, which might limit the generalizability of the study. The difference might have resulted from recruiting participants in the ambulatory care settings and not the hospital settings. The mean age of participants in many HF studies (range 54–64),<sup>25, 55–58</sup> including a few large multi-center trials,<sup>56, 57</sup> was similar to participant age in our study. Regardless, the findings from this study provide important information related to the nature of the relationships among functional status, HRQOL and cardiac events in patients with HF.

## Conclusions

In this study, we found that functional status was a mediator between HRQOL and cardiac event-free survival. Because patients with HF tend to experience poorer HRQOL and poorer functional status, it is vital for clinicians to regularly assess HRQOL and functional status in these patients, and provide tailored, evidence-based interventions to improve HRQOL, functional status, and health outcomes.

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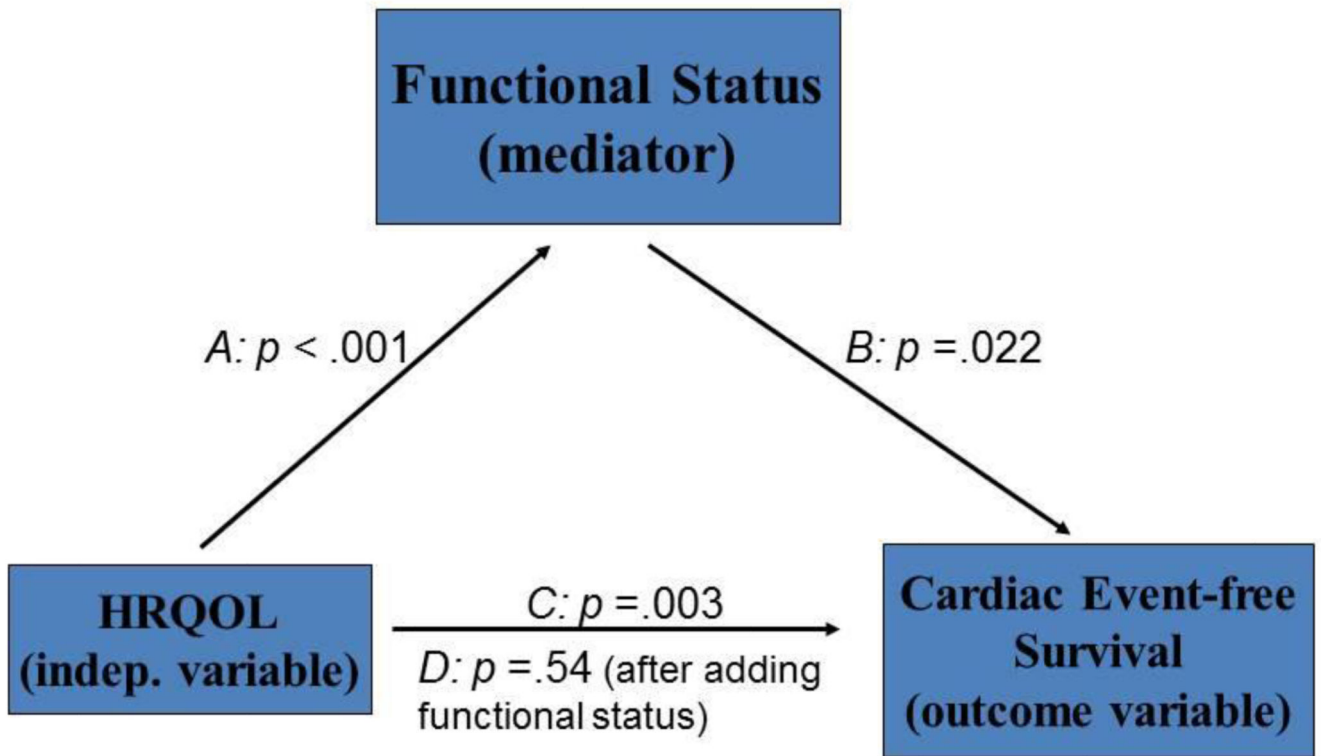
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**Figure 1.**

Functional Status is a mediator

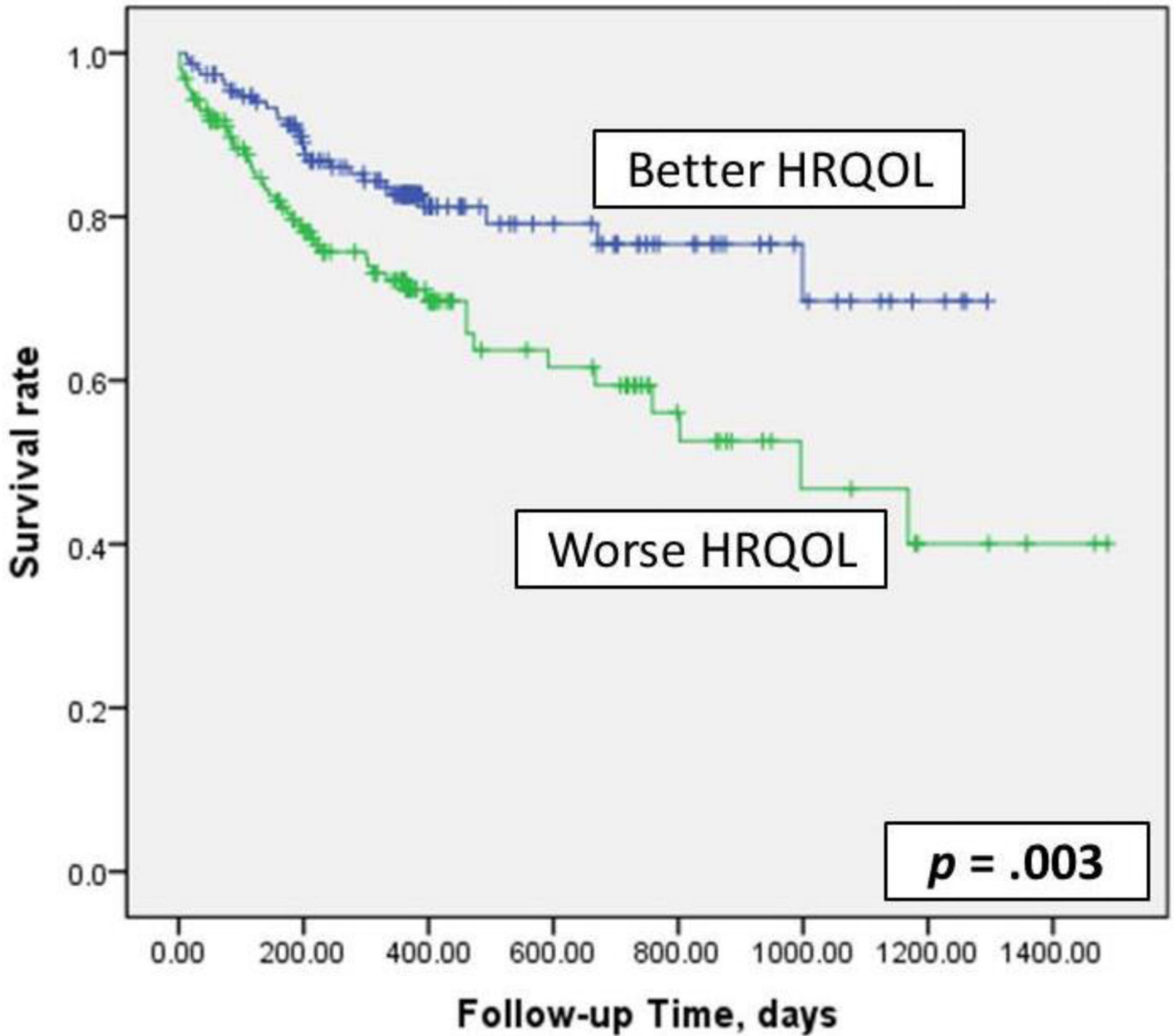
Path A: Test of whether HRQOL is a predictor of functional status

Path B: Test of whether functional status is a predictor of cardiac event-free survival

Path C: Test of whether HRQOL is a predictor of cardiac event-free survival

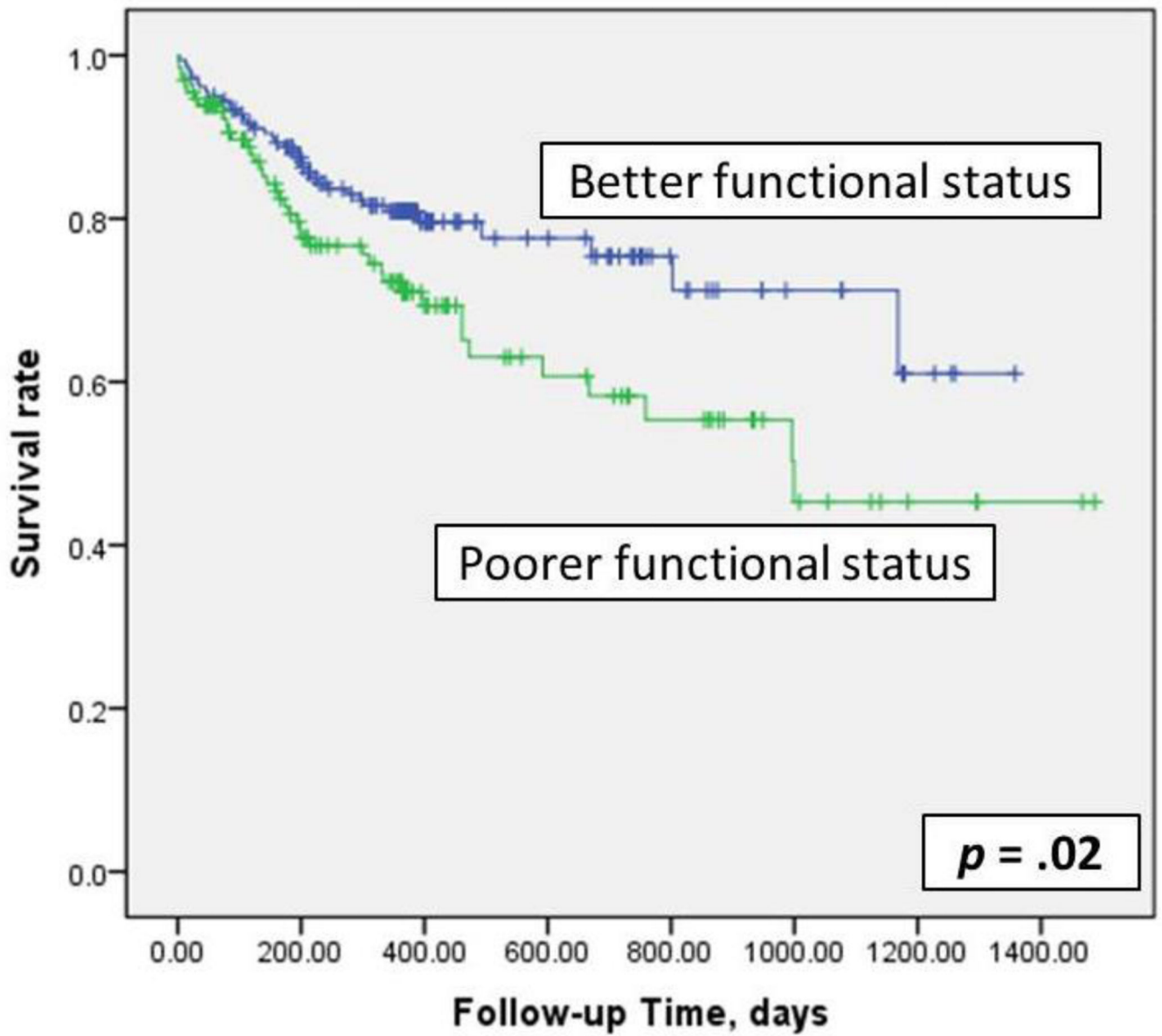
Path D: Test of whether HRQOL and functional status together are predictors of cardiac event-free survival

### Health-related Quality of Life and Cardiac Event-free Survival



**Figure 2.** Kaplan-Meier plots and log-rank test: health-related quality of life and cardiac event-free survival

### Functional Status and Cardiac Event-free Survival



**Figure 3.** Kaplan-Meier plots and log-rank test: functional status and cardiac event –free survival

Sample characteristics and comparison of clinical and demographic characteristics by health-related quality of life and functional status classification

**Table 1**

Characteristics	Overall (N = 313)	Better HRQOL (n = 154)	Worse HRQOL (n = 159)	p	Better Functional status (n = 181)	Poorer Functional status (n = 132)	p
Age, years	62±11	64±11	60±11	.006	62±11	62±11	.961
Male	217 (69)	111 (72)	106 (67)	.328	136 (75)	81 (61)	.013
Living alone	98 (31)	48 (31)	50 (31)	1.00	55 (30)	43 (33)	.712
Caucasian	247 (79)	116 (75)	131 (82)	.130	139 (77)	108 (82)	.327
Education, years	13±3	14±3	12±3	< .001	14±3	13±3	.001
Etiology, ischemic	159 (52)	78 (51)	81 (52)	.467	89 (50)	70 (54)	.636
Body mass index	30.6±7.5	29.7±7.5	31.5±7.4	.062	29.8±7.5	31.6±7.5	.071
Comorbidity	3.2±2.0	3.1±1.8	3.4±2.1	.213	2.9±2.0	3.6±2.0	.002
Prior heart attack	161 (52)	79 (51)	82 (53)	.820	88 (49)	73 (56)	.249
Hypertension	224 (73)	112 (74)	112 (73)	.898	123 (70)	101 (78)	.091
LVEF, %	35±14	34±14	35±15	.938	34±14	35±14	.552
ACEI use	227 (73)	115 (75)	112 (70)	.448	133 (74)	94 (71)	.701
Beta-blocker use	278 (89)	139 (90)	139 (87)	.476	160 (88)	118 (89)	.857
HRQOL	32.3±20.6	14.9±8.4	49.2±13.7	< .001	23.7±17.1	44.2±19.1	< .001
Functional status	16.2±12.9	22.5±14.4	10.2±7.2	< .001	23.5±12.6	6.2±2.5	< .001

Data are presented as means ± SD, or N (%), interval level data compared by independent t-test, categorical by Chi-square; NYHA = New York Heart Association; LVEF = left ventricular ejection fraction; ACEI =angiotensin-converting-enzyme inhibitor.



**Table 2**

Cox Regression Modeling: Health-related Quality of Life on Cardiac Event-free Survival (N = 313)

Variables	Hazard Ratio	Wald	Significance
<b>*Simple Cox Regression HRQOL (MLHFQ score)</b>	<b>1.015</b>	<b>7.054</b>	<b>.008</b>
<b>**Multiple Cox Regression Age</b>	1.018	2.594	.107
Gender	.555	3.876	.049
Ethnicity	1.385	1.179	.277
Education	.983	.242	.622
Living status	.764	1.266	.261
Financial status	.918	.200	.655
LVEF	.985	2.715	.099
Comorbidity	1.055	.870	.351
Taking ACEI	.802	.694	.405
Taking BB	.669	1.339	.247
<b>HRQOL (MLHFQ score)</b>	<b>1.016</b>	<b>6.995</b>	<b>.008</b>

\* $\chi^2 = 7.148, p = 0.008$ ;

\*\* $\chi^2 = 23.977, p = 0.013$

ACEI = angiotensin-converting-enzyme inhibitor; BB = beta blocker; HRQOL = health-related quality of life; LVEF = left ventricular ejection fraction; MLHFQ = Minnesota Living with Heart Failure Questionnaire

**Table 3**

Cox Regression Modeling: Functional Status on Cardiac Event-free Survival (N = 313)

Variables	Hazard Ratio	Wald	Significance
<u>*Simple Cox Regression</u> Functional status (DASI score)	<b>.956</b>	<b>12.213</b>	<b>&lt; .001</b>
<u>**Multiple Cox Regression</u> Age	1.010	868	352
Gender	.512	5.067	.024
Ethnicity	1.423	1.380	.240
Education	.983	.234	.628
Living status	.735	1.613	.204
Financial status	.932	.135	.713
LVEF	.987	2.280	.131
Comorbidity	1.027	.199	.656
Taking ACEI	.801	.696	.404
Taking BB	.630	1.741	.187
<b>Functional status (DASI score)</b>	<b>.956</b>	<b>11.726</b>	<b>.001</b>

\* $\chi^2 = 12.876, p < .001$ ;

\*\* $\chi^2 = 29.015, p = 0.002$

ACEI = angiotensin-converting-enzyme inhibitor; BB = beta blocker; DASI = Duke Activity Status Index; LVEF = left ventricular ejection fraction