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## Weight Management and Physical Activity Throughout the Cancer Care Continuum

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Abstract: Mounting evidence suggests that weight management and physical activity (PA) improve overall health and well being, and reduce the risk of morbidity and mortality among cancer survivors. Although many opportunities exist to include weight management and PA in routine cancer care, several barriers remain. This review summarizes key topics addressed in a recent National Academies of Science, Engineering, and Medicine workshop entitled, "Incorporating Weight Management and Physical Activity Throughout the Cancer Care Continuum." Discussions related to body weight and PA among cancer survivors included: 1) current knowledge and gaps related to health outcomes; 2) effective intervention approaches; 3) addressing the needs of diverse populations of cancer survivors; 4) opportunities and challenges of workforce, care coordination, and technologies for program implementation; 5) models of care; and 6) program coverage. While more discoveries are still needed for the provision of optimal weight-management and PA programs for cancer survivors, obesity and inactivity currently jeopardize their overall health and quality of life. Actionable future directions are presented for research; practice and policy changes required to assure the availability of effective, affordable, and feasible weight management; and PA services for all cancer survivors as a part of their routine cancer care. CA Cancer J Clin 2017;000:000-000. © 2017 American Cancer Society.

Keywords: nutrition, physical activity, supportive care, survivorship, weight management

#### Introduction

With growing evidence of the association between obesity, excess weight, and cancer, the National Cancer Policy Forum (NCPF) of the National Academies of Sciences, Engineering, and Medicine hosted a workshop in 2011 on the "Role of Obesity in Cancer Survival and Recurrence." That workshop examined epidemiologic evidence, biologic mechanisms, preclinical studies, and a limited number of randomized controlled trials (RCTs) of interventions that promoted weight loss via caloric restriction or increased physical activity (PA) in patients with cancer.<sup>1</sup> Two scientific papers emanated from this endeavor—one that was more mechanistic in nature,<sup>2</sup> and the other focused on translational research for patient care.<sup>3</sup> Both emphasized gaps in knowledge.

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#### TABLE 1. Prioritizing Patient Voices

It is essential to hear and prioritize the voices of cancer survivors. Below are select statements from 2 patient advocates who, despite differing backgrounds, diagnoses, and points in their survivorship journey, offer statements with common themes.

Karen Cochrane is a white, 53-year old nurse who was recently diagnosed with early stage breast cancer and is currently receiving chemotherapy. She is concerned about her overall well being and is working toward reaching a healthy weight and being physically active.

Robert Harrison is a black, 72-year old, retired, businessman who was diagnosed with metastatic prostate cancer 14 years ago. Currently, he is actively monitoring his cancer with his urologist. He has played an active role in his care and lost over 60 pounds, and he considers himself a cancer "thriver."

Both voice their thoughts about weight management and physical activity during cancer treatment and beyond:

- "Understand that we are human systems, we are not just a disease to be treated."
- "This isn't just about surviving the treatment of cancer, it's making sure I do everything I can to live a long healthy life."
- "Weight Management and physical activity are essential components of the treatment of cancer patients and can be, should be, and must be integrated
  into and monitored throughout treatment to help achieve the most beneficial treatment outcomes."
- "You know they (patients) are going to gain weight. You know they (patients) are going to lose muscle. From the very beginning, integrate it into treatment, so that we can minimize these negative impacts and maximize the opportunities to keep the body as healthy as possible."
- "People need to be told how to lose weight and followed up. There needs to be a long-standing program and support."
- "It would be most helpful, and dollars well spent, to provide programs and support to teach people how to make healthy food choices and how to be
  physically active."
- "It is not just providing some information or saying that you need to lose weight, it needs to be an ongoing, life-long process."
- "It isn't easy, because if it were we would all be normal weight and very active; but, with help and support, it can be done."
- "What has helped me was to have a plan and follow-up."

Six years later, and after considerable advances in this arena, including a position paper and campaign on obesity and cancer issued by the American Society of Clinical Oncology (ASCO),<sup>4</sup> another NCPF workshop on obesity and cancer was convened-this time focusing on translating research findings into clinical practice and communitybased programs. That workshop, "Incorporating Weight Management and Physical Activity Throughout the Cancer Care Continuum," also drew international experts, but with greater emphasis on behavioral science, clinical research, public policy, dissemination science, and health economics. Workshop presentations and discussions examined the available evidence regarding the value of promoting weight management and PA across the period of cancer survivorship, from diagnosis to end of life,<sup>5,6</sup> along with evaluation of opportunities and challenges in current approaches to promote PA and weight management. To ensure that discussions would be patient-centered, the opening session of the workshop featured 2 cancer survivors who shared their experiences with cancer treatment, weight management, and PA (for key excerpts, see Table 1). Also, throughout the workshop, presentations and discussions illustrated the broad spectrum of diversity among cancer survivors in terms of cancer type, stage, molecular subtype, length of survivorship, comorbidity and functional status, age, race/ethnicity, gender, and geographic location that require consideration in tailoring weight-management and PA interventions and recognition that "one size does not fit all." The workshop culminated in a discussion of whether the strength of evidence warranted the provision and coverage of services for weight management and PA that are specifically directed toward cancer survivors and ways to enhance the delivery of these services to the growing sector of cancer survivors in this nation, who currently number well over 15 million.<sup>7</sup> Herein, we report a summary of the workshop presentations, discussions, and conclusions.

## Body Weight, PA, and Health Outcomes for Cancer Survivors: Knowledge and Gaps

This first section, which provides a foundation for the remainder of the article, is devoted to critically evaluating the state of knowledge regarding the relationship between body weight or PA and health outcomes for cancer survivors. It begins with an overview of the evidence on cancer outcomes and then addresses other outcomes, such as

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quality of life (QOL) and fatigue. Finally, it identifies evidence gaps addressed by ongoing and recent trials, gaps that remain, and the opportunities to fill these gaps.

### **Overview of Obesity and Cancer Outcomes**

Excess weight gain, overweight, and obesity are associated with an increased risk of many cancers; recently, the International Agency for Research on Cancer reported that there is sufficient evidence to conclude that avoidance of excess body fat is associated with a lower risk for cancers of the endometrium, esophagus (adenocarcinoma), gastric cardia, kidney (renal cell), multiple myeloma, meningioma, liver, pancreas, colorectum, gallbladder, breast (postmenopausal), ovary, and thyroid.<sup>8</sup> There also is growing observational evidence that obesity is associated with poorer cancer outcomes among individuals with cancer. The largest body of evidence relates to breast cancer. A meta-analysis of 82 studies involving 213,075 women with breast cancer found a 41% relative increase in all-cause mortality for women with obesity versus those of normal weight (relative increases were 75% in premenopausal women and 34% in postmenopausal women).9 That study also reported increased all-cause mortality for overweight women, albeit the relative increases were smaller. Another meta-analysis found that the risk of mortality associated with overweight and obesity was similar for patients with estrogen receptor (ER)-positive and ERnegative breast cancer, although some (but not all) subsequent individual studies have suggested that the risk may be present only in women with ER-positive disease.<sup>10</sup> Among breast cancer survivors treated with anthracyclines, obesity also is associated with greater cardiotoxicity.<sup>11</sup> Similarly, adverse associations of obesity with survival are reported for endometrial, prostatic, pancreatic, colorectal, and ovarian cancer as well as some hematologic malignancies.<sup>3,4</sup> In contrast, overweight and obesity are associated with somewhat better outcomes in lung, esophageal, and kidney cancercancers in which the morbidity of cachexia and advanced stage at diagnosis are more common.<sup>12</sup>

The association of excess weight gain, overweight, and obesity with cancer is biologically complex. Increased adiposity results in changes in adipose tissue, including death of adipocytes, leading to infiltration of inflammatory cells, as well as secretion of cytokines and other factors that stimulate cancer cell growth, invasion, angiogenesis, and metastasis.<sup>13</sup> Increased adiposity also is associated with changes in systemic physiology, including insulin resistance, dysglycemia, altered adipokines, and increased inflammation; together, these changes enhance signaling through key growth pathways (eg, PI3K, RAS, JAK-STAT) and alter cellular metabolism.<sup>2,14</sup> These obesity-associated effects at the tissue and physiologic levels invoke changes in many of the hallmarks of cancer,<sup>15</sup> including sustained proliferative signaling, activated invasion and metastasis, induced angiogenesis, and resistance to cell death.<sup>2</sup> Overweight and obesity also enable deregulation of cellular energetics and tumor-promoting inflammation.<sup>2,15</sup> While observational data obtained from a multitude of studies, coupled with this strong biologic rationale, provide strong support for an association of obesity with poor cancer outcomes, there is insufficient evidence to conclude that this association is causal.

Studies examining weight-related changes in the transcriptome of breast cancers indicate that cancers developing in women with obesity are biologically different from those in women of normal weight in terms of altered gene regulation and expression.<sup>16</sup> At this time, it is unclear whether reversal of obesity will lead to reversal of these differences or lead to improved cancer outcomes. RCTs of weight loss or pharmacologic interventions that reverse obesity-associated changes related to overweight and obesity are needed.

## Overview of the Evidence on Cancer Outcomes Related to PA

Evidence linking increased PA to improved cancer outcomes is preliminary but promising. A recent systematic review and pooled analysis of 26 observational studies found that cancer survivors who engaged in higher levels of PA (>18 metabolic equivalent hours per week) had a 37% lower risk of dying from cancer, compared with those who engaged in lower levels of PA (<1.5 metabolic equivalent hours per week; hazard ratio [HR], 0.63; 95% confidence interval [95% CI], 0.54-0.73).<sup>17</sup> This risk reduction is remarkably consistent across breast, colorectal, and prostate cancer survivors. There also is growing evidence that the association between PA and cancer mortality varies by specific molecular or genetic markers, suggesting a possible precision medicine approach to exercise oncology (eg, a strong inverse association between PA and colon cancer mortality is noted for survivors whose tumors express p27 [HR, 0.32; 95% CI, 0.12-0.85]). In addition, the link between PA and cancer outcomes has strong biologic plausibility related to sex hormones, cell growth regulators, DNA damage repair, inflammatory markers, immune function, and antioxidant pathways.<sup>18</sup>

RCTs are needed to establish the causal effects of PA on cancer outcomes. The Colon Health and Life-Long Exercise Change (CHALLENGE) Trial is the first phase 3 trial examining the effects of a 3-year structured PA program on disease-free survival in patients with stage II and III colon cancer who have recently completed chemotherapy.<sup>19</sup> To date, the trial has demonstrated feasibility in accrual<sup>20</sup> and PA behavior change<sup>21</sup>; it has randomized over 590 of the planned 962 patients. The Intense Exercise for Survival (INTERVAL) Trial is another phase 3 trial examining the effects of a 2-year structured PA program on overall survival

in 866 men with metastatic, castrate-resistant prostate cancer.<sup>22</sup> These trials, and others like them, will provide the first definitive evidence on the role of PA in improving cancer outcomes.

## Influence of Weight Management and PA on QOL Outcomes

Although obtaining evidence of the impact of weight management and PA on cancer progression and mortality is critical, many cancer survivors experience significant comorbidities, or cancer-related and treatment-related physical and psychosocial problems that compromise their QOL.<sup>23</sup> Healthy eating, regular PA, and maintaining a healthy weight have been recommended for cancer survivors to prevent, mitigate, and manage these downstream sequelae.<sup>24,25</sup>

Weight gain with concomitant loss of muscle (ie, sarcopenic obesity) and bone are common after chemotherapy and hormone therapy, placing cancer survivors at risk for comorbidities, such as cardiovascular disease (CVD), diabetes, second primary cancers, osteoporosis, and functional decline.<sup>25,26</sup> Research indicates that diet-induced and exercise-induced weight management interventions can produce clinically meaningful weight loss in cancer survivors within 6 months, resulting in improved blood lipids and metabolic health and reduced inflammation.<sup>27-31</sup> Also, several studies have reported positive effects of targeted PA on bone health,<sup>32-34</sup> which is important because osteoporosis and the risk of subsequent fractures is increased by 15% to 20% among cancer survivors who receive hormone treatment for breast or prostate cancer.<sup>35</sup>

Growing numbers of studies have examined the effects of PA on CVD in cancer survivors, with a meta-analysis indicating that PA improves cardiorespiratory fitness—a powerful predictor of mortality.<sup>36,37</sup> Growing evidence also suggests that PA may improve cognitive function<sup>38</sup> and lessen peripheral neuropathy,<sup>39</sup> lymphedema,<sup>40</sup> and arthralgia<sup>41</sup> in patients treated for cancer.

In 2010, the American College of Sports Medicine (ACSM) published PA guidelines for cancer survivors based on 85 PA trials conducted during or after treatment.<sup>24</sup> The systematic review and findings from 2 more recent metaanalyses<sup>42,43</sup> demonstrate that PA is safe and effective in improving QOL, cancer-related fatigue, and physical function. While overall effect sizes are small, there is consistent empirical evidence to support PA promotion as part of cancer care.<sup>42,43</sup>

### Evidence Gaps and Ongoing Randomized Weight Management and PA Trials in Cancer Survivors

Although many trials have evaluated the impact of weight management and PA interventions on outcomes, such as body composition, fitness, and QOL in cancer survivors,

not yet available that weight management or increased PA after cancer diagnosis will improve survival or reduce cancer recurrence. The Women's Intervention Nutrition Study, conducted among 2437 women with early stage breast cancer, provides some of the only RCT data and suggests that a mean nonprescribed weight loss of 6 pounds, resulting from a fat-restricted diet, was associated with a significant decrease in subsequent breast cancer events (local, regional, and distant recurrence; ipsilateral breast recurrence after lumpectomy; and contralateral breast cancer) compared with a control arm (ie, 9.8% vs 12.4%; P = .034), a finding driven by women diagnosed with ER-negative disease.<sup>44</sup> However, because the dietary fat intervention also led to weigh loss in that study,<sup>45</sup> it is impossible to disentangle whether the low-fat diet or the weight loss was most responsible for cancer control. Other questions remain about the biologic pathways that underlie the relationship between weight management and PA and malignancy; the relative contribution of body weight, diet, and/or PA to cancer outcomes; the optimal timing, dose, and duration of weight management and PA interventions; and the best ways of implementing weight management and PA interventions in diverse cancer populations.<sup>44</sup> Moreover, the science regarding cancer outcomes and sedentary time also needs to be further developed, as well as the evaluation of potential interventions to limit sedentary time.

critical gaps remain. Most notably, evidence from RCTs is

Several ongoing trials aim to address these evidence gaps (see Table 2).<sup>19,46-48</sup> Each of the ongoing studies examines the impact of weight loss or increased PA, either alone or in combination with improvements in diet quality, on cancer recurrence, cancer-related mortality, or overall survival in individuals diagnosed with a single malignancy. None of the studies compares the effects of different weight management or PA interventions or of different doses or durations of intervention. one-half of the trials enroll breast cancer survivors, and the majority focus on those with no evidence of active disease.

Although these trials will provide critical information regarding the role of weight management and PA in the management of cancer survivors, several gaps will remain. Given that each trial focuses on the effect of a particular weight management or PA intervention in a specific cancer survivor population, it will be difficult to generalize the information gained from these studies across all cancer survivors or to other types of interventions. Moreover, from a feasibility and economic standpoint, it is unlikely that there ever will be trials conducted to evaluate the effect of each type of weight management and PA intervention on every malignancy. So, how do we bridge these evidence gaps and ensure that all cancer survivors have access to weight management and PA interventions that could reduce the risk of

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No.	3136	$\sim$ 1400	1241	1040	962	866
Cancer site	Breast	Breast	Breast	Ovary	Colon	Prostate
Disease stage	III-II	II-II	III-I	II-IV after optimal debulking	II-II	IV, Castrate resistant
Primary endpoint	Invasive disease-free survival	Disease-free survival	Invasive disease-free survival	Progression-free survival	Disease-free survival	Overall survival
Study design	Two-arm RCT: Weight loss intervention + health education vs heath education	Two-arm RCT: General lifestyle + intensive lifestyle intervention vs general lifestyles alone	Two-arm RCT: Weight loss intervention vs general lifestyle guidance	Two-arm RCT: Diet and physical activity intervention vs attention control intervention	Two-arm RCT: Structured physical activity intervention vs general health education materials	Two-arm RCT: Supervised exercise intervention vs supportive care
Intervention target	Weight loss (diet + increased physical activity)	Mediterranean / macrobiotic dietary plan + increased physical activity	Weight loss (diet + physical activity)	Increased physical activity, low fat, increased intake of vegetables and fiber	Increased physical activity	Increased high-intensity physical activity
Intervention duration	2 y	2 y	4 y	2 y	3 y	2 y
Intervention approach	Telephone-based	Telephone-based	Clinic-based group	Telephone-based	Supervised, mixed clinic-based and home-based	Supervised, mixed clinic-based and home-based
Enrollment setting	Cooperative group	Cooperative group	Individual clinics	Cooperative group	Cooperative group	Individual clinics
Correlative specimens	Blood, tumor, and benign tissue	Blood	Blood	Blood	Blood	Blood
Abbreviations: BWEL, Brea:	Abbreviations: BWEL, Breast Cancer Weight Loss; CHALLENGE, Colon Health	Information Health and Lifestyle Lo	Abbreviations: BWEL, Breast Cancer Weight Loss; CHALLENGE, Colon Health and Lifestyle Long Exercise; DIANA-5, Diet and Androgens-5; GAP4, Global Action Plan 4; INTER with Metheotic Controls Devictor Devices (TVES - Historyla Internation for Origins Concord Station, NDC Operation DCT, and onized controls 444	Androgens-5; GAP4, Global Actio	and Lifestyle Long Exercise; DIANA-5, Diet and Androgens-5; GAP4, Global Action Plan 4; INTERVAL, Intense Exercise for Survival Among Men	sise for Survival Among Men

with Metastatic Castrate Resistant Prostate Cancer; LIVES, Lifestyle Intervention for Ovarian Cancer Enhanced Survival; NRG, NRG Oncology; RCT, randomized controlled trial.

recurrence and improve survival after a cancer diagnosis? Part of the key to expanding knowledge gained from ongoing individual trials comes from the correlative science that is embedded in each and the potential to pool data and samples across smaller studies. By evaluating the effect of weight management and PA interventions on blood-based biomarkers (and extant tumor tissue) and determining the relationship between changes in markers, such as insulin and C-reactive protein, and cancer recurrence and survival, intermediate biomarkers could be established to inform future research,<sup>3</sup> akin to research in CVD, in which trials are powered to examine changes in blood pressure or lipid levels rather than on clinical endpoints, such as myocardial infarction.<sup>49</sup> Correlative science also could discover predictive markers of response and determine which cancer survivor populations are most likely to derive benefit from specific interventions.

#### Summary

Ongoing trials will provide vital information on weight management and PA interventions with and without improvements in diet quality on cancer recurrence and survival, but several important knowledge gaps will remain. Biomarker analyses offer the potential to extend the knowledge gained from these trials to other patient populations and could ultimately determine the components of optimal interventions and how they are best applied in a personalized medicine approach to improve cancer outcomes. While more research is needed to elucidate the impact of weight management and PA on cancer-specific outcomes, it is important to note that ASCO now recommends discussion about weight management, including dietary and PA changes, among oncology providers and their patients.<sup>4</sup> This recommendation stems from solid evidence that diet, PA, and reduced adiposity play critical roles in preventing CVD and diabetes and exert a positive influence on QOL, physical function, and fatigue.

## Effective Approaches for Improving Weight Management and PA

Large RCTs related to weight loss and the control of chronic conditions, such as diabetes, have provided sufficient evidence to warrant changes in weight management recommendations from the US Preventive Services Task Force (USPSTF) and from professional organizations.<sup>4,50-53</sup> Research evidence on the benefits of PA (including aerobic, resistance training, and flexibility and coordination) has significantly expanded. Currently, the Physical Activity Guide-lines Advisory Committee is preparing a scientific report with a scheduled release date of early 2018. This evidence has contributed to the development of weight management and PA interventions for cancer survivors and RCTs to

evaluate the impact on important short-term and longer term outcomes in cancer survivors, such as QOL, tolerance for cancer therapy, comorbidity, and disease-free survival. Long-term evidence from previous weight loss and PA trials has demonstrated repeatedly that a high proportion of trial participants have difficulty maintaining behavior changes outside the context of a clinical trial, in part because the current US environment provides little support for being physically active or eating a healthy diet. This recognition has led to an increase in research examining the environmental, policy, and systems changes needed to help individuals adopt and maintain recommended behaviors.

### Interventions for Weight Management in Other Populations That Are Applicable to Cancer Survivors

Lifestyle modifications to alter eating behaviors and increase PA are the cornerstone of treatment for overweight and obesity and have been used successfully in several large-scale trials. The American College of Cardiology (ACC), the American Heart Association (AHA), and The Obesity Society (TOS) reviewed the results from these RCTs and concluded within the 2013 Guidelines for the Treatment of Obesity that adherence to a calorically restricted diet predicts weight loss success, independent of the type of diet or macronutrient composition.54 The guidelines also recommend using body mass index (BMI) and waist circumference to advise patients of their risk of developing other comorbidities and to prescribe a set number of calories (kcal) per day according to the following: 1200 to 1500 kcal per day for women and 1500 to 1800 kcal per day for men to promote a 1-pound to 2-pound weight loss per week.<sup>54</sup> A sustained weight loss of as little as 3% to 5% of initial body weight reduces the risk of type II diabetes and risk factors for CVD.54

The Look AHEAD (Action for Health in Diabetes)<sup>55</sup> and Diabetes Prevention Program (DPP)<sup>56</sup> trials are 2 of the most successful long-term studies to illustrate the ability of lifestyle interventions to reduce and maintain body weight and reduce the risk of chronic diseases; both were instrumental in informing the 2013 AHA/ACC/TOS obesity treatment guidelines. The lifestyle interventions in these 2 trials were similar (ie, low-fat, low-calorie diet with the use of meal replacements and 150-175 minutes per week of moderate-to-vigorous-intensity PA).

Subsequently, some studies have demonstrated that higher protein diets (1.2-1.6 g protein/kg of body weight per day) provide benefits beyond weight loss and may preserve lean body mass (LBM), especially in older men and women.<sup>57,58</sup> Resistance training also has been shown to be particularly beneficial in older adults, including breast and prostate cancer survivors, to preserve LBM and bone health, maintain a higher resting metabolic rate, preserve physical functioning, and reduce falls and injury).<sup>59-61</sup>

Successful long-term weight management requires several behavioral strategies.<sup>62</sup> Tactics for weight success include: maintaining a low-fat, low-calorie dietary pattern; limiting dietary variety; eating breakfast most days of the week; daily to weekly weighing; performing 2500 kcal per week of PA (eg, brisk walking for approximately 1 hour per day); and reducing television watching.

## Interventions for PA in Other Populations That are Applicable to Cancer Survivors

Over the past few decades, numerous studies have clearly shown that PA of sufficient volume and intensity reduces the risk of several chronic diseases and improves physical function.<sup>63</sup> More recently, research has started to examine the impact of physical inactivity on overall morbidity and mortality.<sup>64-66</sup>

Aerobic PA of sufficient volume and intensity ("exercise") to improve cardiorespiratory fitness, which is a potent biomarker of morbidity and all-cause mortality, needs to be frontline care in both healthy and cancer survivor populations.<sup>67-69</sup> Consistent aerobic exercise can delay the onset of disability by more than 10 years and markedly increases survival among older adults with projected lifespans of at least 20 years (the length of survivorship that burgeoning numbers of cancer survivors are now achieving).<sup>7,70</sup> Among cancer survivors, data indicate that high versus low cardiorespiratory fitness reduces the risk of mortality by 45%.<sup>71</sup> Likewise, resistance PA of sufficient volume and intensity ("exercise") to increase neuromuscular fitness (ie, LBM, strength, power, fatigue resistance) is key to frontline care. Low LBM is a major predictor of all-cause mortality and physical disability.<sup>72</sup> Resistance exercise has repeatedly been shown to improve neuromuscular fitness and skeletal health and to reduce the risk of disability.73,74

Although the molecular underpinnings of PA-driven health benefits have not been fully elucidated, significant progress has been made,<sup>75</sup> and more information will be gleaned via the National Institutes of Health (NIH)-funded Molecular Transducers of Physical Activity Consortium (MoTrPAC), which is charged with mapping molecular responses to aerobic and resistance exercise to more fully understand the cellular and molecular signals that drive potential health benefits. This vital step will enable a precision medicine approach and the individualization of exercise prescriptions.<sup>63,75</sup>

## Interventions for Weight Management in Cancer Survivors

Data indicate that weight loss can be promoted among cancer survivors who are overweight or have obesity.<sup>31,76</sup> Sentinel studies of weight management are summarized in Table 3<sup>27-30,77-95</sup> and generally rely on cornerstone elements of weight loss (ie, dietary modification to promote caloric restriction, increased PA, and behavior modification). Nonetheless, there are acknowledged limitations to this research (eg, brief study periods, lack of repeated and objective measures [including body composition outcomes], and overrepresentation of breast cancer survivors who may be "worried, white, and well"). Many questions remain within the context of well-designed and controlled efficacy trials (eg, intervention timing, inclusion of sleep hygiene or stress management components, and discerning the full impact of weight loss across a broad range of symptoms and conditions). Discovery is needed to inform personalized medicine-based approaches and thereby elucidate molecular and metabolic predictors important for tailoring weight loss regimens for individual patients in terms of dose and optimal macronutrient distribution. In addition, there is a need for pragmatic interventions that can overcome well known barriers imposed by distance, economics, co-occurring medical conditions, and culture. The diversity of needs among cancer survivors, many of whom are older, increases the urgency of pragmatic trials to test and compare both hightouch/more-effect approaches and lower-touch/less-burden approaches. Well-designed research across the spectrum requires broad representation by cancer-type, age, sex, and race/ethnicity as well as sufficient sample sizes to conduct subgroup analyses. Ideally, interventions need to be designed with the input of oncologists, dietitians, exercise specialists, behavioral scientists, statisticians, software specialists (if needed), community stakeholders, and, most important of all, cancer survivors. The input of health economists also is key to developing programs that are sustainable and can be widely disseminated.

## Interventions to Improve PA in Cancer Survivors

Various PA interventions have been evaluated in cancer survivors, although the body of evidence is primarily limited to short-term studies of 12 to 16 weeks' duration among breast cancer survivors. Approaches are typically clinic-based or home-based (eg, telephone counseling, print, Web, social media). In general, stronger outcomes are associated with clinic-based programs, whereas greater reach and reduced participant burden are associated with home-based interventions.<sup>96</sup> However, this generalization is affected by the motivation of the cancer survivor; as shown by data from the LEAN (Lifestyle, Exercise, and Nutrition) study, which found no differences in effects between the modes of delivery.<sup>30</sup> On-site, clinic-based programs are generally supervised by exercise professionals and tend to have higher exercise intensity dose and closer supervision and monitoring. Home-based programs tend to promote

TABLE 3. Select Weight	Select Weight-Management Intervention Trials in		er Survivors by	Cancer Survivors by Mode of Delivery		
REFERENCE (NAME)	CANCER TYPE	NO. OF PARTICIPANTS	MEAN AGE, y	INTERVENTION	DURATION	WEIGHT CHANGE, kg
Clinically based/ supervised Thompson 2015 <sup>77</sup> (CHOICE)	Breast, postmenopausal	249	54.9	RD counseling, 42-d cycle menu, low fat or low carbohydrate, 3500 kcal deficit/wk, 10,000 steps/d	6 то	-8.9, -10.5, -0.3 kg: Low fat, low carbohydrate, and control, respectively
Swisher 2015 <sup>78</sup> (GetFit for Fight)	Breast, triple-negative	28	53.7	RD counseling to decrease fat intake by 200 kcal/wk; exercise physiologist-supervised, moderate aerobic PA 3 times/wk/2 unsupervised stretching + resistance training sessions	12 wk	— 3.0 vs —0.4 kg: Intervention vs control
Thomson 2010 <sup>79</sup> (modified Atkins)	Breast, stage I-II	40	56	RD counseling: low fat or low carbohydrate, 500 kcal deficit/d	6 то	-6.3, $-5.9$ kg: Low fat vs low carbohydrate, respectively
McCarroll, 2014 <sup>80</sup> (SUCCEED)	Endometrial, stage I-II	75	Not reported	Physician-led group and individual courseling, 16 sessions; diet, PA and behavior modification	6 то	$-1.5 \text{ kg/m}^2 \text{ vs} + 0.1 \text{ kg/m}^2$ : Intervention vs usual care
Travier 2014 <sup>81</sup>	Breast, stage I-IIIb	42	55.8	Twelve wkly, 1-h group sessions with RD; 1200- 1500 kcal/d; PA: 24 biwkly, 75-min supervised aerobic + strength sessions	12 wk	– 7.8 kg (Completers): Phase 2, single arm
Saxton 2014 <sup>82</sup>	Breast	85	56	Three small-group, supervised exercise sessions/ wk: aerobic + strength; weight loss on a plate Scottish Dietetic Association program/600 kcal/d deficit; wkly small-group nutrition seminars	6 що	-1.1 vs $-0.4$ kg: Intervention vs usual care
Mixed-modalities: Clinic-based and						
terephone contraming Rock 2015 <sup>27</sup> (ENERGY)	Stage I-III, breast	692	56	RD-led, 4 mo of wkly group sessions, tapering to biwkly, then monthly, reinforced by 1:1 telephone/email; deficit 500-1000 kcal/d + PA 60 min/d; tailored print materials	2 y	-3.6 vs -0.9 kg: Group + telephone vs control
von Gruenigan 2008 <sup>83</sup>	Stage I-II, endometrial	45	54.5	RD-led + MD-led wkly, then biwkly, then monthly group sessions (9); telephone or newsletter every nongroup meeting wk; walking 5 d/wk for >45 min	6 що	-3.3 vs +2.1 kg: Group + telephone vs usual care
Sheppard 2016 <sup>84</sup> (Stepping Stone)	Breast	22, Black (analytic)	I	Nutritionist/exercise physiologist-led, individually tailored group sessions twice monthly/alternate wk telephone counseling by trained survivor coach; survivor and interventionist toolkit; ACS guidelines for diet (lower fat + fruits and vegetables) and PA (10,000 steps)	12 wk	-0.8 vs +0.2 kg: Intervention vs control
Telephone counseling Harrigan 2016 <sup>30</sup> (LEAN)	Breast	100	20	RD/exercise physiologist-delivered; 1:1 wkly $\times$ 4 wks, biwkly $\times$ 8 wks, monthly $\times$ 3 mo; DPP adaptation with 500 kcals/25% fat kcals = 150 min PA/wk (walking, lower sedentary time) + mindfulness; self-monitoring	6 m o	Three-arm RCT (in-person, telephone, control), – 5.6, – 4.8, – 1.7 kg; in-person, telephone, and usual care, respectively

Herefore, numberMercy matrix mercy matrix mercy matrixMercy matrixMercy matrixMercy matrixMercy matrix mercy matrixMercy matrix <th>TABLE 3. Continued</th> <th></th> <th></th> <th></th> <th></th> <th></th> <th></th>	TABLE 3. Continued						
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SA     Breast, 11-13M01- 13M00 on letrozole     338     61     Trained lifestyle coaches: 19 calls + workbook or end of a low leads; 159-200 min PAWk; behavior nonfriction     24 mo       Breast, stage HII     91     589     Trained lifestyle coaches: 19 calls + workbook or end of starts; 250 min PAWk; behavior and resistance; 581 min PAWk; behavior and resistance; 581 movies are resistance; 5	Harris 2013 <sup>85</sup> (CASTLE)	Breast, stage I-IIIa	52	52.8	Health professionals employed by Trestle-Tree, Inc (San Marcos, TX) provided 15-min to 60-min telephone coaching wkly $\times$ 25 wk, then monthly $\times$ 6 mo; behavioral targets, diet and PA	12 mo	—4.0 vs —3.3 kg: Telephone vs in-person, respectively
Break, stage HI         01         58.9         Trained group leads, 34 conference calls in PANA, captors of 14, kcal effort, mad with pand kcalks(it mad inelacements, 255 min PANA; (172 plase 2)         6 mo calls mad inelacements, 255 min PANA; erobic and resistance; self-montoning         6 mo calls mad inelacements, 255 min PANA; erobic and resistance; self-montoning         6 mo calls mad inelacements, 255 min PANA; erobic and resistance; self-montoning         6 mo           Breast, stage         70         56.6         Phase 1, se abover, maintenance: new kell         6 % 12 mo           Breast, stage         70         56.6         Poh, heath educators, trailers delivered; anield revolaterer         12 wk           Duil, Hispanic         70         56.6         Poh, heath educators, trailers delivered; anield revolaterer         12 wk           Densat         23 African         51.4         Study calsers + 2 mo contrables + three drono education         12 wk           Breast, stage Ivil         48         36.70         Nuth meetings vs 11 wh P0 wky × 3         12 mo           Breast, stage Ivil         42. Hispanic, black         50.6         New Min eterings vs 11 wh P0 wky × 3         12 mo           Breast, stage Ivil         42. Hispanic, black         50.6         New Min eterings vs 11 wh P0 wky × 3.6         12 mo           Breast, stage Ivil         43. Hispanic, black         50.6         New Min eterings vs 11 wh P0 wky × 3.6	Goodwin 2014 <sup>28</sup> (LISA)	Breast, T1-T3N01 N3M0 on letrozole	338	61	Trained lifestyle coaches; 19 calls + workbook- directed call content adapted from DPP; -500 to -1000 kcal/d; 150-200 min PA/wk; behavior modification	24 mo	-3.1 vs $-0.3$ kg: Telephone vs print alone (all received general health educational print material)
Breast, stage I-II         Rurdi: 210         58         Phase 1, as above, maintenance: new kell         6 & 12 mo           (172 phase 2)         0ail: 2 min Pakki;         continued bukky goup conference cals on maintenance: new kell         6 & 12 mo           Breast, stage         70         56.6         RD. MD, heith educators, trialist delivered;         12 wk           Breast, stage         70         56.6         RD. MD, heith educators, trialist delivered;         12 wk           Breast, stage         70         56.6         RD. MD, heith educators, trialist delivered;         12 wk           Breast         23. African         51.4         Study bass with         6 mo           Breast, stage I-II         48         36-70         NW Wm retrugtum         6 mo           Breast, stage I-II         48         36-70         NW WM metings v1: with RD wkly × 3         12 mo           Breast, stage I-II         48         36-70         NW WM metings v1: with RD wkly × 3         12 mo           Breast, stage I-II         48         36-70         NW WM metings v1: with RD wkly × 3         12 mo           Breast, stage I-II         48         36-70         NW WM metings v1: with RD wkly × 3         12 mo           Breast, stage I-II         48         36-70         NW WM wetings v1: with RD wkly × 3	Befort 2012 <sup>86</sup>	Breast, stage I-III	6	58.9	Trained group leads; 24 conference calls in groups of 8-14; kcal deficit, -500 to -1000 kcals/d; meal replacements; 225 min PA/wk, aerobic and resistance; self-monitoring	6 то	— 12.5: Single arm, group telephone
Breast, stage         70         56.6         RU, Mi, health education, trialists delivered; out, Hispanic         12 wk           0-III, Hispanic         7         55 h cooking sessions + 2 food shopping field trips over 12 wk (24 h), cuturally tailored cook foor Your Life curriculum         12 wk           Breast         23. African         51.4         Stwd educators, trialists delivered; trips over 12 wk (24 h), cuturally tailored cook         12 wk           Breast         23. African         51.4         Stwd educators, trialists delivered; trips over 12 wk (24 h), cuturally tailored cook         6 mo           Breast, stage I-II         48         36-70         Wky WW meetings vs 1:1 with RD wkly × 3         12 mo           Breast, stage I-II         48         36-70         Wky WW meetings vs 1:1 with RD wkly × 3         12 mo           Breast, stage I-II         48         36-70         Wky WW meetings vs 1:1 with RD wkly × 3         12 mo           Breast, stage I-II         48         36-70         Wky WW meetings vs 1:1 with RD wkly × 3         12 mo           Breast, stage I-II         42, Hispanic, black         50.6         Curves staffed Curves weight-management         6 mo           Breast, stage I-II         42, Hispanic, black         50.6         Curves staffed Curves weight-management         6 mo	Befort 2016 <sup>29</sup>	Breast, stage I-III	Rural: 210 (172 phase 2)	58	Phase 1, as above; maintenance: new kcal goal; 2 meal replacements/d; 225 min PA/wk; continued biwkly group conference calls or mailed newsletter	6 & 12 mo	-12.2 vs 13.2: Regain +3.3 vs +4.9 kg; group telephone vs newsletter
Breast23, African51.4Study lead and certified exercise instructor from the community, 24 wkly classes with predetermined topics: diet, PA, and behavior modification; food shopping field trip, menu planning6 moBreast, stage I-II4836-70Wkly WW metrings vs 1:1 with RD wkly × 3 mo, binkly × 3 mo, monthly × 6 mo, binkly self-monitoring12 moBreast, stage O-IIIa42, Hispanic, black50.6Curves staff-led Curves weight-management self-monitoring6 moBreast, stage O-IIIa42, Hispanic, black50.6Curves staff-led Curves weight-management stength training + low-impact areobic PA; 60% increasing to 75% maximal heart rate; six 1-h wkly untrition classes, 1200 kcals/d increasing to to block classes	Community based Greenlee 2015 <sup>87</sup> (Cocinar Para Su Salud)	Breast, stage 0-III, Hispanic	70	56.6	RD, MD, health educators, trialists delivered; four 2-h nutrition education roundtables + three 3.5-h cooking sessions + 2 food shopping field trips over 12 wk (24 h); culturally tailored Cook for Your Life curriculum	12 wk	-2.5 vs $+3.8$ : Intervention vs control
Breast, stage I-II     48     36-70     Wkly WW meetings vs 1:1 with RD wkly × 3     12 mo       Reast, stage I-II     42, Hispanic, black     50.6     Wkly WW meetings vs 1:1 with RD wkly × 3     12 mo       Breast, stage 0-IIIa     42, Hispanic, black     50.6     Curves staff-led Curves weight-management     6 mo       Strength training + low-impact aerobic PA; 60%     Strength training + low-impact aerobic PA; 60%     6 mo       Mky nutrition classes, 1200 kcals/d increasing to 15% maximal heart rate; six 1-h     9 wkly nutrition classes, 1200 kcals/d increasing to 1600 kcals/d + wkly MI telephone calls	Stolley 2009 <sup>88</sup> (Moving Forward)	Breast	23, African American	51.4	Study lead and certified exercise instructor from the community; 24 wkly classes with predetermined topics: diet, PA, and behavior modification; food shopping field trip, menu planning	6 то	-2.52: Single arm
2013 <sup>90</sup> Breast, stage 0-Illa 42, Hispanic, black 50.6 Curves staff-led Curves weight-management 6 mo program curriculum; exercise at Curves 30 min, 3 times/wk + 2 d at home; bidirectional strength training + low-impact aerobic PA; 60% increasing to 75% maximal heart rate; six 1-h wkly nutrition classes, 1200 kcals/d increasing to 1600 kcals/d + wkly MI telephone calls	Commercial program Djuric 2002 <sup>89</sup> (NVV)	Breast, stage I-II	48	36-70	Wkly WW meetings vs 1:1 with RD wkly × 3 mo, biwkly × 3 mo, monthly × 6 mo; both promoted low fat, 500-1000 kcal deficit/d + self-monitoring	12 mo	-2.6, -8.0, -9.4, and -0.85; Control, WW alone, individual (RD counseling), and WW + individual, respectively
	Greenlee 2013 <sup>90</sup> (Curves)	Breast, stage 0-Illa		50.6	Curves staff-led Curves weight-management program curriculum; exercise at Curves 30 min, 3 times/wk + 2 d at home; bidirectional strength training + low-impact aerobic PA; 60% increasing to 75% maximal heart rate; six 1-h wkly nutrition classes, 1200 kcals/d + wkly MI telephone calls 1600 kcals/d + wkly MI telephone calls	6 mo	-2.87 vs -1.42: Curves program vs waitlist control

REFERENCE (NAME)	CANCER TYPE	NO. OF PARTICIPANTS	MEAN AGE, y	INTERVENTION	DURATION	WEIGHT CHANGE, kg
Home-based, print materials Morey 2009 <sup>91</sup> (RENEW)	Breast, prostate, colorectal	641, Older	73	Health counselor-delivered intervention; person- ally trailored workbook + quarterly newsletters +15 telephone counseling sessions and 8 prompts; PA: 15 min strength training twice wkly + 30 min aerobic PA/d; high fruit/ vegetable + low saturated and total fat; kcal restriction to promote up to 1 lb/wk	12 mo	— 2.06 vs — 0.92, Intervention vs control
Demark-Wahnefried 2014 <sup>92</sup> (DAMES)	Breast, stage 0-III	N = 136; 68 mothers, 68 daughters	I	Tailored print materials; ACS and US Dietary Guidelines; Kcal deficit of 500-1000/d, remove or substitute 3 major diet recall-identified caloric sources; 150 min/wk PA; self-monitoring	12 mo	-3.77 vs $-0.87$ In mothers (individual vs control)
Technology-based Huang 2014 <sup>93</sup> (Fit4Life)	ALL	80 M	13	Professional, caregiver, and parent + survivor input to material development; 4-mo Web-based and text and telephone counseling; ACS and Children's Oncology Group guidelines for healthy weight; kcal deficit, 60 min moderate-vigorous PA/d; 15,000 steps/d; self-monitoring	4 mo	-0.1 vs +1.4 Web SMS, telephone vs control
McCarroll 2015 <sup>94</sup> (Loseit! App)	Stage I-II, endometrial and breast	50	58.4	Study personnel-assisted application (App) orien- tation (30-60 min); Loseit! App for daily self- monitoring of diet and PA; low carbohydrate, high fiber; 150 min PA/wk, vigorous > 40 min/wk	4 wk	-2.3, Single arm
Haggerty 2016 <sup>95</sup> (Text4Diet)	Endometrial, pre and stage 1	20 y	60.5	Telemedicine-adapted DPP delivered by MD and MS clinician; weekly telephone and Wi-Fi weigh- ins daily $\times$ 6 mo; SMS received Text4Diet with 3-5 personalized SMS daily with monthly themes; 2-way communications, wkly Wi-Fi weigh-ins; all educated to consume 1200-1500 kcals/d; self- monitoring	о Ш	-9.7 vs -3.9, Telemedicine vs SMS

moderate-to-light PA, reach individuals who cannot travel or meet the scheduling requirements of on-site programs, are more likely to be theory-based, and are less costly.<sup>97</sup> However, supervision for home-based programs may be minimal, so individuals with significant comorbidity or safety issues are generally excluded. More recently, hybrid programs that are able to support sustained PA have emerged. Generally, these programs begin with an on-site, supervised phase and then taper to an off-site phase (eg, the CHALLENGE trial).<sup>21</sup>

A review of behavior change studies, including a metaanalyses of 14 RCTs among breast cancer survivors,98 found that the key elements of effective interventions are selfmonitoring of PA, individualized guidance or coaching, and setting clear goals and expectations. Because PA maintenance may be particularly challenging with long-term (eg, fatigue) or later effects of cancer treatment (eg, arthralgia), attention to symptom management (which ideally starts as prehabilitation and continues across the cancer survivorship trajectory) may be an important consideration to optimize PA uptake and adherence long term.<sup>21,99</sup> Likewise, there is potential for interventions (especially PA) to reduce these symptoms and thus contribute to the survivor's ability to maintain healthy behavior changes over time. Given that obesity may adversely affect adherence to PA regimens, there is a need to determine the relative timing or sequence of PA in relation to caloric restriction within the context of weight management.<sup>73,100</sup> As indicated above, research is needed to determine what type of PA works (modality [eg, aerobic, strength], intensity, frequency, and duration) to achieve which outcomes and for whom,<sup>101</sup> as well as when in the course of the cancer continuum programs should be offered. Identifying the minimal PA dose for QOL improvements, weight and symptom management, and survival will assist in developing pragmatic programs that target outcomes relevant to patients' needs (eg, management of fatigue and pain).

Looking ahead, improvements in the recovery and functioning of the growing numbers of cancer survivors may emerge from use of behavior change theories to inform intervention development, use of information technology and mHealth technologies to widen the reach of programs, plans for maintenance of behavior change, and assessment of program costs. There are numerous missed opportunities for health care professionals in oncology and primary care settings to promote PA for their patients at various points across the cancer care continuum. Addressing the barriers faced by providers<sup>96,102</sup> and providing guidelines to help triage patients to effective programs (eg, clinic-based, community-based, or home-based) are sorely needed.

#### Summary

Research in the general population has demonstrated the benefits of weight management and PA for the prevention

and management of diabetes and other chronic diseases, reducing disability, and delaying mortality. As research on weight management, PA, and cancer survivorship moves beyond small, early clinical trials that evaluate the effects of these interventions on biomarkers and QOL, larger trials are needed to test the effects of the interventions on disease-free and overall survival, especially in disease sites other than breast cancer, and to include adequate representation of population subgroups defined by comorbidity/ functional status, race, ethnicity, or age. To ensure that clinical trials are appropriately designed to provide more definitive answers, the NIH has recently established new guidance related to funding applications submitted to the NIH for all clinical trials.<sup>103</sup> Specific institutes, such as the National Cancer Institute, have used expert working groups to discuss trial design issues within their clinical trial networks, including those related to behavioral interventions.

Within the NIH Obesity Research Task Force, the National Heart Lung and Blood Institute is leading an effort to identify additional factors that may predict successful response to weight management interventions. The Accumulating Data to Optimally Predict obesity Treatment (ADOPT) Core Measures Project was developed in response to the well documented individual variability in response.<sup>104</sup> ADOPT is designed to provide investigators with tools to generate an evidence base that may advance understanding of the behavioral, psychosocial, environmental, and biologic sources of this variability. Working with an expert panel of investigators, a trans-NIH group identified an initial core set of high-priority measures that, when consistently used in trials, may facilitate the prediction of treatment response. The NIH is now exploring approaches for increasing the use of consistent measures across trials so that data can be pooled and used to identify reliable predictors, mediators, and moderators of response. This accumulation of efficacy evidence will likely spur the translation of effective interventions into clinical practice, although research in implementation science also is needed to best adapt interventions to enhance their reach, scope, and uptake among populations and settings that may not be representative of clinical trials.

### Addressing the Weight Management and PA Needs of Diverse Populations of Cancer Survivors

Low-income, minority populations, particularly African Americans and Hispanics, as well as those who are older and live in rural areas bear a disproportionate burden of cancer.<sup>7,105-107</sup> Moreover, these populations also are more likely to be overweight or obese, physically inactive, and to manifest health conditions that are affected by these factors, such as metabolic syndrome—all of which are associated with greater comorbidity and reduced survival

DOMAIN	ACTION
Peripheral	Design study materials to appear culturally appropriate (ie, logo, recruitment materials)
Evidential	Enhance relevance of targeted health issues by presenting evidence of their impact (ie, cancer disparities, impact of obesity, comorbidity burden)
Constituent-involving	Draw directly on the experiences of the target group (ie, staff represent target group; inform intervention using qualitative data from target population; engage advisory group to provide feedback on study materials and procedures)
Sociocultural	Discuss health-related issues in the context of broader social and/or cultural values (ie, role of God and faith in one's daily life, woman's central role in families, cancer fatalism and stigma, body image ideals, and the traditional roles of food)
Linguistic	Make health education programs and materials more accessible by providing them in the dominant or native language of target group

(overall and cancer-specific). This section addresses weight management and PA among diverse populations.

## Meeting the Needs of Diverse Survivors in Terms of Race/Ethnicity, Culture, and Language

Among cancer survivors, prevalence of overweight and obesity is high, especially among non-Hispanic blacks and Hispanics compared with non-Hispanic whites.<sup>108</sup> Concordantly, cancer survivors who are members of minority groups have lower adherence to diet and PA guidelines and are more likely to report poorer health status compared with nonminority cancer survivors or racial/ethnic minorities without cancer.<sup>109-115</sup> Thus, there is a critical need to develop and examine weight management and PA interventions among cancer survivors of racial/ethnic minority status to enhance outcomes and reduce disparities. Consideration of patients' environmental, cultural, and survivorship context is critical to the success of these efforts.<sup>116</sup> Racial and ethnic minority survivors are more likely to live in areas characterized by high segregation, traffic density, and crime and by low neighborhood socioeconomic status and access to fullservice supermarkets and PA resources.<sup>117-119</sup> Despite this, most communities, including lower socioeconomic neighborhoods, also have some assets, such as farmers' markets, public recreation systems, and community gardens that support PA, healthy eating, and reduction of chronic disease.<sup>120-122</sup> Partnering with community organizations to bring interventions to under-resourced neighborhoods provides opportunities to build social capital, reach more cancer survivors, and increase potential for sustainability.<sup>123</sup>

Consideration of cultural norms is important. Culture varies among and within racial and ethnic groups, influencing beliefs, behaviors, and patient-provider interactions related to cancer, obesity, and lifestyle. The conceptual framework of Kreuter et al<sup>124</sup> can inform cultural tailoring and structure formative work, thereby enhancing the relevance of an intervention approach and content to a particular population (see Table 4).<sup>124</sup> Similarly, because cancer survivors report greater interest in programs that acknowledge their cancer journey and concerns, it is important for programs to address these issues.<sup>125</sup> In addition to context, biopsychosocial approaches to research are needed to understand and address the multilevel factors (from cells to society) that affect weight status and behavior and influence cancer control, overall health, and QOL.<sup>107</sup>

To date, weight management and PA interventions among racial/ethnic minority cancer survivors have established feasibility and safety and report positive, albeit modest, results, including weight loss, behavioral changes, improved QOL and biomarker status, and decreased cancer-related anxiety.<sup>84,87,88,90,126-133</sup> Limitations of many studies include quasi-experimental designs, small sample sizes, a focus on behavioral outcomes, and sole inclusion of breast cancer survivors. Only 3 studies assessed biomarkers,<sup>87,90,128</sup> and only one targeted a cancer other than breast<sup>134</sup>—none included men. Recent efforts address some of these limitations.<sup>135,136</sup> Important steps to advance the science of obesity and lifestyle interventions in diverse populations include using more rigorous methodologies, addressing multilevel mediators and moderators of change, examining biologic mechanisms related to energy balance and cancer, and addressing more diverse cancer survivor populations.

## Meeting the Needs of Cancer Survivors Across the Lifespan

Currently, 62% of cancer survivors are age 65 years and older—a subpopulation that will continue to grow with the aging of the population along with earlier diagnosis and improvements in treatment.<sup>137</sup> However, cancer also affects the young. In the United States, there are almost 400,000 childhood cancer survivors and 70,000 adolescent and young adult cancer survivors, many of whom could have long lives ahead of them.<sup>137</sup> Lifestyle interventions are sorely needed by survivor populations of all age groups, because suboptimal diets and insufficient PA are noted in 40% to 70% and 54% to 84% of younger cancer survivors, respectively,<sup>138-141</sup> and in 52% to 85% and 53% to 70% of older cancer survivors, respectively.<sup>142-144</sup> In addition, up to 71% of older cancer survivors are overweight or have obesity, and these conditions also are prevalent in children, adolescents, and young adults diagnosed with acute lymphocytic leukemia and some forms of brain cancer.<sup>140,142</sup>

A unifying theme shared by both young and old cancer survivors is that of the long-term and late effects of cancer and their treatment, many of which are influenced by nutritional status and PA,<sup>145</sup> such as increased risk of CVD, second cancers, osteoporosis, metabolic syndrome, fatigue, cognitive changes, and sarcopenia. Underlying many of these conditions is the process of accelerated aging and frailty among cancer survivors, which occurs across all age groups.

Frailty, or an "insufficient reserve to recover," is generally preceded by diminished function,<sup>146</sup> both of which increase with age. However, illnesses and injury, as occur with cancer and its treatment, accelerate this course, especially among females.<sup>145</sup> Other factors, such as a poor diet, physical inactivity, and obesity, further exacerbate functional decline and the onset of frailty.<sup>147</sup> Current data indicate that the odds of frailty are significantly increased among individuals with a BMI of 30 kg/m<sup>2</sup> or greater (odds ratio, 1.12; 95% CI, 1.01-1.19 [P = .003]).<sup>148</sup> Weight management and PA interventions can potentially reorient neuromuscular control, increase muscle strength, and reduce frailty.<sup>33,60,61,149</sup>

Despite the potential benefits of weight management to forestall frailty and common comorbidities, caution is needed in pursuing weight loss. Until more data are available specifically on cancer survivors, the AHA/ACC/TOS and National Heart Lung and Blood Institute guidelines can inform best practices.<sup>54,150</sup> Among these guidelines is the recommendation for a rate of weight loss of 1 to 2 pounds per week. Because sarcopenia is a common condition that accompanies cancer and its treatment and one that accelerates aging, slower rates of weight loss that minimize LBM loss, concurrent with strength training, are recommended.<sup>151</sup> For older adults, weight loss guidelines suggest an energy deficit of 500 to 750 kcal daily to promote a weight loss of up to 1.5 pounds per week<sup>152</sup>; for childhood and adolescent cancer survivors, energy deficits of up to 250 kcal per day to invoke a maximum weight loss of 0.5 pounds per week are recommended.<sup>153</sup> Also, behavioral approaches, such as substituting higher energy with lower energy density foods, avoiding distracted eating, and adopting slower rates of eating, are commonly used tactics to help prevent weight gain in adults and allow children to "grow into their weight."

Regular PA is important for cancer survivors of any age to achieve optimal health. Thus the avoidance of inactivity is key, and adaptations need to be made to accommodate limitations or comorbidities because of cancer or its treatment.<sup>24</sup> Guidelines suggest that children pursue 300 minutes per week of moderate to vigorous PA, whereas the guidelines for adults (including older adults) suggest 150 minutes of moderate PA or 75 minutes of vigorous PA per week.<sup>25,154</sup> Strength training 2 or 3 times per week is recommended across the lifespan<sup>25,154</sup>; although, for children, this recommendation is made within the context of a sports curriculum and with adequate supervision.<sup>155</sup> The benefits of low-intensity PA have recently been reported as well; however, to date, there are no guidelines in this area.<sup>156</sup> The means by which weight management and PA are promoted in younger versus older cancer survivors differs.<sup>157-161</sup> For example, children have preferences for game-based or playbased interventions, whereas older adults favor holistic interventions (eg, gardening, dancing) that have personal meaning and/or involve others. Because of prevalent functional and sensory deficits among the young and the old, it is critical that interventions use large font (and screen size), volume control, module brevity, pretraining, and support for new technologies, especially those that allow for homebased delivery.<sup>162</sup> Exemplar interventions are featured in Table 3.91,93,163 In addition, given the key role that caregivers play in the lives of children and older adults, an unexplored area with potential is the use of dyadic approaches.

## Meeting the Needs of Rural Cancer Survivors

Rural cancer survivors (ie, those residing in nonmetropolitan counties, as defined by the Office of Management and Budget)<sup>164</sup> have higher cancer mortality rates compared with urban residents across all regions of the United States. Moreover, death rates from cancer have declined at a slower rate in rural compared with urban counties.<sup>165</sup> Among cancer survivors, those from rural areas report poorer health status, more psychological distress,<sup>166</sup> higher rates of depression and anxiety,<sup>167</sup> and greater knowledge gaps related to their cancer and its treatment effects. Rural cancer survivors also report high levels of unmet support needs.<sup>168</sup> These disparities are compounded by higher rates of comorbidities, obesity, and physical inactivity among rural compared with urban residents.<sup>169-172</sup>

Rural cancer disparities affect a significant proportion of our population. Nearly 20% of Americans,<sup>173</sup> and an estimated 21% of cancer survivors, reside in a rural area, representing roughly 2.8 million survivors.<sup>166</sup> Rural residents are a diverse group. Nationwide, 78% of rural residents are non-Hispanic white; however, higher proportions of African Americans and Hispanics reside in the south and southwest, respectively. Despite their diversity, rural residents often share common cultural elements, such as conventional attitudes, self-reliance, and orientation toward work, family, and religion.<sup>174</sup> Rural residents of all racial/ethnic groups are also older, poorer, and have less education than their urban counterparts.<sup>161</sup> These demographic differences, in addition to the contextual, cultural, and access factors stemming from place of residence, contribute to rural cancer disparities.<sup>175</sup> All of these factors need to be considered when designing lifestyle interventions for rural cancer survivors.

Effective, remote-based interventions are essential to maximize reach into rural communities because of challenges with access to health care services (including specialized services for weight management and PA), travel time, and financial barriers. With approximately 3% of medical oncologists<sup>176</sup> and very few specialized psycho-oncology providers practicing in rural communities, there is a gap in services for supportive care and lifestyle interventions.<sup>177</sup> Travel time and transportation costs pose barriers for inperson lifestyle interventions, particularly in frontier regions.

Survey results among rural breast cancer survivors in Kansas and Illinois found that the vast majority did not met PA guidelines<sup>178</sup> but rated PA and weight management programs as a top need.<sup>179</sup> To date, there is only one published trial of a full-scale lifestyle intervention done exclusively in a rural setting. Befort et al<sup>180</sup> enrolled 210 breast cancer survivors into a 6-month telephone-based intervention delivered by a weekly conference calls. The study demonstrated feasibility and achieved a 12.9% weight loss-with a 10.6% net loss maintained at 18 months-through continued, but scaled back, conference calls. Lessons learned from that study are needs for: 1) direct patient recruitment through cancer registries (direct mailing yielded 84% of participants, whereas physician referrals yielded only 4%); 2) clinical integration (because of high levels of comorbid conditions); and 3) group support among rural women.<sup>181</sup> While the intervention was exclusively home based, many participants arranged to meet in person with one another; therefore, some face-to-face contact may enhance intervention efficacy for some and needs to be considered in future programming. Additional research also is needed to better understand environmental determinants of diet and PA in rural areas and contextual factors influencing successful implementation across various health care and community settings.

#### Summary

Overall, there are limited data available on the effectiveness of weight management and PA interventions for diverse populations, although feasibility and safety have been established. In each unique population, various factors must be addressed to ensure that weight management, PA, and behavioral modification elements address physiologic needs and health issues (eg, promoting slower weight loss among pediatric and geriatric cancer survivors who are at greater risk for stunting and sarcopenia, respectively), individual preferences (eg, home-based delivery to overcome travel barriers), and community-based resources (eg, partnering with community-based organizations for program implementation). Given the higher prevalence of overweight and obesity and suboptimal lifestyle practices among certain subpopulations of cancer survivors (eg, racial/ethnic minorities, those residing in rural areas, pediatric and older cancer survivors) there is a need to target interventions to these diverse populations that are currently more likely to have poorer outcomes and shorter years of survival.

## Opportunities and Challenges for the Workforce, Care Coordination, and Technologies to Support Weight Management and PA in Cancer Survivors

Several factors currently limit the ability to deliver weight management and PA programs to all cancer survivors who need them. These barriers exist at multiple levels. Barriers at the level of the cancer survivor and family have been covered in the sections above and include factors such as high costs, lack of geographic access to these programs, or lack of knowledge or motivation of how to change health behaviors. These may be compounded by barriers at the level of the clinician, such as lack of clinician comfort with discussing weight with patients or lack of knowledge of which intervention to refer or prescribe, as well as competing demands for time in the clinical encounter. Finally, barriers at the level of the health care system and the environment present further challenges; for example, a lack of prioritization of PA, weight management, or disease prevention in general; a lack of insurance coverage for lifestyle change programs; or the obesogenic environment. These challenges and current strategies to overcome them are described below.

## Weight Management and PA: Clinical Care Opportunities and Challenges

A health care professional's recommendation to exercise significantly improves PA engagement,96,182 yet many providers do not counsel patients who might benefit on the need for PA or weight management. Research shows that providers are more likely to encourage a change in health behavior if they have established a positive patient-physician relationship, have available referral resources to facilitate health behavior change, and believe that health behavior engagement will benefit cancer outcomes or overall health and well being.<sup>183</sup> Several individual and systems barriers need to be overcome to promote PA and weight management in the delivery of survivorship care. Competing time demands during oncology visits dictate that oncology care providers help patients make difficult choices about therapy, monitor side effects, promote adherence to oral medications, administer screening evaluations, and help patients cope with the psychological effects of cancer diagnosis and

ORGANIZATION	RESOURCES AVAILABLE	WEB SITE	TELEPHONE
American Cancer Society	Survivorship guidelines (nutrition and physical activity; cancer-specific)	cancer.org	(800) 227-2345
American College of Sports Medicine	Physical activity guidelines for exercise professionals	acsm.org	(317) 637-9200
American Institute for Cancer Research	Health behavior information and recommendations	aicr.org	(800) 843-8114
American Physical Therapy Association/ The Oncology Section	Physical activity and safety consider- ations for the cancer survivor	apta.org	(800) 999-2782
American Society of Clinical Oncology	Survivorship Compendium, Obesity Toolkit	asco.org	(571) 483-1300
Cancer Nutrition Consortium	Nutrition Guidance	cancernutrition.org	(857) 301-8495
LIVESTRONG	Health behavior tools, LIVESTRONG at the YMCA	livestrong.org	(855) 220-7777
National Cancer Institute/Office of Cancer Survivorship	Facing Forward series, general recommendations, workshops and conferences	cancercontrol.cancer.gov/ocs	(800) 422-6237
National Center for Health Promotion & Disease Prevention/VHA	Weight management resources	move.va.gov	(844) 698-2311
National Comprehensive Cancer Network	Survivorship and disease-specific guidelines for health care providers	nccn.org	(215) 690-0300
National Heart, Lung, and Blood Institute	Weight management resources	nhlbi.nih.gov	(301) 592-8573
Silver Sneakers	Physical activity for older adults	silversneakers.com	(866) 584-7389

TABLE 5. Resources for Weight Management and Physical A
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treatment, presenting significant obstacles to discussions of PA and weight management.<sup>183-186</sup> Moreover, in the current age of electronic medical records, health care professionals in general spend only one-half of their scheduled clinic visit time talking with the patient and over one-third of their time on documentation.<sup>187</sup> Furthermore, most health care professionals do not receive adequate training in how to operationalize health behavior recommendations at the level of an individual patient,<sup>185</sup> so a generalized lack of expertise or competency in knowing what and how to recommend changes is a major barrier. Providers also need resources for appropriate services (eg, dietitians, exercise specialists, or programs)<sup>183,188,189</sup>; but a lack of supportive infrastructure, including lack of access to appropriate referrals or effective strategies as well as limited insurance coverage, can inhibit provider recommendations.<sup>184,185</sup> Care providers also may be skeptical that a cancer survivor can change behavior or may worry that engaging in weight management or PA during or after treatment may be risky to the survivor's overall health.<sup>183</sup> Finally, a lack of knowledge about the benefits of weight management and PA can result in a lack of motivation to focus on this topic.<sup>183</sup> Fortunately, there are several resources available for cancer survivors and health care professionals to help promote weight management and PA (see Table 5).

## Overcoming Workforce Issues and Establishing Common Competencies

The contribution of excess weight and physical inactivity to cancer and overall health outcomes emphasizes the need for health professional education and adoption of appropriate competencies. The lack of a standard of care for overweight and obesity and its associated lifestyle factors, the mismatch of disease burden with care provider capacity, and the lack of integrated clinical and community services constitute major barriers to effective care. Behavior change is the cornerstone of therapy. For the care of pediatric patients with obesity, the USPSTF recommends moderate-to-highintensity behavioral interventions, including nutrition, PA, and behavioral counseling for a minimum of 26 contact hours.<sup>190</sup> For adults with obesity, the USPSTF recommends obesity behavioral interventions that include selfmonitoring and from 12 to 26 visits over the course of a year.<sup>51</sup> However, few providers have been trained in the delivery of behavior change therapies; and, currently, few major insurance plans provide reimbursement for the duration of care recommended by the USPSTF.

A second gap is the lack of understanding of the most fundamental elements of obesity care. For example, less than 50% of internists, family practitioners, obstetricians/ gynecologists, and nurse practitioners surveyed knew the

#### TABLE 6. Competencies for Health Care Professionals in Obesity Prevention and Control (Bipartisan Policy Center 2017<sup>192</sup>)

Understanding the framework of obesity as a medical condition Knowledge of epidemiology and key drivers of the epidemic Knowledge of disparities and inequities in obesity prevention and care Providing interprofessional obesity care Integration of clinical and community care for obesity Use of patient-centered communication Recognition and mitigation of weight bias and stigma Accommodation of people with obesity Use of strategies for patient care related to obesity Recognition of acute warning signs of obesity complications

recommended level of PA for adults, and even fewer knew the USPSTF guidelines for treatment of obesity. Similar surveys have not been administered to oncology care providers; however, their knowledge in these areas is not likely to be better.

To address these gaps, 24 organizations involved in the care of obesity convened to develop common competencies for the prevention and treatment of obesity.<sup>191</sup> This effort, funded by the Robert Wood Johnson Foundation, led to the development of 10 major competencies, which are shown in an abbreviated form in Table 6.<sup>192</sup> The consensus-derived competencies are not intended to be an obesity curriculum. Rather, the expectation is that each of the groups involved in their development will adapt them to the needs of their specific profession; however, a limitation is that oncology was not represented among these groups.<sup>191</sup>

A final issue is the need for providers to be sensitive to the issue of stigma and bias. The stigma associated with obesity may be secondary only to race.<sup>193,194</sup> Because obesity is so highly stigmatized, providers who are uncomfortable with the topic and unaware of how to discuss it with their patients often add to this burden.<sup>193</sup> Consequently, patients with obesity may not receive the care they need. Therefore, the competencies include understanding terms that are acceptable to patients in discussions about weight and the need for joint decision making with respect to care.

## Opportunities and Challenges Posed by New Technologies

Self-monitoring is a strong predictor of weight management success,<sup>195</sup> but engagement with self-tracking declines over time.<sup>196</sup> New technologies have improved adherence over traditional paper modes, and pairing feedback with tracking optimizes behavior change.<sup>196</sup> However, the challenges of maintaining self-monitoring of diet and PA remain. Daily weighing, via an internet-connected scale paired with text message feedback, has been found to promote clinically meaningful weight loss of 6% within 6 months in healthy populations.<sup>197</sup> These data suggest that self-monitoring strategies that are both discrete and simple achieve high

engagement and desired clinical outcomes. However, mobile application (app) abandonment rates have been well documented in commercial and research settings.<sup>198,199</sup> Such data indicate that multiple strategies (including Web, e-mail, interactive voice response, and text messaging) are needed to keep users connected to feedback or coaching and are necessary to complement self-monitoring strategies-strategies that may vary by population subgroup. Current research shows that feedback strategies that are real-time responsive (eg, app messages) are better positioned than those that are delayed (eg, weekly coaching calls), but there is a need to better understand the reasons why users disengage from technology. Moreover, while technology is currently used to address data collection, analytics, and integration with a goal of providing actionable feedback to users, the integration with health care professionals to provide health care decision support has yet to be made and is a needed leap.

### Summary

The considerable body of research on weight management and PA interventions has documented myriad positive effects during and after cancer care. Despite the numerous challenges in delivering weight management and PA programs for cancer survivors, this is a time of unprecedented opportunity to include these programs as part of standard cancer and follow-up care. Several national trends and changes in health care are contributing to this opportunity while addressing some of the multilevel barriers to delivering these programs. At the level of the survivor and family, it is clear that engaged and activated patients participate more fully in their health care. Thus, efforts by clinical and public health groups have focused on patient activation and education about the importance of weight management and PA. Examples of this are patient education materials available on the websites of ASCO, the American Cancer Society (ACS), and the National Cancer Institute, as well as the Springboard Beyond Cancer mobile health tool (survivorship.cancer.gov) and SurvivorSHINE (survivorshine.org) websites, which help survivors change their lifestyle behaviors along with managing their ongoing symptoms and selfmanaging their health. The latter are examples of another ongoing trend discussed above: using technology to increase the feasibility of delivering interventions to survivors regardless of location. Technological solutions also may help reduce financial barriers to these programs for survivors who have limited economic resources.

At the level of the health care professional and clinical practice, there are now multiple guidelines from the ACS, ACSM, ASCO, NCCN, and others delineating weight management and PA as part of overall cancer care and follow-up care. The work described above, to establish weight management competencies for clinicians, will help increase

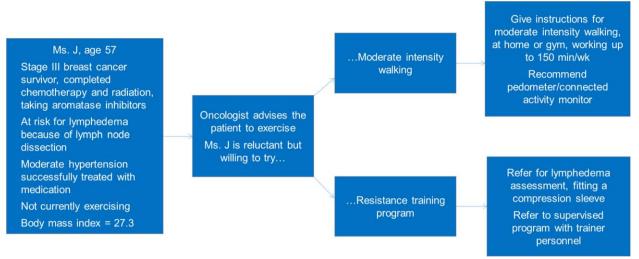


FIGURE 1. Example of Tailoring an Exercise Program to the Patient's Physical Needs and Preferences.

the ability of the clinical workforce to deliver appropriate care. In addition, health care professionals need safe and efficacious programs available for patient referral and time during the clinical encounter to discuss these issues and resources with patients and survivors. Interventions in community settings, such as LIVESTRONG at the YMCA, the availability of ACSM-certified Cancer Exercise Trainers in gyms across the country, and other programs for cancer survivors are helping to provide sources for appropriate referral.

Finally, changes in American health care present an opportunity to integrate weight management and PA programs into cancer and follow-up care. The transition from fee-forservice to value-based payment models is also bringing attention to identifying interventions that are "good buys" (ie, that result in positive effects in multiple ways, in terms of patient outcomes and cost savings). The multiple positive effects of weight management and PA on overall health, well being, and QOL are prompting health care funders and decision makers look more seriously at these interventions.

## Weight Management and PA Programs for Cancer Survivors: Models of Care

Successful incorporation of PA and weight management services into cancer survivorship care requires effective models of care delivery services. Because survivors come with a range of needs and preferences for services, oncology health care professionals need algorithms for assessing these factors to triage survivors and refer them to appropriate programs. Guidelines for providing smoking-cessation assistance in primary care settings use a schema that may prove useful in conceptualizing this process; the 5 As (Ask if a patient is smoking, Advise quitting, Assess patient motivation for making a quit attempt, Assist with counseling referral and pharmacotherapy, and Arrange follow-up within a week of quit date).<sup>200</sup>

## Ask/Advise

Referral of cancer survivors to PA and weight management services starts with a conversation between the health care professional and their patient about these issues. A survey of 15,254 cancer survivors in the United Kingdom found that survivors who recalled a conversation with their care provider about exercise were 88% more likely to be physically active and to meet PA recommendations<sup>201</sup>; however, only 31% of the respondents recalled such a conversation. In the US, discussion about PA is documented in only 35% of patientoncologist encounters.<sup>202</sup> Therefore, provider prompts may enhance discussions on this topic within the workflow and need to be considered in models of care delivery.

#### Assess

Cancer survivors have a range of needs and limitations regarding their symptoms, physical limitations, and comorbid health conditions that need to be considered. A process for assessing these conditions is necessary to determine optimal programs for weight loss or PA to ensure appropriate supervision and patient safety. At the same time, the cancer survivors' goals, preferences, and prior experience with PA and weight management need to be considered. The ACSM health screening guidelines, which take into account the patient's current activity level, signs and symptoms of disease, and desired intensity of exercise, are used to assess capacity for PA in the general population and may be useful to implement for cancer survivor populations.<sup>203</sup> Figure 1 provides an example of how survivor goals and preferences might interact in recommending an exercise program for a breast cancer survivor at risk for lymphedema.

## Assist (or Refer)

Few clinical practices providing oncology/survivorship care are able to provide PA and weight management services and

often require referral to outside programs. There are growing numbers of programs available; some are cancer specific (eg, LIVESTRONG at the YMCA), but programs intended for the general population may also be appropriate (eg, Silver Sneakers). In addition, home-based or self-directed programs that rely on books and other print material, websites, mobile apps, and wearable devices (like pedometers and connected activity monitors and scales) can assist cancer survivors with PA and weight management. Cancer survivorspecific versions of such programs have shown efficacy in research studies but are not widely available outside of the research setting. Cancer survivors who have physical limitations or comorbid medical conditions may need to start with a clinic-based program, such as a cancer rehabilitation program or a medical weight-loss program. A challenge for health care professionals is knowing which programs are reliable and of high quality for cancer survivors. There is a need to develop and refine program standards and staff certification to optimize safety and effectiveness.

#### Connect

Even if effective programs are available, there often is a need to motivate cancer survivors to follow up on recommendations to access services. Research on smoking cessation shows that placing the onus of contact on the service provider (assuming that the patients' permission is obtained to share their contact information) can boost enrollment by 13-fold to 30-fold.<sup>204,205</sup> A similar connection strategy could be used by an oncology clinic referring patients to programs like LIVESTRONG at the YMCA or cancer rehabilitation.

Cancer survivors need access to a variety of safe and effective programs and clinical services to assist them with increasing PA and managing their weight. In addition, there also is a need to increase provider competencies; develop tools to assist providers; and build capacity in workflow and procedures that increase the likelihood that the patient-provider discussion about weight management and PA transpire, that referral to appropriate programs that are aligned with survivors' needs and preferences occurs, and that follow-through takes place.

### **Exemplar Programs**

Over the past 2 decades, and with increasing recognition of the heightened needs of cancer survivors for supportive care to improve both emotional and physical health, several PA and weight management interventions have been developed and tested. Brief descriptions of some of these exemplars follow:

#### MOVE!

In 2006, the Veterans Health Administration implemented the MOVE! Weight Management Program for Veterans,<sup>206</sup> an evidence-based comprehensive lifestyle intervention available to veterans receiving care at all Veterans Health Administration

or have obesity and an obesity-associated condition, such as cancer or CVD, to achieve clinically significant weight loss.<sup>206</sup> The program adheres to evidence-based recommendations from the Department of Veterans Affairs/Department of Defense Clinical Practice Guideline for Screening and Management of Overweight and Obesity,<sup>207</sup> which provides flexibility in addressing the unique needs of cancer survivors. The Department of Veterans Affairs/Department of Defense guideline recommendations include: yearly screening and documentation of overweight/obesity; pharmacotherapy and bariatric surgery as adjuncts to comprehensive lifestyle intervention; shared decision-making among providers and patients to support patient engagement based on individual values and preferences; and repeated assessment of response to treatment, with adjustments as needed to ensure clinically meaningful weight change. MOVE! and weight management care are embedded within the health care system while also supporting care coordination across primary and specialty care settings (including oncology) and inpatient and outpatient care.

medical centers. MOVE! assists veterans who are overweight

### LIVESTRONG at the YMCA

Over the past decade the YMCA, which consists of a national resource center (YMCA of the USA, or Y-USA) and over 2700 local YMCAs, has partnered with many public health and health care stakeholders to transform its network to better serve the health of the nation. Local YMCAs are trained to build their capacity and to develop competencies needed to become a strong partner. Thereafter, they are licensed to deliver standardized programs and services for those with special health needs (eg, the YMCA's DPP for people with prediabetes, Healthy Weight and Your Child for families with children who are challenged by obesity, etc). To address the challenges faced by cancer survivors, Y-USA partnered with the LIVESTRONG Foundation to develop and scale a 12-week PA program.

The program model was nested within a 6-month organizational change process modeled after the Institute for Health Care Improvement's "Plan, Do, Study, Act." Local YMCAs that have an interest in serving cancer survivors within their communities are encouraged to submit a readiness assessment to Y-USA. The LIVESTRONG Foundation, the local YMCA leadership, and, more recently, the Centers for Disease Control and Prevention have supported YMCAs that demonstrate the highest levels of commitment to go through the organizational change process and become trained and authorized to deliver the LIVE-STRONG at the YMCA program. Through the organizational change process, YMCA staff build competencies, such as those related to understanding cancer survivors' needs, providing welcoming environments, and developing partnerships with local cancer centers and oncology health care professionals.

Data indicate that the program leads to significant improvements in PA, stamina, and QOL<sup>208</sup>; and it has been scaled to over 500 locations in 39 states, serving nearly 50,000 cancer survivors. Currently, philanthropy is required to make this program free of charge to all cancer survivors, and wait lists have been established in cases where demand exceeds availability.

In 2016, Y-USA completed a claims-based cost-savings demonstration of the YMCA's DPP that led to Medicare coverage of that program.<sup>209</sup> A partnership between Y-USA and the Robert Wood Johnson Foundation was instrumental in transforming the systems behind scaling and sustainability that produced health outcomes and value within the health care system. Connections were made between the electronic medical record system of health care organizations and local YMCA business units. Health care providers are increasingly referring patients to programs like LIVE-STRONG at the YMCA under alternative payment models (eg, Accountable Care Organization [ACO] structures or bundled payment models) in which the value and savings of these programs are reinvested in partnerships with the YMCA, and payers are providing coverage for the YMCA's DPP under claims-based reimbursements.

### The Physical Activity and Lymphedema and Strength After Breast Cancer trials

The challenge of knowing which cancer survivors can be safely referred to home-based and community-based exercise remains unresolved,<sup>210-212</sup> although initiating exercise programming within the context of outpatient rehabilitation clinics is a potential approach. The Physical Activity and Lymphedema (PAL) trial was revised to use this approach among breast cancer survivors after establishing the safety and efficacy of a weight-lifting intervention for lymphedema and other side effects of cancer and its treatment. 32,41,213-220 The PAL intervention was originally implemented in select YMCAs in which fitness staff received training (including preintervention safety evaluations) and ongoing support from PAL investigators. Because of high staff turnover at the YMCAs, concern about implementing safety evaluations, and an unwillingness of participants to continue to pay for YMCA memberships, this program led to the creation of "Strength After Breast Cancer"-a dissemination and implementation study to translate PAL into the setting of outpatient rehabilitation.<sup>221</sup>

As noted in other effective interventions, the "Strength After Breast Cancer" intervention optimally begins with a referral by the oncology/primary care provider to the outpatient rehabilitation clinic, which then contacts the patient. Clinic staff implement a safety evaluation, then deliver an educational session on lymphedema, as well as 4 sessions of weight-lifting instruction, before release to a home-based program. Favorable comparison of the efficacy of this revised program with the original PAL trial led to an online continuing-education course that targets outpatient rehabilitation specialists (klosetraining.com/course/online/strengthabc).<sup>221</sup> Over 400 outpatient rehabilitation clinicians have completed the course, and no difficulties have been reported in obtaining third-party payer reimbursement.

#### Healthy Living after Cancer

Healthy Living after Cancer (HLaC) is a partnership project among 4 Australian state-based Cancer Councils, which are funded by the Australian National Health and Medical Research Council. It is evaluating the implementation of an evidence-based, 6-month, telephone-delivered lifestyle program, delivered by the Cancer Councils through their national cancer information and support service. HLaC is provided free of charge to cancer survivors with any type cancer after treatment with curative intent. It provides behaviorally based support to achieve internationally agreed recommendations for PA, healthy eating, and healthy weight. In this phase 4 dissemination study (single-group, pre-post design with assessments at baseline and 6 months), primary outcomes relate to program implementation: adoption (referral sources); reach (number of participants) and retention; fidelity of implementation; participant and staff satisfaction; and fixed and recurrent program costs. Secondary outcomes are patient-reported and validated measures of weight, PA, dietary intake/behavior, QOL, cancer-related side effects, and fear of recurrence. To date, 500 patients have enrolled (89% women; mean age [± standard deviation],  $55 \pm 11$  years; average BMI [ $\pm$  standard deviation],  $29 \pm 6 \text{ kg/m}^2$ ), with a wide range of cancers. The retention (program completion) rate is 57%. Among the first 200 program completers, significant (P < .05) and clinically meaningful improvements have been seen in all secondary, patient-reported outcomes. This collaborative undertaking provides an opportunity for national dissemination of an evidence-based intervention to support healthy living among cancer survivors. Rigorous evaluation of service-level and patient-reported outcomes will provide the practicebased evidence needed to achieve sustained support.

## Summary

MOVE!, Strength After Breast Cancer, LIVESTRONG at the Y, and HLaC all serve as model programs that effectively address the PA and weight management needs of cancer survivors. These programs, and many others like them, have established feasibility, safety, and efficacy. Although they are successful, common challenges, such as referral, training, triage, support, and reimbursement, remain as barriers.

# Insurance Coverage of Weight Management and PA in Cancer Care

Obesity presents unique challenges in patients with cancer; however, because of the high prevalence of obesity within the United States and its association with a constellation of chronic diseases,<sup>222</sup> most payers consider it a general concern across all of their health plan membership. Although evidence of what works in the treatment of obesity is growing,<sup>54,223,224</sup> the services that are covered and how the coverage is implemented in health benefit plans remain highly variable. Private and governmental payers of health insurance/benefit plans consider multiple factors as they decide what to cover and how to implement coverage. While many consumers and health care professionals assume that coverage decisions are based solely on cost, payers generally consider several factors in coverage decisions, including consumer/employer demand for a service, evidence for effectiveness and efficiency of the service, the ability to administer the benefit consistently and fairly, the presence of state/federal governmental mandates for a service, and how the benefit will affect the marketability/adverse selection of a health plan.

In its role as a fiduciary agent for its members, and for taxpayers in the case of the Centers for Medicare and Medicaid Services (CMS), insurers are faced with balancing their fiscal responsibilities with the mission and values of their organizations. An insurer first determines whether a service is effective, and then assesses the impact of that service on the overall cost of care for the members using that service. For example, coverage of the previously mentioned Y-USA DPP was based on relatively short-term data from a CMS Innovation Center demonstration project that indicated modest weight loss and significant savings in total costs of care. However, because the program was projected to reduce premature mortality, actuarial evaluation suggested that the program could ultimately lead to higher costs for Medicare patients who had care needs for a longer period of time. In response, CMS determined that longer life would not be considered as a cost (ie, care costs over a longer life span were zeroed-out). As a result, CMS announced that, in 2018, Medicare would begin reimbursing all DPP programs that meet CDC requirements.<sup>225</sup>

Interestingly, as value-based payment systems become more prevalent for both government and private payers, issues historically considered within the insurer's purview will shift to health care providers who share the financial risks (eg, next-generation ACOs).<sup>226</sup> These payment models provide payment based on a defined population of consumers and include incentives and potential penalties for both quality and cost metrics. Private payers, including multistate Anthem Blue Cross and Blue Shield and most national health plans, have developed similar value-based payment models. The expectation is that ACOs, or groups of physicians, hospitals, and other health professionals who come together voluntarily to provide coordinated highquality care, will provide services that improve the overall health and well being for a defined set of patients, including both clinical and community services for the prevention and treatment of obesity.

Although ACOs currently account for approximately 10% of the private health insurance market, insurers such as Anthem indicate that up to 60% of their fully insured membership may be covered under value-based payment models. The US Department of Health and Human Services has set a goal of tying 90% of all Medicare fee-for-service payments to quality or value metrics by 2018.<sup>227</sup>

Value-based metrics, however, are evolving and incomplete. The National Quality Forum has only endorsed 4 screening metrics related to obesity and has yet to reach consensus on any outcome measures,<sup>228</sup> but promising developments have transpired with Medicare coverage of the DPP, which includes an outcomes-based payment tied to both short-term and long-term patient outcomes.

The Bipartisan Policy Center has convened both commercial and government payers to discuss coverage for services related to obesity, including an effort to develop a shared-benefit design. Their discussions have focused on benchmarks of efficacy and cost effectiveness, in addition to issues related to member retention, return on investment, community partnerships, senior leadership support, and data tracking. Thus health professionals interested in working with payers to improve coverage for obesity-related services need to understand not only the needs of their patients but also the context in which payers make their coverage determinations.

## Advancing Progress in Tertiary Prevention: Stakeholder Insights and Recommendations

As cancer treatment advances and survivors live significantly longer, enhancing health and QOL for cancer survivors has become a major public health goal—one that also has widespread implications for the financial well being of survivors and families and of the health care system. Evidence continues to accumulate strongly suggesting that weight management and PA can improve the management of cancer, comorbid conditions, and QOL. However, there are 3 urgent challenges that must be overcome to connect cancer survivors with interventions that can ideally help them.

The first challenge is identifying the optimal type of intervention for a given survivor (eg, specific tumor type, cultural factors, comorbidities, functional status) and a specific goal (eg, fatigue management or decreasing risk of recurrence). This involves research to test varied types of interventions and to capture multiple types of patient data (tumor subtypes, clinical laboratory values, patient-reported symptom data) in multiple settings (identifying who needs medically supervised programs vs who benefits from community programs vs home-based interventions).

The second challenge is identifying how to deliver evidence-based interventions to support weight management and PA, not only in medical settings but also in the community or through the use of technology. Effective examples of model programs exist (eg, MOVE!, Strength After Breast Cancer, LIVESTRONG, and HLaC). Research is needed to continue to test clinic-based, community-based, and technology-delivered interventions and to identify how to facilitate referrals from oncologists and other health care professionals to these interventions. Educating health care providers about the importance of these interventions is a valuable but insufficient step. Future efforts need to address how to integrate weight management and PA interventions into standard cancer care.

The third challenge is accumulating the right data about weight management and PA programs to inform health care payer decisions to cover these interventions. Reimbursement by insurers will help make these interventions more affordable for survivors and drive widespread availability of these services. Part of this evidence will come from ongoing trials that will provide evidence regarding the benefits of postdiagnosis weight management and PA on recurrence and survival. However, even if these interventions do not specifically affect recurrence, coverage decisions may be made based on comorbidity management or effects on downstream health care utilization. Future research needs to test the cost effectiveness of these interventions on these important outcomes to inform health care coverage decisions.

The key to our success in implementing weight management and PA programs will be to bridge the silos of expertise, as represented in the NCPF workshop in 2017 in cancer biology, epidemiology, survivorship, nutrition, PA, weight management, and economics, as well as health care systems (eg, ACOs, payers, hospitals, oncology practices), care providers (eg, oncologists, primary care providers, allied health professionals, cancer rehabilitation, behavioral medicine) and the community (eg, advocacy organizations, YMCAs). The efforts to implement effective programs will need to address individual, provider/workforce, and systemic barriers, which include barriers specific to cancer survivors (symptoms and treatment side effects), as well as individual and cultural differences. The challenge is great. The opportunities and benefits of collaboration across disciplines and key stakeholders have significant potential to enhance outcomes for the growing number of cancer survivors in the United States and beyond.

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