) original contributions

Exercise Recommendation for People With Bone Metastases: Expert Consensus for Health Care Providers and Exercise Professionals

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QUESTION ASKED: What are the best practice recommendations for exercise programming for people with bone metastases?

SUMMARY ANSWER: People with bone metastases should be supported and encouraged to engage in regular physical activity, including structured exercise, to obtain the well-established general health benefits and as a strategy to manage side effects related to cancer and treatments. Perceived risks of skeletal complications should be weighed against the potential health benefits in consultation between the person with bone metatases, health care team, and exercise professional.

WHAT WE DID: The International Bone Metastases Exercise Working Group (IBMEWG) developed best practice recommendations on the basis of published research, clinical experience, and expert opinion using (1) *modified Delphi survey*, (2) *systematic review*, (3) *cross-sectional survey* to physicians and nurse practitioners, (4) *inperson meeting*, and (5) *stakeholder engagement*.

WHAT WE FOUND: On the basis of the review of available evidence and expert consensus, the IBMEWG developed five key recommendations: (1) Before exercise testing or training, perform a risk assessment to inform the likelihood of a skeletal complication from exercise. (2) Consultation with the medical team is strongly encouraged to obtain key medical information and establish bidirectional communication. (3) Exercise professionals best suited for this population are physical therapists and clinical exercise physiologists (or equivalent) who have additional cancer exercise training. (4) Professional judgement should be used to consider if exercise testing is necessary. (5) Exercise prescription

should follow the standard exercise recommendations as outlined by the International Exercise Guidelines for Cancer Survivors, with greater emphasis on postural alignment, controlled movement, proper technique, and consideration of the bone lesion location and presentation.

BIAS, CONFOUNDING FACTORS: Some authors of this paper are employed by entities that provide physical activity programming for people with cancer, are involved in professional organizations that promote the role of exercise professionals within health care settings, and/or were authors for the International Exercise Guidelines for Cancer Survivors. Although these are strengths in terms of expertise, they may also lead to bias.

REAL-LIFE IMPLICATIONS: Exercise has been underutilized in people with advanced or incurable cancer despite the potential to improve physical function and reduce psychosocial morbidity, especially for people with bone metastases because of concerns over skeletal complications. These recommendations provide a framework and starting point for members of the medical team and exercise professionals to improve the integration of physical activity into the care of people with bone metastases. It is anticipated that the recommendations provided here will continue to evolve as more literature is available. Fundamental limitations remain around a lack of definitive literature on standardized approaches to predict the risk of skeletal complications and a paucity of data on the safety and efficacy of exercise for specific people that may be at increased risk of skeletal complications with exercise (ie, elderly individuals with multiple myeloma).

ASSOCIATED CONTENT

Appendix

Author affiliations and disclosures are available with the complete article at ascopubs.org/ iournal(cz

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PURPOSE Exercise has been underutilized in people with advanced or incurable cancer despite the potential to improve physical function and reduce psychosocial morbidity, especially for people with bone metastases because of concerns over skeletal complications. The International Bone Metastases Exercise Working Group (IBMEWG) was formed to develop best practice recommendations for exercise programming for people with bone metastases on the basis of published research, clinical experience, and expert opinion.

METHODS The IBMEWG undertook sequential steps to inform the recommendations: (1) *modified Delphi survey*, (2) *systematic review*, (3) *cross-sectional survey* to physicians and nurse practitioners, (4) *in-person meeting* of IBMEWG to review evidence from steps 1-3 to develop draft recommendations, and (5) *stakeholder engagement*.

RESULTS Recommendations emerged from the contributing evidence and IBMEWG discussion for pre-exercise screening, exercise testing, exercise prescription, and monitoring of exercise response. Identification of individuals who are potentially at higher risk of exercise-related skeletal complication is a complex interplay of these factors: (1) lesion-related, (2) cancer and cancer treatment-related, and (3) the person-related. Exercise assessment and prescription requires consideration of the location and presentation of bone lesion(s) and should be delivered by qualified exercise professionals with oncology education and exercise prescription experience. Emphasis on postural alignment, controlled movement, and proper technique is essential.

CONCLUSION Ultimately, the perceived risk of skeletal complications should be weighed against potential health benefits on the basis of consultation between the person, health care team, and exercise professionals. These recommendations provide an initial framework to improve the integration of exercise programming into clinical care for people with bone metastases.

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INTRODUCTION

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Accepted on November 18, 2021 and published at ascopubs.org/journal/ op on January 6, 2022: D0I https://doi. org/10.1200/0P.21. 00454 People living with advanced or incurable cancers are treated with sequential cancer therapies to improve survival and quality of life, but treatment burden can be high, including fatigue, reduced physical function, and psychosocial morbidity.¹ In this context, people with advanced or incurable cancer highlight that maintaining functional independence and managing symptom burden are top priorities.^{2,3}

Exercise (ie, planned and structured physical activity aimed to improve health) has been shown to improve fatigue, physical function, and psychosocial morbidity

in people with early-stage cancers.⁴ In the advanced or incurable cancer setting, the presence of bone metastases complicates prescribing exercise and promoting physical activity because of concerns about potential skeletal complications such as pathologic fracture, hypercalcemia, or spinal cord compression.⁵ Although people with bone metastases regularly express an interest in information on exercise, ⁶⁻⁸ health care professionals report uncertainties regarding risk management.⁹⁻¹¹ Recent systematic reviews in advanced or incurable cancer generally show exercise to be safe, feasible, and beneficial for quality of life, physical function, and fatigue.¹²⁻¹⁸ However, these





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systematic reviews have not focused specifically on the safety or efficacy of exercise for people with bone metastases.

The International Bone Metastases Exercise Working Group (IBMEWG) was formed on the basis of the recognition that people with bone metastases could potentially benefit from exercise and are not receiving direction from the health care team, as they may be reticent because of lack of guidance in the literature or clinical practice. This group aimed to develop best practice recommendations on the basis of existing research evidence, clinical experience, and expert opinion.

METHODS

The IBMEWG is a multidisciplinary, international panel of physicians, physical therapists, clinical exercise physiologists, and researchers. Sequential steps (Fig 1) were taken to inform the recommendations: (1) modified Delphi survey to ascertain the practical considerations of implementing exercise for people with bone metastases in the outpatient oncology setting¹⁹ (all participants provided informed consent, Research Ethics Board approval, University of British Columbia; H19-00379); (2) systematic review to assess the safety, feasibility, and efficacy of exercise in controlled trials that included people with bone metastases (International Prospective Register of Systematic Reviews: CRD42019121958)²⁰; (3) cross-sectional survey to physicians and nurse practitioners to understand the attitudes toward exercise for people with bone metastases and components required to promote exercise referral²¹ (all participants provided informed consent, Research Ethics Board approval, University of British Columbia; H19-00379); (4) in-person meeting of IBMEWG to review evidence from steps 1-3 to develop recommendations; and (5) stakeholder engagement, first with people with bone metastases (draft recommendations) and second with clinician peers (penultimate recommendations).

RESULTS

Recommendation 1

Before exercise testing or training, perform a risk assessment to inform the likelihood of a skeletal complication from exercise.

The Delphi identified consensus on information that respondents deemed key to determine exercise suitability (Table 1). However, reliance on an individual's *selfreported* bone lesion details did not reach consensus. The systematic review showed that randomized controlled trials of exercise involving people with bone metastases commonly exclude people deemed to have structurally unstable bone lesion(s) or pain associated with bone lesion(s).²⁰ Determination of lesion stability was either undefined^{22,23} or two randomized controlled trials used Taneichi's subtype A-C scoring for osteolytic thoracic and lumbar spine lesions, with computed tomography or magnetic resonance imaging assessed independently by a radiologist and orthopedic surgeon.^{24,25} In the survey, most respondents agreed that exercise for people with bone metastases was safe (82%) and these individuals look to their health care providers to provide exercise referral and recommendations (74%). To facilitate respondents providing an exercise recommendation to individuals, the most frequently cited resource identified was a set of clinical guidelines for exercise in this population (25%), an easy-to-use screening scale (20%), and a consult with a qualified exercise professional (13%).²¹

The IBMEWG acknowledged that in the absence of literature, all people with bone metastases can be at risk of an exercise-related skeletal complication, but this should not preclude prescribing exercise. Identification of individuals who are potentially at higher risk of exercise-related skeletal complication is a complex interplay of these factors: (1) lesion-related, (2) cancer and cancer treatment-related, and (3) person-related (Table 2). Furthermore, it was acknowledged that bone scan reports may not consistently provide information on lesion volume and that a computed tomography and magnetic resonance imaging may be more reliable for structural qualities of the bone and preferable for people deemed higher risk. Finally, more research is required to confirm the utility of pre-exercise skeletal assessment scales to guide risk assessment. Stakeholders with bone metastases reported that they were keen to exercise but fearful, lacking in confidence for self-guidance, and uncertain where to look for exercise advice. They confirmed that discussion with the health care team on how to manage risk was of key interest to people with bone metastases.

Recommendation 2

Consultation with the medical team is strongly encouraged before an exercise professional provides structured exercise for a person with bone metastases, to obtain key medical information and establish bidirectional communication for initial assessment and exercise training throughout care.

The primary reason reported by exercise professionals to seek medical clearance or guidance was to obtain key medical information, such as results of bone scan reports to guide exercise prescription and to establish two-way communication (ie, to allow for information sharing with the goal of better safety and overall care). When asked if a primary reason for seeking medical clearance or guidance was a requirement for professional liability (ie, signed approval by a physician), only 39% of exercise professionals strongly agreed or agreed (Table 1). In the systematic review, 24% of trials required medical clearance by a physician. For the remainder, exclusion criteria specific to lesion characteristics of bone metastases were used, such as excluding people with unstable bone metastases (24%) or pain associated with the bone lesion (41%).²⁰ Most

(held as informal side meeting of attende	Prelimina ees at the an	r y planning meeting nual meeting 2018 American College of Sports Medicine; n = 16)		
Modified Delphi consensus process (participants, n = 73 [R1], 59 [R2], 57 [R3])	>	Inclusion criteria Involved with research that used a mode of exercise and involved participants with bone metastases; OR Exercise or health care professional who has worked with a minimum of 10 people with bone metastases per year; OR Considered expert in exercise and bone metastases through clinical or research experience		
Systematic review (No. of trials included, n = 17)	>	Inclusion criteria Randomized controlled or controlled clinical trials Adults (≥ 18 years) with cancer diagnosis and included people with bone metastases Exercise intervention (> 1 session structured exercise) Outcome: physical function, functional capacity, muscular strength, or treatment side effect		
Physician and nurse practitioner survey (participants and physicians, n =109; nurse practitioners, n = 15)	>	Inclusion criteria Providing care for people with metastatic cancer in publicly funded provincial cancer service in British Columbia, Canada AND Oncologists (medical, radiation, and surgeons) OR General practitioners in oncology OR Nurse practitioner		
In-person meeting of the International Bone Metastases Exercise Working Group and recommendation development (participants = 21)	>	Inclusion criteria Previous research expertise in exercise oncology, particularly in advanced cancer or bone metastases; OR Clinicians engaged in clinical oncology care; AND Willingness to attend 2-day meeting in-person or virtually.		
Draft recommendation statements by International Bone Metastases Exercise Working Group				
Stakeholder engagement: People with bone metastases (participants = 3)	>	Inclusion criteria Person living with bone metastases > 1 year Experience in engaging in exercise Able to attend in-person focus group		
Update of recommendations statements and draft of rationale statements by smaller working group (D.S.M., J.G., K.A.B., K.L.C., K.M.W.S., M.L.M., N.H.H., P.C., S.W.)				
Stakeholder engagement: clinician peers (participants = 5)	>	Inclusion criteria Exercise or health care professional who has experience in working with people with bone metastases		
Finalization of recommendations by Interna	tional Bone	Metastases Exercise Working Group		

FIG 1. Information gathering and recommendation development process.

survey respondents agreed that people with bone metastases expected their physician to provide physical activity recommendations (74%), and these recommendations would be well received (66%) and followed (58%). However, less than half of the physicians and nurse practitioners (43%) felt confident to recommend exercise to people with bone metastases.²¹

The IBMEWG agreed that the term medical consultation versus clearance was the most appropriate term for communication between the exercise professional and health care provider. The goal of such medical consultation is to establish two-way communication to assess the cost to benefit-risk of an exercise prescription, as it can be difficult for health care providers to feel that they have enough understanding of exercise to provide medical clearance and for exercise professionals to obtain or interpret all the desired clinical information to assess risk, especially without access to the medical chart (Appendix Table A1, online only).

Recommendation 3

Exercise professionals best suited to prescribe exercise to people with bone metastases are physical therapists and clinical exercise physiologists (or equivalent), who have additional cancer exercise training and appropriate experience in working with people with a cancer diagnosis.

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DR	Strongly Agree	Agree	Neutral	Disagree	Strongly Disagree	Consensus (%)
		A. Pre-	exercise screening	g questions (Recom	mendation 1)	
DR2	When performing a uses the following lesion location, typ bone pain within	pre-exercise scre g information to d pe, progression, a the past 3 month	ening on an indivi etermine exercise nd history; (2) bone is; (4) symptoms a	dual with MBD, it is suitability: (1) bone pain details (during associated with the l	STRONGLY RECOMMENDE lesion details, ideally from a rest, ADLs, or physical activity bone lesions	D that an exercise professional diagnostic report that includes y); (3) any medical treatment for
	91%	9%	0%	0%	0%	100% agree
DR2	In addition to the above items, when performing a pre-exercise screening on an individual with MBD, it is STRONGLY RECOMMENDE that an exercise professional also uses the following information to determine exercise suitability: (1) detailed medical history, (2) currer medications or treatments specific to MBD or pain, (3) history of fractures and falls, (4) bone mineral density report, (5) the individual physical activity goals, and (6) current physical activity levels					
	86%	12%	2%	0%	0%	98% agree
DR2	In the absence of a b bone lesion(s) AF	oone scan report (RE SUFFICIENT	or a similar diagnos	stic report or letter fro	om medical doctor), an indivic	lual's self-reported details of the
	2%	14%	23%	35%	26%	No consensus reached
		B. Medical	clearance and Me	edical guidance (Re	commendation 2)	
DR3	As an exercise professional, the primary objective when seeking a medical clearance or medical guidance from a physician, in relation to an individual with MBD is					
	a. Medical inform	nation (eg, reports	from recent scan	s or tests and any ι	pcoming treatments)	
	65%	28%	5%	0%	2%	93% agree
	b. Establish two-v	vay communicati	on			
	53%	23%	7%	16%	2%	76% agree
	c. Recommendat	ions for exercise	prescription			
	10%	23%	23%	33%	11%	No consensus reached
	d. Professional lia	ability				
	12%	27%	12%	33%	16%	No consensus reached
DR3	Individuals with MBD who meet any of the following conditions REQUIRE medical guidance from the individual's medical professional pr to commencing a structured exercise program with a clinical exercise professional: (1) bone lesions that are unstable or of unknow stability, (2) bone pain or medical treatment of bone pain in the past 3 months, and (3) history of disease-related fractures within the pa 12 months					
	63%	26%	2%	5%	4%	89% agree
		C. Su	itable exercise pro	fessionals (Recomm	nendation 3)	
DR2	Please select all the disease who is de	e exercise profess eemed HIGHER F	ionals that you fee RISK OF FRACTUF	el are suitable to saf RE (eg, multiple bor	ely prescribe exercise to an i ne lesion locations, unknown	individual with metastatic bone stability, and symptomatic)
	a. Physical therap	oist				39
	b. Physical therap	pist with cancer e	exercise training			95
	c. Exercise physic	ologist				20
	d. Exercise physic	plogist with cance	er exercise training	5		85
	e. Kinesiologist or	r exercise or spor	ts scientist or exer	cise therapist		13
	f. Kinesiologist or	exercise sports s	cientist or exercise	e therapist with can	cer exercise training	52
	g. Personal traine	er				0
	h. Personal traine	er with cancer exe	ercise training			13
	i. Fitness instruct	or				0
	j. Fitness instruct	or with cancer ex	ercise training			7
			(continued	on following page)		

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DR	Strongly Agree	Agree	Neutral	Disagree	Strongly Disagree	Consensus (%)
			D. Exercise testir	ng (Recommendatio	on 4)	
DR2	In an individual with but there are son	n MBD who is at ne tests that are o	a LOWER risk of fi contraindicated	racture, I recommer	nd exercise testing,	
	70%				30%	70% agree
	In an individual with contraindicated	n MBD who is at	a HIGHER risk of	fracture, I recomme	end exercise testing, but there	are some tests that are
	88%				12%	88% agree
DR3	When considering s	trength testing fo	r a person with bo	one metastases, I wo	ould	
	a. Avoid any strer	ngth testing (7%)				
	b. Avoid any strei	ngth testing that	places stress on th	ne lesion site (44%)		
	c. Use caution wi	ith any testing tha	at places stress on	the lesion site (37%	%)	
	d. Not make any	considerations (7	7%)			
	e. I am not an ex	ercise profession	al and do not feel	I can answer this q	uestion (2%)	
		E	E. Exercise prescrip	ption (Recommenda	ation 5)	
DR2	If an individual is dee education or mov individuals' medio	emed UNSAFE TO vement prescriptio cal team	DEXERCISE (aka "Hon that is modified	high" risk), it is appro to specifically suit t	opriate for an exercise profession the individual and is provided	onal to provide physical activity in collaboration with the
	65%	26%	2%	4%	4%	91% agree
DR3	Once an individual v Exercise Physiolo Instructor) may b established by th	with MBD has rec gist with cancer e e appropriate. Th e Physiotherapist	ceived an exercise exercise training), r e less qualified exe or Exercise Physic	prescription from a referral to a less qua ercise professional v plogist	highly qualified exercise profe alified exercise professional (e vould administer and monitor	ssional (eg, Physiotherapist or g, Personal Trainer or Fitness the specific exercise program
	16%	49%	12%	16%	7%	No consensus reached
DR3	For individuals with I that place stress	ower risk MBD (eg on the bone lesic	g, stable and asymp on site	otomatic), exercise p	rescription is normally safe if it	USES CAUTION with exercises
	54%	39%	5%	0%	2%	93% agree
DR3	What does the term	"use caution" m	iean to you, in the	context of the ques	stion above?	
	a. Start by avoiding	ng any exercises	that place stress o	n the lesion site (14	1%)	
	b. For exercises the exacerbation of	hat target the lesion f pain or adverse	on site, start with a symptoms (47%)	ctive movement and	I no load or weight and progre	ss slowly, provided there is no
	c. For exercises th exacerbation of	at target the lesior f pain or symptor	n site, start with low ns (28%)	loads (eg, theraband	d or light dumbbells) and progr	ess slowly, provided there is no
	d. Other (10%)					
		F. Direction for	r future exercise re	esearch for people v	vith bone metastases	
	Clinical prediction n	nodels for adverse	e event risk during	g physical activity		
	In exercise studies	of people with ad	vanced cancer, re	port in detail for pe	ople with bone metastases	
	a. Numbers of pa	articipants				
	b. Lesion(s) detai	ls: type, location,	size, and treatment	nts		
	c. Screening proc	cedures, tools, or	decision-making p	process		
	d. Adaptations to	testing and exerc	cise protocols			
	e. Adverse events	s related and unre	elated to exercise			
	Conduct studies of	safety and efficad	cy that test			
	a. Different exerc	ise prescription a	pproaches			
	b. Use of establis	hed scales or too	ols to screen or info	orm exercise prescr	iption	
	c. Exercise dose	that optimizes ma	aintenance or impr	rovement on target o	outcome	
	d. Exercise for inc	dividuals with uns	stable or painful bo	one lesion(s)		
	Qualitative analysis	of neonle with bo	ne metastases nei	rcentions of exercise	e particularly safety benefit	and worny

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Abbreviations: ADL, activity of daily living; DR, Delphi Round; MBD, metastatic bone disease.

TABLE 2. Recommendation for Team-Based Approach to Provision of Exercise Programming to People Living With Bone Metastases

Medical communication					
Communication with current medical team (eg, Oncologist) is recommended to	Obtain any necessary medical guidance or additional information (eg, specific bone lesion details, previous and current cancer therapies, etc)				
	Establish and maintain a two-way communication pathway to share information on				
	New presentation of pain				
	Other symptoms				
	Functional concerns				
Access to information on lesion-specific factors	Most recent bone scan report or other diagnostic report that describes the following:				
	Type of lesion (osteolytic, sclerotic, or mixed)				
	Number of lesion (s)				
	Location of lesion(s) (ie, load-bearing areas such as femur, spine, pelvic ring, acetabulum, and tibial plateau)				
	Size of lesion(s) (ie, small v large)				
	Series of reports to determine status of the lesion(s), rate of progression, and structural stability of bone (ie, any evidence of vertebral compression)				
	Pre-exercise history				
Cancer and cancer treatment-related factors to consider					
Cancer diagnosis	Prognosis or progression of disease				
	Metastases to other organ systems (ie, brain and lung)				
	History of fractures since cancer diagnosis				
Management of bone metastases	Planned or prior treatment approach to management of bone metastases (ie, surgical, radiation, and medical)				
	Planned or prior treatment approach to management or for SSE (ie, surgical, radiation, and medical)				
	Current use of bone agents (eg, denosumab and bisphosphonates > 6 months)				
	Current use of agents that may increase risk of SSE (ie, corticosteroids, hormonal therapy, and anticoagulants)				
General bone health	Bone health risk factors (ie, smoking and family history of osteoporosis)				
Diagnosis and severity of osteoporosis					
	Pharmaceutical management of osteoporosis (eg, bisphosphonates)				
Pain	Severity				
	Location				
	Use of pain medication related to site of lesion(s)				
	Triggers (eg, functional pain, during ADLs, loading, at rest, and at night)				
	Change in quality or location of pain (ie, new pain with weight bearing, transitional pain when performing ADLs, and pain worsening at night)				
Neurologic	Sudden or recent muscle weakness in any region				
	Sudden, new or recent change in bowel and/or bladder control (ie, progressive urinary retention, bladder or bowel incontinence)				
	New or progressive gait or balance impairment				
	Loss of sensation or reflexes in any region				
Falls	Occurrence within the past 12 months				
	If positive, number of falls				
	If positive, injuries consequent to fall				
Person-related factors to consider					
Overall medical and symptom profile (eg, comorbid con	ndition, fatigue, and cachexia)				
Worse ECOG PS					
Limitations in ADLs or instrumental ADLs					
Cognitive impairment (eg, making adherence to precau	itions unreliable)				

Exercise history

Abbreviations: ADL, activity of daily living; ECOG PS, Eastern Cooperative Oncology Group Performance Status; SSE, symptomatic skeletal event.

The Delphi showed consensus that the exercise professionals best suited to prescribe exercise to people with bone metastases were university qualified with clinical expertise, namely, physical therapists (95% agree) and clinical exercise physiologists (85% agree), provided that each has additional cancer exercise education and training (Table 1). In the systematic review, exercise supervision was provided predominantly by universitytrained exercise professionals, including physical therapists and clinical exercise physiologists. All but one trial included at least one session of in-person supervised exercise instruction (ie, individualized demonstration and practice).²⁰

The IBMEWG recommended that all people are assessed by a physical therapist or clinical exercise physiologist, with additional cancer exercise education and ideally experience in working with people who have bone metastases (Appendix Table A1). This may necessitate that exercise professionals initially are advised or supervised by a more experienced colleague, and these recommendations may need to be adapted in different jurisdictions and clinical settings. Stakeholders with bone metastases endorsed the importance of members of the health care team being able to guide people to qualified professionals. The IBMEWG highlighted that considerable work is needed to establish a referral pathway to improve equitable access to qualified exercise professionals. Each exercise professional must consider their own training and experience, as well as the setting in which they work, to determine if it is appropriate to work with that client or to refer. Finally, the results of the systematic review suggest that an element of supervised exercise instruction should be initially included before the addition of unsupervised exercise for individuals with bone metastases. Although this supports the potential feasibility of delivering supervised virtual exercise programs in light of the COVID-19 pandemic, research is needed to determine the overall safety profile of virtual exercise for individuals with bone metastases.²⁶

Recommendation 4

Professional judgment should be used to consider if exercise testing at baseline and follow-up is necessary by weighing the risks and benefits of including the test or if the testing protocols may need to be modified.

In the Delphi, for lower and higher risk examples, the statement that "exercise testing was recommended but there were some tests that were contraindicated" met consensus. Specific to strength testing, there was consensus to avoid tests or use caution with a test that places stress on a lesion site (Table 1). In the systematic review, few studies provided sufficient details on which participants completed the tests outlined in the methods and why a test may or may not be used for select participants. For the four studies exclusively in people with bone metastases, all

outlined specific adjustments to the testing protocol.²⁰ For example, Galvao et al²⁷ excluded people with metastatic bone lesion(s) in the proximal femur from completing the 1-RM leg press and 400-m walk tests, and people with lesions in ribs, thoracic spine, or humerus were excluded from completing the 1-RM chest press and 1-RM seated row.

The IBMEWG assessed that there was insufficient evidence to guide specific recommendations on a standardized approach for safe exercise testing. To provide additional expert guidance, the recommendation is to focus on the goal of exercise testing and to follow the testing guidance in the International Exercise Guidelines for Cancer Survivors⁴ until further evidence is available to update this approach (Appendix Table A1). In addition, exercise practitioners should use professional judgment to consider if a test is necessary and consistent with the person's goals, while providing written justification for inclusion of the test as part of the treatment plan in an appropriate charting location.

Recommendation 5

Exercise prescription should follow the standard exercise recommendations as outlined by the International Exercise Guidelines for Cancer Survivors, with greater emphasis on postural alignment, controlled movement, and proper technique, as well as consideration given to the location and presentation of the bone lesion(s). Formal monitoring of exercise response and adjustment of exercise prescription should be ongoing.

The Delphi shows a 91% consensus that education and advice on safe movement patterns for activities of daily living is important for all people with bone metastases (Table 1). Specific to prescribing an exercise that could directly place stress on the lesion site, most respondents agreed that the correct approach was to use caution. Respondents defined this as "start with active movement and no load or weight and progress slowly, provided there is no exacerbation of pain or adverse symptoms" (47%) or "start with low loads and progress slowly, provided there is no exacerbation of pain or adverse symptoms (28%)." This is a notable departure from approaches outlined originally by Cormie et al²⁸ and Galvao et al,²⁷ in which exercise involving the affected bone region was avoided altogether. In the systematic review, exercise prescription modifications specific to the presence of bone metastases were included in 41% of trials, whereas the remainder of trials did not report exercise modifications specific to bone metastases. How a participant's response to exercise was monitored and the related adjustments made to the exercise prescription were not uniformly reported.²⁰

The IBMEWG discussed the available literature that could be used to inform specific guidance around exercise prescription approaches. Exercise prescription for resistance exercises that prevent direct stress on the site of bone lesions has been published by IBMEWG members (P.C.,

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N.H.H., and R.U.N.).^{27,28} It was acknowledged that these previously published approaches were conservative and provided a starting point, while evidence regarding safety continued to emerge. The challenge in applying these generic prescription approaches is that individuals with widespread metastatic disease may be recommended to engage in limited exercise options that do not target the overall function or the goals of the individual. The approved movements can also be more restrictive than many activities of daily living that place considerable load on the body, including sites of bone metastases, such as descending stairs.

The IBMEWG recommendation aims to move beyond the initial and restrictive approaches to exercise testing and prescription with the goal better informing the development of efficacious exercise prescriptions for people with bone metastases in the clinical setting. The standard approach to exercise prescription for people living with and beyond cancer as outlined by the International Exercise Guidelines for Cancer Survivors from the American College of Sports Medicine⁴ should be used, with ongoing monitoring of individual response to adjust the exercise prescription or to send the person for further review. It was also deemed important to take additional guidance from exercise literature in osteoporosis that has sought to reduce the risk of skeletal-related complications, especially in vertebral bodies and hip.^{29,30} This includes an emphasis on correct technique and postural alignment with all exercises; avoidance of rapid or loaded end-range movements, such as rotation, flexion, or extension movements that involve the area of the lesions; consideration of impact loading of an exercise along with the type of movement (ie. concentric. eccentric, and plyometric) in the area of the bone lesion(s); and education and precautions to minimize the risk of falls. An additional consideration is that people with bone metastases may be deconditioned, because of treatment or higher symptom burden, and exercise volume should be adjusted accordingly.

Overarching Recommendation

Regular exercise has the potential to maintain or improve physical function and health-related quality of life in people with bone metastases, and the perceived risk of skeletal complication should be weighed against the potential health benefits.

On the basis of the review of available evidence and expert consensus, the IBMEWG has concluded that people with bone metastases should be supported and encouraged to engage in physical activity, including structured exercise, to obtain the well-established general health benefits, as a strategy to manage side effects related to cancer and treatments.⁴ Overall, the perceived risk of skeletal complications should be weighed against the potential health benefits of regular physical activity, including prevention of further loss of functional capacity. Exercise professionals

should communicate with the health care team to minimize participant risk while providing sufficient exercise stimulus to improve or maintain function or to slow decline, while prioritizing the goals of the person. The IBMWG also identified research priorities to allow for the continuous evolution of evidence-based guidelines (Table 1).

This recommendation was strongly endorsed by our stakeholders, both people with bone metastases and clinician peers, who felt it essential for members of the health care team to initiate a conversation about exercise, explain why exercise is important for people with bone metastases, and frame exercise as a range of activities, spanning from activities of daily living to sports. Stakeholders with bone metastases also highlighted the need to increase the availability of exercise services led by qualified exercise professionals and the provision of more information on how to access services.

DISCUSSION

These recommendations aim to provide a framework for improved integration of exercise programming into the care of people with bone metastases. The overarching recommendation is that routine exercise has the potential to improve or maintain physical function and quality of life and reduce treatment side effects, while also potentially increasing resilience for future treatments. To achieve provision of exercise programming, a collaborative relationship between the health care team and qualified exercise professionals is optimal. This approach is intended to improve the health care team's confidence in encouraging people with bone metastases to engage in regular exercise and provide navigation on how to access qualified exercise professionals who can develop appropriate and individualized exercise programming.

Determining the ideal approaches to pre-exercise screening, exercise testing, and exercise programming related to safety and skeletal-related events (SREs) remains a challenge. The IBMEWG recommends that exercise programming for people with bone metastases is administered by qualified exercise professionals, namely, physical therapists and clinical exercise physiologists, who have additional training and experience in working with people with a cancer diagnosis. These exercise professionals have university-level training (or equivalent) in most countries and a scope of practice that includes working with individuals with complex medical conditions.

A specific risk stratification tool to evaluate the appropriateness of exercise participation for an individual is desired by the health care team, exercise professionals, and people with bone metastases. However, the utility to apply existing screening tools used to predict the risk of fracture or guide decisions on prophylactic surgical fixation (ie, Taneichi, Mirels, and Spinal Instability Neoplastic Score³¹) within the exercise screening process for people with bone metastases is unknown. These have not been trialed extensively as part of the pre-exercise screening process to date and cannot be scored independently by an exercise professional. More research into the practical use of such a screening tool or newly developed tool is needed to determine if a standardized approach is feasible. Confounding this issue, the initial detection of bone metastasis is evolving with the advent of new imaging modalities. Unlike soft tissue tumors where Response Evaluation Criteria in Solid Tumors are clear, the response of bone metastasis to cancer treatment remains poorly understood.³² A recently proposed algorithm for the investigation of bone metastasis was proposed by the European Society for Medical Oncology in 2020, and this could be considered for future exercise studies.³²

The impact of cancer therapy and bone-modifying agents (BMA) on bone remodeling remains an area of intense study. With respect to BMA, zoledronic acid remains to date the only bisphosphonate to show broad efficacy in reducing SREs in people with bone metastases from almost all malignancies (including lung carcinoma and other solid tumor types: prostate, breast, thyroid, head and neck, thyroid, and renal cell).³² Data also show decreased SREs with the use of RANK ligand inhibitors (denosumab) for cancers of the prostate, breast, and renal cell, non-smallcell lung cancer, and multiple myeloma.³² In the future, the use of BMA should be clearly documented in exercise studies to develop a better understanding of their potential influence on bone remodeling. Furthermore, future studies should explore the use of bone biomarkers and imaging, especially positron emission tomography in exploring

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potential risk indicators and treatment response of exercise interventions.

A key strength of the IBMEWG was the process to merge a systematic understanding of the current literature with a formalized Delphi process and stakeholder input. The IBMEWG recommendations are consistent with two existing publications. Sheill et al³³ published a narrative review of considerations for exercise prescription for people with bone metastases, and the support organization Macmillan Cancer Support in the United Kingdom developed a user-friendly guidance document for health care professionals.³⁴ The IBMEWG aimed to address gaps in these documents by documenting the specific information gathering and decision-making processes used to generate the recommendations.

A fundamental limitation is a lack of definitive literature on standardized approaches to safety screening, exercise testing and prescription, and safety reporting of minor or major complications with exercise, including between exercise sessions (ie, pain, disability, and need for analgesia). In addition, there is a paucity of data on specific individuals that may be at increased risk of fracture with exercise (ie, elderly individuals with multiple myeloma).

In conclusion, these recommendations provide a framework and starting point for exercise professionals and members of the health care team to improved integration of physical activity into the care of individuals with bone metastases. It is anticipated that the recommendations provided here will continue to evolve as more literature is available.

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Exercise Recommendation for People With Bone Metastases: Expert Consensus for Health Care Providers and Exercise Professionals

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TABLE A1. Guidance for Exercise Testing Approach and Monitoring Exercise Response With Examples	of Qualified Exercise Professionals
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	Guidance for exercise testing approac	h			
Is the test necessary?	Consideration should be given to why the test is discussion with patient and healthcare team	being conducted, weighing the risks against the benefits, in			
Criteria for specific test selection:	or specific test selection: Testing should inform clinical decision making and exercise prescription				
	Testing should be adapted according to patier	nt goals			
	Individuals must be informed and feel comfort	table to perform the test safely			
Practical considerations	Avoid tests that place high loads on site(s) of bo proximal femur or vertebrae) ⁴	ne metastasis (ie, 1-RM leg press for individual with lesion in			
	Consider potential of fall risk when using tread available and blood pressure monitoring shou	Imill or other ambulatory machines (ie, handrails must be Ild be done with consideration of not compromising balance)			
	Consider forces testing approach on site(s) of including open <i>v</i> closed chain)	bone metastasis (ie, eccentric concentric, or isometric,			
	Consider the compressive forces of testing app press machine or repeated chair rise causir	proach (ie, pressure on lumbar spine from backrest of leg ng rapid/forceful impact onto the chair)			
	Be mindful of movement and forces needed to get into position for tests not just the test itself (ie, range of hip flexion needed to get into position for leg press in a leg press machine) as this can inadvertently cause unsafe movement related to risk of skeletal complication				
	Guidance for monitoring response to exercise	e training			
Observation	Exercise professionals should continuously monitor overall response to each exercise prescribed and adjust prescription as appropriate to reduce potential risk of exercise-related adverse event				
Self-reported response	Exercise professionals must ask the individual before and after each session about				
	Pain ^a	New or increased pain; or Change in quality or location of pain; or Unexplained pain; or Change in pain medication (dose and/or type) prior to or following an exercise session Ideally using a standardized tool, such as a visual analog scale			
	Neurological symptoms ^a	New symptom or change in symptom (eg, muscle weakness, loss of sensation or change in bladder/bowel function, balance or gait)			
	ADL	New limitation or change in limitations			
Exam	ples of qualified exercise professionals and can	cer certifications			
Professional title ^b	Regulatory or professional body	Examples of additional cancer training or certifications			
Licensed or registered physical therapist or physiotherapist	Colleges of Physical Therapy (State, Province, or National)	Specialist Certification in Oncology from the American Physical Therapy Association ³⁵			
		Clinical Specialist from Canadian Physiotherapy Association ³⁶			
ACSM-CEP	ACSM	ACSM Cancer Exercise Trainer ³⁷			
		CanRehab ³⁸			
CSEP-CEP	CSEP	Thrive Cancer and Exercise Training ³⁹			
ESSA-AEP	ESSA	ExMedCancer ⁴⁰			
		Exercise Medicine Research Institute ⁴¹			

Or equivalent in a given jurisdiction

Abbreviations: ACSM, American College of Sports Medicine; ADL, activity of daily living; AEP, Accredited Exercise Physiologist; CEP, Clinical Exercise Physiologist; CSEP, Canadian Society for Exercise Physiology; ESSA, Exercise and Sport Science Australia; RM, repetition maximum.

^aFor individuals with complex presentation, exercise professional should consider asking the individual before and after each exercise performed, especially if it is a new exercise.

^bThere is currently no single standardized licensing examination or number of clinic hours identified as optimal credentials for exercise professionals, specific to working with people with bone metastases.

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