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# Weight Lifting for Women at Risk for Breast Cancer–Related Lymphedema

## A Randomized Trial

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**M**ORE THAN 2.4 MILLION breast cancer survivors live in the United States.<sup>1</sup> Lymphedema ranks high among their concerns because it causes swelling and discomfort, impairing arm function and quality of life<sup>2,3</sup> and increasing health care costs.<sup>4</sup> Lymphedema remains a frequent complication among survivors, despite lymphatic-sparing procedures such as sentinel lymph node biopsy. Of the 61% of patients who undergo sentinel lymph node biopsy, 5% to 7% develop breast cancer–related lymphedema.<sup>5,6</sup> However, one-third of patients with breast cancer require complete axillary dissection,<sup>5</sup> which is associated with 13% to 47% incident lymphedema.<sup>7,8</sup>

Breast cancer survivors at risk for lymphedema alter activity, limit activity, or both from fear and uncertainty about their personal risk level, and upon guidance advising them to avoid lift-

**Context** Clinical guidelines for breast cancer survivors without lymphedema advise against upper body exercise, preventing them from obtaining established health benefits of weight lifting.

**Objective** To evaluate lymphedema onset after a 1-year weight lifting intervention vs no exercise (control) among survivors at risk for breast cancer–related lymphedema (BCRL).

**Design, Setting, and Participants** A randomized controlled equivalence trial (Physical Activity and Lymphedema trial) in the Philadelphia metropolitan area of 154 breast cancer survivors 1 to 5 years postunilateral breast cancer, with at least 2 lymph nodes removed and without clinical signs of BCRL at study entry. Participants were recruited between October 1, 2005, and February 2007, with data collection ending in August 2008.

**Intervention** Weight lifting intervention included a gym membership and 13 weeks of supervised instruction, with the remaining 9 months unsupervised, vs no exercise.

**Main Outcome Measures** Incident BCRL determined by increased arm swelling during 12 months ( $\geq 5\%$  increase in interlimb difference). Clinician-defined BCRL onset was also evaluated. Equivalence margin was defined as doubling of lymphedema incidence.

**Results** A total of 134 participants completed follow-up measures at 1 year. The proportion of women who experienced incident BCRL onset was 11% (8 of 72) in the weight lifting intervention group and 17% (13 of 75) in the control group (cumulative incidence difference [CID],  $-6.0\%$ ; 95% confidence interval [CI],  $-17.2\%$  to  $5.2\%$ ;  $P$  for equivalence = .04). Among women with 5 or more lymph nodes removed, the proportion who experienced incident BCRL onset was 7% (3 of 45) in the weight lifting intervention group and 22% (11 of 49) in the control group (CID,  $-15.0\%$ ; 95% CI,  $-18.6\%$  to  $-11.4\%$ ;  $P$  for equivalence = .003). Clinician-defined BCRL onset occurred in 1 woman in the weight lifting intervention group and 3 women in the control group (1.5% vs 4.4%,  $P$  for equivalence = .12).

**Conclusion** In breast cancer survivors at risk for lymphedema, a program of slowly progressive weight lifting compared with no exercise did not result in increased incidence of lymphedema.

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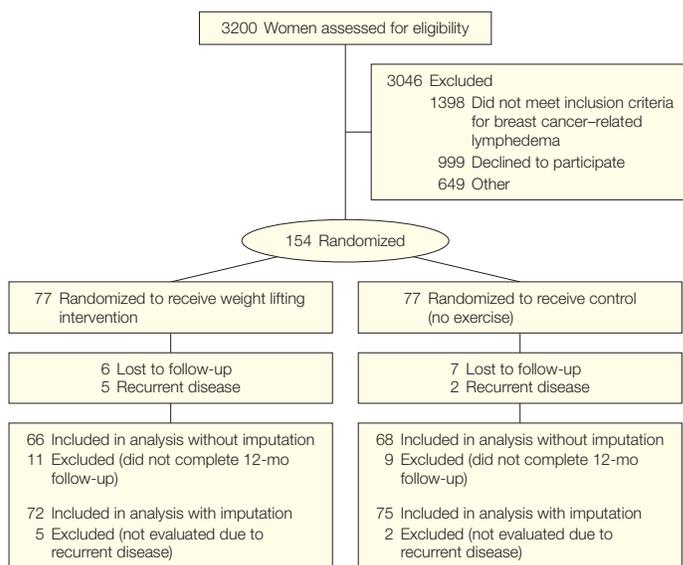
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**Figure.** Flow of Participants Through the Physical Activity and Lymphedema Trial

ing children, heavy bags, or other objects with the at-risk arm.<sup>9,10</sup> Such guidance is often interpreted in a manner that deconditions the arm, increasing the potential for injury, overuse, and, ironically, lymphedema onset.<sup>11</sup> Adherence to these precautions may limit physical recovery after breast cancer and, for some women, result in lost employment. Furthermore, activity avoidance may deter survivors from performing regular exercise, which may prevent cancer recurrence and improve survival.<sup>12,13</sup> By contrast, controlled physiological stress through progressive weight lifting may increase the maximal physical work capacity of the affected arm, protecting it from injury.

In a pilot study, we found no evidence that slowly progressive weight lifting precipitated lymphedema among breast cancer survivors,<sup>14</sup> although that study had limited statistical power and follow-up. The Physical Activity and Lymphedema (PAL) trial was conducted among breast cancer survivors to determine whether exercise is safe for women at risk for lymphedema. Women were randomized to a 1-year weight lifting intervention group or a 1-year non-intervention group. The PAL trial was a single study statistically powered to ad-

dress 2 distinct primary goals. We previously published the findings of the first primary goal, which was to assess the effects of weight lifting on lymphedema worsening.<sup>15</sup> Herein, we report the results of the second primary goal, which was to evaluate incident lymphedema from weight lifting from a distinct pool of PAL participants.

## METHODS

### Study Participants

Breast cancer survivors with and at risk for lymphedema were recruited throughout the Philadelphia metropolitan area. Participants were recruited between October 1, 2005, and February 2007, with data collection ending in August 2008. Recruitment methods included letters sent by Pennsylvania and New Jersey state cancer registries, media advertisements and interviews, and flyers at support groups. After baseline measurements that confirmed whether women had lymphedema, participants were randomized into the trial about lymphedema worsening (results of which have already been published),<sup>15</sup> or into the trial described herein, which evaluated incident lymphedema from weight lifting. Eligibility requirements for the trial included female sex, history of unilat-

eral nonmetastatic breast cancer diagnosis between 1 and 5 years before study entry, body mass index (calculated as weight in kilograms divided by height in meters squared) of 50 or less, currently cancer free, no medical conditions that would limit participation in exercise, no weight lifting in the year before study entry, no plans for surgery or to be away for at least 1 month during the study, currently weight stable and not actively trying to lose weight, at least 2 lymph nodes removed, no prior lymphedema diagnosis, and no evidence of current lymphedema. For the purpose of eligibility, lymphedema was defined as an interlimb difference of at least 10% as measured by water volumetry, greatest circumferential difference, or, per the Common Toxicity Criteria version 3.0 adverse events criteria,<sup>16</sup> swelling or obscuration of anatomic architecture or pitting edema. Women with suspected lymphedema were sent for evaluation with a certified lymphedema therapist (CLT) to verify eligibility.<sup>17</sup> The FIGURE shows the 154 participants who entered the PAL trial at risk for lymphedema.

Women were placed into 2 equally sized groups through a computerized process called minimization<sup>18,19</sup> in a manner that was unpredictable and concealed from research staff who determined eligibility. This approach balanced important potential confounders at baseline: age (<54 vs ≥54 years), number of lymph nodes removed (<6 vs ≥6), obesity (body mass index <30 vs ≥30), and history of radiation treatment (yes vs no). The study was approved by the University of Pennsylvania institutional review board. Women provided written informed consent and written clearance from a physician before participation.

### Intervention

Participants in the weight lifting intervention group received a 1-year membership to a community fitness center (usually a YMCA) near their homes. For the first 13 weeks, women were instructed twice weekly on safe performance of exercises in groups of 2 to 6

survivors. Certified fitness professionals employed by the fitness centers led these 90-minute sessions. Upper body exercises (seated row, supine dumbbell press, lateral or front raises, bicep curls, and triceps pushdowns) were performed with dumbbells or variable resistance machines. Lower body exercises (leg press, back extension, leg extension, and leg curl) were performed with variable resistance machines. The specific equipment used varied across the fitness centers at which the intervention was delivered. Three sets of each exercise were performed at each session, 10 repetitions per set. After 13 weeks, participants continued twice weekly unsupervised exercise to 1 year. Weight was increased for each exercise by the smallest possible increment after 2 sessions of completing 3 sets of 10 repetitions with no change in arm symptoms. Fitness trainers called women who missed more than 1 session per week throughout the year. Participants who missed more than 2 consecutive sessions were asked to reduce resistance and rebuild as per protocol above. Participants in the control group were asked to not change baseline level of exercise during study participation and were offered a 1-year fitness center membership with 13 weeks of supervised instruction following study completion. Further details of the intervention are provided elsewhere.<sup>20</sup>

All trainers who worked with participants underwent a 3-day training course including the exercise protocol and an overview of lymphedema prevention, symptoms, and treatment.<sup>21-23</sup> An intervention coordinator met with trainers weekly during the first 13 weeks, then monthly to ensure protocol fidelity. All participants (weight lifting intervention and control groups) who developed lymphedema were provided a custom-fitted compression garment (Jobst, BSN Medical, Charlotte, North Carolina) and were required to wear these garments during weight lifting sessions. Trainers asked about changes in symptoms weekly and took circumference and water volume measurements monthly to ensure arm

swelling changes were detected and treated promptly. In addition, all participants (weight lifting intervention and control groups) were required to attend a 1-hour educational lecture about lymphedema risk reduction, treatment, and exercise safety based on position stands from the National Lymphedema Network.<sup>17,22,24</sup>

### Measurement

Measurements of all participants at baseline and 12 months were completed by trained staff using standardized methods. Measurement staff (including CLTs) were blinded to treatment allocation. Participants were reminded to not reveal their group assignment before measurement and evaluation sessions.

Demographic characteristics (age, education, race, occupation) were self-reported at baseline. Cancer stage was taken from the state cancer registry, surgical pathology report, or self-report, according to data availability. Treatment history was self-reported for radiation and chemotherapy. The number of lymph nodes removed was collected from surgical pathology reports. Anthropometry measures included weight, height (baseline only), and whole-body dual-energy x-ray absorptiometry scan (Hologic Discovery, software version 12.4, Bedford, Massachusetts). Percentage of body fat is presented without bone mass to avoid misrepresenting changes in relative fat mass due to changes in bone density. Physical activity outside of weight lifting was assessed using the International Physical Activity Questionnaire.<sup>25</sup> Diet was assessed using the Diet History Questionnaire.<sup>26</sup>

The primary outcome was lymphedema onset defined as a 5% or more increase in arm swelling, which was defined by interlimb water volume difference [(affected arm volume-unaffected arm volume)/unaffected arm volume].<sup>3</sup> Water volume displacement was used to measure arm volumes at baseline and 12 months.<sup>27</sup> Water volume is accurate by raters to 1% and was taken once per side.<sup>28</sup> For clinician-defined onset, CLTs<sup>17</sup> at Penn Therapy and Fitness used a standardized clinical evalu-

ation based on the Common Toxicity Criteria version 3.0 criteria,<sup>16</sup> including interlimb differences, and changes in tissue tone or texture, as well as symptoms. Participants were sent for evaluation of possible onset upon report of a change in symptoms lasting 1 week or longer or if monthly preexercise safety measurements by fitness trainers or 3-month interval measurements by measurement staff indicated a change in treated arm volume of at least 5% and at least 5% interlimb difference. Lymphedema-related arm symptom presence and severity were reported using a validated and reliable survey for detecting prevalent lymphedema.<sup>29</sup>

Strength measurements at baseline and 12 months provided physiological evidence of intervention adherence and strength gains. The maximum amount of weight that can be lifted once (1 repetition maximum=1-RM) was assessed for the bench press and leg press. One-RM tests, the standard for evaluating increases in muscular strength,<sup>30</sup> are safe for most populations when properly supervised.<sup>30-32</sup> Methods for the strength measurements have been reported elsewhere.<sup>20</sup> Intervention adherence was also evaluated by attendance logs completed by fitness trainers.

### Statistical Analysis

All analyses were conducted using SAS version 9.2 (SAS Institute, Cary, North Carolina). Descriptive statistics for baseline variables included rates for binary variables and means, medians, and SDs for continuous variables. The rates of occurrence of lymphedema and other binary outcomes were compared between the exercise and control groups using Fisher exact test and continuous outcomes were compared using the Wilcoxon rank sum test, with 2-sided  $P < .05$ . Cumulative incidence ratios (relative risks) of outcomes are shown with 95% confidence intervals (CIs). Because clinician-defined onset required follow-up for 12 months, patients who were not evaluated due to recurrent cancer (5 in the weight lifting intervention group and 2 in the control group) or patients who dropped out of the trial ( $n=13$ ) were ex-

cluded from this analysis. Simple imputation-based sensitivity analyses were conducted to examine the potential effect of these missing data on results. First, all dropouts were assumed to have had the event in question and then to not have had the event. For continuous lymphedema outcomes (eg, arm swelling and symptoms), data were imputed using a regression model, incorporating baseline covariates that predicted the outcome at  $P \leq .25$ , and properly incorporated patient-specific variability.

Sample size calculations were based on the primary comparisons of interest. The PAL trial had 2 primary comparisons of interest. The PAL trial assessed outcomes in 2 independent subgroups of women, those with and

without lymphedema at baseline; each substudy was designed to demonstrate safety of the intervention using a type I error rate of .05. Herein, we reported on women at risk for lymphedema, in whom the study was designed with 80% power to show equivalent lymphedema onset between the weight lifting intervention and control groups, allowing up to a 20% loss to follow-up. Given these parameters, the trial sought to recruit at least 144 women at risk for lymphedema to detect more than a doubling of the rate of lymphedema onset, with an assumption that the background rate among the control group would be 6%.<sup>6</sup> Furthermore, we planned a subgroup analysis among women who had 5 or more

nodes removed. This threshold was chosen to be consistent with our prior work<sup>14</sup> and published accounts that the majority of sentinel lymph node biopsies involve removal of 1 to 4 nodes.<sup>33</sup>

## RESULTS

TABLE 1 shows the 154 randomized PAL trial participants at risk for lymphedema at baseline, including the 7 (4.5%) who had recurrent cancer and the 13 (8.4%) who were lost to follow-up. Participants were aged 36 to 75 years at baseline and diverse regarding education, race/ethnicity (29% nonwhite), and occupation. The number of lymph nodes removed ranged between 2 and 26, with a mean of 8 in the weight lifting intervention group and 9 in the control group; 94 women had at least 5 nodes removed. Interlimb differences in arm volume ranged between  $-11\%$  and  $13\%$  (comparing affected with unaffected limb), with a mean of  $0.13\%$  and  $-0.27\%$ , respectively, in weight lifting intervention and control group women.

TABLE 2 shows baseline and 12-month data for strength, anthropometry, and diet and physical activity. At baseline, the range for the 1-RM bench press test was 0 to 80 lb and the range for the leg press was 65 to 345 lb. Participants in both groups were well-balanced at baseline on strength and anthropometrics. Women in the weight lifting intervention group became stronger compared with the no exercise group. Percentage body fat was lower among the weight lifting participants at 12 months. Median attendance was 79% among the 77 women in the weight lifting intervention group, including the 13 lost to follow-up. No between-group differences were noted in dietary intake or self-reported physical activity outside of weight lifting at 12 months.

TABLE 3 shows lymphedema onset outcomes at 12 months. The proportion of women who experienced a 5% or more increase in interlimb volume difference during the 12 months was 17% (13 of 75) in the control group and 11% (8 of 72) in the weight lifting in-

**Table 1.** Baseline Characteristics of the 154 Study Participants at Risk for Lymphedema<sup>a</sup>

Characteristic	Weight Lifting Intervention (n = 77)	Control (n = 77)	P Value
Age, mean (SD), y	54 (8)	56 (8)	.36
Education			
≤High school	7 (9)	11 (14)	.51
Some college	28 (36)	23 (30)	
≥College	42 (55)	43 (56)	
Race/ethnicity			
White	50 (65)	59 (76)	.04
Black	19 (25)	17 (22)	
Other <sup>b</sup>	8 (10)	1 (<1)	
Occupation			
Professional	32 (42)	34 (44)	.95
Clerical or service	18 (23)	16 (21)	
Homemaker, student, or unemployed	6 (8)	8 (10)	
Other or unknown	8 (10)	6 (8)	
Retired	13 (17)	13 (17)	
Months since cancer diagnosis, mean (SD)	39 (15)	42 (16)	.26
Cancer stage <sup>c</sup>			
Ductal carcinoma in situ	1 (1)	0	.31
1	43 (56)	43 (56)	
2	8 (10)	6 (8)	
3	25 (33)	28 (36)	
No. of nodes removed, mean (SD)	8 (6)	9 (6)	.50
Chemotherapy	56 (73)	53 (69)	.72
Radiation	59 (77)	58 (75)	>.99
Current receipt of drugs			
Tamoxifen	27 (21)	25 (19)	>.99
Aromatase inhibitors	0	1 (1)	>.99
Arm volume difference, mean (SD) [median], %	0.13 (5.09) [−0.34]	−0.27 (4.93) [−0.43]	.61

<sup>a</sup>Data are presented as No. (%) unless otherwise specified. Percentages may not sum to 100 due to rounding. Control group included no exercise.

<sup>b</sup>Includes American Indian or Alaskan Native, Asian, Hispanic, Latino, Portuguese, or Cape Verdean, Native Hawaiian or other Pacific Islander.

<sup>c</sup>The staging system used was from the American Joint Commission on Cancer and refers to the extent of the cancer in the body, with ductal carcinoma in situ as the least and stage 3 as the greatest extent of cancer in the study participants.

intervention group (cumulative incidence difference [CID], -6.0%; 95% CI, -17.2% to 5.2%; *P* for equivalence = .04; cumulative incidence ratio [CIR], 0.64; 95% CI, 0.28-1.45; *P* for equivalence = .003; the upper limit of the CI is below the a priori equivalence boundary of 2 for the CIR). These results are based on imputed data for intention-to-treat analyses; findings were robust

with repeated analysis without imputation. Among the 134 women at risk for lymphedema who had no new or recurrent cancers and not lost to follow-up, there were 4 incident cases of clinician-defined lymphedema (1 in the weight lifting group and 3 in the control group), producing a CIR of 0.34 (95% CI, 0.04-3.22; *P* for equivalence = .12). Sensitivity analyses (as-

suming the women lost to follow-up all had no events or all had events) resulted in CIRs that did not substantively alter the results. No notable differences in the number or severity of symptoms were observed in the weight lifting and control groups.

In planned secondary analyses limited to women with 5 or more nodes removed, the proportion of women who

**Table 2.** Strength, Anthropometry, and Diet and Physical Activity at Baseline and 12 Months

Variables	Baseline					12 Months				
	Weight Lifting Intervention		Control		<i>P</i> Value <sup>a</sup>	Weight Lifting Intervention		Control		<i>P</i> Value <sup>a</sup>
	No.	Mean (SD)	No.	Mean (SD)		No.	Mean (SD)	No.	Mean (SD)	
Strength										
Bench press, lb	77	41 (13)	75	41 (13)	.93	59	54 (12)	63	43 (11)	<.001
Leg press, lb	77	170 (48)	76	181 (54)	.23	61	213 (50)	63	192 (53)	.02
Anthropometry										
Weight, kg	77	73.87 (15.21)	77	76.76 (17.16)	.27	66	72.36 (14.88)	68	75.46 (17.07)	.27
BMI	77	27.52 (5.09)	77	28.55 (6.17)	.26	66	26.94 (4.99)	68	28.03 (5.95)	.25
Body fat, %	77	37.71 (5.60)	77	39.26 (6.38)	.11	65	37.34 (5.35)	68	39.59 (6.45)	.03
Fat mass, kg	77	28.11 (9.10)	77	30.56 (10.69)	.13	65	27.18 (8.48)	68	30.3 (10.57)	.06
Lean mass, kg	77	46.84 (7.05)	77	47.30 (7.50)	.70	65	46.25 (7.42)	68	46.3 (7.58)	.97
Diet and physical activity										
Dietary intake, kcal	77	1637 (1139)	77	1691 (1446)	.79	63	1492 (798.8)	65	1535 (844.2)	.77
Self-reported physical activity (MET-min/wk) <sup>b</sup>	70	2670.4 (2.34)	73	2079.7 (3.06)	.14	58	3041.2 (2.29)	60	2440.6 (3.10)	.46

Abbreviations: BMI, body mass index, calculated as weight in kilograms divided by height in meters squared; MET, metabolic equivalent.

<sup>a</sup>Wilcoxon rank sum 2-sample *t* tests.

<sup>b</sup>Data reported are geometric means at baseline and log-transformed physical activity levels to improve normality of distribution at 12 months.

**Table 3.** Lymphedema Onset Outcomes at 12 Months<sup>a</sup>

	Weight Lifting Intervention		Control		Cumulative Incidence Ratio (95% CI)	<i>P</i> Value <sup>b</sup>
	No./Total No. (%)	Mean (SD)	No./Total No. (%)	Mean (SD)		
All participants						
Defined by ≥5% increase in arm swelling <sup>c</sup>	8/72 (11)		13/75 (17)		0.64 (0.28-1.45)	.003
Clinician-defined onset	1/66 (1.5)		3/68 (4.4)		0.34 (0.04-3.22)	.12
Participants who had ≥5 lymph nodes removed						
Defined by ≥5% increase in arm swelling <sup>c</sup>	3/45 (7)		11/49 (22)		0.30 (0.09-1.00)	.001
Clinician-defined onset	1/42 (2.4)		3/46 (6.5)		0.37 (0.04-3.38)	.13
	<b>Total No.</b>		<b>Total No.</b>		<b>Mean (SD) Difference</b>	
All participants						
Δ in No. of symptoms reported	72	-0.51 (1.57)	75	-0.42 (2.26)	-0.10 (0.32)	.77
Δ in symptom severity <sup>d</sup>	72	-0.27 (0.97)	75	-0.28 (0.86)	0.003 (0.15)	.99
Participants who had ≥5 lymph nodes removed						
Δ in No. of symptoms reported	45	-0.63 (1.86)	49	-0.83 (1.52)	0.21 (0.35)	.55
Δ in symptom severity <sup>d</sup>	45	-0.30 (1.06)	49	-0.41 (0.88)	0.12 (0.20)	.56

Abbreviation: CI, confidence interval.

<sup>a</sup>Results for arm swelling and symptoms include imputed data.

<sup>b</sup>Test for equivalence, using Fisher exact test for arm volume changes and Wilcoxon rank sum 2-sample test for change in symptoms.

<sup>c</sup>Arm swelling = [(affected arm volume - unaffected arm volume) / unaffected arm volume] (eg, interlimb volume difference).

<sup>d</sup>Possible values were 0 (did not have symptom) to 4 (very severe) for each item; outcomes reported are average changes in symptom severity across all 14 possible symptoms (rings too tight, watch too tight, bracelets too tight, clothing too tight, puffiness, could not see knuckles, could not see veins, skin felt leathery, arm felt tired, pain, pitting, swelling after exercise, difficulty writing, or other).

experienced a 5% or more increase in interlimb volume during the 12 months was 7% (3 of 45) in the weight lifting group and 22% (11 of 49) in the control group (CID,  $-15.0\%$ ; 95% CI,  $-18.6\%$  to  $-11.4\%$ ;  $P$  for equivalence = .003; CIR, 0.30; 95% CI, 0.09–1.00;  $P$  for equivalence = .001). No between-group differences were observed in clinician-defined lymphedema onset or symptoms in secondary analysis limited to women with 5 or more nodes removed.

## COMMENT

Breast cancer survivors who performed slowly progressive weight lifting twice weekly for 1 year were less likely to experience clinically significant increases in arm swelling than women in the control group. The majority of breast cancer survivors do not have lymphedema; however, they alter the use of their arms and upper body activities out of fear of developing lymphedema. The findings from our trial should help clarify clinical advice to patients who have completed breast cancer treatment regarding the safety of resuming or beginning a weight lifting program. These results are consistent with the well-defined hormetic effect of exercise training—small, slowly progressive increases in physiological stress buffer the body's ability to respond to infection, inflammation, and injury through gradual adaptations to muscle mass, metabolic demand on tissues, altered microcirculation, reduced oxidative stress, and improved inflammatory profile.<sup>34</sup>

Prior randomized trials of weight lifting safety among breast cancer survivors, all of which agree with the current findings, have been smaller, shorter, and some have included lymphedema as a secondary outcome.<sup>14,35–37</sup> Studies in Norway and Spain have demonstrated that when upper body exercise is combined with other lymphedema therapeutic modalities, no increased risk of onset is conferred<sup>8</sup> or lymphedema may be prevented.<sup>38</sup> Our study is the first well-powered clinical trial to our knowledge to demonstrate no increased risk of

lymphedema onset with weight lifting alone, with the possibility of reduced likelihood of increased arm swelling among higher risk women with 5 or more nodes removed.

Strengths of the PAL trial include the large sample size and the delivery in community fitness centers, primarily YMCAs, by trainers employed by these fitness centers. This approach was purposeful, with a goal of increasing the likelihood of broad dissemination. Additional strengths include the participant diversity (29% nonwhite participants), long intervention (1 year), and minimal loss to follow-up (8.4% of women did not have recurrent cancers). Limitations included marginal significance of a treatment effect on lean mass. Changes in lean mass were more favorable in women in the weight lifting group vs the control group during the 12-month trial ( $-0.45$  vs  $-1.47$  kg,  $P = .06$ ), but it is unclear why there were decreases in lean mass on average given the notable increases in strength (and in contrast with findings from the pilot study).<sup>14</sup>

Multiple elements of the PAL trial intervention were specifically designed to reduce the risk of lymphedema onset. First, breast cancer survivors started with 13 weeks of supervision to learn to perform the exercises properly and to progress the resistance appropriately. Second, participants started training at a low weight (1 or 2 lb) and progressed resistance according to symptom response. If there was a break in exercise that lasted 1 week or more, the protocol specified that the resistance should be reduced and increased gradually. It was vital to the participants' sense of safety that there were CLTs<sup>17</sup> available to whom they could be referred. These intervention elements are in keeping with the National Lymphedema Network position statement on exercise and the American College of Sports Medicine guidance for exercise in breast cancer survivors.<sup>39,40</sup> Third, all fitness trainers were certified by a national organization and underwent training about the specific exercises used, adapta-

tions that might be required, and when and whom to call if there were symptom or measurement changes that might require medical evaluation.

In conclusion, the findings of our study remove concerns that slowly progressive weight lifting will increase risk of lymphedema onset in breast cancer survivors. In a preplanned secondary analysis limited to women with 5 or more nodes removed, the incidence of 5% increase in arm swelling was reduced by 70% among women in the weight lifting intervention group compared with no exercise. No between-group differences were noted for clinically defined lymphedema onset or symptom changes in the total cohort or this higher-risk subset. The primary goal was to test safety of weight lifting, not superiority; therefore, additional research is needed before concluding that weight lifting prevents lymphedema. However, even with the finding of no harm, our results combined with previously published results for women with breast cancer–related lymphedema<sup>15</sup> suggest that the many health benefits of weight lifting should now become available to all breast cancer survivors.

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**Study concept and design:** Schmitz, Ahmed, Cheville. **Acquisition of data:** Schmitz, Lewis-Grant, Bryan. **Analysis and interpretation of data:** Schmitz, Troxel, Cheville, Smith, Williams-Smith, Chittams.

**Drafting of the manuscript:** Schmitz, Cheville. **Critical revision of the manuscript for important intellectual content:** Schmitz, Ahmed, Troxel, Cheville, Lewis-Grant, Smith, Bryan, Williams-Smith, Chittams. **Statistical analysis:** Troxel, Williams-Smith, Chittams. **Obtained funding:** Schmitz, Ahmed.

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