ORIGINAL ARTICLE

Adherence to prescribed exercise time and intensity declines as the exercise program proceeds: findings from women under treatment for breast cancer

Hsiang-Ping Huang • Fur-Hsing Wen • Jen-Chen Tsai • Yung-Chang Lin • Shiow-Ching Shun • Hsien-Kun Chang • Jong-Shyan Wang • Sui-Whi Jane • Min-Chi Chen • Mei-Ling Chen

Received: 10 September 2014 / Accepted: 7 December 2014 / Published online: 21 December 2014 © Springer-Verlag Berlin Heidelberg 2014

Abstract

Purpose Adherence to prescribed exercise is a challenge for cancer patients undergoing treatment. The changing pattern of exercise adherence over time cannot be fully understood by an overall measure of adherence. This study was aimed to identify the trajectory of exercise adherence and its predictors for women with breast cancer during their chemotherapy.

H.-P. Huang · S.-W. Jane

Department of Nursing, Chang Gung University of Science and Technology, Tao-Yuan, Taiwan

F.-H. Wen Department of International Business, Soochow University, Taipei, Taiwan

J.-C. Tsai School of Nursing, National Yang-Ming University, Taipei, Taiwan

Y.-C. Lin · H.-K. Chang Division of Hematology-Oncology, Department of Internal Medicine, Chang Gung Medical Foundation, Tao-Yuan, Taiwan

S.-C. Shun Department of Nursing in College of Medicine, National Taiwan University, Taipei, Taiwan

J.-S. Wang

Graduate Institute of Rehabilitation Science, Chang Gung University, Tao-Yuan, Taiwan

M.-C. Chen

Department of Public Health and Biostatistics Consulting Center, Chang Gung University, Tao-Yuan, Taiwan

M.-L. Chen (🖂)

Graduate Institute of Nursing, College of Medicine, Chang Gung University, 259 Wen-Hwa 1st Road, Kwei-Shan, Tao-Yuan, Taiwan e-mail: mechenl@mail.cgu.edu.tw *Methods* Participants were 78 women with breast cancer assigned to the exercise arm of a randomized control trial. Based on the weekly adherence rates in time and intensity, patients were classified as good (>100 %), acceptable (80–100 %), and poor (<80 %) adherents. Data were analyzed using ordinal logistic hierarchical linear modeling.

Results The trajectories for both time and intensity adherence declined significantly. The decline in exercise-time adherence was significantly slower in women who reported higher interest in exercise. Women with higher perceived importance of exercise, early disease stage, and employed were more likely to be classified as good intensity adherents. Poorer weekly adherence for both exercise time and intensity was associated with higher fatigue level for that week.

Conclusions Adherence to exercise adherence in breast cancer patients declined as the dose of exercise prescription increased. Factors influencing overall adherence and adherence trend were identified.

Keywords Adherence \cdot Home-based exercise \cdot Breast cancer \cdot Chemotherapy \cdot HLM

Introduction

For cancer patients undergoing treatment, exercise or physical activity has been shown to reduce symptoms such as fatigue [1], mood disorders [2], sleep disturbance [3], and physical function [4] and to improve quality of life [2]. Furthermore, women who exercised moderately after a breast cancer diagnosis were found to have a better survival rate than sedentary women [5]. A moderate-to-high intensity aerobic training is safe for breast cancer patients during chemotherapy [6].

Indeed, exercise has been shown in systematic reviews to benefit cancer patients during and after treatment [7–9].

Despite the benefits of exercise, many people do not adhere to recommended exercise prescriptions [10]. Among cancer survivors, the exercise adherence rate for both home-based and supervised exercise programs was 60 to 85 % [11]. Nonadherence to exercise interventions is a challenge for evaluating their efficacy [12–14].

Exercise adherence has been measured by two approaches: to determine overall performance or to average repeated measures of exercise-related behaviors over an exercise program. The most commonly used indicator of overall performance is the ratio of sessions attended/ completed to sessions expected/prescribed [13, 15–20]. Exercise-related behaviors include exercise frequency, exercise duration, pedometer steps, intensity, etc. [21, 22]. For both the overall and averaged approaches, exercise adherence is treated as a time-invariant variable. Since exercise training is often prescribed with progressively increasing doses in both intensity and time, exercise adherence may change during the training period. Furthermore, for cancer patients under treatment, adherence to exercise may fluctuate with treatment side effects.

Changes in patterns of exercise adherence and its predictors in breast cancer survivors have been examined in only a few studies [23, 24]. For example, breast cancer survivors' adherence to a 12-week home-based exercise program was assessed by three indicators: total weekly exercise (minutes/week), number of steps/week, and meeting a negotiated weekly goal [23]. Participants' first two outcomes significantly increased during the program, but the percentage of those meeting the negotiated goal was >80 % only in the first 4 weeks [23]. In the second study, the adherence of female breast cancer patients receiving chemotherapy to the prescribed regimen of 10, 000 steps/day was measured in two ways: total number of steps/week and mean number of steps/day (on days with any steps recorded) [24]. During the 12-month study, the adherence rate was lower in the first 6 weeks, but increased at 3, 6, and 9 months before slightly dropping at 12 months [24].

Adherence has been defined as the extent to which a person's behavior corresponds with the agreed recommendation [25]. Therefore, the absolute amount of exercise activity, either steps or exercise time, does not reflect the concept of adherence. Only percentage of actual exercise activity over an expected exercise activity can better represent the essence of adherence. To develop effective strategies to improve breast cancer patients' exercise adherence, those at risk for poor adherence must be identified. This study examined female breast cancer patients' adherence to a 12-week home-based exercise intervention while undergoing chemotherapy. Specifically, the study had two purposes: (1) to examine the trajectory of exercise adherence in terms of time and intensity and (2) to explore predictors of the exercise adherence trajectory, in terms of intercept and slope.

Materials and methods

Sample and setting

The study sample was from the intervention arm of a randomized control study examining the effect of homebased walking-exercise program on sleep disturbance and other symptoms for women with breast cancer receiving chemotherapy. Participants for the original study were recruited from women treated at Chang Gung Memorial Hospital in northern Taiwan who met the following criteria: (1) diagnosed with stage I–III breast cancer, (2) reporting insomnia, fatigue, pain, or depressive symptoms after the first cycle of chemotherapy, and (3) willing to be randomly assigned to an intervention or control group. Women were excluded if they had diagnosed sleep disorders, a psychiatric diagnosis, cardiovascular problems, muscle-skeletal problems, or conditions judged by their treating physicians as not suitable for exercise.

Among 492 patients assessed for eligibility in the original study, 221 were disqualified and 112 refused to participate. Of the remaining 159 patients, 81 were randomly allocated to the exercise group, but 3 dropped out before the intervention, leaving 78 who received the exercise intervention (Fig. 1). Among these 78 patients, 64 completed the 12-week exercise program and 14 dropped out for various reasons: not feeling well physically (n=8), unwilling to exercise (n=2), dislike of brisk walking (n=1), and unknown reasons (n=3). Those who withdrew and those who completed the 12-week program did not differ significantly at baseline on fatigue, performance status, and distance walked in 6 min. The sample for the current analysis included the 78 women who participated in the exercise intervention for at least 1 week. The study was approved by the hospital's Institutional Review Board, and all participants provided their written informed consents.

Exercise intervention

The exercise intervention was a 12-week home-based aerobicwalking program that followed ACSM guidelines of progressive increase of exercise prescription through three stages: initial, improvement, and maintenance stage [26]. Since the participants in this study were cancer patients under treatment, a moderate exercise intensity was set with 30–70 % heart rate reserved (HRR), which was 10–15 % lower than suggested by ACSM. The suggested exercise frequency was 3 times/week Fig. 1 Flow chart of subject recruitment



for the first 6 weeks and 5 times/week for the second 6 weeks. Exercise duration was also increased gradually, from 15 to 25 min/session for weeks 1 to 4, to 25–35 min/session for weeks 5 to 8, and 35–40 min/session for weeks 9 to 12 (Table 1).

All exercise instructions were provided by a coach with professional training on sports. The coach first instructed patients about the benefit of exercise followed by a demonstration of brisk walking. An exercise manual using both simple wording and pictures was used to assist the instruction. Content of this exercise manual includes purposes and potential benefits of exercise, the step-by-step procedure of the brisk walking with pictures, methods of monitoring heart-rate, indicators for stopping exercise, and contact information. A progressive weekly goal based on individual patient's age and rest heart rate was planned for

 Table 1
 Exercise frequency, intensity, and duration by stage

Stage		Week	Frequency (sessions/week)	Intensity (% of HRR)	Duration (minutes/session)
I. Initial	Ι	1-4	3	30–40	15–25
II. Improvement	IIa	5-6	3	40–50	25–35
	IIb	7	5	40–50	25–35
	IIc	8	5	50-60	25–35
III. Maintenance	III	9–12	5	60–70	35–40

HRR heart rate reserve

each participants. Participants were taught to record their walking time, frequency, duration, HR before and after walking, maximum HR during walking, exertion level and any injuries or adverse events on a diary. During the weekly follow-up phone call, the same coach would usually remind patient to do exercise and make sure if the goal for that week was reached. Patients would be asked if they had any difficulty adhering to the prescribed exercise dose and if they experienced any uncomfortable situation during the exercise. The instructor often provided positive reinforcement for those who reached the goal and encouragement for those who did not reach the goal. To ensure the feasibility and standardization of the instruction, the coach performed several mock instructions under the guidance of the second author.

Measures

The primary outcomes were exercise adherence rates on two elements: time and intensity. We conceptualized four types of predictors: demographics, disease/treatment characteristics, exercise behavior and attitude factors, and physical factors (fatigue and physical status).

Exercise time and intensity

Data of actual exercise time, heart rate during exercise were self-recorded by participants using a researcher-developed exercise diary. This diary included each participant's weekly prescription for exercise frequency, duration, and target heart rate. Participants monitored their heart rate using a FDA-approved sports pulse ring that detects heart rates of 30–250 beats/min. In this study, this sports pulse ring had an error <3 %.

Exercise behavior and attitudes

Data were collected at baseline on previous regular exercise habits (yes/no), degree of interest in exercise, and perceived importance of exercise using a researcherdeveloped questionnaire. Interest in exercise and perceived importance of exercise were measured on a 0 (not at all) to 10 (extremely interested/important).

Fatigue

Baseline fatigue was measured by the Brief Fatigue Inventory (BFI) [27]. The first part of the BFI assesses fatigue intensity at the time of assessment, as well as average and worst during the past 24 h. The second part of the BFI assesses fatigue interference with six aspects of life. Both intensity and interference items are measured on a 0 (no fatigue/interference) to 10 (worst fatigue imaginable/extreme interference) scale. The BFI has demonstrated satisfactory validity and reliability in cancer patients [27]. Average fatigue intensity was used in the current study. In each exercise day during the 12-week intervention, patients recorded their fatigue level on a 0 (not at all) to 10 (extremely) scale in a researcher-developed exercise diary. Average weekly fatigue was used in the analysis.

Physical status

Physical status was assessed by performance on two measures: the Karnofsky Performance Scale (KPS) [20] and the six-minute walking test (6MWT) [28]. The KPS score ranges from 0 % (dead) to 100 % (normal, no complaints; no evidence of disease). The 6WMT measures the distance (in meters) that the subject can walk in 6 min. Both methods have been widely used in cancer patients with satisfactory reliability and validity [20, 28, 29].

Demographic and disease/treatment characteristics

Demographic (age, marital status, educational level, religious affiliation, menopause status, and employment status) and disease/treatment (disease stage, surgery type, chemotherapy regimen, and other adjuvant therapy) characteristics were collected from patients or medical records using a researcher-developed questionnaire.

Procedure

At the second chemotherapy cycle, eligible and consented patients were assessed for baseline data. At the third chemotherapy cycle, the exercise coach instructed patients how to perform their home-based walking and how to use the pulse ring to monitor heart rate during exercise. The exercise coach called patients weekly to encourage adherence to the exercise program and to help them deal with exercise-related problems, if any. Patients were asked to record exercise-related information in the exercise diary throughout the 12-week exercise intervention period.

Statistical analysis

Exercise adherence was evaluated by two elements of the exercise prescription: time and intensity. The weekly total exercise time was obtained from the diary information of exercise duration/session and frequency/week. Adherence to the prescribed exercise time was defined by the ratio of actual exercise time/week to the prescribed minimal exercise time/week. Adherence to exercise intensity was defined by the ratio of the highest heart rate during exercise to the target heart rate. The target heart rate for each week

was calculated for each patient based on their age using the following equation:

$$HR_{target} = ([HR_{max} - HR_{rest}] \times percent intensity) + HR_{rest}$$

where

HR _{target}	target heart rate
HR _{max}	maximal heart rate=200-age in years
HR _{rest}	heart rate at rest
HR _{max} -HR _{rest}	heart rate reserve (HRR)
Percent intensity	30 to 70 %

The obtained adherence ratios (rates) for time and intensity were further categorized into three ordinal groups: "good adherence" (>100 %), "acceptable adherence" (80–100 %), and "poor adherence" (<80 %).

The trajectory of exercise adherence and its predictors was examined by ordinal logistic hierarchical linear modeling (HLM). In this study, repeated measures of adherence were considered as being nested within individuals. Therefore, we used a multilevel analysis. Level I data contained information of within-person changes over time, while level II data comprised information about individual characteristics that could be used to predict the parameters estimated from level I. The analytic process was divided into three steps. First, an unconditional model was specified to determine the pattern of change in adherence over time. Second, univariate analysis was used to explore potential factors predicting adherence. The predictors explored in this step included demographic variables, disease and treatment variables, exercise behavior and attitude variables, and physical variables including fatigue. Third, significant factors identified were entered into multivariate analysis and final models with significant predictors of adherence, for both time and intensity, were formed. In these analyses, the weekly time- and intensity-adherence categories were time-variant dependent variables. Time (weeks) and weekly mean fatigue severity were considered level-I time-variant predictors. Personal and disease/treatment characteristics, exercise behavior and attitude, and physical variables were level-II time-invariant variables. For each week after the first week, patients with missing adherence data were compared to patients with valid data for their previous week's adherence rates to determine whether the data were missing at random.

Results

Sample characteristics

Participants' mean age was 48.27 years (SD=8.03). Most of the women were married (85.9 %) and were diagnosed

Variable	Mean (SD)	n (%)
Age (years)	48.27 (8.03)	
Height (cm)	155.58 (4.70)	
Weight (kg)	57.43 (10.66)	
Interest in exercise $(0-10)^a$	5.83 (2.65)	
Perceived importance of	8.92 (1.68)	
Baseline 6 min-walking	434.1 (78.76)	
Karnofsky performance	91.13 (4.23)	
Fatigue severity	3.90 (2.22)	
(BFI score, 0–10)		
Regular exercise before		
diagnosis Ves		35 (44 9)
No		43 (55 1)
Provious exercise frequency		45 (55.1)
(days/week) (n=35)		2(57)
≥ 2		2(3.7)
3-4		10 (28.6)
_⊃ Marital status		25 (55.7)
Marian status		0(115)
Never married		9 (11.5)
Diversed/widewed		67 (83.9) 2 (2.6)
Divorced/widowed		2 (2.6)
Vee		22(42.2)
Yes		33 (42.3)
N0		45 (57.7)
Religion		
None		22 (28.2)
Buddhism		38 (48.7)
Christianity/Catholicism		3 (3.9)
Taoism		15 (19.2)
Employment		
None		20 (25.6)
Full time		55 (70.5)
Part time		3 (3.8)
Working status after diagnosis		
Same		39 (50.0)
Shift to part time		3 (3.8)
Quit		19 (24.4)
Leave job temporarily		12 (15.4)
Other		5 (6.4)
Education level		
≤Elementary school		19 (24.4)
Junior high school		11 (14.1)
Senior high school		32 (41.0)
≥College		16 (20.6)
AJCC disease stage		
Ι		31 (39.8)
IIa		28 (36.4)

Table 2 (continued)

Variable	Mean (SD)	n (%)
IIb		9 (11.7)
IIIa		5 (6.5)
IIIc		5 (6.5)
Surgery type		
Mastectomy		34 (43.6)
Breast-conserving surger	ry	44 (56.4)
Lymph node dissection		
Yes		41 (52.6)
No		37 (47.4)
Comorbidity		
Yes		30 (38.5)
No		48 (61.5)
Chemotherapy regimen		
Cyclophosphamide, methotrexate, fluoroura	acil	21 (26.9)
CEF		26 (33.3)
CEF + taxane		30 (38.5)
Taxane only		1 (1.3)
Radiotherapy		
Yes		36 (46.2)
No		42 (53.8)
Hormone therapy		
Yes		48 (61.5)
No		30 (38.5)

BFI Brief Fatigue Inventory, *AJCC* American Joint Committee on Cancer, *CEF* cyclophosphamide, epirubicin, 5-fluorouracil

^aNumbers in parentheses indicate possible range of scores

with stage I or IIa breast cancer (76.2 %). The average scores of interest in exercise and perceived importance of exercise were5.83 (SD=2.65) and 8.92 (SD=1.68), respectively. Detailed information regarding cancer treatment and physical functioning are presented in Table 2. No injury due to exercise was reported in the period of exercise intervention. Some adverse events occurred with low incidence, such as cold sweating (0.07 %), palpitation (0.07 %), dizziness (0.34 %), and shortness of breath (0.14 %).

Exercise-time adherence

The mean weekly total exercise time was 185.91 min, with the longest time in weeks 5 and 9 (206.58 and 207.21 min, respectively) and the shortest in week 1 and 4 (149.83 and 149.85 min, respectively) (Table 3). The mean time-adherence rate was 87.1 %, with the highest adherence in week 3 (99.4 %) and the lowest in week 11 (74.0 %). The proportion of good adherents (adherence rate>100 %) decreased over the exercise program (mean=69.0 %, range=

40.8–92.3). In the last stage of the program (week 9–12), only 40.8–50.0 % of participants fully adhered to the prescribed exercise time (Fig. 2). The missing of exercise-time adherence was not associated with the time adherence rate at the previous week.

Exercise-intensity adherence

Participants' exercise intensity was represented by the highest HR achieved during exercise. The mean highest HR during exercise was 128.81 beats/min (range=78–186). Overall, the weekly highest HR during exercise showed an increasing trend, except for a slight drop in week 4 (Table 3). The overall mean intensity-adherence rate was high, 97.59 % (range=95.14–99.18). The proportion of good adherents (adherence rate>100 %) decreased over the exercise program (mean=73.1 %, range=45.8–94.2). The proportion of good adherents decreased notably after week 8, although most participants still had an 80 % adherence rate (Fig. 3). The missing of exercise-intensity adherence rate at the previous week.

Trajectory and predictors of adherence

Changes in adherence over time and factors predicting adherence were explored using ordinal adherence categories (poor, acceptable, and good) of time and intensity as the dependent variables. For time adherence, unconditional HLM results indicated that adherence declined as the exercise program progressed (coefficient=-0.342, p < 0.001) (Table 4). Over time, the probability of being categorized as a good adherent decreased (odds=0.710, 95 % CI=0.662-0.762). However, the probability of this adherence declining with time was shown to be lower in multivariate analysis for patients with higher perceived interest in exercise (odds=1.038, 95 % CI=1.006-1.072). In addition, the probability of being categorized as a good adherent was lower in the corresponding week that patients had higher weekly fatigue severity (odds= 0.770, 95 % CI=0.673-0.882) (Table 4). In other words, the weekly exercise-time adherence fluctuated with patients' level of fatigue in that week.

Intensity adherence was also shown by the unconditional model to decline as the program progressed (coefficient=-0.360, p < 0.001). Overtime, the probability of being categorized as a good adherent decreased (odds= 0.698, 95 % CI=0.627-0.776) (Table 5). The probability of being categorized as a good adherent was greater for patients who attached higher importance to exercise (odds=1.593, 95 % CI=1.013-2.504), were employed (odds=6.103, 95 % CI=1.037-35.925), and with early-stage disease (odds=0.426, 95 % CI=0.270-0.788).

 Table 3
 Weekly exercise time and intensity and corresponding adherence rate over 12 weeks

	Exercise	time										
Week (n)	1 (48)	2 (52)	3 (53)	4 (46)	5 (55)	6 (56)	7 (49)	8 (50)	9 (53)	10 (51)	11 (49)	12 (52)
Exercise time (min) ^a												
Mean (SD)	149.83 (105.82)	188.02 (116.08)	184.92 (105.07)	149.85 (118.62)	206.58 (164.86)	190.54 (134.59)	184.08 (115.11)	182.28 (126.88)	207.21 (158.3)	199.24 (136.16)	185.02 (139.72)	203.37 (122.80)
Time adherence (%)												
Mean (SD)	97.82 (9.90)	97.01 (11.67)	99.37 (4.58)	92.08 (18.61)	91.42 (21.87)	96.07 (11.75)	82.68 (27.35)	81.94 (25.52)	75.68 (30.67)	76.57 (28.46)	74.02 (30.55)	80.00 (26.41)
	Exercise	intensity										
Week (<i>n</i>) Highest HR ^b	1 (47)	2 (51)	3 (52)	4 (46)	5 (54)	6 (50)	7 (47)	8 (49)	9 (48)	10 (47)	11 (45)	12 (48)
Mean (SD)	126.73 (18.63)	127.42 (15.93)	129.22 (15.83)	124.75 (15.43)	126.40 (14.66)	128.37 (14.55)	127.98 (12.27)	127.90 (12.70)	131.79 (12.68)	131.45 (11.36)	131.83 (10.10)	131.83 (12.38)
Intensity adherence (%)											
Mean (SD)	98.53 (4.52)	98.89 (3.68)	98.90 (4.08)	98.81 (3.63)	97.67 (5.27)	98.72 (4.15)	99.18 (1.94)	97.16 (4.13)	95.14 (6.25)	95.79 (4.78)	96.44 (4.29)	95.86 (5.54)

^a Exercise time = the weekly total exercise minutes

^b Highest HR = the highest heart rate during exercise

Additionally, the probability of being categorized as a good adherent was lower in the corresponding week that a patient had higher weekly fatigue (odds=0.836, 95 % CI=0.710-0.983) (Table 5). Thus, weekly exercise-intensity adherence also fluctuated with patients' level of fatigue that week.

Discussion

This study found that two aspects of exercise adherence (i.e., time and intensity) to a home-based walking-exercise program decreased over the 12-week program. Predictors of time adherence were fatigue and interest in exercise, and predictors of



Fig. 2 Percentage of good, acceptable, and poor time adherence over 12 weeks

Fig. 3 Percentage of good, acceptable, and poor intensity adherence over 12 weeks



Table 4 Models of time adherence

Unconditional model					
Fixed effect	Coefficient	Odds ratio	95.0 % con Lower	fidence interval Upper	р
For intercept					
Intercept	3.266	26.218	14.158	48.553	< 0.001
For week slope, B1					
Intercept	-0.342	0.710	0.662	0.762	< 0.001
For threshold					
<i>d</i> (2)	0.614	1.848	1.588	2.151	< 0.001
Multivariate prediction m	odel				
Fixed effect	Coefficient	Odds ratio	95.0 % con	fidence interval	р
			Lower	Upper	
For intercept					
Intercept	4.244	69.690	1.669	2909.777	0.027
Education (years)	0.119	1.127	0.931	1.364	0.217
Exercise preference	-0.146	0.864	0.645	1.158	0.323
Employment	-1.359	0.257	0.043	1.524	0.132
Disease stage	0.022	1.023	0.636	1.644	0.925
Type of surgery	1.361	3.899	0.775	19.622	0.097
For week slope					
Intercept	-0.426	0.653	0.441	0.968	0.034
Education (years)	-0.011	0.990	0.970	1.010	0.300
Interest in exercise	0.037	1.038	1.006	1.072	0.023
Employment	0.066	1.068	0.860	1.327	0.543
Disease stage	-0.026	0.974	0.930	1.020	0.261
Type of surgery	-0.109	0.897	0.743	1.083	0.253
For weekly fatigue severi	ty				
Intercept	-0.261	0.770	0.673	0.882	0.000
For threshold					
<i>d</i> (2)	0.695	2.003	1.653	2.427	0.000

Table 5 Models of intensity adherence

Unconditional model					
Fixed effect	Coefficient	Odds ratio	95.0 % cont Lower	idence interval Upper	р
For intercept					
Intercept	3.998	54.515	22.001	135.079	< 0.001
For week slope, B1					
Intercept	-0.360	0.698	0.627	0.776	< 0.001
For threshold					
<i>d</i> (2)	4.727	112.915	36.039	353.778	< 0.001
Multivariate prediction me	odel				
Fixed effect	Coefficient	Odds ratio	95.0 % conf	idence interval	р
			Lower	Upper	
For intercept					
Intercept	1.746	5.732	0.241	136.340	0.275
Exercise importance	0.465	1.593	1.013	2.504	0.044
Employment	1.809	6.103	1.037	35.925	0.046
Disease stage	-0.854	0.426	0.230	0.788	0.008
For week slope					
Intercept	-0.354	0.702	0.454	1.085	0.109
Exercise importance	-0.012	0.988	0.938	1.041	0.651
Employment	-0.189	0.828	0.654	1.048	0.115
Disease stage	0.058	1.060	0.994	1.129	0.073
For weekly fatigue severit	y				
Intercept	-0.179	0.836	0.710	0.983	0.031
For threshold					
<i>d</i> (2)	5.165	175.123	43.595	703.475	0.000

intensity adherence were fatigue, perceived importance of exercise, employment status, and disease stage.

The dropout rate in this study (18 %) was higher than the 8.8 to 10.3 % rates reported in previous studies of exercise adherence with cancer patients [6, 30-32]. These differences may be due to exercise mode (supervised exercise) [6, 30], treatment phase (treatment completed) [31], and combining dietary consultation with exercise intervention and more frequent telephone reminders [32]. Supervised exercise often has a higher adherence rate than home-based exercise [33]. Treatment-related side effects may prevent patients from engaging in physical activity. Meta-analysis has shown that the effect of exercise in reducing fatigue was only significant for patients who completed or who were completing adjuvant treatment [8].

The decreasing exercise-time and exercise-intensity adherence reflects the exercise prescription, which gradually increased the frequency, duration, and percent of HRR over the 12-week program. Our results are consistent with previous reports that adherence decreased for sedentary adults [34] and breast cancer survivors [23] as prescription intensity increased. A higher requirement of exercise time was also found to be associated with a lower adherence [14]. The simultaneous increase of exercise intensity, frequency, and duration

2069

along the exercise program may have an additive effect in decreasing exercise adherence. Simultaneous increase of different aspects of exercise prescription has been used in newly diagnosed breast cancer patients with low non-adherence rate [35]. However, no detailed longitudinal adherence data were reported in that study. Future study may consider increase one exercise parameter at a time. Since the majority of our participants had good adherence during the first stage of the 12week exercise program, the initial prescription for exercise time and HRR percentage may have been too weak and could have been stronger. In contrast, the decreased proportions of good adherents during the last stage of the exercise program suggest that the prescription was too strong for patients under chemotherapy.

We also found that changes in weekly time adherence and intensity adherence were associated with weekly fatigue. Participants who felt tired may not have been motivated to exercise or may have exercised less frequently for shorter durations and at lower intensity, as previously reported for patients receiving cancer treatment [36-38]. On the other hand, those with poor adherence may have experienced greater fatigue due to not exercising. The exercise intervention in the original trial was designed to reduce cancer-related symptoms, including fatigue. It is reasonable to hypothesize that participants with good exercise adherence experienced more benefit in symptom relief, in this case reduced fatigue. The causal relationship between fatigue level and exercise adherence needs to be further studied.

Exercise-time and exercise-intensity adherence were significantly predicted by participants' degree of interest in exercise and perceived importance of exercise, respectively. This finding is consistent with the theory of planned behavior where personal attitudes (e.g., interest in exercise) and subjective norm (e.g., perceived importance of exercise) toward a specific behavior (i.e., exercise adherence) influence the actual performance of that behavior [39]. An interesting note is that employment status was associated with better intensity adherence, but not with time adherence. One possible explanation is that employed patients may be in better physical condition, allowing them to have better adherence to the intensity requirement. However, employed patients may lack enough time to exercise at the required frequency or duration [13]. Disease stage was also found to predict intensity adherence, but not time adherence. Patients with early-stage disease may have better physical functioning, allowing them to have better intensity adherence. However, patients with better physical functioning do not necessarily have enough interest in exercise which, as shown in this study, is an important factor for time adherence.

Clinical implications

Our longitudinal adherence data can inform the design of future exercise programs for breast cancer patients undergoing chemotherapy. Specifically, the prescribed exercise time and intensity may be elevated during the initial stage and reduced during the last stage of the exercise program.

Limitations

First, our participants were breast cancer patients with selfreported insomnia, fatigue, pain, or depressive symptoms after the first chemotherapy cycle and sampled from a single hospital in Taiwan. This limitation prevents the generalizability of our findings to all women with breast cancer in Taiwan. Second, potential factors predicting adherence, such as change stage of behavior and self-efficacy, were not measured. Third, the psychometric properties of the single-item measurement of exercise belief and attitude need to be tested in the future study. Finally, non-adherence does not always represent performing less than the prescribed exercise; over-exercise can be a mode of non-adherence [40]. Future studies should sample breast cancer patients from hospitals throughout Taiwan, examine other potential predictors of exercise adherence, and over-exercise.

Conclusions

Both exercise-time and exercise-intensity adherence decreased as the prescription requirements increased. Poor exercise-time and exercise-intensity adherence was associated with chemotherapy-related fatigue. Greater exercise-intensity adherence was more likely in employed patients with earlystage disease. Exercise adherence was predicted by attitudes toward exercise.

Acknowledgments This study was supported by a research grant from the National Science Council of Taiwan to Dr. Mei-Ling Chen (grant number: NSC 97-2314-B-182-035-MY3)

Conflict of interest There is no financial relationship to be claimed in this study. The authors have full control of all primary data and they agree to allow the journal to review their data if requested.

Ethical approval The study was approved by the Institutional Review Board of Chang Gung Memorial Hospital.

References

- Mock V, Frangakis C, Davidson NE, Ropka ME, Pickett M, Poniatowski B, Stewart KJ, Cameron L, Zawacki K, Podewils LJ, Cohen G, McCorkle R (2005) Exercise manages fatigue during breast cancer treatment: a randomized controlled trial. Psychooncology 14(6):464–477. doi:10.1002/pon.863
- Mock V, Pickett M, Ropka ME, Muscari Lin E, Stewart KJ, Rhodes VA, McDaniel R, Grimm PM, Krumm S, McCorkle R (2001) Fatigue and quality of life outcomes of exercise during cancer treatment. Cancer Pract 9(3):119–127
- Payne JK, Held J, Thorpe J, Shaw H (2008) Effect of exercise on biomarkers, fatigue, sleep disturbances, and depressive symptoms in older women with breast cancer receiving hormonal therapy. Oncol Nurs Forum 35(4):635–642
- Markes M, Brockow T, Resch KL (2006) Exercise for women receiving adjuvant therapy for breast cancer. Cochrane Database Systematic Reviews (Online) (4):CD005001. doi:10.1002/ 14651858.CD005001.pub2
- Holmes MD, Chen WY, Feskanich D, Kroenke CH, Colditz GA (2005) Physical activity and survival after breast cancer diagnosis. JAMA 293(20):2479–2486. doi:10.1001/jama.293.20.2479
- 6. Hornsby WE, Douglas PS, West MJ, Kenjale AA, Lane AR, Schwitzer ER, Ray KA, Herndon JE 2nd, Coan A, Gutierrez A, Hornsby KP, Hamilton E, Wilke LG, Kimmick GG, Peppercorn JM, Jones LW (2014) Safety and efficacy of aerobic training in operable breast cancer patients receiving neoadjuvant chemotherapy: a phase II randomized trial. Acta Oncol 53(1):65–74. doi:10.3109/ 0284186X.2013.781673
- Pinto BM, Floyd A (2007) Methodologic issues in exercise intervention research in oncology. Semin Oncol Nurs 23(4):297–304
- McNeely ML, Campbell KL, Rowe BH, Klassen TP, Mackey JR, Courneya KS (2006) Effects of exercise on breast cancer patients and survivors: a systematic review and meta-analysis. Can Med Assoc J 175(1):34–41
- Knols R, Aaronson NK, Uebelhart D, Fransen J, Aufdemkampe G (2005) Physical exercise in cancer patients during and after medical treatment: a systematic review of randomized and controlled clinical trials. J Clin Oncol 23(16):3830–3842

- Martin KA, Sinden AR (2001) Who will stay and who will go? A review of older adults' adherence to randomized controlled trials of exercise. J Aging Phys Act 2(2):91–114
- Courneya KS, Segal RJ, Reid RD, Jones LW, Malone SC, Venner PM, Parliament MB, Scott CG, Quinney HA, Wells GA (2004) Three independent factors predicted adherence in a randomized controlled trial of resistance exercise training among prostate cancer survivors. J Clin Epidemiol 57(6):571–579. doi:10.1016/j.jclinepi.2003.11. 010S0895435603004293
- Pickett M, Mock V, Ropka ME, Cameron L, Coleman M, Podewils L, Pickett M, Mock V, Ropka ME, Cameron L, Coleman M, Podewils L (2002) Adherence to moderate-intensity exercise during breast cancer therapy. Cancer Pract 10(6):284–292
- Courneya KS, Friedenreich CM, Quinney HA, Fields AL, Jones LW, Fairey AS (2004) Predictors of adherence and contamination in a randomized trial of exercise in colorectal cancer survivors. Psychooncology 13(12):857–866. doi:10.1002/pon.802
- Husebo AM, Dyrstad SM, Mjaaland I, Soreide JA, Bru E (2014) Effects of scheduled exercise on cancer-related fatigue in women with early breast cancer. Sci World J 2014:271828. doi:10.1155/ 2014/271828
- Courneya KS, Blanchard CM, Laing DM (2001) Exercise adherence in breast cancer survivors training for a dragon boat race competition: a preliminary investigation. Psychooncology 10(5):444–452. doi:10. 1002/pon.524
- Courneya KS, Segal RJ, Gelmon K, Reid RD, Mackey JR, Friedenreich CM, Proulx C, Lane K, Ladha AB, Vallance JK, McKenzie DC (2008) Predictors of supervised exercise adherence during breast cancer chemotherapy. Med Sci Sports Exerc 40(6): 1180–1187. doi:10.1249/MSS.0b013e318168da45
- Kim CJ, Kang DH, Smith BA, Landers KA (2006) Cardiopulmonary responses and adherence to exercise in women newly diagnosed with breast cancer undergoing adjuvant therapy. Cancer Nurs 29(2):156– 165
- Milne HM, Wallman KE, Gordon S, Courneya KS (2008) Effects of a combined aerobic and resistance exercise program in breast cancer survivors: a randomized controlled trial. Breast Cancer Res Treat 108(2):279–288
- Peddle CJ, Jones LW, Eves ND, Reiman T, Sellar CM, Winton T, Courneya KS (2009) Correlates of adherence to supervised exercise in patients awaiting surgical removal of malignant lung lesions: results of a pilot study. Oncol Nurs Forum 36(3):287–295. doi:10. 1188/09.ONF.287-295
- Schag CC, Heinrich RL, Ganz PA (1984) Karnofsky performance status revisited: reliability, validity, and guidelines. J Clin Oncol 2(3): 187–193
- Daley AJ, Crank H, Mutrie N, Saxton JM, Coleman R (2007) Determinants of adherence to exercise in women treated for breast cancer. Eur J Oncol Nurs 11(5):392–399
- 22. Latka RN, Alvarez-Reeves M, Cadmus L, Irwin ML (2009) Adherence to a randomized controlled trial of aerobic exercise in breast cancer survivors: the Yale exercise and survivorship study. J Cancer Surviv 3(3):148–157. doi:10.1007/s11764-009-0088-z
- Pinto BM, Rabin C, Dunsiger S (2009) Home-based exercise among cancer survivors: adherence and its predictors. Psychooncology 18(4):369–376. doi:10.1002/pon.1465
- Swenson KK, Nissen MJ, Henly SJ (2010) Physical activity in women receiving chemotherapy for breast cancer: adherence to a walking intervention. Oncol Nurs Forum 37(3):321–330. doi:10. 1188/10.ONF.321-330
- WHO (2003) Adherence to long-term therapies. Evidence for Action. http://apps.who.int/medicinedocs/en/d/Js4883e/6.html#Js4883e.6.1.1

- ACSM (2006) ACSM's Guidelines for Exercise Testing and Prescription In: Medicine ACoS (ed). 7th edn. Lippincott William & Wilkins, Philadelphia, pp 149–151
- Mendoza TR, Wang XS, Cleeland CS, Morrissey M, Johnson BA, Wendt JK, Huber SL (1999) The rapid assessment of fatigue severity in cancer patients: use of the Brief Fatigue Inventory. Cancer 85(5): 1186–1196. doi:10.1002/(SICI)1097-0142(19990301) 85:5<1186::AID-CNCR24>3.0.CO;2-N
- Kervio G, Carre F, Ville NS (2003) Reliability and intensity of the six-minute walk test in healthy elderly subjects. Med Sci Sports Exerc 35(1):169–174. doi:10.1249/01.MSS.0000043545. 02712.A7
- Yates JW, Chalmer B, McKegney FP (1980) Evaluation of patients with advanced cancer using the Karnofsky performance status. Cancer 45(8):2220–2224
- Andersen C, Adamsen L, Moeller T, Midtgaard J, Quist M, Tveteraas A, Rorth M (2006) The effect of a multidimensional exercise programme on symptoms and side-effects in cancer patients undergoing chemotherapy—the use of semi-structured diaries. Eur J Oncol Nurs 10(4):247–262. doi:10.1016/j.ejon.2005.12.007
- Vallance JK, Courneya KS, Plotnikoff RC, Yasui Y, Mackey JR (2007) Randomized controlled trial of the effects of print materials and step pedometers on physical activity and quality of life in breast cancer survivors. J Clin Oncol 25(17):2352–2359
- 32. Demark-Wahnefried W, Case LD, Blackwell K, Marcom PK, Kraus W, Aziz N, Snyder DC, Giguere JK, Shaw E (2008) Results of a diet/exercise feasibility trial to prevent adverse body composition change in breast cancer patients on adjuvant chemotherapy. Clin Breast Cancer 8(1):70–79
- 33. Irwin ML, Cadmus L, Alvarez-Reeves M, O'Neil M, Mierzejewski E, Latka R, Yu H, Dipietro L, Jones B, Knobf MT, Chung GG, Mayne ST (2008) Recruiting and retaining breast cancer survivors into a randomized controlled exercise trial: the Yale Exercise and Survivorship Study. Cancer 112(11 Suppl):2593–2606. doi:10.1002/cncr.23446
- 34. Perri MG, Anton SD, Durning PE, Ketterson TU, Sydeman SJ, Berlant NE, Kanasky WF Jr, Newton RL Jr, Limacher MC, Martin AD (2002) Adherence to exercise prescriptions: effects of prescribing moderate versus higher levels of intensity and frequency. Health Psychol 21(5):452–458
- 35. Wang YJ, Boehmke M, Wu YW, Dickerson SS, Fisher N (2011) Effects of a 6-week walking program on Taiwanese women newly diagnosed with early-stage breast cancer. Cancer Nurs 34(2):E1–E13. doi:10.1097/NCC.0b013e3181e4588d
- 36. Courneya KS, McKenzie DC, Reid RD, MacKey JR, Gelmon K, Friedenreich CM, Ladha AB, Proulx C, Lane K, Vallance JK, Segal RJ (2008) Barriers to supervised exercise training in a randomized controlled trial of breast cancer patients receiving chemotherapy. Ann Behav Med 35(1):116–122
- 37. Shang J, Wenzel J, Krumm S, Griffith K, Stewart K (2012) Who will drop out and who will drop in: exercise adherence in a randomized clinical trial among patients receiving active cancer treatment. Cancer Nurs 35(4):312–322
- Schwartz AL (2000) Daily fatigue patterns and effect of exercise in women with breast cancer. Cancer Pract 8(1):16–24
- 39. Ajzen I (2011) The theory of planned behaviour: reactions and reflections. Psychol Health 26(9):1113–1127. doi:10.1080/ 08870446.2011.613995
- Martin KA, Bowen DJ, Dunbar-Jacob J, Perri MG (2000) Who will adhere? Key issues in the study and prediction of adherence in randomized controlled trials. Control Clin Trials 21(5 Suppl): 195S–199S