

available at [www.sciencedirect.com](http://www.sciencedirect.com)journal homepage: [www.eu-openscience.europeanurology.com](http://www.eu-openscience.europeanurology.com)

European Association of Urology



## Brief Correspondence

# Can Exercise Adaptations Be Maintained in Men with Prostate Cancer Following Supervised Programmes? Implications to the COVID-19 Landscape of Urology and Clinical Exercise

Pedro Lopez<sup>a,\*</sup>, Dennis R. Taaffe<sup>a,b,c</sup>, Robert U. Newton<sup>a,b,c</sup>, Nigel Spry<sup>a,b,d</sup>, Tom Shannon<sup>e</sup>, Mark Frydenberg<sup>f</sup>, Fred Saad<sup>g</sup>, Daniel A. Galvão<sup>a,b</sup>

Exercise medicine clinics have had to change their services including those for men with prostate cancer (PCa) due to the recent coronavirus disease 2019 (COVID-19) where face-to-face supervised exercise programmes are required to cease. Unsupervised home-based (ie, off-site) interventions are an alternative method to improve physical activity behaviour in different clinical populations; however, despite being superior to usual care [1], benefits derived from these programmes are modest compared with face-to-face clinic-based programmes and not sustained for prolonged periods in men with PCa at different stages of disease [1,2]. Moreover, it is undetermined whether a change in programme delivery (ie, face-to-face clinic-based to unsupervised home-based programmes) would preserve clinical outcomes. These outcomes are relevant as the gains achieved with supervised periods of exercise could be significantly reduced, resulting in loss of exercise adaptations and decline in psychological and metabolic health.

In this brief correspondence, we analyse the effects of change from face-to-face supervised to unsupervised home-based exercise programmes on fatigue, quality of life (QoL), and body composition in men with PCa. Although none of the studies were performed during a viral pandemic, we highlight and contextualise our findings to their application for the COVID-19 pandemic, where face-to-face interventions have been restricted. This information will help provide a rationale for the delivery of exercise medicine to PCa patients in clinical practice in the current COVID-19 landscape.

A systematic review was undertaken in randomised controlled trials (RCTs) evaluating the effects of supervised resistance-based exercise programmes with subsequent change to a nonsupervised exercise intervention in PCa patients. Data were extracted from four manuscripts [3–6] describing three RCTs in men with PCa on androgen

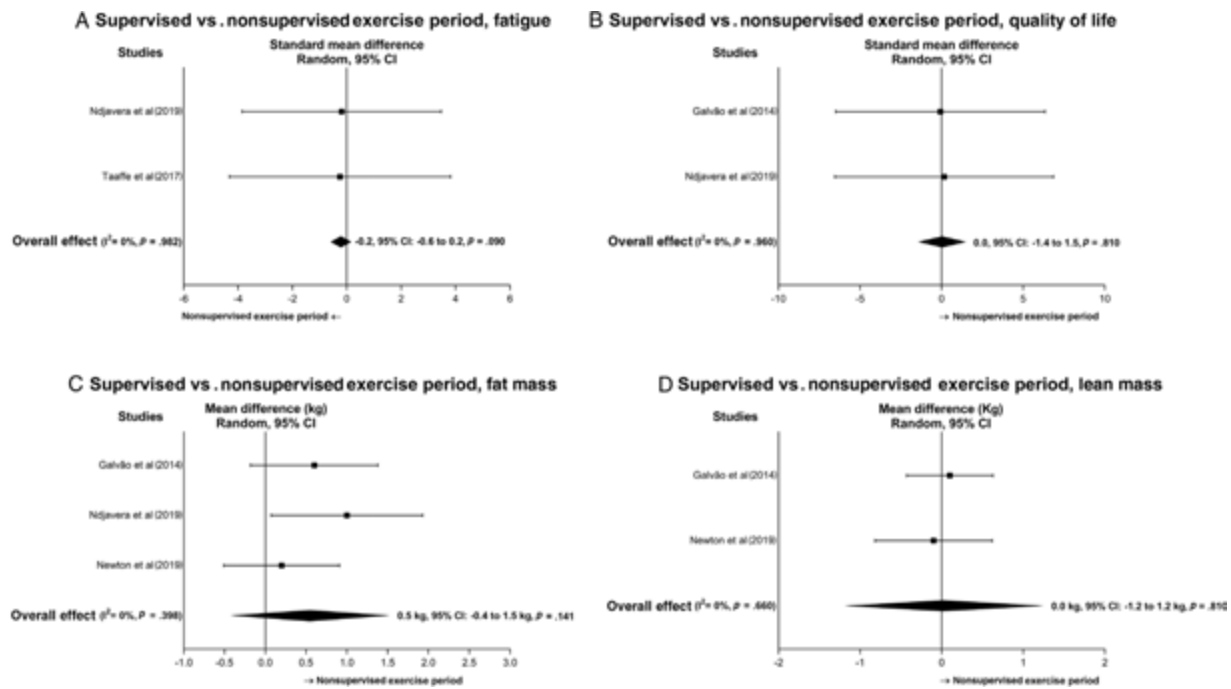
suppression therapy (AST) or previously treated with AST, which included fatigue, QoL, and body composition (ie, fat and lean mass) from the completion of face-to-face supervised and unsupervised home-based periods. The study selection procedure and results, and the main characteristics are described in the Supplementary material and provided in Table 1. We undertook a meta-analysis using a random-effect model and the Hartung-Knapp-Sidik-Jonkman method. Pooled-effect estimates were obtained from within-group values. Statistical heterogeneity was assessed using the Cochran Q test and expressed by  $I^2$ .

One-third of men undergoing AST experience fatigue due to associated depression, anxiety, pain, and insomnia, resulting in reduced QoL. Despite the maintenance of patient-reported outcomes during unsupervised home-based programmes (Fig. 1A and 1B), issues related to the participants' baseline levels and programme design may affect the interpretation of our findings. In the included studies, participants mostly presented with low baseline levels of fatigue and high levels of QoL, and this may have attenuated further change from the exercise programme during the unsupervised period. Thus, we may need to consider that fatigue and QoL are likely to be affected during the pandemic period in patients with higher fatigue levels and poor QoL. This concern is not only because of the distress associated with time on AST or fears related to tumour recurrence, but also because of distancing from family and friends, and fears associated with the COVID-19 pandemic. Moreover, most unsupervised home-based programmes require participants to have a self-controlled physical activity habit. Although this might work for those with regular exercise habits, it is unlikely to be generalised to all patients. Thus, current unsupervised models based on a usual weekly exercise volume (eg, 150 min/wk) or self-directed physical activity may not fit patient needs during

<http://dx.doi.org/10.1016/j.euros.2020.09.002>

2666-1683/© 2020 The Author(s). Published by Elsevier B.V. on behalf of European Association of Urology. This is an open access article under the CC BY-NC-ND license (<http://creativecommons.org/licenses/by-nc-nd/4.0/>).

Please cite this article in press as: Lopez P, et al. Can Exercise Adaptations Be Maintained in Men with Prostate Cancer Following Supervised Programmes? Implications to the COVID-19 Landscape of Urology and Clinical Exercise. Eur Urol Open Sci (2020), <http://dx.doi.org/10.1016/j.euros.2020.09.002>



**Q1** Fig. 1 – Mean difference and standard mean difference effects of face-to-face exercise programmes compared with unsupervised home-based exercise programmes on (A) fatigue, (B) quality of life, (C) fat mass, and (D) lean mass. Squares represent study-specific estimates and diamonds represent pooled estimates of random-effect meta-analysis. CI = confidence interval.

the COVID-19 pandemic. A redesigned home-based exercise programme leveraging the rapidly developing technology resources may help facilitate and assist participants in overcoming barriers related to exercise practice, such as lack of self-discipline, safety and monitoring, time, and treatment-related fatigue. Digital health facilitating technology might include the following:

- 1 Wearable biosensors (eg, heart rate, physical activity, steps, and distance)
- 2 Digital exercise prescription platforms providing the exercise programme and instructions, and recording exercises completed on the patient's smart device or computer
- 3 Video chat with qualified exercise professional to monitor and support the patient

During face-to-face supervised exercise, studies reported positive effects on body fat (range:  $-2.6$  to  $-0.6$  kg) and lean mass (range:  $0.1$ – $0.7$  kg) [3,5,6]. However, our analysis reveals that PCa patients were more likely to increase fat mass during follow-ups with unsupervised programmes, although lean mass was preserved (Fig. 1C and 1D). Considering the maintenance of physical activity levels and nutritional status reported in those studies [3,6], our findings suggest that body fat is likely to be increased during COVID-19 restriction, potentially adversely affecting metabolic health and disease prognosis [7]. Although there is evidence that patients on AST may be less likely to develop COVID-19 than PCa patients not receiving AST [8],

poor body composition and reduced physical activity levels can contribute to poorer disease prognosis, altering systemic and cellular factors and increasing the incidence of comorbidities such as diabetes or hypertension [9]. Thus, our results highlight the need for different exercise strategies such as an adequate increase of exercise stimulus (volume or intensity) to balance energy expenditure during physical activity restrictions or for facilitating intervention delivery as outlined above using different resources such as exercise smartphone Apps and online programmes (eg, telehealth). This may help provide instructions and feedback to help maintain patient motivation during the outbreak and avoid physical inactivity. Therefore, a higher exercise stimulus and increased contact with patients are likely to help maintain exercise adherence and counteract expected weight gain during self-quarantine [10].

In summary, despite the relatively small number of studies and patients in our meta-analysis, the direction of our results and the change of habits during the lockdown [10] are a concern when viewed in the context of the current worldwide situation. Changing from face-to-face to unsupervised self-directed home-based exercise programmes is unlikely to provide further benefits on fatigue, QoL, and body composition in patients with PCa during physical distancing restrictions, but may help with maintenance. Therefore, use of various technologies to keep patients motivated during self-quarantine and an increase in exercise stimulus to counteract physical distancing restrictions are some of our suggestions for the current COVID-19 landscape to avoid physical inactivity. These measures are

**Table 1 – Study characteristics: cancer therapy duration, demographic and clinical characteristics, sample size, supervised and unsupervised exercise prescription, and outcomes assessed.**

Author (year)	Cancer therapy duration	Demographical and clinical characteristics	Face-to-face supervised exercise period	Unsupervised home-based exercise period	Outcomes
Galvão et al (2014) [3]	Previous AST duration of ~12 mo with time since its cessation of 38 mo	Age: 71.4 yr	Combined resistance and aerobic training:	24 wk; booklet with detailed information about a home exercise prescription including resistance, aerobic, and flexibility exercises	Fat mass, lean mass, SF-36 <sup>a</sup>
		II–IV	n = 50, 2 sessions per week for 24 wk; RT: 2–4 sets of 6–12 RM; AT: 20–30 min at 70–85% HR		
		Previous AST and radiotherapy			
Taaffe et al (2017) [4] <sup>b</sup>	Minimum exposure to AST of 2 mo and anticipated to receive AST for the subsequent 12 mo	Age: 68.8 yr	Combined resistance and aerobic training:	24 wk; home-based programme that recommended 150 min of aerobic exercise per week and resistance exercise using body weight and resistance bands	EORTC QLQ-C30 <sup>Fatigue</sup> <sup>c</sup>
		Localised and nodal metastases	n = 54, 2 sessions per week for 24 wk; AT: 20–30 min at 60–85% HR; RT: 2–4 sets of 6–12 RM		
		Gleason score: 7.8			
		AST			
		AST plus radiotherapy			
		AST plus antiandrogen			
		AST plus surgery			
Ndjavera et al (2020) [5]	Patients with newly diagnosed prostate cancer and beginning AST treatment	Age: 72.0 yr	Combined resistance and aerobic training:	12 wk; patients were instructed to continue exercising and maintain self-directed levels of physical activity	Fat mass, FACT-P <sup>a</sup> , FACIT-Fatigue <sup>c</sup>
		Locally advanced and metastatic patients	n = 24, 2 sessions per week for 12 wk; AT: 6 bouts of 5 min at 55–85% HR; RT: 2–4 sets of 10 reps at 11–15 RPE		
		Gleason score range from 6 to 10			
		AST			
		AST plus radiotherapy			
Newton et al (2019) [6] <sup>b</sup>	Minimum exposure to AST of 2 mo and anticipated to receive AST for the subsequent 12 mo	Age: 69.0 yr	Combined resistance and aerobic training:	24 wk; home-based programme that recommended 150 min of aerobic exercise per week and resistance exercise using body weight and resistance bands	Fat mass, lean mass
		Localised and nodal metastases	n = 50, 2 sessions per week for 24 wk; AT: 20–30 min at 60–85% HR; RT: 2–4 sets of 6–12 RM		
		Gleason score: 7.8			
		AST			
		AST plus radiotherapy			
		AST plus antiandrogen			

AST = androgen suppression therapy; AT = aerobic training; EORTC QLQ-C30 = European Organisation for Research and Treatment of Cancer Quality of Life Questionnaire C30; FACIT = Functional Assessment of Chronic Illness Therapy; FACT-P = Functional Assessment of Cancer Therapy–Prostate; HR = heart rate; RM = repetitions maximum; RPE = rate of perceived exertion; RT = resistance training; SF-36 = 36-Item Short Form Survey.

<sup>a</sup> Included in quality-of-life meta-analysis.

<sup>b</sup> Papers derived from the same trial.

<sup>c</sup> Included in fatigue meta-analysis.

139 necessary to guarantee the continuation of appropriate and  
140 targeted exercise medicine delivery to PCa patients.

141 **Author contributions:** Pedro Lopez had full access to all the data in the  
142 study and takes responsibility for the integrity of the data and the  
143 accuracy of the data analysis.

144 **Study concept and design:** Lopez, Taaffe, Newton, Galvão.

145 **Acquisition of data:** Lopez.

146 **Analysis and interpretation of data:** Lopez, Taaffe, Newton, Spry, Shannon,  
147 Frydenberg, Saad, Galvão.

148 **Drafting of the manuscript:** Lopez, Taaffe, Newton, Spry, Shannon,  
149 Frydenberg, Saad, Galvão.

150 **Critical revision of the manuscript for important intellectual content:**  
151 Lopez, Taaffe, Newton, Spry, Shannon, Frydenberg, Saad, Galvão.

152 **Statistical analysis:** Lopez.

153 **Obtaining funding:** None.

154 **Administrative, technical, or material support:** None.

155 **Supervision:** None.

156 **Other:** None.

157 **Financial disclosures:** Pedro Lopez certifies that all conflicts of interest,  
158 including specific financial interests and relationships and affiliations  
159 relevant to the subject matter or materials discussed in the manuscript  
160 (eg, employment/affiliation, grants or funding, consultancies, honoraria,  
161 stock ownership or options, expert testimony, royalties, or patents filed,  
162 received, or pending), are the following: None.

163 **Funding/Support and role of the sponsor:** Pedro Lopez is supported by  
164 the National Health and Medical Research Council (NHMRC) Centre of  
165 Research Excellence (CRE) in Prostate Cancer Survivorship Scholarship.  
166 Daniel A. Galvão and Robert U. Newton are funded by an NHMRC CRE in  
167 Prostate Cancer Survivorship. The results of the study are presented  
168 clearly, honestly, without fabrication, falsification, or inappropriate data  
169 manipulation. Sponsors were not involved in the study design, analysis  
170 or interpretation of data, manuscript writing, and decision to submit the  
171 Q4 manuscript for publication.

## 172 Appendix A. Supplementary data

173 Supplementary material related to this article can be  
174 found, in the online version, at doi:[https://doi.org/10.1016/j.  
175 euros.2020.09.002](https://doi.org/10.1016/j.euros.2020.09.002).

## 176 References

177 [1] Buffart Lm, Kalter J, Sweegers Mg, et al. Effects and moderators of  
178 exercise on quality of life and physical function in patients with  
cancer: an individual patient data meta-analysis of 34 RCTs. *Cancer  
Treat Rev* 2017;52:91–104.

- [2] Galvão DA, Newton RU, Girgis A, et al. Randomized controlled trial  
of a peer led multimodal intervention for men with prostate cancer  
to increase exercise participation. *Psychooncology* 2018;27:199–  
207. 179 180
- [3] Galvão DA, Spry N, Denham J, et al. A multicentre year-long ran-  
domised controlled trial of exercise training targeting physical  
functioning in men with prostate cancer previously treated with  
androgen suppression and radiation from TROG 03.04 RADAR. *Eur  
Urol* 2014;65:856–64. 181 182 183
- [4] Taaffe DR, Newton RU, Spry N, et al. Effects of different exercise  
modalities on fatigue in prostate cancer patients undergoing an-  
drogen deprivation therapy: a year-long randomised controlled  
trial. *Eur Urol* 2017;72:293–9. 184 185 186
- [5] Ndjaverwa W, Orange ST, O'Doherty AF, et al. Exercise-induced at-  
tenuation of treatment side-effects in patients with newly diag-  
nosed prostate cancer beginning androgen-deprivation therapy: a  
randomised controlled trial. *BJU Int* 2020;125:28–37. 187 188 189
- [6] Newton RU, Galvão DA, Spry N, et al. Exercise mode specificity for  
preserving spine and hip bone mineral density in prostate cancer  
patients. *Med Sci Sports Exerc* 2019;51:607–14. 190 191
- [7] King AJ, Burke LM, Halson SL, Hawley JA. The challenge of main-  
taining metabolic health during a global pandemic. *Sports Med*  
2020;50:1233–41. 192 193
- [8] Montopoli M, Zumerle S, Vettor R, et al. Androgen-deprivation  
therapies for prostate cancer and risk of infection by SARS-CoV-  
2: a population-based study (N = 4532). *Ann Oncol* 2020;31:1040–5. 194 195
- [9] Smith MR, Saad F, Egerdie B, et al. Sarcopenia during androgen-  
deprivation therapy for prostate cancer. *J Clin Oncol* 2012;30:3271–  
6. 196
- [10] Zachary Z, Brianna F, Brianna L, et al. Self-quarantine and weight  
gain related risk factors during the COVID-19 pandemic. *Obes Res  
Clin Pract* 2020;14:210–6. 197 198

<sup>a</sup>Exercise Medicine Research Institute, Edith Cowan University, Perth,  
Western Australia, Australia

<sup>b</sup>School of Medical and Health Sciences, Edith Cowan University, Perth,  
Western Australia, Australia

<sup>c</sup>School of Human Movement and Nutrition Sciences, University of  
Queensland, St. Lucia, Queensland, Australia

<sup>d</sup>Faculty of Medicine, University of Western Australia, Nedlands, Western  
Australia, Australia

<sup>e</sup>The Prostate Clinic, Hollywood Private Hospital, Nedlands, Western  
Australia, Australia

<sup>f</sup>Department of Surgery, Faculty of Medicine, Nursing and Health Sciences,  
Monash University, Clayton, Victoria, Australia

<sup>g</sup>Division of Urology and Urologic Oncology, Centre Hospitalier de  
l'Université de Montréal, Montreal, Quebec, Canada

\*Corresponding author. Exercise Medicine Research Institute, Edith  
Cowan University, 270 Joondalup Drive, Joondalup WA 6027, Australia.  
Tel. +61 416463228.

E-mail address: [plopezda@our.ecu.edu.au](mailto:plopezda@our.ecu.edu.au) (P. Lopez).