# original article

Wien Klin Wochenschr (2020) 132:124–131 https://doi.org/10.1007/s00508-019-01582-z

# Wiener klinische Wochenschrift

The Central European Journal of Medicine



# Relevant parameters for recommendations of physical activity in patients suffering from multiple myeloma

# A pilot study

Fadime Cenik · Mohammad Keilani · Timothy Hasenöhrl · Dominikus Huber · Bianca Stuhlpfarrer · Anna Pataraia · Richard Crevenna

Received: 13 September 2019 / Accepted: 6 November 2019 / Published online: 29 November 2019 © The Author(s) 2019

Summary

*Purpose* This pilot study aimed to describe physical performance, self-reported physical activity, health-related quality of life, anxiety and depression in patients who were assigned from Austrian self-help groups for multiple myeloma patients. These parameters were then discussed in the context of clinical decision-making concerning the recommended type of regular physical activity and exercise.

Methods Members of the self-help groups were invited to participate. Physical performance and physical activity were assessed with the 6 min walk test (6MWT), handgrip strength test, timed up and go test (TUG), Tinetti performance oriented mobility assessment (POMA), falls efficacy scale (FES), international physical activity questionnaire (IPAQ), health-related quality of life (EORTC QLQ-C30) and the hospital anxiety and depression scale (HADS).

Results A total of 40 patients (female:male=15:25, mean age: 63.8±9.0 years, range 41–80 years) were identified. In total 20 (50%) reached the performance of healthy peers in the tests 6MWT, handgrip strength, TUG and POMA, while 50% showed at least 1 result below the reference value or cut-off-point for each test. Self-reported activity levels were high. Patients showed a tendency to overestimate the risk of falling but a case by case analysis revealed a tendency for underestimating the actual performance in the respective tests (TUG, POMA).

F. Cenik·M. Keilani·T. Hasenöhrl·
D. Huber·B. Stuhlpfarrer·A. Pataraia·
Univ. Prof. Dr. R. Crevenna, MBA, MMSc (⋈)
Department of Physical Medicine, Rehabilitation and
Occupational Medicine, Medical University of Vienna,
Waehringer Gürtel 18–20, 1090 Vienna, Austria
richard.crevenna@meduniwien.ac.at

Conclusion The performance of healthy peers was reached by a substantial number of the participants in tests of physical performance and they reported high levels of physical activity. Nevertheless, they tended to overestimate the specific risk of falling. Patients with notably impaired physical performance might be suitable to perform regular physical activity and exercise in an individual therapy, whereas those with good physical performance are suited for training in exercise groups; however, individual contraindications and clinical considerations should be noted in a multiprofessional and interdisciplinary setting.

**Keywords** Plasmacytoma  $\cdot$  Exercise  $\cdot$  Self-help groups  $\cdot$  Bone lesion

#### Introduction

Multiple myeloma is the second most common hematologic malignancy [1]. Patients suffering from multiple myeloma show an increase in cancer-specific survival rate due to modern cancer treatment [2, 3]. Regular physical activity and exercise are important measures to improve health-related quality of life and physical performance in cancer patients [4–7]. Recommendations for regular physical activity and exercise consist of 150 min/week of moderately intense or 75 min/week of vigorously intense activity or an equivalent combination, muscle strengthening activities of at least moderate intensity at least 2 days/week for each major muscle group and stretching of major muscle groups and tendons [2, 4–12].

In Austria approximately 2200 patients suffer from multiple myeloma [13]. There are two main self-help groups in Austria, the so-called *Multiples Myelom* and *Myelom- und Lymphomhilfe* offering regular meetings and conferences for patients [14, 15]. Nevertheless, the opportunity to be involved in myeloma exer-



cise groups does not yet exist in Austria. Due to bone manifestation of multiple myeloma and special clinical features (especially hypercalcemia and monoclonal gammopathy), a multiprofessional and interdisciplinary approach is required for planning and prescribing regular physical activity and exercise [2, 8–12]. Hereby, physical performance seems to be an important factor to identify which patients should receive an individual therapy or a therapy in an exercise group [11, 16–19]. Physical performance depends on endurance capacity, muscle strength and sensorimotor function.

This pilot study aimed to describe physical performance, self-reported physical activity, health-related quality of life, anxiety and depression in patients who were assigned from two Austrian self-help groups. These parameters were then discussed in the context of clinical decision-making, concerning the recommended type of regular physical activity and exercise.

#### Patients, materials and methods

The two main self-help groups (Myelom- und Lymphomhilfe and Multiples Myelom) invited their members with multiple myeloma (aged between 18 and 99 years) to participate in this cross-sectional study. The patients were invited via conferences, emails and the webpage and also via Facebook. After agreeing to participate and checking out the inclusion criteria via telephone, a medical examination was carried out to survey status and demographic data before assessing the parameters presented below. Patients with preliminary stages, such as monoclonal gammopathy of undetermined significance (MGUS) or asymptomatic (smoldering/indolent) myeloma without targeted cancer treatment were excluded from this study. Furthermore, the patients who did not fully understand the study measures or could not perform them, did not speak German sufficiently well or were unable to cognitively or physically follow the course of study were excluded. Illiteracy and refusal to participate were other exclusion criteria. To give the patients from all regions of Austria the best opportunity to participate in this study travel expenses were remunerated.

This pilot study took place at the Department of Physical Medicine, Rehabilitation and Occupational Medicine of the Medical University of Vienna, Austria. The ethics committee of the Medical University of Vienna, Austria approved the present study (EK.-Nr.: 1725/2018). All patients signed a written informed consent before enrolment. The participants were assessed from February 2019 to July 2019.

### Assessment

To assess physical performance, the 6-min walk test (6MWT), the handgrip strength test (HGST by JAMAR®

dynamometry [Patterson Medical, Warrenville, IL, USA]), the timed up and go test (TUG) and the Tinetti performance oriented mobility assessment (POMA) were performed. Additionally, the patients were asked to fill in a series of standardized questionnaires and scales:

- Health-related quality of life of the participants (EORTC QLQ-C30)
- Clinically significant states of anxiety and depression (hospital anxiety and depression scale, HADS)
- Self-reported physical activity (international physical activity questionnaire, IPAQ)
- Fall-related self-efficacy (falls efficacy scale, FES)

The TUG is a commonly used screening tool to assist clinicians to evaluate gait and balance functions, and to identify patients at risk of falling [20]. There are different procedures for the TUG, namely with normal and fast walking speeds [20]. In the present study the normal walking speed and cut-off-point for impaired mobility were used (>10 s) based on the recommendations of the Austrian Society for Geriatrics and Gerontology [21] as well as Podsiadlo and Richardson [22]. Concerning the POMA, the assessment was based on Tinetti with the following classification: <19 points for high fall risk, 19–23 points for moderate fall risk and 24–28 points for minimal fall risk [23].

The IPAO was used to indirectly evaluate an individual's extent of sedentary behavior and moderate to vigorous physical activity throughout the last 7-day week. The questionnaire was scored in terms of estimated metabolic equivalent task minutes per week (MET-min/week) [24]. For the FES the cut-off-points established by Delbaere et al. were used, who classified low concern about falling with 16–19 points, moderate concern with 20-27 points and high concern with 28-64 points [25]. The HADS evaluation was based on Zigmond and Snaith with the following classification and definition: 0-7 points as normal case, 8-10 as doubtful case, ≥11 as definitive case for each sub-scale [26]. The age-matched normative data published by Breeman et al. were used for analysis, which were available only for people younger than 65 years old [27]. Therefore, all older participants were classified as being 65 years old in this study. The 6MWT is widely used as a test of general physical performance, mobility and submaximal exercise capacity [28]. For the 6MWT, the following formula was used to calculate the predicated walking distance in meters for each participant:  $(7.57 \times height_{cm}) - (5.02 \times age_{vear}) - (1.76 \times weight_{kg}) - 309$  $(2.11\times height_{cm}) - (2.29\times age_{vear})$ and  $(5.78 \times \text{weight}_{kg}) + 667$  for female participants [28]. For the HGST mean values of each side were used to compare with the JAMAR® age and sex-matched reference values [29–31]. Existing reference values with multiple myeloma patients from the EORTC group were taken from Scott et al. (2008) [32]. Continuous variables are presented as means and standard deviations, medi-

# original article

ans and ranges and nominal variables as percentages. The Kolmogorov-Smirnov test was used to test each variable for normal distribution. In case the null-hypothesis from this test was retained, non-parametric solutions like Wilcoxon-signed rank test and Spearman's correlation were employed to test the related samples for difference in means and correlation, respectively. Due to the retrospective character of the study, no risk ratios could be calculated. Nonetheless, odds ratios (OR, corrected via Haldane-Anscombe for low sample size) were calculated for a pathological test result in one of the primary outcome parameters (columns) given a specific patient characteristic (rows).

#### Results

In total, 55 patients agreed to participate of whom 9 were excluded due to exclusion criteria (suffering from preliminary stages; had not received any cancer treatment), 4 were not available after self-registration for this study, 1 patient did not want to travel to the study location and 1 patient was recommended not to participate by the family physician. Finally, 40 patients (female:male=15:25) fulfilled the inclusion criteria. The demographic data of the participants are presented in Table 1.

Each patient had received (n=8) or was still receiving (n=32) chemotherapy, new substances and/or immunotherapy (with or without stem cell transplantation). The results of the 6MWT, HGST, POMA, TUG, HADS, IPAQ and EORTC QLQ-C30 are presented in Table 2.

On average the participants showed a moderate subjective risk of falling (moderate concern in the FES) and a low to moderate fall risk in the objective fall risk assessments (POMA and TUG test, Table 3). In more detail, 20 patients (50%) reported a moderate or high concern about falling (FES >10) while in fact only 10 patients (25%) showed an elevated risk in the TUG test, of whom 9 patients (22.5%) were deemed at high risk by the POMA. Among those patients who reported a moderate to high subjective risk of falling (in the FES), 70% performed well in the remaining objective risk assessment tools. A case by case analysis therefore revealed a tendency of patients to overestimate the fall risk and underestimate the actual performance in the respective tests.

The third objective test for physical performance was the handgrip strength test. Overall, 6 (15%) participants showed a reduced handgrip strength in comparison with age and sex-matched reference values. The 6MWT was employed as fourth objective evaluation of physical performance and 12 patients showed results below the individual age, sex, height and weight-predicted values (see methods). Results from this test are the only data in this study that followed a normal distribution (Kolmogorov-Smirnov test retains the null hypothesis of normal distribution

Table 1 Demographic data

Table T Demographic data			_		
n=40	Mean	SD	Range		
Age (years)	63.8	9.0	41–80		
Female	64.5	7.7	52–80		
Male	63.3	9.8	41–80		
Family status	п				
Married	23				
Cohabitation	6				
Divorced	4				
Single	3				
Widowed	3				
n.s.	1				
Employment					
Yes	7				
No (retired)	32				
No (housewife)	1				
Highest level of education					
University	13				
Business/high school	15				
Obligatory education/apprenticeship	12				
Household income/year (€)					
10,000–25,000	10				
25,000–50,000	17				
>50,000	3				
n. s.	10				
Smoker	3				
Time since initial diagnosis					
<1 year	5				
1–2 years	5				
3–5 years	10				
5–10 years	12				
>10 years	8				
Stem cell transplantation					
Yes	28				
No	12				
Pathologic fracture/risk					
Yes	21				
No	19				
Radiation due to fracture/risk	10				
No	21				
Yes	19				
Operation due to fracture/risk	13				
No	31				
Yes	9				
Polyneuropathy	22				
Cancer rehabilitation performed	22				
·					
No Voc	20				
Yes	19				
n.s.	1				
SD standard deviation, n. s. not specifie	u				



Table 2 Results of all parameters

Idabio = Tiocdito of all	paramotoro		
Parameter	Total (n=40) mean ± SD	Median	Range
6-MWT in m	548 ± 128	573	225-835
HGST in kg	$32.4 \pm 9.6$	32.5	17–51
POMA (<19 = high fall risk) (points)	$23.5 \pm 4.8$	26	12–28
Balance	$13.4 \pm 3.0$	15	5–16
Gait	$10.1 \pm 2.1$	11	6–12
TUG (s)	$9.5 \pm 3.1$	9	5.5-21.0
FES	$21.2 \pm 6.0$	20	16–39
HADS (points)	$10.5 \pm 5.0$	10	1–23
Anxiety	$5.7 \pm 3.2$	6	0–12
Depression	$4.9 \pm 2.7$	4	1–11
MET-min/week	Total (n=35) mean ± SD		
Vigorous	$888.0 \pm 1873.3$	0	0-10,080
Moderate	$830 \pm 1275.3$	240	0-5040
Walk	1110.7 ± 1127.8	792	0-2772
Total	$2829.5 \pm 3041.5$	1584	0-16,506
EORTC QLQ-C30 (points)	Total (n=40) mean ± SD		Reference
Global health status/QOL	$60.0 \pm 21.5$	62.5	$55.7 \pm 22.8$
Physical functioning	75.3 ± 18.1	80.0	$67.7 \pm 23.4$
Role functioning	$65.0 \pm 23.5$	66.7	$60.1 \pm 33.4$
Emotional functioning	$67.3 \pm 24.7$	66.7	$71.3 \pm 22.7$
Cognitive functioning	$70.8 \pm 24.7$	66.7	$78.1 \pm 23.8$
Social functioning	$64.6 \pm 29.0$	66.7	$63.2 \pm 31.0$
Fatigue	$47.8 \pm 22.3$	44.4	$48.7 \pm 26.7$
Nausea and vomiting	$7.1 \pm 13.5$	0	$10.5 \pm 19.2$
Pain	$36.7 \pm 30.5$	33.3	$47.1 \pm 33.9$
Dyspnea	$30.0 \pm 25.9$	33.3	$26.0 \pm 27.3$
Insomnia	$42.5 \pm 33.7$	33.3	$28.9 \pm 30.6$
Appetite loss	12.5 ± 24.7	0	$23.2 \pm 30.2$
Constipation	13.3 ± 27.0	0	$23.2 \pm 29.9$
Diarrhea	$20.0 \pm 30.0$	0	$9.6 \pm 19.4$
Financial difficulties	$20.0 \pm 29.0$	0	$16.1 \pm 26.6$
SD standard deviation. QOI	quality of life. 6MI	<i>NT</i> 6-min w	alk test.

SD standard deviation, QOL quality of life, 6MWT 6-min walk test, HGST handgrip strength test, POMA Tinetti performance oriented mobility assessment, TUG timed up and go test, FES falls efficacy scale, HADS hospital anxiety and depression scale, MET metabolic equivalent of task, EORTC QLQ-C30 European Organization for Research and Treatment of Cancer Quality of Life Questionnaire-Core 30

at p>0.20). Patients walked on average 547.9 m in 6 min and the standard deviation was 127.8 m. Patients who did not reach the predicted value (n=12) walked on average 419 m and those who did reach the predicted values (n=28) 603 m. Despite the low sample size, both groups followed a normal distribution (Kolmogorov-Smirnov test p>0.20) and the means in these two groups are highly significantly different (2-sided Wilcoxon-signed-rank test p<0.01). On average, patients who reached the predicted values walked 43% further in 6 min than the peers who did not reach their predicted values. When combining the objective tests for hand grip strength, exercise

**Table 3** Patients underwent four different objective measures of physical performance (6MWT, POMA, TUG, HGST)

Sex	Age (years)	6MWT (m)	HGST (kg)	POMA (points)	TUG (s)		
m	59	600	38	28	8		
m	54	643	44	28	7		
m	61	640	41	27	7		
f	67	530	26	25	9		
m	70	550	38	27	8		
m	65	750	42	28	6		
f	64	475	26	27	9		
f	59	560	26	27	9		
f	63	580	34	28	9		
m	67	602	51	28	8		
f	52	625	30	27	8		
f	66	605	22	26	8		
f	66	517	32	26	7		
m	51	720	45	27	9		
m	60	605	43	26	9		
m	55	835	40	26	8		
m	70	620	30	26	7		
m	57	660	50	28	9		
m	70	680	34	27	6		
f	61	575	26	28	7		
m	69	470	36	26	7		
m	59	600	50	20	8		
f	66	460	22	22	8		
m	79	500	26	16	9		
m	61	500	32	25	10		
f	62	590	15	24	9		
m	53	650	40	27	8		
f	57	570	26	24	13		
f	71	478	18	21	9		
m	78	525	30	18	10		
m	55	660	20	25	13		
m	72	520	35	18	11		
m	41	650	36	21	9		
m	59	358	30	22	12		
f	78	360	20	16	14		
f	55	305	17	15	12		
m	61	425	38	13	11		
m	80	427	50	18	11		
m	77	225	37	12	21		
f	80	270	18	15	19		
N-20 nationts showed a nathological result in at least one test (hold) while							

N= 20 patients showed a pathological result in at least one test (*bold*) while the other 20 patients reached the individual cut-off-points of the healthy peers (*italic*)

m male, f female, 6MWT 6-min walk test in meters, POMA Tinetti performance oriented mobility assessment, TUG timed up and go test, HGST handgrip strength test of the dominant hand in kilograms

## original article

**Table 4** Comparison of the patients below (n = 12) the individual age, sex, height and weight-predicted values, with the remaining patients

	6MWT (m)	Age	POMA (points)	TUG (s)	HGST (kg)	HADS (points)	FES (points)	MET-min/week	QOL (points)	Pain (points)
Patients be	low their re	eference va	alues (n =	12)						
Mean	419.8	65.8	19.0	12.2	29.8	11.4	25.6	2902.7	57.6	37.5
SD	123.9	12.1	4.8	4.1	10.3	4.9	8.4	2513.1	25.5	29.4
Median	426.0	65.0	19.5	11.5	31	13.0	22.5	1413.0	58.3	33.3
Remaining patients (n = 28)										
Mean	602.8	62.9	25.4	8.3	34.1	10.1	19.3	2791.4	61.0	36.3
SD	83.2	7.3	3.4	1.5	9.7	5.1	3.3	3336.9	20.0	31.4
Median	601.0	62.5	27.0	8.2	34.0	10.0	18.0	1653.0	66.7	33.3

SD standard deviation, m male, f female, 6MWT 6-min walk test, POMA Tinetti performance oriented mobility assessment, TUG timed up and go test, HGST handgrip strength test of the dominant hand, HADS hospital anxiety and depression scale, FES falls efficacy scale, MET metabolic equivalent of task, QOL quality of life

**Table 5** Odds ratios for pathologic test results

	TUG (s)	POMA (points)	6MWT (m)	HGST (kg)	
Rehabilitation	1.07 (0.26; 4.10)	1.48 (0.28; 4.93)	0.73 (0.23; 3.27)	2.26 (0.26; 7.68)	
Stem cell therapy	0.10 (0.05; 2.54)	0.17 (0.02; 8.98)	0.11 (0.02; 7.43)	1.19 (0.14; 7.96)	
PNP	1.55 (0.30; 4.78)	1.01 (0.24; 4.19)	1.18 (0.24; 3.47)	1.61 (0.23; 6.61)	
Smoking	1.57 (0.14; 10.4)	0.43 (0.03; 14.6)	4.37 (0.22; 16.2)	12.4 (0.31; 28.3)	
TUG timed up and go test, POMA performance oriented mobility assessment, 6MWT 6-min walk test, HGST handgrip strength test, PNP polyneuropathy					

Table 6 Effect size on failing an outcome parameter cutoff

	TUG (s)	POMA (points)	6MWT (m)	HGST (kg)
Rehabilitation	-0.02	0.09	-0.07	0.20
Stem cell therapy	-0.57	-0.42	-0.52	0.04
PNP	0.17	0.00	0.04	0.11
Smoking	0.13	-0.20	0.35	0.60

TUG Timed up and Go test, POMA Performance Oriented Mobility Assessment, 6MWT 6-minute walk test, HGST Handgrip strength test, PNP Polyneuropathy

capacity (6MWT) and risk of falling (TUG test and POMA), 20 (50%) patients in the sample of 40 multiple myeloma patients showed a pathologic result in at least 1 test, while the other 50% performed above the cut-off points in all tests (Table 3).

Self-reported physical activity measured by the IPAQ showed high activity levels among the patients in this study. During the scoring procedure, 5 participants were excluded from the analysis (due to either missing information regarding duration of physical activity or being an outlier: self-reported total activity >960 min). In total, 8 patients (23%) showed total MET-min values equivalent of low activity, while 77% reported at least moderate activity levels. In terms of exercise intensity, 8 patients reported only walking activity during the last week. The remaining patients performed at least moderate if not vigorous activity. The 12 participants who showed results below the individual age, sex, height and weight-predicted values in the 6MWT also seemed to have impairments in mobility, self-reported physical activity and healthrelated quality of life. Furthermore, they had higher levels regarding depression and anxiety (Table 4).

With respect to the small sample size of n=40, no further statistical evaluation was performed.

The ORs are presented in Table 5. All ORs showed only very weak evidence of impacting any of the outcomes (OR of 1 contained in the 95% confidence interval). In terms of effect size, an approximation of Cohen's d was calculated from the OR (Table 6). Among all recorded disease-modifying factors, only stem cell therapy in addition to chemotherapy showed a strong effect on three of the four primary outcome parameters (reducing the odds of a pathological outcome). Oddly, the positive patient characteristics had a positive effect size on not reaching the cut-off in the primary outcomes.

## **Discussion**

The present study was performed by analyzing members of two self-help groups in Austria. Self-help groups are an essential measure of patient involvement and play an important part in the healthcare system. Members of self-help groups are often wellinformed about the management of the disease and highly motivated to participate in regular physical activity. The finding that most patients reported at least moderate to high levels of physical activity supports this observation. Moreover, a substantial amount of study participants reached the performance of the healthy peers in tests of physical performance, such as the 6MWT (n = 28, 70%), the TUG test (n = 29, 73%), the POMA (n=26, 65%) and handgrip strength test (n=34, 85%). While this performance seems to be underestimated by the patients themselves, they tend to overestimate theispecific risk of falling (FES). This discrepancy is of particular interest as a higher subjective



concern of falling can lead in the long run to a vicious circle between impaired physical performance and physical activity. This consequently leads to loss of muscle and bone mass lowering bone density and therefore increasing risk of bone fractures. This is in line with the results as the 12 participants with results in the 6MWT below the individual age, sex, height and weight-predicted values had lower selfreported physical activity and health-related quality of life as well as higher levels of depression and anxiety. Patients with impaired subjective and objective walking performance should therefore be encouraged to at least maintain physical activity levels for as long as possible and perform suitable and supervised individual exercise to improve sensorimotor function, mobility, muscle strength, endurance capacity and independence from others' help. In comparison, exercise studies with multiple myeloma patients usually exclude patients with pathological fractures or increased risk of fractures [11, 16, 17]. Therefore, a multidisciplinary and interdisciplinary approach as described is necessary to manage the actual condition and to evaluate bone load-bearing capacity of this patient group [11, 18, 19].

Most patients (75%) who did not reach normal values in the 6MWT, TUG, POMA or handgrip strength tests did not reach their goal in at least two of these tests. Data only allow a differentiation between patients who reached the reference values and those who did not. More than half (n=22, 55%) of the patients reported having symptoms of polyneuropathy, which could influence the fear of falling and fall risk in theory; however, the performance in the four objective physical performance tests (6MWT, TUG, POMA, HGST) did not seem to differ between patients with PNP and those without. Overall 20 (50%) patients in the sample of 40 multiple myeloma patients did not reach the reference values in at least 1 of the tests for handgrip strength, exercise capacity (6MWT) and risk of falling (TUG test and POMA), while the other 50% performed above cut-off values in all tests (Table 3). Clinical recommendations for group therapy could safely be given to the latter patient sample, while the former group would be assigned to single therapy sessions. Additionally, a patient's medical history, clinical examination, laboratory parameters, electrocardiography, echocardiography, exercise testing, spirometry, and radiographic findings are required for planning and prescribing regular physical activity and exercise for patients suffering from multiple myeloma in an interdisciplinary setting [11, 18, 19]. Special clinical features that clinicians have to consider are bone manifestation, hypercalcemia (with risk of cardiac arrhythmia or kidney failure) and monoclonal gammopathy [11, 18, 19]. Additionally, due to the natural history of multiple myeloma with changing clinical status over time, exercise should be performed under supervision of physicians. Besides these objective measures there are also subjective measures, such

as pain which was assessed in this study via EORTC QLQ-C30; however, even the patients in the non-fit group (6MWT values below the individual predicted values) reported pain intensity similar to the reference values. Moreover, besides radiographic findings pain is an important factor for evaluation of spinal stability of cancer lesions [33] and pathological fractures [34, 35]. As pain will always be a limiting factor for physical activity and exercise, the symptom pain should be considered when planning exercise groups and treated in a multidisciplinary manner by use of a multimodal approach [19].

None of the recorded patient characteristics (oncological rehabilitation, additional stem cell therapy, polyneuropathy, and smoking) could establish statistical evidence of modifying the outcome of the primary parameters (POMA, TUG, 6MWT, HGST). Stem cell therapy, however, showed a strong effect size on the balance tests and walking distance. A future study on multiple myeloma patients should therefore bolster this finding with greater sample size. Moreover, more confounding variables from the patient's medical records, such as bone metastases, fractures, precise chemotherapy regimen and a more detailed history of physical therapy should be taken into account.

The present study has some limitations. The majority of the mobility tests were developed for use in geriatric patients in the original context. For the POMA and TUG there are various existing cut-off points, classifications and approaches for the test procedure in the literature but mostly researched in older people [20]. Moreover, due to non-existing reference groups in patients older than 65 years, the oldest participants were classified as if they were 65 years old. Despite the low sample size of n=40 patients, statistical inference was possible to a certain degree; however, a larger sample would have been beneficial and would have allowed a stronger generalizability. When comparing subgroups, however, statistical analysis is less reliable and warrants great caution. Another weakness of the assessment battery was its length, which might have discouraged some participants from participating in this study in the first place. Finally, the recruitment of this study sample was performed via invitations. Therefore, a selection bias regarding motivation to participate is to be expected. Clinical experience shows that self-help group members are especially motivated, educated and active, which is a clear bias towards a generally healthier patient sample. Moreover, assessing fracture risk from the existing patients' diagnostics, which were incomplete in this respect and not including radiographic assessments during the study protocol made it impossible to categorize the participants into high, moderate, and low fracture risk as a possible basis for more precise recommendations for regular physical activity and exercise in patients with multiple myeloma.

#### Conclusion

A substantial number of participants reached the performance of healthy peers in tests of physical performance and reported high levels of physical activity. Nevertheless, they tended to overestimate the specific risk of falling. Patients with notably impaired physical performance might be suitable to perform regular physical activity and exercise in an individual therapy, whereas those with good physical performance are suited for training in exercise groups. To ensure safety and effectiveness of these interventions in patients with multiple myeloma, however, individual contraindications and clinical considerations should be noted in a multiprofessional and interdisciplinary setting.

**Funding** Open access funding provided by Medical University of Vienna.

**Conflict of interest** F. Cenik, M. Keilani, T. Hasenöhrl, D. Huber, B. Stuhlpfarrer, A. Pataraia, and R. Crevenna declare that they have no competing interests.

**Open Access** This article is distributed under the terms of the Creative Commons Attribution 4.0 International License (http://creativecommons.org/licenses/by/4.0/), which permits unrestricted use, distribution, and reproduction in any medium, provided you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons license, and indicate if changes were made.

#### References

- 1. Radocha J, Hájek R, Brožová L, et al. Simplified novel prognostic score for real-life older adults with multiple myeloma-registry-based analysis. Ann Hematol. 2019;98(4):951–62.
- 2. Warren JL, Harlan LC, Stevens J, Little RF, Abel GA. Multiple myeloma treatment transformed: a population-based study of changes in initial management approaches in the United States. J Clin Oncol. 2013;31:1984–9.
- 3. Kumar SK, Dispenzieri A, Lacy MQ, et al. Continued improvement in survival in multiple myeloma: changes in early mortality and outcomes in older patients. Leukemia. 2014;28:1122–8.
- 4. Mock V, Pickett M, Ropka M, et al. Fatigue and quality of life outcome of exercise during cancer treatment. Cancer Pract. 2001;9(3):119–27.
- 5. Crevenna R. Aspects of cancer rehabilitation: an Austrian perspective. Disabil Rehabil. 2019;27:1.
- 6. Crevenna R. Cancer rehabilitation and palliative care-two important parts of comprehensive cancer care. Support Care Cancer. 2015;23(12):3407–8.
- 7. Palma S, Keilani M, Hasenoehrl T, Crevenna R. Impact of supportive therapy modalities on heart rate variability in cancer patients—a systematic review. Disabil Rehabil. 2018;4:1–8.
- 8. American College of Sports Medicine, Schmitz KH, Courneya KS, Matthews C, et al. American College of Sports Medicine roundtable on exercise guidelines for cancer survivors. Med Sci Sports Exerc. 2010;42(7):1409–26. https://doi.org/10.1249/MSS.0b013e3181e0c112. Erratum in: Med Sci Sports Exerc. 2011 Jan;43(1):195..

- 9. Dimeo F, Schwartz S, Fietz T, Wanjura T, Boning D, Thiel E. Effects of endurance training on the physical performance of patients with hematological malignancies during chemotherapy. Support Care Cancer. 2003;11:623–8.
- 10. ChaS, KimI, LeeSU, Seo KS. Effect of an inpatient rehabilitation program for recovery of deconditioning in hematologic cancer patients after chemotherapy. Ann Rehabil Med. 2018;42(6):838–45.
- Keilani M, Kainberger F, Pataraia A, et al. Typical aspects in the rehabilitation of cancer patients suffering from metastatic bone disease or multiple myeloma. Wien Klin Wochenschr. 2019; https://doi.org/10.1007/s00508-019-1524-3.
- 12. Kumar SK, Rajkumar V, Kyle RA, et al. Multiple myeloma. Nat Rev Dis Primers. 2017;3:17046.
- Statistik Austria. Plasmozytom, myelom. 2019. https://www.statistik.at/web\_de/statistiken/menschen\_und\_gesellschaft/gesundheit/krebserkrankungen/plasmozytom\_myelom/index.html. Accessed 7 Aug 2019.
- 14. Multiples Myelom. 2019. http://www.multiplesmyelom.at. Accessed 7th Aug 2019
- 15. Myelom-Lymphom. 2019. http://www.myelomlymphom.at. Accessed 7th Aug 2019
- 16. Coleman EA, Coon S, Hall-Barrow J, Richards K, Gaylor D, Stewart B. Feasibility of exercise during treatment for multiple myeloma. Cancer Nurs. 2003;26(5):410–9.
- 17. Groeneveldt L, Mein G, Garrod R, et al. A mixed exercise training programme is feasible and safe and may improve quality of life and muscle strength in multiple myeloma survivors. BMC Cancer. 2013;13:31.
- 18. Crevenna R, Kainberger F, Wiltschke C, et al. Cancer rehabilitation: current trends and practices within an Austrian university hospital center. Disabil Rehabil. 2018; https://doi.org/10.1080/09638288.2018.1514665.
- 19. Crevenna R. From neuromuscular electrical stimulation and biofeedback-assisted exercise up to triathlon competitions-regular physical activity for cancer patients in Austria. Eur Rev Aging Phys Act. 2013;10:53–5. https://doi.org/10.1007/s11556-012-0110-8.
- Schädler S. Grenzwerte (Cut-off), Sensitivität und Spezifität. 2010. https://www.stefan-schaedler.ch/wp-content/uploads/2014/11/Sturzrisikoassessments.pdf. Accessed 7 Aug 2019.
- 21. Österreichische Gesellschaft für Geriatrie und Gerontologie. Geriatrisches Basisassessment. 2019. https://www.geriatrie-online.at/publikationen/basisassessment-aneinem-strang-ziehen/. Accessed 7 Aug 2019.
- 22. Podsiadlo D, Richardson S. The timed "Up & Go": a test of basic functional mobility for frail elderly persons. J Am Geriatr Soc. 1991;39(2):142–8.
- 23. Tinetti ME. Performance-oriented assessment of mobility problems in elderly patients. J Am Geriatr Soc. 1986;34(2):119–26.
- 24. Craig CL, Marshall AL, Sjöström M, et al. International physical activity questionnaire: 12-country reliability and validity. Med Sci Sports Exerc. 2003;35(8):1381–95.
- 25. Delbaere K, Close JC, Mikolaizak AS, Sachdev PS, Brodaty H, Lord SR. The Falls Efficacy Scale International (FES-I). A comprehensive longitudinal validation study. Age Ageing. 2010;39(2):210–6.
- 26. Zigmond AS, Snaith RP. The hospital anxiety and depression scale. Acta Psychiatr Scand. 1983;67(6):361–70.
- 27. Breeman S, Cotton S, Fielding S, Jones GT. Normative data for the Hospital Anxiety and Depression Scale. Qual Life Res. 2015;24(2):391–8.



- 28. Enright PL, Sherrill DL. Reference equations for the six-minute walk in healthy adults. Am J Respir Crit Care Med. 1998;158(5 Pt 1):1384–7.
- 29. Mathiowetz V, Weber K, Volland G, Kashman N. Reliability and validity of grip and pinch strength evaluations. J Hand Surg Am. 1984;9(2):222–6.
- 30. Mathiowetz V, Kashman N, Volland G, Weber K, Dowe M, Rogers S. Grip and pinch strength: normative data for adults. Arch Phys Med Rehabil. 1985;66(2):69–74.
- 31. Mathiowetz V, Wiemer DM, Federman SM. Grip and pinch strength: norms for 6- to 19-years-old. Am J Occup Ther. 1986;40(10):705–11.
- 32. Scott NW, Fayers PM, Neil KA, et al. EORTC QLQ-C30 Reference Values. EORTC Quality of Life Group. 2008. https://www.eortc.org/app/uploads/sites/2/2018/

- 02/reference\_values\_manual2008.pdf. Accessed 7 Aug 2019.
- 33. Mirels H. Metastatic disease in long bones: a proposed scoring system for diagnosing impending pathologic fractures. Clin Orthop Relat Res. 2003;415 (Suppl): S4–S13.
- 34. Macedo F, Ladeira K, Pinho F, et al. Bone metastases: an overview. Oncol Rev. 2017;11(1):321.
- 35. Knips L, Bergenthal N, Streckmann F, Monsef I, Elter T, Skoetz N. Aerobic physical exercise for adult patients with haematological malignancies. Cochrane Database Syst Rev. 2019; https://doi.org/10.1002/14651858.CD009075.pub3.

**Publisher's Note** Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

