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EDITORIAL COMMENT

The Continued Importance of Promoting Exercise as Part of Oncology Care for Breast Cancer Patients*

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he benefits of exercise during cancer treatment are multifaceted and welldocumented. Most studies demonstrate that exercise is not only safe for cancer patients undergoing treatment, but also highly beneficial for physical and mental health during and after cancer treatment.¹⁻³ Exercise prescriptions are beneficial in the management of several disease- and treatment-related side effects including anxiety, depression, physical function, and lymphedema,^{2,4} maintenance of cardiac health,⁵ as well as improved overall survival, cardiovascular disease survival, and cancer-specific survival.⁶ Despite the known benefits and call to incorporate exercise during oncology treatment and care, third-party payers generally provide coverage for exercise programs to cancer patients at higher risk for cardiovascular events.

The benefits of exercise during breast cancer treatment are among the most extensively studied of any cancer type. Studies in breast and other cancer patients have demonstrated the benefits of exercise in improving cardiorespiratory fitness (CRF) in cancer patients during treatment.^{7,8} CRF often declines during treatment due to the high cardiac toxicity of many chemotherapies, particularly those used in the treatment of HER2⁺ breast cancer,^{9,10} thus strategies to mitigate any cardiac effects in cancer patients would have tremendous JACC: CardioOncology, Peck et al¹² examine whether self-reported moderate-to-vigorous intensity physical activity (MVPA) is associated with better quality of life, cardiac function, and post-treatment CRF in 88 women with HER2⁺ breast cancer. MVPA and outcomes were measured at 6 time points over the course of approximately 1 year: near the time of diagnosis before treatment initiation; following anthracyclines, but before initiating trastuzumab; every 3 months during trastuzumab treatment; and at the completion of trastuzumab treatment. They found that higher MVPA was associated with better quality of life, measures of cardiac function, and post-treatment CRF. The findings around cardiac function are of particular importance, given the documented negative effects of anthracyclines and trastuzumab on cardiac function for women with HER2⁺ breast cancer.^{13,14} The combination of these drugs can lead to severe cardiac damage¹⁰; thus, these findings offer additional evidence of the benefits of exercise in mitigating these cardiac effects specifically for HER2⁺ breast cancer patients receiving sequential anthracyclines and trastuzumab. These findings are consistent with other prior

potential in improving longer-term outcomes

including mortality.¹¹ To that end, in this issue of

studies of breast cancer patients and support the continued need to promote exercise as part of oncology care, but there are some limitations of this study that should be acknowledged. Many of these limitations were acknowledged by the investigators including the small sample size and use of selfreported MVPA data. The investigators stated that pre-existing cardiovascular disease risk factors contribute to poor cardiac outcomes, but they had a limited measure of important potential confounders such as prediagnosis levels of alcohol, diet quality,

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and excess body fatness. The investigators were also unable to explore the role of prediagnosis MVPA, which may be a limitation given the documented association between MVPA before diagnosis and improved cardiovascular disease outcomes among breast cancer survivors.¹⁵ Finally, the lack of activity type, strength training, and sedentary time are additional limitations. While acknowledging these limitations, it is important to note the high-quality, robust CRF measures in the present study are a major strength.

Future studies addressing these limitations will be important and should, in particular, include: 1) measures of quality of life and CRF before treatment initiation to examine treatment-related changes; and 2) more detailed measures of physical activity measured before and during treatment. Although the Godin physical activity survey instrument used in the present study has shown moderate comparability with other self-reported measures of MVPA, the lack of detail on MVPA type, strength training, and sedentary time prevent analyses on of the roles of these distinct domains. As cancer survivor-specific recommendations for physical activity include aerobic exercise, resistance training, or a combination of both for expected patient benefits, future studies relying on self-report should consider newer, more

detailed, validated survey instruments that capture all of these activity domains.^{16,17}

This study reinforces the benefits of MVPA during treatment for breast cancer, here particularly for HER2⁺ breast cancer patients receiving anthracyclines followed by trastuzumab. Given the wellestablished benefits of exercise on physical and mental health, various patient-reported outcomes, improved survival, and other endpoints among cancer survivors, and the mounting evidence for improving CRF during treatment, guidelines are increasingly calling for the incorporation of exercise into oncology care.^{1,3,18} Most studies support that exercise is generally safe for individuals undergoing cancer treatment. Oncology care teams should encourage breast cancer patients to incorporate exercise as part of their treatment plan.

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REFERENCES

1. Campbell KL, Winters-Stone KM, Wiskemann J, et al. Exercise guidelines for cancer survivors: consensus statement from international multidisciplinary roundtable. *Med Sci Sports Exerc*. 2019;51(11):2375-2390. https://doi.org/10.1249/ MSS.00000000002116

2. Stout NL, Santa Mina D, Lyons KD, Robb K, Silver JK. A systematic review of rehabilitation and exercise recommendations in oncology guidelines. *CA Cancer J Clin.* 2021;71(2):149-175. https://doi. org/10.3322/caac.21639

3. Rock CL, Thomson CA, Sullivan KR, et al. American Cancer Society nutrition and physical activity guideline for cancer survivors. *CA Cancer J Clin.* 2022;72(3):230–262. https://doi.org/10. 3322/caac.21719

4. Rees-Punia E, Patel AV, Nocera JR, et al. Selfreported physical activity, sitting time, and mental and physical health among older cancer survivors compared with adults without a history of cancer. *Cancer.* 2020;127(1):115-123. https://doi.org/10. 1002/cncr.33257

5. Gilchrist SC, Barac A, Ades PA, et al. American Heart Association Exercise, Cardiac Rehabilitation, and Secondary Prevention Committee of the Council on Clinical Cardiology; Council on Cardiovascular and Stroke Nursing; and Council on Peripheral Vascular Disease. Cardio-oncology rehabilitation to manage cardiovascular outcomes in cancer patients and survivors: a scientific statement from the American Heart Association. *Circulation*. 2019;139(21):e997-e1012. https://doi. org/10.1161/CIR.000000000000679

6. Patel AV, Friedenreich CM, Moore SC, et al. American College of Sports Medicine Roundtable Report on physical activity, sedentary behavior, and cancer prevention and control. *Med Sci Sports Exerc.* 2019;51(11):2391-2402. https://doi.org/10. 1249/MSS.00000000002117

7. Naaktgeboren WR, Groen WG, Jacobse JN, et al. Physical activity and cardiac function in long-term breast cancer survivors: a crosssectional study. J Am Coll Cardiol CardioOnc. 2022;4(2):183-191. https://doi.org/10.1016/j.jaccao.2022.02.007

8. Fakhraei R, Peck BKin SS, Abdel-Qadir H, et al. Research quality and impact of cardiac rehabilitation in cancer survivors: a systematic review and meta-analysis. *J Am Coll Cardiol CardioOnc*. 2022;4(2):195-206. https://doi.org/10.1016/j.jaccao.2022.03.003

9. Foulkes SJ, Howden EJ, Bigaran A, et al. Persistent impairment in cardiopulmonary fitness after breast cancer chemotherapy. *Med Sci Sports* Exerc. 2019;51(8):1573-1581. https://doi.org/10. 1249/MSS.0000000000001970

10. Mohan N, Jiang J, Dokmanovic M, Wu WJ. Trastuzumab-mediated cardiotoxicity: current understanding, challenges, and frontiers. *Antib Ther.* 2018;1(1):13-17. https://doi.org/10.1093/abt/ tby003

11. Lavín-Pérez AM, Collado-Mateo D, Mayo X, et al. High-intensity exercise to improve cardiorespiratory fitness in cancer patients and survivors: a systematic review and meta-analysis. *Scand J Med Sci Sports*. 2021;31(2):265-294. https://doi. org/10.1111/sms.13861

12. Peck SS, Esmaeilzadeh M, Rankin K, et al. Selfreported physical activity, quality-of-life, cardiac function, and cardiorespiratory fitness in women with HER2+ breast cancer. *J Am Coll Cardiol CardioOnc.* 2022;4:387-400.

13. Chen JJ, Wu PT, Middlekauff HR, Nguyen KL. Aerobic exercise in anthracyclineinduced cardiotoxicity: a systematic review of current evidence and future directions. *Am J Physiol Heart Circ Physiol.* 2017;312(2):H213-H222. https://doi.org/10.1152/ajpheart.00646. 2016

14. Mclaughlin M, Florida-James G, Ross M. Breast cancer chemotherapy vascular toxicity: a review of

mediating mechanisms and exercise as a potential therapeutic. *Vasc Biol.* 2021;3(1):R106-R120. https://doi.org/10.1530/VB-21-0013

15. Okwuosa TM, Ray RM, Palomo A, et al. Prediagnosis exercise and cardiovascular events in primary breast cancer: Women's Health Initiative. *J Am Coll Cardiol CardioOnc*. 2019;1(1):41-50. https://doi.org/10.1016/j.jaccao.2019.08. 014 **16.** Rees-Punia E, et al. Reliability and validity of the cancer prevention study-3 physical activity survey items. *J Meas Phys Behav.* 2019;2(3):157-165.

17. Rees-Punia E, Matthews CE, Evans EM, et al. Demographic-specific validity of the Cancer Prevention Study-3 sedentary time survey. *Med Sci Sports Exerc.* 2019;51(1):41–48. https://doi.org/10. 1249/MSS.00000000001743 **18.** Hayes SC, Newton RU, Spence RR, Galvão DA. The Exercise and Sports Science Australia position statement: exercise medicine in cancer management. *J Sci Med Sport.* 2019;22(11):1175-1199. https://doi.org/10.1016/j. jsams.2019.05.003

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