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Relationship of Race With Functional and Clinical Outcomes With the REHAB-HF Multidomain Physical Rehabilitation Intervention for Older Patients With Acute Heart Failure

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ORIGINAL RESEARCH

Relationship of Race With Functional and Clinical Outcomes With the REHAB-HF Multidomain Physical Rehabilitation Intervention for Older Patients With Acute Heart Failure

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BACKGROUND: The REHAB-HF (Rehabilitation Therapy in Older Acute Heart Failure Patients) randomized trial demonstrated that a 3-month transitional, tailored, progressive, multidomain physical rehabilitation intervention improves physical function, frailty, depression, and health-related quality of life among older adults with acute decompensated heart failure. Whether there is differential intervention efficacy by race is unknown.

METHODS AND RESULTS: In this prespecified analysis, differential intervention effects by race were explored at 3 months for physical function (Short Physical Performance Battery [primary outcome], 6-Minute Walk Distance), cognition, depression, frailty, health-related quality of life (Kansas City Cardiomyopathy Questionnaire, EuroQoL 5-Dimension-5-Level Questionnaire) and at 6 months for hospitalizations and death. Significance level for interactions was $P \le 0.1$. Participants (N=337, 97% of trial population) self-identified in near equal proportions as either Black (48%) or White (52%). The Short Physical Performance Battery intervention effect size was large, with values of 1.3 (95% Cl, 0.4–2.1; P=0.003]) and 1.6 (95% Cl, 0.8–2.4; P<0.001) in Black and White participants, respectively, and without significant interaction by race (P=0.56). Beneficial effects were also demonstrated in 6-Minute Walk Distance, gait speed, and health-related quality of life scores without significant interactions by race. There was an association between intervention and reduced all-cause rehospitalizations in White participants (rate ratio, 0.73 [95% Cl, 0.55–0.98]; P=0.034) that appears attenuated in Black participants (rate ratio, 1.06 [95% Cl, 0.81–1.41]; P=0.66; interaction P=0.067).

CONCLUSIONS: The intervention produced similarly large improvements in physical function and health-related quality of life in both older Black and White patients with acute decompensated heart failure. A future study powered to determine how the intervention impacts clinical events is required.

REGISTRATION: URL: https://www.clinicaltrials.gov. Identifier: NCT02196038.

Key Words: frailty
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CLINICAL PERSPECTIVE

What Is New?

 In this prespecified analysis of the REHAB-HF (Rehabilitation Therapy in Older Acute Heart Failure Patients) trial, the 3-month transitional, tailored, progressive multidomain physical rehabilitation intervention produced similarly large improvements in physical function and healthrelated quality of life in older Black and White patients with acute decompensated heart failure.

What Are the Clinical Implications?

- This analysis demonstrates the potential generalizability and robustness of the intervention and also suggests that the protocol, inclusive of comprehensive adherence and retention strategies, was successful in supporting both Black and White patients with varying baseline health determinants.
- Though the trial was not powered for definitive conclusions from clinical events, the intervention appeared possibly associated with a differential effect on 6-month all-cause rehospitalization rates by race, suggesting the need for a larger follow-up study with robust racial and ethnic representation.

Nonstandard Abbreviations and Acronyms

ADHF HRQOL	acute decompensated heart failure health-related quality of life
REHAB-HF	Rehabilitation Therapy in Older
	Acute Heart Failure Patients
SPPB	Short Physical Performance Battery

Severe impairments in physical function, which are infrequently identified or addressed in routine heart failure (HF) clinical care, may contribute to the persistently poor outcomes associated with acute decompensated heart failure (ADHF). Prior interventions have primarily focused on pharmaceutical-based targets. The REHAB-HF (Rehabilitation Therapy in Older Acute Heart Failure Patients) trial demonstrated that a transitional, tailored, progressive multidomain physical rehabilitation intervention improved physical function, health-related quality of life (HRQOL), and depression in older adults hospitalized with ADHF, regardless of ejection fraction (EF) phenotype.^{1,2} Whether there is differential efficacy of the intervention by race has not yet been explored.

Groups that have been historically marginalized in clinical practice and underrepresented in clinical studies are at increased risk for poor HF-related health outcomes. For example, across all age groups, Black individuals are more likely than White individuals to have an ADHF event, and death related to HF is significantly higher for Black compared with White individuals.^{3–5} A prior REHAB-HF subgroup analysis stratified by race and sex found substantial differences in many baseline characteristics, notably that White men had less severely impaired walking function than 3 other groups (Black men, Black women, and White women) despite adjustments for EF category and educational level and traditional cardiovascular disease risk factors (age, body mass index, comorbidity).⁶ In addition, White individuals were more likely to have the social support of living with a spouse or family member than Black individuals. Taken together, the differences found across these studies likely stem in part from inequities in social determinants of health such as health care policy and access, income level and economic stability, and neighborhood and built-environment guality. In turn, these disparities impact health-sustaining behaviors, health care usage, and HF self-management regimens and likely contribute to variable health outcomes. Related to the current study, there is the question of whether there is a variable response by race, a proxy for the constellation of socialized experiences, to the REHAB-HF intervention.⁷

There is currently limited literature exploring the effects of physical rehabilitation and exercise interventions by race in ADHF. In contrast, in patients who have established cardiovascular disease or are recovering from an acute cardiovascular disease event, non-White individuals have lower enrollment in and adherence to outpatient cardiac rehabilitation than White individuals, despite the well-described benefits associated with cardiac rehabilitation on long-term health outcomes.^{8–11} Thus, evaluating whether the effects of physical rehabilitation interventions vary across racial groups in ADHF is a critical first step toward the end goal of appropriately personalizing care to reduce disparities and optimize outcomes.

METHODS

Study Design

Details regarding trial design,^{12,13} interventional fidelity,¹⁴ cross-sectional baseline characteristics by race and sex as well as EF phenotype,^{6,15} overall trial results,¹ and EF subgroup analysis results² have been previously published. In brief, REHAB-HF was a randomized, controlled trial that examined a 3-month (12-week) transitional, tailored, progressive, one-onone (interventionist:participant), multidomain physical

Functional and Clinical Status by Race in REHAB-HF

rehabilitation intervention (focused on strength, balance, mobility, and endurance) in a population of older individuals following hospitalization for ADHF that met its primary end point of improving the Short Physical Performance Battery (SPPB) at 3 months in the intervention group. When possible, intervention sessions began on an inpatient basis and continued to outpatient for 12 weeks, 3 days weekly, totaling 36 sessions. Of the 7 participating hospitals in North Carolina and Pennsylvania, 4 were academic and 3 were community based. All centers approved study enrollment through their respective institutional review boards. All participants provided informed consent. Data requests will be considered on a case-by-case basis by the REHAB-HF steering committee.

Participants

The study included 349 individuals aged \geq 60 years hospitalized for \geq 24 hours for ADHF, including both HF with preserved EF and HF with reduced EF. Additional inclusion criteria included the ability to ambulate 4 meters (with or without an assistive device), functional independence before admission, and an expectation of being discharged home. Key exclusion criteria included acute myocardial infarction; end-stage HF, including those on inotropic agents or who may be considered for advanced HF therapies in the next 6 months; severe valvular disease; advanced kidney disease defined by an estimated glomerular filtration rate \leq 20 mL/min/1.73 m² or requiring dialysis; current enrollment in cardiac rehabilitation; or limitations with participation due to stroke, dementia, or other conditions.

In accordance with the US Office of Management and Budget minimum reporting standards for federally funded research, race and biological sex were selfreported. Due to small numbers and, as previously described,⁶ individuals who participated in the study and self-reported a race other than White or Black were included in the trial but not in race-specific analyses (n=12): Asian (n=4, 1%), American Indian or Alaska Native (n=7, 2%), and no racial identification (n=1, <1%). Among participants selecting multiple races, individuals who selected Black were categorized as Black (n=6), whereas individuals who selected White and any race other than Black were classified as White (n=1). The Consolidated Standards of Reporting Trials checklist for the primary trial and diagrams for the primary trial and current analysis are included in Figures S1 and S2.

Study Variables

Data collection was performed by individuals unaware of trial-group assignments before hospital discharge and at 3 months. The primary end point was physical function as measured by the SPPB, which has been validated in frail, older populations to predict numerous outcomes, including death.¹⁶ It comprises 3 components, including static standing balance; 4-meter gait speed; and time to perform 5 chair stands scored from 0 to 4 with a total score of 0 to 12, with higher scores indicating better physical function. Scores <10 are considered at risk for physical disability.¹⁷

Secondary outcomes captured at 3 months included physical function as assessed by 6-Minute Walk Distance¹⁸; heart failure symptoms and quality of life by Kansas City Cardiomyopathy Questionnaire¹⁹; and general quality of life by the EuroQol 5-Dimension-5-Level Questionnaire; depression by the Geriatric Depression Scale-15²⁰; cognitive function by the Montreal Cognitive Assessment²¹; and Frailty by the Fried Frailty Phenotype.²² See Table S1 for functional outcome variables, numerical ranges, and associated prognostic cutoffs. Additionally, rehospitalization, rehospitalization days, death, and falls were collected at 6 months through interviews and electronic health record review.

Statistical Analysis

Baseline characteristics were reported as mean (SD) or median (interguartile range) for continuous variables and frequency (percentage) for categorical variables. Differences in characteristics between White and Black participants were compared using t-tests and chisquare tests for continuous and categorical variables, respectively. To evaluate the potential moderating effect of race on the effect of the intervention on 3-month outcomes (SPPB total and component scores. Montreal Cognitive Assessment, Geriatric Depression Scale-15, Fried Frailty Criteria, 6-Minute Walk Distance, gait speed, Kansas City Cardiomyopathy Questionnaire, and EuroQol 5-Dimension-5-Level Questionnaire), general linear models that included indicator variables for intervention, race, and their interaction were used. All analyses were adjusted for 2 stratification factors used for randomization, EF category of <45% or $\ge 45\%$ and clinical site as well as 2 other established independent predictors of HF outcomes-baseline age and sex-as in other analyses of REHAB-HF trial data. Furthermore, as baseline age and sex differed by race and EF trended to differ by race, they could be confounders in the association of race with outcomes. We used least square means to estimate the effects of the intervention in race groups, and effect sizes were reported with 95% Cls.

The moderating effect of race on the effect of the intervention on 6-month clinical outcomes was assessed using Poisson regression for all-cause rehospitalizations, HF rehospitalizations, and deaths; negative binomial regression for facility-free days and hospitalized days due to overdispersion; and logistic regression for the presence of any falls during the study. All analyses were adjusted for age, sex, EF category, and clinical

	White				Black				
Characteristics	All (n=175)	Rehabilitation (n=93)	Attention control (n=82)	Within-group P value	All (n=162)	Rehabilitation (n=76)	Attention control (n=86)	Within- group <i>P</i> value	Between- race <i>P</i> value
Age, y	74.6 (8.3)	74.9 (8.6)	74.3 (8.1)	0.61	70.4 (7.2)	70.8 (7.5)	70.2 (7.0)	0.61	<0.0001
Female	76 (43.4)	37 (39.8)	39 (47.6)	0.30	103 (63.6)	47 (61.8)	56 (65.1)	0.67	<0.001
Ejection fraction ≥45%	101 (57.7)	52 (55.9)	49 (59.8)	0.61	77 (47.5)	39 (51.3)	38 (44.2)	0.06	0.061
NYHA class									
	32 (18.3)	16 (17.2)	16 (19.5)	0.69	31 (19.1)	16 (21.1)	15 (17.4)	0.67	0.24
=	102 (58.3)	57 (61.3)	45 (54.9)		81 (50)	39 (51.3)	42 (48.8)	1	
2	41 (23.4)	20 (21.5)	21 (25.6)		50 (30.9)	21 (27.6)	29 (33.7)		
BMI, kg/m ²	31.9 (8.6)	32.3 (8.5)	31.4 (8.7)	0.50	33.8 (8.2)	33.6 (7.8)	33.9 (8.6)	0.83	0.040
BNP, pg/mL (n=197)*	516 (278–1041)	583 (306–1038)	500 (193–1104)	0.27	669 (374–1292)	689 (430–1089)	642 (304–1381)	0.85	0.25
N-terminal proBNP, pg/mL (n=116)*	3549 (1939–6983)	3688.0 (1969–9074)	3464.0 (1793–4547)	0.49	2796 (1425–5358)	3131 (1492–6640)	2488 (1357–5174)	0.27	0.37
Index hospital LOS, days	5.0 (3.0-7.0)	5.0 (3.0–7.0)	5.0 (3.0–7.0)	0.37	4.0 (3.0–7.0)	4.0 (3.0-7.0)	5.0 (3.0–7.0)	0.19	0.17
Hospitalization in past 6 mo	69 (39.4)	34 (36.6)	35 (42.7)	0.41	80 (49.4)	38 (50.0)	42 (48.8)	0.88	0.066
HF Hospitalization in past 6 mo	40 (22.9)	20 (21.5)	20 (24.4)	0.70	45 (27.8)	20 (26.3)	25 (29.1)	0.82	0.18
Fall in past 3 mo [‡]	22 (14.8)	10 (12.8)	12 (16.9)	0.49	21 (16.2)	13 (20.6)	8 (11.9)	0.18	0.75
Comorbidities									
Total number	5.4 (2.0)	5.5 (2.2)	5.2 (1.8)	0.37	5.0 (1.8)	5.2 (1.9)	4.8 (1.8)	0.13	0.043
Hypertension	155 (88.6)	81 (87.1)	74 (90.2)	0.51	155 (95.7)	73 (96.1)	82 (95.3)	0.83	0.016
History of MI	34 (19.4)	16 (17.2)	18 (22.0)	0.43	25 (15.4)	12 (15.8)	13 (15.1)	0.91	0.33
History of coronary revascularization	70 (40)	38 (40.9)	32 (39.0)	0.80	28 (17.3)	15 (19.7)	13 (15.1)	0.44	<0.0001
Atrial fibrillation	109 (62.3)	55 (59.1)	54 (65.9)	0.36	61 (37.7)	30 (39.5)	31 (36.0)	0.65	<0.0001
Diabetes	84 (48)	52 (55.9)	32 (39.0)	0.026	94 (58)	48 (63.2)	46 (53.5)	0.21	0.066
Hyperlipidemia	114 (65.1)	58 (62.4)	56 (68.3)	0.41	109 (67.3)	48 (63.2)	61 (70.9)	0.29	0.68
COPD	52 (29.7)	30 (32.3)	22 (26.8)	0.43	43 (26.5)	23 (30.3)	20 (23.3)	0.31	0.52
CKD	55 (31.4)	28 (30.1)	27 (32.9)	0.69	57 (35.2)	29 (38.2)	28 (32.6)	0.46	0.46
Stroke	21 (12)	9 (9.7)	12 (14.6)	0.31	27 (16.7)	14 (18.4)	13 (15.1)	0.57	0.22
PVD	23 (13.1)	16 (17.2)	7 (8.5)	0.09	17 (10.5)	11 (14.5)	6 (7.0)	0.12	0.45
Arthritis, muscle/joint pain, CTD	81 (46.3)	46 (49.5)	35 (42.7)	0.37	68 (42)	36 (47.4)	32 (37.2)	0.19	0.44
History of cancer	44 (25.1)	28 (30.1)	16 (19.5)	0.11	28 (17.3)	11 (14.5)	17 (19.8)	0.37	0.079
Sleep-disordered breathing	57 (32.6)	35 (37.6)	22 (26.8)	0.13	65 (40.1)	32 (42.1)	33 (38.4)	0.63	0.15
Depression	44 (25.1)	22 (23.7)	22 (26.8)	0.63	17 (10.5)	7 (9.2)	10 (11 6)	0.61	50001

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	White				Black				
Characteristics	All (n=175)	Rehabilitation (n=93)	Attention control (n=82)	Within-group P value	All (n=162)	Rehabilitation (n=76)	Attention control (n=86)	Within- group <i>P</i> value	Between- race <i>P</i> value
Dementia or cognitive impairment	4 (2.3)	2 (2.2)	2 (2.4)	0.90	5 (3.1)	3 (3.9)	2 (2.3)	0.55	0.65
Urinary incontinence [†]	22 (15)	12 (15.8)	10 (14.1)	0.77	17 (13.2)	6 (9.5)	11 (16.7)	0.23	0.67
BMI indicates body mass index; BNP, B-type natriuretic peptide; CKD, chronic kidney disease; COPD, chronic obstructive pulmonary disease; CTD, connective tissue disease; HF, heart failure; LOS, length of stay; Mi, myocardial infarction; N-terminal pro-BNP, N-terminal pro-B-type natriuretic peptide; NYHA, New York Heart Association; and PVD, peripheral vascular disease. *Presented as n (%), mean (SD), or median (interquartile range). *Data collected in Attention Control=137. Rehabilitation Intervention=139.	B-type natriuretic pepti BNP, N-terminal pro-B-t dian (interquartile range 37. Rehabilitation Interve	ide; CKD, chronic kidne. :ype natriuretic peptide; .).	y disease; COPD, chro NYHA, New York Hear	nic obstructive pul t Association; and I	monary disease; PVD, peripheral v	CTD, connective tiss ascular disease.	ue disease; HF, h	eart failure; LOS	3, length of sta

site. All-cause rehospitalization was also adjusted for baseline SPPB score as prespecified. Effect sizes for White and Black participants were summarized as rate ratio for count-based outcomes and odds ratio for binary outcomes. We also performed a sensitivity analysis to investigate the effect of baseline hospitalizations on the 6-month all-cause rehospitalizations. A stratified analysis was conducted to evaluate the moderating effect of race on the effect of the intervention in patients with and without any baseline hospitalization.

A *P* value of <0.05 was prespecified as statistically significant for overall comparisons. The interaction between race and the intervention was prespecified as significant if P<0.10. Due to the hypothesis-generating nature of the analysis, there was no correction for multiple comparisons.

RESULTS

Baseline Characteristics

Of the 349 participants included in the REHAB-HF trial, 337 self-identified as Black (48%; 76 intervention, 86 control) or White (52%; 93 intervention, 82 control) individuals. Tables 1 and 2 present the baseline characteristics by race as well as treatment arm. Black participants tended to be younger than White participants (mean age, 70.8 ± 7.5 versus 74.6 ± 8.3 years; *P*<0.001) and to have a higher body mass index (33.6 ± 8.2 versus 31.9 ± 8.6 kg/m²; *P*=0.040). Black participants were more likely to be women (64% versus 43%; *P*<0.001).

Overall, comorbidity burden was high across both races. Black participants had a lower total comorbidity count than White participants (5.0 versus 5.4; P=0.043; Table 1). Black participants were more likely to have hypertension (96% versus 87%; P=0.016) and were less likely to have atrial fibrillation (38% versus 62%; P<0.001) and to have a diagnosis of depression (11%) versus 25%; P<0.001). Though Black and White participants had a similar prevalence of myocardial infarction, Black participants were less likely to have prior revascularization (17% versus 40%; P<0.001). Medical and device therapies for HF were evenly distributed between both races (Table 2). There was a nonsignificant trend for Black participants to have had a hospitalization in the preceding 6 months compared with White participants (49% versus 39%; P=0.066; Table 1). This appeared to be due to more non-HF hospitalizations as there was no difference in baseline HF hospitalizations between Black and White individuals (17% versus 22%; P=0.18; Table 1).

Adherence and Retention Metrics

Of the 169 participants randomized to intervention, 11 died before completing the intervention (6 Black participants, 5 White participants), 13 dropped from the

Continued

able 1.

Data collected in Attention Control=138, Rehabilitation Intervention=141.

	White				Black				
Characteristics	All (n=175)	Rehabilitation (n=93)	Attention control (n=82)	Within- group <i>P</i> value	All (n=162)	Rehabilitation (n=76)	Attention control (n=86)	Within- group <i>P</i> value	Between- race <i>P</i> value
Loop diuretic	164 (94.3)	88 (94.6)	76 (93.8)	0.83	150 (92.6)	68 (89.5)	82 (95.3)	0.15	0.54
Beta blocker	134 (77)	73 (78.5)	61 (75.3)	0.62	133 (82.1)	61 (80.3)	72 (83.7)	0.25	0.25
Angiotensin-converting enzyme inhibitors	61 (35.1)	32 (34.4)	29 (35.8)	0.85	65 (40.1)	31 (40.8)	34 (39.5)	0.34	0.34
Angiotensin II receptor blockers	36 (20.7)	19 (20.4)	17 (21.0)	0.93	37 (22.8)	18 (23.7)	19 (22.1)	0.63	0.63
Aldosterone antagonists	34 (19.5)	19 (20.4)	15 (18.5)	0.75	28 (17.3)	9 (11.8)	19 (22.1)	0.09	0.59
Digoxin	10 (5.7)	6 (6.5)	4 (4.9)	0.67	8 (4.9)	1 (1.3)	7 (8.1)	0.74	0.74
Insulin	47 (27)	27 (29.0)	20 (24.7)	0.52	47 (29)	25 (32.9)	22 (25.6)	0.31	0.68
Oral diabetic agents	41 (23.6)	28 (30.1)	13 (16.0)	0.029	41 (25.3)	22 (28.9)	19 (22.1)	0.32	0.71
Implantable cardioverter-defibrillator	28 (16)	16 (17.2)	12 (14.6)	0.64	29 (17.9)	13 (17.1)	16 (18.6)	0.80	0.64
Biventricular pacemaker	16 (9.1)	9 (9.7)	7 (8.5)	0.79	8 (4.9)	3 (3.9)	5 (5.8)	0.58	0.13

*Presented as n (%), mean (SD), or median (interquartile range).

study (6 Black participants, 7 White participants), and 16 discontinued the intervention but contributed to the primary outcome assessment (10 Black participants, 6 White participants). Overall intervention retention was 82% (ie, participants who did not drop from the study or die). There were no significant differences between Black and White participants in adherence to the intervention. Of all participants retained in the intervention, Black and White participants completed 24.8 ± 1.5 sessions ($69 \pm 4\%$ adherence) and 27.0 ± 1.1 sessions ($75 \pm 3\%$ adherence), respectively (P=0.24). After accounting for sessions missed due to illness and medical appointments, intervention adherence rates were $76 \pm 4\%$ and $82 \pm 3\%$ for Black and White participants, respectively (P=0.26).

Three-Month Functional Outcomes

Functional outcomes at 3 months are delineated by race in Figure 1 and Table 3. Improvements in the primary trial outcome, SPPB, were similarly large in both Black and White participants with respective differences between mean intervention and control of 1.3 (95% CI, 0.4–2.1) units and 1.6 (95% CI, 0.8–2.4) units (both P<0.01) and no significant interaction (P=0.56). Additionally, there were large improvements in gait speed in both Black and White participants with mean intervention and control value differences of 0.12 m/s (95% CI, 0.06–0.18; P<0.001) and 0.12m/s (95% CI, 0.06–0.18; P<0.001), respectively, and no significant interaction (P=0.96).

There were no significant interactions in other 3month outcomes (Figure 1, Table 3). These included 6-Minute Walk Distance with mean differences between study groups in respective Black and White

participants with gains of 31.1 m (95% CI, -1.7 to 64.0; P=0.06) versus 37.6 m (95% Cl, 6.1-69.1; P=0.020; interaction P=0.78). There were no significant differences in HRQOL as captured by the Kansas City Cardiomyopathy Questionnaire-Overall Summary Score with a difference in mean intervention and control values of 7.6 (95% Cl, 0.1-15.2, P=0.05) for Black participants and 6.3 (95% Cl, -1.0 to 13.6; P=0.09) for White participants (interaction P=0.81). A similar pattern was observed for EuroQol 5-Dimension-5-Level Questionnaire utility weights with a difference of mean intervention and control values of 6.3 (95% Cl, -0.7 to 13.3; P=0.08) for Black participants and 7.5 (95% Cl. 0.8-14.3; P=0.029) for White participants (interaction P=0.81). Differences between mean values for the intervention and control groups among Black and White patients on cognition, depression, and frailty are reported in Table 3.

Six-Month Clinical Events

There were no significant interactions between study groups and race in 6-month HF rehospitalization, all-cause death, rehospitalization and death, or injurious falls (Figure 1, Table 4). There was a significant interaction (P=0.067) in the rate ratios of all-cause rehospitalization in Black versus White participants with respective rate ratios of 1.06 (95% CI, 0.81–1.41; P=0.66) versus 0.73 (95% CI, 0.55–0.98; P=0.034). All-cause rehospitalization rates were similar among Black participants in both the intervention and control groups (1.38 and 1.24) as well as the White participants in the control group (1.33), while the rate was lower among White participants in the intervention group (0.98). Given that the percentage of Black participants



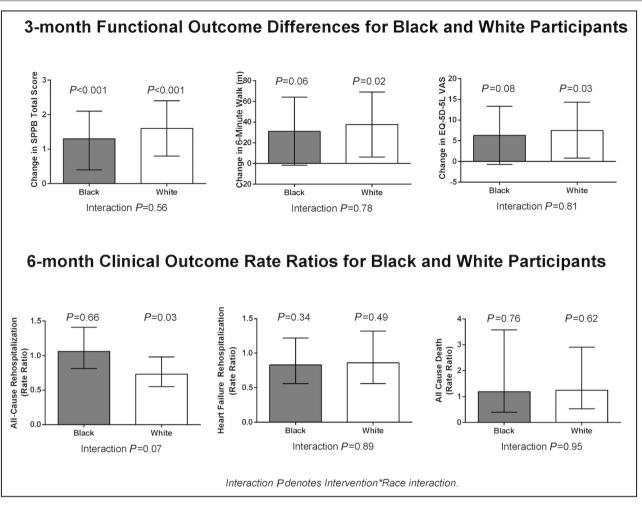


Figure 1. Functional outcome differences for Black and White participants.

Three-month functional outcomes and 6-month clinical outcomes for Black and White participants. Bars represent intervention vs control effect sizes (95% CI error bars). Interaction *P* values below each panel represent intervention*race interaction.

hospitalized within 6 months before baseline showed a nonsignificant trend to be higher than White participants (49% versus 39%; P=0.066), an additional analysis of all-cause rehospitalizations was performed examining the effects of recent baseline hospitalization. The results did not reveal significant variations in intervention effects on those with prior hospitalization for Black (Rate Ratio, 0.77 [95% Cl, 0.52–1.14]; P=0.20) or White participants (RR, 1.15 [95% Cl, 0.81–1.62]; P=0.43) or an interaction between the study intervention and race (P=0.14). Similar patterns were observed in participants without a hospitalization at baseline.

DISCUSSION

Key Findings

In this prespecified secondary analysis of the multicenter REHAB-HF trial, the innovative, 3-month transitional, tailored, progressive multidomain physical rehabilitation intervention in older adults with ADHF had similar adherence rates and produced similarly large improvements across multiple measures of physical function, including the primary trial outcome SPPB, and HRQOL for Black and White individuals, despite variable baseline characteristics. Although the trial was not sufficiently powered to draw definitive conclusions from clinical events, the intervention appeared potentially associated with lower 6-month all-cause rehospitalization rates in White participants, supporting the need for larger follow-up studies with robust recruitment of racial and ethnic diversity. Overall, this analysis demonstrates the potential generalizability and robustness of the intervention regardless of race and also suggests that the protocol, inclusive of comprehensive effective adherence and retention strategies across races, was successful in overcoming baseline demographic, social, and physical health-determinant differences between Black and White participants.

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J Am Heart Assoc.	2023;12:e030588.	DOI: 10.1161/JAH/	A.123.030588

Difference Rehab intervention (95% Cl) Rehab prevention (1=0.08 to 2.4) Rehab intervention (n=76) Attention (n=76) Difference (95% Cl) P value 1.6 (0.8 to 2.4) <0.001 8.0 (0.3) 6.7 (0.3) 1.3 (0.4 to 2.1) 0.003 0.4 (0.1 to 0.8) 0.024 3.2 (0.2) 3.0 (0.2) 0.3 (-0.1 to 0.7) 0.15 0.5 (0.1 to 0.8) 0.007 2.1 (0.2) 1.3 (0.1) 0.3 (-0.1 to 0.7) 0.15 0.5 (0.1 to 0.8) 0.007 2.1 (0.2) 1.3 (0.1) 0.3 (-0.1 to 0.7) 0.15 0.5 (0.1 to 0.8) 0.007 2.1 (0.2) 1.3 (0.1) 0.3 (-0.1 to 0.7) 0.15 0.5 (0.1 to 0.8) 0.007 2.1 (0.2) 1.3 (0.1) 0.3 (-0.1 to 0.6) 0.36 0.5 (0.1 to 6.8) 0.10 0.10 0.3 (0.2) 0.3 (-1.7 to 0.6) 0.36 0.12 (0.06 to 0.18) 0.10 0.12 (0.02) 0.12 (0.02) 0.16 0.16 0.12 (0.08 to 0.18) 0.10 0.12 (0.02) 0.12 (0.02) 0.12 0.06 0.12 (0.08 to 1.42) 0.09 0.12 (0.02)		White				Black				latonion tion
me 8.1 (0.3) 6.5 (0.3) 1.6 (0.8 to 2.4) <0.001		Rehabilitation intervention (n=93)	Attention control (n=82)	Difference (95% CI)	P value	Rehab intervention (n=76)	Attention control (n=86)	Difference (95% CI)	P value	by race interaction, <i>P</i> value
8.1 (0.3) $6.5 (0.3)$ $1.6 (0.8 tr 2.4)$ < 0.001 $8.0 (0.3)$ $6.7 (0.3)$ $1.3 (0.4 to 2.1)$ 0.003 $3.1 (0.1)$ $2.7 (0.1)$ $2.7 (0.1)$ $2.7 (0.1)$ $0.4 (0.1 to 0.8)$ 0.224 $3.2 (0.2)$ $3.0 (0.2)$ $0.3 (-0.1 to 0.7)$ 0.15 $5core$ $3.1 (0.1)$ $2.5 (0.1)$ $2.6 (0.1)$ $0.6 (0.3 to 0.8)$ 0.024 $3.2 (0.2)$ $3.0 (0.2)$ $0.3 (0.1 to 0.7)$ 0.15 $5core$ $1.9 (0.1)$ $1.4 (0.1)$ $0.5 (0.1 to 0.8)$ 0.007 $2.1 (0.2)$ $1.3 (0.1)$ $0.3 (0.4 to 1.2)$ 0.08 $5.3 (0.5)$ $2.3.4 (0.5)$ $0.5 (0.1 to 0.8)$ 0.007 $2.1 (0.2)$ $1.3 (0.1)$ $0.8 (0.4 to 1.2)$ 0.06 $5.3 (0.5)$ $2.3.4 (0.5)$ $0.7 (-1.2 to 1.1)$ 0.87 $2.1 (0.2)$ $2.1 (0.5)$ $0.8 (0.4 to 1.2)$ $0.06 (-1.7 to 0.6)$ $5.3 (0.5)$ $2.3.3 (0.5)$ $2.3.4 (0.5)$ $-0.1 (-1.2 to 1.1)$ 0.87 $2.1 (0.5)$ $2.2.1 (0.5)$ $0.8 (-1.7 to 0.6)$ $2.3.3 (0.5)$ $3.7 (0.3)$ $0.7 (-1.5 to 0.1)$ 0.87 $0.8 (0.2)$ $0.6 (-1.7 to 0.6)$ 0.36 $7.0 (0.3)$ $2.3.6 (0.3)$ $0.7 (0.3)$ 0.10 $1.3 (0.1)$ $0.06 (-1.7 to 0.6)$ 0.16 $5.0 (0.3)$ $0.10 (0.2)$ $0.10 (0.2)$ $0.10 (0.2)$ $0.10 (-1.2 to 0.2)$ $0.11 (-1.2 to 0.2)$ $5.0 (0.3)$ $0.23 (0.7 to 0.2)$ $0.21 (0.5)$ $0.21 (0.5)$ $0.21 (0.5)$ $0.11 (-1.7 to 0.6)$ $0.10 (0.2)$ $0.24 (0.2)$ $0.72 (0.23)$ $0.12 (0.03)$ <	Primary outcome									
31 (0.1) 2.7 (0.1) 0.4 (0.1 to 0.8) 0.024 3.2 (0.2) 3.0 (0.2) 0.3 (-0.1 to 0.7) 0.15 score 3.1 (0.1) 2.5 (0.1) 2.5 (0.1) 0.3 (0.0 to 0.6) 0.08 0.08 $score$ 3.1 (0.1) 1.4 (0.1) 0.5 (0.1 to 0.8) 0.007 2.1 (0.2) 1.3 (0.1) 0.3 (0.0 to 0.6) 0.08 s 1.9 (0.1) 1.4 (0.1) 0.5 (0.1 to 0.8) 0.007 2.1 (0.2) 1.3 (0.1) 0.3 (0.4 to 1.2) <0.001 s $2.3.3$ (0.5) $2.3.4$ (0.5) 0.5 (0.1 to 0.8) 0.007 2.1 (0.2) 1.3 (0.1) 0.8 (0.4 to 1.2) <0.001 s $2.3.3$ (0.5) $2.3.4$ (0.5) 0.5 (0.1 to 0.8) 0.07 2.1 (0.5) $2.2.1$ (0.5) 0.6 (1.7 to 0.6) 0.35 s 0.03 3.7 (0.3) -0.1 (-1.2 to 1.1) 0.87 216 (0.5) 2.21 (0.5) 0.6 (1.7 to 0.6) 0.35 s 0.03 3.7 (0.3) 0.7 (0.3) 0.7 (1.5 to 0.1) 0.07 3.6 (0.3) 0.6 (1.7 to 0.6) 0.35 s 0.03 3.7 (0.3) 0.7 (1.5 to 0.1) 0.07 0.07 0.36 (1.7 to 0.6) 0.35 s 0.03 0.35 0.34 (1.2) 0.36 (1.1 to 0.6) 0.35 0.35 0.35 0.35 s 0.02 0.29 0.02 2.21 (0.5) 0.36 (1.3) 0.11 (1.7 to 0.6) 0.05 s 0.12 (0.03) 0.12 (0.01) 0.02 0.29 (1.12) 0.12 (0.02) 0.12 (0.0	SPPB	8.1 (0.3)	6.5 (0.3)	1.6 (0.8 to 2.4)	<0.001	8.0 (0.3)	6.7 (0.3)	1.3 (0.4 to 2.1)	0.003	0.56
31 (0.1) 2.5 (0.1) 0.6 (0.3 to 0.9) <0.001 2.8 (0.1) 2.5 (0.1) 0.3 (0.0 to 0.6) 0.08 1.9 (0.1) 1.4 (0.1) 0.5 (0.1 to 0.8) 0.007 2.1 (0.2) 1.3 (0.1) 0.3 (0.4 to 1.2) <0.001	Balance score	3.1 (0.1)	2.7 (0.1)	0.4 (0.1 to 0.8)	0.024	3.2 (0.2)	3.0 (0.2)	0.3 (-0.1 to 0.7)	0.15	0.61
1:9(0.1) 1.4(0.1) 0.5(0.1 to 0.8) 0.007 2.1(0.2) 1.3(0.1) 0.8(0.4 to 1.2) <0.001 23.3(0.5) 23.4(0.5) 23.4(0.5) -0.1(-1.2 to 1.1) 0.87 21.6(0.5) 22.1(0.5) -0.6(-1.7 to 0.6) 0.35 3.0(0.3) 3.7(0.3) -0.7(-1.5 to 0.1) 0.10 3.6(0.3) 22.1(0.5) -0.6(-1.7 to 0.6) 0.35 1.4(0.1) 1.7(0.1) 0.23(-0.6 to 0.1) 0.10 1.5(0.1) 1.8(0.2) -0.3(-0.6 to 0.1) 0.13 286.9(12.7) 249.3(13.8) 37.6(6.1 to 68.1) 0.10 1.5(0.1) 1.8(0.2) 0.31(-1.7 to 64.0) 0.13 0.84(0.02) 0.72(0.03) 0.12 0.001 0.79(0.03) 0.67(0.02) 0.12 0.06 1.3(2.9) 66.2(3.1) 0.17 66.1(3.1) 66.7(3.1) 0.17 61.02) 0.13 0.13 0.13 1.1.3(2.9) 66.2(3.1) 61.10 1.5(0.1) 1.1(2.9) 0.67 0.12 0.13 0.13 0.13 0.13 0.13 0.13 0.13	4-m gait speed score	3.1 (0.1)	2.5 (0.1)	0.6 (0.3 to 0.9)	<0.001	2.8 (0.1)	2.5 (0.1)	0.3 (0.0 to 0.6)	0.08	0.16
23.3 (0.5) 23.4 (0.5) -0.1 (-1.2 to 1.1) 0.87 21.6 (0.5) 22.1 (0.5) -0.6 (-1.7 to 0.6) 0.35 3.0 (0.3) 3.7 (0.3) 3.7 (0.3) -0.7 (-1.5 to 0.1) 0.87 21.6 (0.5) 22.1 (0.5) -0.6 (-1.7 to 0.6) 0.35 3.0 (0.3) 3.7 (0.3) -0.7 (-1.5 to 0.1) 0.10 3.6 (0.3) 4.3 (0.3) -0.6 (-1.5 to 0.2) 0.15 1.4 (0.1) 1.7 (0.1) -0.3 (-0.6 to 0.1) 0.10 1.5 (0.1) 1.8 (0.2) -0.3 (-0.6 to 0.1) 0.13 286.9 (12.7) 249.3 (13.8) 37.6 (6.1 to 69.1) 0.10 1.5 (0.1) 1.8 (0.2) 0.31 (-1.7 to 64.0) 0.13 0.84 (0.02) 0.72 (0.03) 0.72 (0.03) 0.12 (1.0 to 13.6) 0.07 0.07 (0.03) 0.67 (0.02) 0.12 (0.6 to 0.18) 0.01 71.3 (2.9) 66.2 (3.1) 61.3 (13.1) 0.17 (14.2) 66.0 (13.9) 0.12 (0.02) 0.13 0.13 0.14 0.16 71.3 (2.9) 66.2 (3.1) 61.3 (13.9) 0.56 (1.0 co 13.6) 0.12 (0.02) 0.12 (0.02) 0.12 (0.02) 0	Chair rise score	1.9 (0.1)	1.4 (0.1)	0.5 (0.1 to 0.8)	0.007	2.1 (0.2)	1.3 (0.1)	0.8 (0.4 to 1.2)	<0.001	0.21
23.3 (0.5) 23.4 (0.5) -0.1 (-1.2 to 1.1) 0.87 21.6 (0.5) 22.1 (0.5) -0.6 (-1.7 to 0.6) 0.35 3.0 (0.3) 3.7 (0.3) 3.7 (0.3) -0.7 (-1.5 to 0.1) 0.10 3.6 (0.3) 4.3 (0.3) -0.6 (-1.7 to 0.6) 0.35 1.4 (0.1) 1.7 (0.1) -0.7 (-1.5 to 0.1) 0.10 1.5 (0.1) 1.8 (0.2) 0.6 (-1.5 to 0.2) 0.15 286.9 (12.7) 249.3 (13.8) 37.6 (6.1 to 69.1) 0.10 1.5 (0.1) 1.8 (0.2) 20.6 (1.7) 0.13 286.9 (12.7) 249.3 (13.8) 37.6 (6.1 to 69.1) 0.020 292.0 (14.2) 260.8 (13.9) 31.1 (-1.7 to 64.0) 0.13 0.84 (0.02) 0.72 (0.03) 0.12 (0.06 to 0.18) <0.001	Secondary outcomes									
3.0 (0.3) 3.7 (0.3) -0.7 (-1.5 to 0.1) 0.10 3.6 (0.3) 4.3 (0.3) -0.6 (-1.5 to 0.2) 0.15 1.4 (0.1) 1.7 (0.1) -0.3 (-0.6 to 0.1) 0.10 1.5 (0.1) 1.8 (0.2) -0.3 (-0.6 to 0.1) 0.13 286.9 (12.7) 249.3 (13.8) 37.6 (6.1 to 69.1) 0.10 1.5 (0.1) 1.8 (0.2) -0.3 (-0.6 to 0.1) 0.13 0.84 (0.02) 0.72 (0.03) 0.72 (0.06 to 0.18) <0.001	MoCA	23.3 (0.5)	23.4 (0.5)	-0.1 (-1.2 to 1.1)	0.87	21.6 (0.5)	22.1 (0.5)	-0.6 (-1.7 to 0.6)	0.35	0.58
1.4 (0.1) 1.7 (0.1) -0.3 (-0.6 to 0.1) 0.10 1.5 (0.1) 1.8 (0.2) -0.3 (-0.6 to 0.1) 0.13 286.9 (12.7) 249.3 (13.8) 37.6 (6.1 to 69.1) 0.020 292.0 (14.2) 260.8 (13.9) 31.1 (-1.7 to 64.0) 0.06 0.84 (0.02) 0.72 (0.03) 0.12 (0.06 to 0.18) <0.001	GDS-15	3.0 (0.3)	3.7 (0.3)	-0.7 (-1.5 to 0.1)	0.10	3.6 (0.3)	4.3 (0.3)	-0.6 (-1.5 to 0.2)	0.15	0.91
286.9 (12.7) 249.3 (13.8) 37.6 (6.1 to 69.1) 0.020 292.0 (14.2) 260.8 (13.9) 31.1 (-1.7 to 64.0) 0.06 0.84 (0.02) 0.72 (0.03) 0.12 (0.06 to 0.18) <0.001	Fried Frailty Criteria	1.4 (0.1)	1.7 (0.1)	-0.3 (-0.6 to 0.1)	0.10	1.5 (0.1)	1.8 (0.2)		0.13	0.98
0.84 (0.02) 0.72 (0.03) 0.12 (0.06 to 0.18) <0.001 0.79 (0.03) 0.67 (0.02) 0.12 (0.06 to 0.18) <0.001 71.3 (2.9) 66.2 (3.1) 5.1 (-2.2 to 12.4) 0.17 66.1 (3.1) 60.2 (3.0) 5.9 (-1.7 to 13.5) 0.13 71.6 (2.9) 65.3 (3.0) 6.3 (-1.0 to 13.6) 0.09 66.5 (3.1) 58.9 (3.0) 7.6 (0.1 to 15.2) 0.05 72.3 (2.7) 64.8 (2.8) 7.5 (0.8 to 14.3) 0.029 71.1 (2.9) 64.7 (2.8) 6.3 (-0.7 to 13.3) 0.08	6MWD	286.9 (12.7)	249.3 (13.8)	37.6 (6.1 to 69.1)	0.020	292.0 (14.2)	260.8 (13.9)	31.1 (-1.7 to 64.0)	0.06	0.78
S 71.3 (2.9) 66.2 (3.1) 5.1 (-2.2 to 12.4) 0.17 66.1 (3.1) 60.2 (3.0) 5.9 (-1.7 to 13.5) 0.13 S 71.6 (2.9) 65.3 (3.0) 6.3 (-1.0 to 13.6) 0.09 66.5 (3.1) 58.9 (3.0) 7.6 (0.1 to 15.2) 0.05 72.3 (2.7) 64.8 (2.8) 7.5 (0.8 to 14.3) 0.029 71.1 (2.9) 64.7 (2.8) 6.3 (-0.7 to 13.3) 0.08	Gait speed, m/sec	0.84 (0.02)	0.72 (0.03)	0.12 (0.06 to 0.18)	<0.001	0.79 (0.03)	0.67 (0.02)	0.12 (0.06 to 0.18)	<0.001	0.96
S 71.6 (2.9) 65.3 (3.0) 6.3 (-1.0 to 13.6) 0.09 66.5 (3.1) 58.9 (3.0) 7.6 (0.1 to 15.2) 0.05 72.3 (2.7) 64.8 (2.8) 7.5 (0.8 to 14.3) 0.029 71.1 (2.9) 64.7 (2.8) 6.3 (-0.7 to 13.3) 0.08	KCCQ-CSS	71.3 (2.9)	66.2 (3.1)	5.1 (-2.2 to 12.4)	0.17	66.1 (3.1)	60.2 (3.0)	5.9 (-1.7 to 13.5)	0.13	0.88
72.3 (2.7) 64.8 (2.8) 7.5 (0.8 to 14.3) 0.029 71.1 (2.9) 64.7 (2.8) 6.3 (-0.7 to 13.3) 0.08	KCCQ-OSS	71.6 (2.9)	65.3 (3.0)	6.3 (-1.0 to 13.6)	0.09	66.5 (3.1)	58.9 (3.0)	7.6 (0.1 to 15.2)	0.05	0.81
	EQ-5D-5L	72.3 (2.7)	64.8 (2.8)	7.5 (0.8 to 14.3)	0.029	71.1 (2.9)	64.7 (2.8)	6.3 (-0.7 to 13.3)	0.08	0.81

Table 3. Analysis of 3-Month Functional Outcomes by Race

6MWD, 6-Minute Walk Distance; CSS, Clinical Summary Score; EQ-5D-5L, EuroQol 5-Dimension 5-Level Questionnaire GDS-15, Geriatric Depression Scale-15; HRQOL, Health-Related Quality of Life; KCCQ, Kansas City Cardiomyopathy Questionnaire; MoCA, Montreal Cognitive Assessment; OSS, Overall Summary Score; SBBP, Short Physical Performance Battery.

	White				Black				Internetion
	Rehabilitation intervention (n=93)	Attention control (n=82)	Rate ratio (95% CI)	P value	Rehabilitation intervention (n=76)	Attention control (n=86)	Rate ratio (95% CI)	P value	by race interaction, <i>P</i> value
Secondary outcomes									
All-cause rehospitalization	84 (0.98)	104 (1.33)	0.73 (0.55–0.98)	0.034	100 (1.38)	102 (1.24)	1.06 (0.81–1.41)	0.66	0.067
HF rehospitalization	41 (0.48)	44 (0.56)	0.86 (0.56–1.32)	0.49	45 (0.62)	59 (0.72)	0.83 (0.56–1.22)	0.34	0.89
All-cause death	13 (0.15)	9 (0.11)	1.24 (0.53–2.91)	0.62	7 (0.10)	6 (0.07)	1.19 (0.40–3.58)	0.76	0.95
Rehospitalizationand death	97 (1.13)	113 (1.44)	0.79 (0.60–1.04)	0.09	107 (1.48)	108 (1.31)	1.06 (0.81–1.38)	0.69	0.14
Falls*	25 (0.27)	32 (0.39)	0.59 (0.31–1.13)	0.11	21 (0.28)	29 (0.34)	0.72 (0.36–1.41)	0.34	0.70

Analvsis of 6-Month Clinical Outcomes by Race Table 4.

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sex, ejection fraction category, and clinical site. All-cause All analyses were adjusted for age, Ô. *count (proportion) and odds ratio (95% (95% CI) except ehospitalization was also adjusted for baseline SPPB score as prespecified. Data are presented as counts (6-mile rate) and rate ratio

Short Physical Performance Battery failure; and SBBP, HF indicates heart

Impact of Personalized Care on **Outcomes**

Population health level data indicate that both sociodemographic and physical characteristics can predispose Black individuals to a higher incidence and prevalence of HF as well as a worse prognosis compared with White individuals.²³ One of the most significant risk factors for HF is hypertension, which is more common in the Black population and was reflected in our sample.³ Likewise, in this trial. Black participants were younger and had higher body mass index, on average, and were less likely to have received coronary revascularization compared with White participants.^{24,25} Furthermore, we previously reported variations in the social living situation and the severity of baseline functional impairments across race and sex.^{6,26-28} However, despite the differences in baseline characteristics, there were similar intervention adherence rates, and the intervention produced similarly large improvements in physical function and HRQOL in both Black and White participants, corroborating and extending findings from exercise studies in chronic stable HF (Figure 2).8,29

Social and functional determinants of health are increasingly acknowledged for their impact on cardiovascular outcomes.³⁰ Indeed, 1 study examining Black individuals attending cardiac rehabilitation demonstrated improved coping efficacy and health benefits in those with greater social support.³¹ The REHAB-HF protocol provided personalization of care by using tailored adherence strategies designed to address the specific social and functional determinants of health needs of the participants that would reduce barriers to trial participation and self-management of exercise.^{12,14,32} For instance, a home and builtenvironment assessment was performed to support exercise safety and independence in the participant's home and community. Transportation support was provided to those who otherwise were unable to reach the outpatient facility. Strategies such as these led to similarly high intervention session adherence and fidelity across race, resulting in correspondingly large improvements in functional outcomes.³² Our results not only underscore opportunities for the intervention to improve physical function and HRQOL but also to address sociodemographic variables that limit optimal participation in care and contribute to variable health outcomes for Black versus White individuals. However, the difference in 6-month all-cause rehospitalization between Black and White individuals (49% in Black versus 39% in White; P=0.066) was not explained by adherence rates or by variations in past hospitalizations at baseline, suggesting potential contributions of unmeasured or unaddressed social determinants of health.

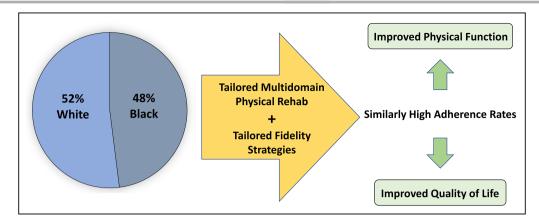


Figure 2. Relationship of race with outcomes in the REHAB-HF population.

Despite differences in baseline characteristics across a diverse sampling of Black and White participants, multidomain physical rehabilitation and intervention fidelity strategies personally tailored to the participant's physical and social health determinants resulted in similarly high intervention adherence rates and produced large improvements in physical function and health-related quality of life. REHAB-HF indicates Rehabilitation Therapy in Older Acute Heart Failure Patients.

Strengths and Limitations

Strengths of the current study include the randomized, controlled trial design; assessment of an innovative, early, transitional, tailored, and progressive multidomain physical rehabilitation intervention for vulnerable, high-risk older patients with ADHF; and strong representation of Black participants (48% of participants). Contextually, Black people comprise 9% of those aged ≥65 years in the United States, suggesting a robust sampling in the current study.³³ Additionally, enrollment was evenly distributed, and intervention session adherence was similar by race, enhancing our ability to test for heterogeneity of intervention effect. We acknowledge there are inherent statistical power limitations for subgroup analyses assessing both the primary outcome and also secondary outcomes.³⁴ Accordingly, confirmation of results in the larger recently launched, National Institutes of Health-funded phase III clinical trial will be crucial for confirming the results (https:// www.clinicaltrials.gov; Identifier NCT05525663).

CONCLUSIONS

In this diverse population of older patients hospitalized for ADHF, the intervention produced large improvements in physical function and HRQOL compared with attention control, irrespective of race. The clinical event data suggest that there may be a differential effect of the intervention by race on all-cause rehospitalization rates, highlighting the need for larger follow-up studies with robust recruitment across racial and ethnic groups.

ARTICLE INFORMATION

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Disclosures

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Supplemental Material

Data S1

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