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Daily Physical Activity and Hot Flashes in the Study of Women's Health Across the Nation FLASHES Study

Carolyn Gibson, MPH, MS^{1,*}, Karen Matthews, PhD², and Rebecca Thurston, PhD²

¹Department of Psychology, University of Pittsburgh

²Departments of Psychiatry, Epidemiology, and Psychology, University of Pittsburgh

Abstract

Objective—To examine the role of physical activity in menopausal hot flashes. Competing models conceptualize physical activity as a risk or protective factor for hot flashes. Few studies have examined this relationship prospectively using physiologic measures of hot flashes and physical activity.

Design—Over two 48 hour-periods, 51 participants wore a physiologic hot flash monitor and activity monitor, and reported their hot flashes in an electronic diary. Physiologic hot flashes, reported hot flashes and reported hot flashes without physiological corroboration were related to activity changes using hierarchical generalized linear modeling, adjusting for potential confounders.

Setting—Community.

Patients—Midlife women.

Interventions—None.

Main Outcome Measures—Physiologically-detected hot flashes and reported hot flashes with and without physiologic corroboration.

Results—Hot flash reports without physiologic corroboration were more likely after activity increases (OR 1.04, 95% CI: 1.00-1.10, $p=.01$), particularly among women with higher levels of depressive symptoms (interaction $p=.02$). No other types of hot flashes were related to physical activity.

Conclusion—Acute increases in physical activity were associated with increased reporting of hot flashes lacking physiologic corroboration, particularly among women with depressive symptoms. Clinicians should consider the role of symptom perception and reporting in relations between physical activity and hot flashes.

Keywords

Menopause; hot flashes; physical activity

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*Corresponding author: Carolyn Gibson, MPH, MS, 201 N. Craig Street, Suite 200, Room 202, Pittsburgh, PA 15214, Phone: 412-648-9697, gibsoncj@upmc.edu.

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Introduction

An estimated 70 to 80% of women report hot flashes during the menopausal transition (1). Despite the common occurrence of these symptoms, the physiology of hot flashes is only partially understood. The role of physical activity as an acute trigger of hot flashes has been anecdotally reported, but there has been little rigorous investigation of this relationship. The current literature is mixed, with findings alternately suggesting positive, negative, and null relationships between physical activity and hot flashes (2). Differences in results may stem in part from varied assessments of both physical activity and hot flashes.

Most studies of physical activity and hot flashes have examined self-reported hot flashes in relation to habitual physical activity, with the hypothesis that higher levels of habitual physical activity and cardiorespiratory fitness may be protective against hot flashes (3). While some studies have found that women who report being physically active (4,5) or increase their habitual physical activity levels through exercise interventions (6,7), are less likely to report severe hot flashes than their more sedentary counterparts, most have found no association between reported physical activity levels (8–14) or increasing habitual physical activity over time (15–18) and self-reported hot flashes. Further research using objective measures of activity and fitness is needed to clarify the potentially protective role of habitual physical activity and fitness against hot flashes.

While habitual physical activity is posited to be protective, acute bouts of physical activity are anecdotally reported as a trigger of hot flashes. This hypothesized relationship aligns with theories of hot flash physiology that frame hot flashes as an acute heat dissipation event occurring in the context of a narrowed thermoneutral zone (19). According to this model, acute physical activity would trigger hot flashes by increasing core body temperature, leading to the heat dissipation event of a hot flash. However, few studies have rigorously examined the acute effects of physical activity on hot flash occurrence. Freedman (19) found that hot flashes physiologically-detected by a skin conductance monitor were more likely to occur following a laboratory-based exercise protocol, and Thurston et al. (20) found an increased likelihood of physiologically-detected hot flashes following diary-reported physical exertion. In contrast, other studies found that increases in aerobic exercise were associated with a decrease in reported hot flashes over the subsequent minutes or day (3,8). Some of this work suggests that relationships between hot flashes and physical activity differ by method of hot flash assessment, and that weight status and fitness level may modify these associations (3). Thus, the role of acute physical activity as a trigger for hot flashes remains to be clarified, and may vary by factors such as participant characteristics and the method of hot flash assessment utilized.

Differences in methods of hot flash assessment contribute to the challenge of interpreting this limited literature. Most studies of hot flashes utilize self-report measures, which may be subject to error, retrospective reporting bias, and the influence of current affective states on symptom perception and reporting (21). Methods of physiologically detecting hot flashes have rarely been used in the literature examining physical activity and hot flashes, though it is known that correlates of these and self-reported hot flashes differ (20). Associations between acute physical activity and self-reported hot flashes may be more indicative of a role of physical activity in symptom perception and interpretation. Alternatively, associations between acute physical activity and physiologically-detected hot flashes may point to a physiological mechanism linking physical activity and hot flashes, such as increased core body temperature coupled with the hypothalamic thermoregulatory dysregulation.

In the current study, we sought to examine acute changes in daily physical activity as a trigger of self-reported and physiologically-detected hot flashes. We expected to find that increases in physical activity would be more likely prior to hot flashes compared to control times. Differences in this relationship by type of hot flash (e.g., self-reported, physiologic) were also explored. Given the important relationships between negative affect and both hot flash reporting (21) and physical activity (22), we also tested whether associations between acute physical activity and hot flashes would vary by baseline levels of negative affect and habitual physical activity.

Materials and Methods

Participants

The study sample was a subcohort of participants (N = 52) of the Pittsburgh site of the Study of Women's Health across the Nation (SWAN). SWAN is a cohort study designed to characterize the menopausal transition, conducted at seven sites across the United States. Details of SWAN procedures have been reported previously (23). At enrollment (1996-1997), SWAN participants (N = 3,302) were aged 42 to 52 years, had an intact uterus and at least one ovary, were not pregnant or breast feeding, had menstruated within 3 months, and were not using oral contraceptives or hormone therapy.

A subcohort of participants at the Pittsburgh SWAN site participated in SWAN FLASHES, an ancillary study using physiologic measures of hot flashes. SWAN FLASHES assessments occurred from 2008 to 2009, most closely corresponding to SWAN's 10th annual visit. By design, the Pittsburgh site recruited only Caucasian and African-American women. SWAN FLASHES inclusion criteria included reporting any hot flashes or night sweats in the past 2 weeks, having a uterus and both ovaries, not being pregnant, not using hormone therapy or selective serotonin reuptake inhibitors/serotonin norepinephrine uptake inhibitors for 3 months, and not currently undergoing chemotherapy. SWAN FLASHES enrolled 52 women. One woman was excluded from analysis because of missing sternal skin conductance data, for a final sample of 51 women (25 African American, 26 Caucasian women). Procedures were approved by the University of Pittsburgh Institutional Review Board. Participants provided written informed consent.

Design and procedures

Participants in SWAN FLASHES wore physiological hot flash monitors and carried an electronic hot flash diary. The monitoring period totaled approximately 96 hours, conducted in the context of two separate 48-hour ambulatory monitoring sessions within approximately 4 weeks. Participants were instructed to continue their regular routines, but to refrain from vigorous exercise while wearing the monitoring equipment. They also completed questionnaires and provided demographic and anthropometric data during a laboratory visit.

Hot Flashes—Both physiologically-detected and self-reported hot flashes were assessed. Physiologic hot flashes were detected with Biolog sternal skin conductance monitors (model 3991/2-SCL; UFI, Morro Bay, CA). The Biolog measures sternal skin conductance sampled at 1 Hz from the sternum via a 0.5-volt constant voltage circuit passed between two Ag/AgCl electrodes (UFI) filled with 0.05 M KCL Velvachol/glycol paste. Physiologic hot flashes were classified via standard methods, with skin conductance rise of 2 μ mho in 30 seconds flagged automatically by UFI software (DPSv3.6) and edited for artifact. Given that some women show submaximal hot flashes failing to reach the 2 μ mho criterion, all potential hot flash events were also visually inspected. Events showing the characteristic hot flash pattern but less than 2 μ mho/30 sec rise were coded as hot flashes. This coding has been shown to be reliable (24,25). A 20-minute hot flash lockout period was implemented

after the start of the flash. Physiologically-detected hot flashes were considered reported if met by a self-report of a hot flash within 5 minutes. Participants reported hot flashes as they occurred during waking hours by completing a portable electronic diary (Palm Z22; Palm, Inc., Sunnyvale, CA) and pressing event mark buttons on the hot flash monitor. Hot flashes reported by either method were categorized as a self-report. Four categories were considered for these analyses: physiologic hot flashes, self-reported hot flashes, self-reported hot flashes not corroborated by a physiologic hot flash, and physiologically-detected hot flashes not met by a self-report.

Physical activity—Daily physical activity was assessed continuously over the monitoring period with accelerometer-derived activity counts from the Biolog monitor. The mean activity count in the 10 minutes prior to a hot flash were calculated for each type of hot flash, and categorized as pre-flash activity. Activity during each type of flash and for the 10 minutes following a hot flash was not included in the analysis, as it was expected that women may be likely modify their physical activity pattern during and immediately following a hot flash. The mean activity count during all other waking hours not preceding hot flashes was calculated and categorized as control activity. For the purposes of these analyses, only activity counts during waking hours were used.

Covariates—All covariates were selected on the basis of previously documented associations with vasomotor symptoms and physical activity, and included education, habitual physical activity, race/ethnicity, body mass index, negative affect, age, menopausal status, and time of day of hot flash occurrence. Race/ethnicity and educational level were self-reported in the SWAN screening interview. Habitual physical activity was assessed with the Kaiser Physical Activity Survey, an adaptation of the Baecke physical activity questionnaire (9). A sum score for total activity was derived from responses to questions about physical activity during sports/exercise, household/caregiving tasks, and daily routine in the previous year. BMI was calculated using weight (kg) and height (m)², and treated as a continuous variable for this analysis. Depressive symptoms were self-reported with the Center for Epidemiological Studies Depression Scale (CESD), a 20-item depression symptom scale (26). Anxiety symptoms were self-reported with the Spielberger State-Trait Anxiety scale (27), a 40-item scale that assesses current anxiety (state anxiety) and typical anxiety levels (trait anxiety). Age was calculated from the participant's date of birth and date of study visit. Menopausal status was defined as perimenopausal (bleeding in the last 12 months, some change in cycle regularity) and postmenopausal (no bleeding in the last 12 months). Time of day was categorized as morning (4 am to noon), afternoon (noon to 4 pm) and evening (4 pm until 4 am). As only those symptoms that occurred during waking hours were including in analysis, morning encompassed symptoms during waking until noon, while evening encompassed symptoms from 4 pm until participants reported going to sleep.

Statistical analyses—Separate hierarchical generalized linear models were used to examine relationships between physical activity and each category of hot flashes (physiologic hot flashes, self-reported hot flashes, self-reported hot flashes without physiologic hot flash, and physiologically-detected hot flashes without self-report). Hierarchical generalized linear modeling utilizes logistic regression methods that adjust for and model the non-independence of repeated measures within individuals, making it appropriate for this assessment of continuous and repeated factors over several days within a sample of women. Associations between mean accelerometer-assessed activity counts in the 10 minute period prior to hot flashes during waking hours was compared to the mean activity counts at all waking non-flash times. Separate non-flash control times were calculated for all hot flash types. Odds ratios of hot flash reporting by each standard deviation increase in activity count were then determined. Final models were adjusted by

education, habitual physical activity, race/ethnicity, body mass index, depressive symptoms, age, menopausal status, and time of day hot flashes were reported/measured. Covariates were initially selected *a priori* due to previously-documented associations between with hot flashes, and subsequently retained in the model with significant univariate associations with hot flash type ($p < .05$). Depressive symptoms and anxiety were examined separately as covariates given their high correlation (depressive symptoms and trait anxiety $r: .84, p < .001$) and associated concerns about multicollinearity. The cross-product of negative affect variables and activity count, as well as habitual physical activity and activity count, were also examined in separate final, fully adjusted models investigating the role of negative affect and habitual physical activity on the relationships found between acute increases in physical activity and each hot flash type.

Results

Characteristics of sample

Participants were relatively highly educated, postmenopausal, and overweight (Table 1). By design, the sample was half Caucasian and half African American. All participants had at least one self-reported or physiologically-detected hot flash during the observed period. Participants reported fewer hot flashes than were detected by the physiologic hot flash monitors (mean of 3.41 self-reported hot flashes per participant per day; mean of 5.09 physiological hot flashes per participant per day). During waking hours a total of 1,067 hot flashes were reported, and 1,587 physiologic hot flashes were detected.

Increased activity as a predictor of hot flashes

We tested the likelihood of each of the four types of hot flashes following increases in physical activity. We observed no relationships between physiologic hot flashes (OR: 1.00, 95% CI: 1.00-1.00, $p = .06$), self-reported hot flashes (OR: 1.00, 95% CI: 1.00-1.00, $p = .50$), or physiologically-monitored hot flashes not met by self-report (OR: 0.99, 95% CI: 0.99-1.00, $p = .66$) and physical activity in multivariable models. However, self-reported hot flashes not corroborated by a physiologic hot flash were more likely following increases in physical activity (OR: 1.04, 95% CI: 1.00-1.10, $p = .02$) in multivariable models. Higher habitual physical activity (OR: 1.27, 95% CI: 1.13-1.41, $p < .001$), higher BMI (OR: 1.1, 95% CI: 1.06-1.17, $p < .001$) more depressive symptoms (OR: 1.05, 95% CI: 1.01-1.06, $p = .02$), and STAI state anxiety (OR: 1.03, 95% CI: 1.01-1.06, $p < .01$) were also associated with self-reported hot flashes not corroborated by a physiologic hot flash (table 2).

The relationship between physical activity and self-reported hot flashes lacking physiologic evidence varied by levels of negative affect, such that the relationship between increased activity and self-reported hot flashes lacking physiologic evidence was stronger among women with higher depressive symptom scores (OR: 1.005, 95% CI: 1.001-1.009, $p = .02$) or higher trait anxiety (OR: 1.003, 95% CI: 1.000-1.007, $p = .05$). Stratified analyses confirmed that the increased likelihood of self-reported hot flashes lacking physiologic evidence following physical activity was most apparent among women with higher CESD scores or trait anxiety. Habitual physical activity did not modify relationships between acute increases in activity and any type of hot flash (data not shown).

Additional findings deserve mention. In multivariable models, African-American women (vs. Caucasian, OR: 1.56, 95% CI: 1.18-2.07, $p < .001$), and postmenopausal women (vs. perimenopausal OR: 1.66, 95% CI: 1.22-2.44, $p < .01$) were most likely to show physiologically monitored hot flashes. Higher depressive symptoms were associated with increased likelihood of self-reported hot flashes (CESD OR: 1.03, 95% CI: 1.01-1.05, $p < .01$). Younger (OR: 0.92, 95% CI: 0.85-0.99, $p = .03$) and postmenopausal women (OR: 1.49,

95% CI: 1.20-1.85, $p < .01$) had a higher likelihood of physiologic hot flashes without self-report.

Discussion

In this investigation of daily physical activity and hot flashes in midlife women, modest increases in physical activity were associated with subsequent self-reported hot flashes lacking physiologic evidence. This relationship was modified by negative affect, such that women with higher levels of depressive symptoms and anxiety showed the highest propensity to report hot flashes lacking physiologic evidence following increases in physical activity. In contrast, acute increases in daily physical activity were not associated with physiologically-monitored hot flashes. Therefore, rather than acting as a trigger of physiologically-detected hot flashes, acute increases in physical activity were linked to a subsequent reporting of hot flashes lacking physiologic evidence.

These findings add to the limited existing literature on physical activity and hot flashes. Physical activity is frequently discussed as both a protective factor and an acute trigger of hot flashes, but there has been relatively little rigorous empirical investigation of this association. Few published studies have utilized both self-report and physiologic measures of both hot flashes and physical activity to test the commonly posited association between physical activity and hot flashes. This addition is important, as self-reported and physiologically-detected assessments of both hot flashes (20) and physical activity (28) are not highly correlated. Further, the risk and protective factors associated with these self-reported and physiologically-detected variables often differ. Some existing studies using physiologic measures of hot flashes and/or physical activity have shown a similar relationship between aerobic exercise (3) and diary-reported physical exertion (20) with increases in hot flashes without physiologic evidence. Limitations of these studies include self-reported physical activity measures (20) and an ethnically homogenous, relatively normal-weight sample with limited generalizability (3). Thus, this study adds importantly to the literature, indicating that the association between acute increases in physical activity may be most linked to the reporting of hot flashes, specifically those lacking physiologic evidence. Taken as a whole, these findings indicate that symptom perception and interpretation likely plays a key role in the relationship between physical activity and hot flash reporting.

This study underscores the potential importance of depressive and anxiety symptoms in the association between acute physical activity and hot flash reporting. Other work has shown that self-reported and physiologically monitored hot flashes may differ, and that women with high levels of negative affect (e.g., depressive symptoms, anxiety) are more likely to report physiologically-undetected hot flashes (20,24). Another study showed that self-reported hot flashes decreased following increases in habitual physical activity over time among women with a history of depression (12). However, previous work has not examined negative affect as a modifier of relationships between physical activity and hot flashes, and our study is the first to examine the role of negative affect on the relationship between acute physical activity and the occurrence of hot flashes. The current study contributes to this small literature with additional evidence that negative affective states are influential in the occurrence of hot flash reporting following acute increases in physical activity.

This study also examined factors commonly associated with hot flashes as selected covariates, with the expectation that the mechanisms contributing to physiologically-detected and self-reported hot flashes would differ. Supporting this concept, race/ethnicity and menopausal status were associated with physiologically-detected hot flashes, but not with self-reported hot flashes. In contrast, negative affect, known to influence symptom

perception, was associated with self-reported hot flashes, but not with physiologically-detected hot flashes. While habitual physical activity was related to self-reported hot flashes not corroborated by physiologic hot flashes, physiologically-detected and self-reported hot flashes were not associated with habitual physical activity in this study. The latter relationships are consistent with most studies of habitual physical activity. While other studies have supported a role of cardiorespiratory fitness in modifying the relationship between hot flashes and physical activity (3), levels of habitual physical activity did not influence relationships between acute activity and any type of hot flash. However, our measure of habitual physical activity cannot be considered a proxy for fitness. In a closer comparison of differences in associations by method of hot flash assessment, covariates associated with physiologic hot flashes without corroborating self-report mirror those of physiologic hot flashes in general, while self-reported hot flashes that were not corroborated by a physiologic hot flash were also associated with higher habitual physical activity, higher BMI, and higher levels of negative affect. Overall, this pattern of findings supports the concept that different methods of measuring hot flashes capture varied information that can contribute to our understanding of these common symptoms.

This study had several strengths. The thorough measurement of hot flashes is particularly notable, and differs from the larger literature that primarily utilizes retrospective self-report of these common symptoms. Diary and event marker self-reporting of hot flashes throughout the observed period reduces the retrospective reporting biases common to typical measurement. The inclusion of physiologic hot flash monitoring allows for the assessment and comparison of self-reported and physiologically-detected hot flashes. This comparison enables an examination of the mechanisms contributing both to physiological changes and the interpretation and subjective experience of symptoms. The reliability and external validity of these findings was enhanced by ambulatory monitoring during participants' normal daily activities. Finally, this study was conducted with a well-characterized sample of African-American and Caucasian women who have been followed throughout the menopausal transition.

Several limitations of this study should also be noted. Intensity of physical activity and changes in energy expenditure cannot be interpreted from accelerometer-derived activity counts, the objective measure of physical activity utilized in this study. Future study would benefit from detailed assessments of activity that could identify these aspects of activity change, in order to clarify the level and intensity of acute activity necessary to exhibit effects on symptom reporting. Overall, women in this study exhibited low levels of physical activity during the observed period, consistent with protocol recommendations to reduce artifact in hot flash measures. These levels may not reflect participants' typical activity levels, which may have masked relations that would be observed with more intense bouts of physical activity. Further, these low levels of physical activity, along with the frequent measurement of small units of time assessed, may have contributed to the small point estimates seen in this analysis. While established parameters were followed in coding hot flashes, these methods are subject to error. Finally, this sample is relatively small, and includes only women who typically report hot flashes and are willing and able to comply with the protocol. This may limit the generalizability of these findings to other samples.

Conclusions

The relationship between daily, low-intensity physical activity and hot flashes seems to be limited to self-reported hot flashes that are not corroborated by a physiologic hot flash. This is particularly true among women with higher levels of negative affect, who are also known to be more likely to report more frequent, severe, and bothersome hot flashes in midlife. The commonly posited relationship between acute physical activity and hot flashes may

therefore be more tied to symptom perception and reporting than to any physiological mechanism. Women in midlife reporting hot flashes often seek treatment options and recommendations to manage their symptoms, but may be wary of behavioral change thought to worsen their symptom experience. Clinicians should consider the potentially modifiable role of affect and the perception of hot flashes in the relationship between physical activity and hot flashes among women when making treatment recommendations for their symptomatic patients.

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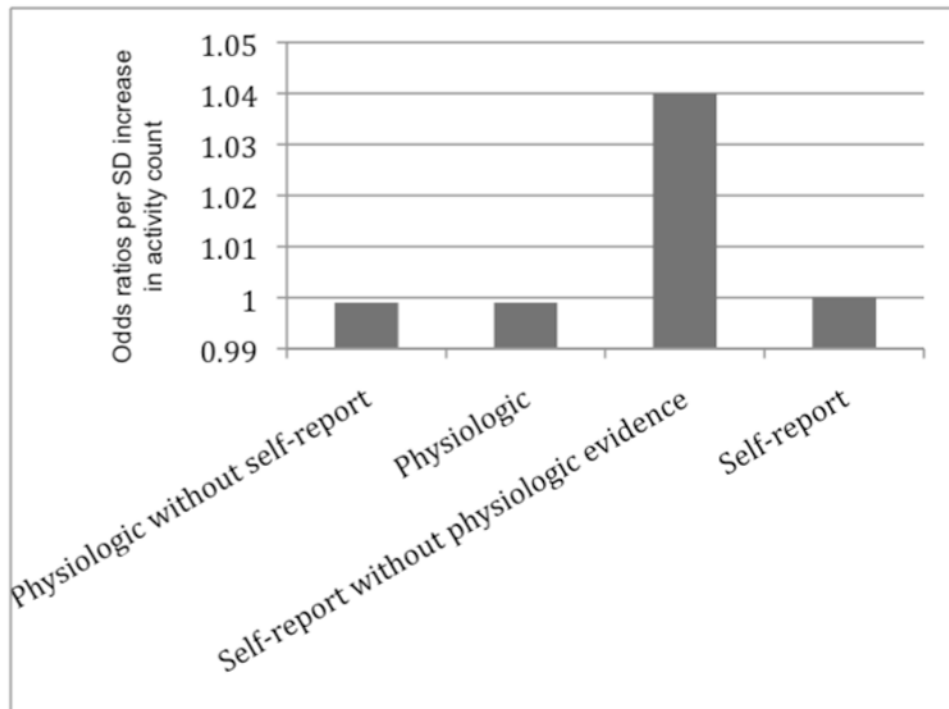


Figure 1. Odds of reporting different types of hot flashes with increases in physical activity.

Table 1

Characteristics of the sample (n=51).

	N (%)
Education	
High school or less	10 (19.6)
Some college or more	41 (80.4)
Race/ethnicity	
Caucasian	26 (51)
African American	25 (49)
Menopausal status	
Perimenopausal	5 (9.8)
Postmenopausal	46 (90.2)
	Mean (SD)
Hot flashes[±]	
Self-reported hot flashes	20.92 (13.60)
Physiologic hot flashes	31.12 (18.83)
Self-reported hot flashes without physiologic evidence	4.71 (7.38)
Physiologic hot flashes without self-report	14.90 (12.34)
Activity count[*]	24.6 (96.7)
Habitual physical activity (KPAS total score)	6.8 (1.3)
Body mass index	29.8 (5.0)
Depressive symptoms (CESD)	7.5 (6.5)
Trait anxiety (STAI trait)	35.1 (8.1)
State anxiety (STAI state)	31.1 (8.6)

[±] Mean number of daytime hot flashes over monitoring period

^{*} Activity count derived from physiologic monitor (range 1 - 3,623).

KPAS: Kaiser Physical Activity Survey

CESD: Center for Epidemiological Studies Depression Survey

STAI: State-Trait Anxiety Inventory

Table 2

Self-reported hot flashes without physiologic evidence (n=240) predicted by 1 standard deviation increase in activity count.

	Odds Ratios (95% Confidence Intervals)	p-values
Education		
Some college or more (compared to high school or less)	1.41 (.91-1.99)	0.11
Habitual physical activity (KPAS total score)	1.27 (1.13-1.41)	<.001
Race/ethnicity		
African-American (compared to Caucasian)	0.83 (.49-1.25)	0.32
Body mass index	1.10 (1.06-1.17)	<.001
Depressive symptoms (CESD)	1.05 (1.01-1.06)	0.02
Age	0.99 (.90-1.06)	0.80
Menopausal status	0.64 (.32-1.67)	0.19
Postmenopausal (compared to perimenopausal)		
Time hot flash reported		
Morning (compared to evening) [#]	1.05 (.82-1.35)	0.66
Afternoon (compared to evening) [#]	0.73 (.58-.91)	<.01
Activity count (change per standard deviation)[*]	1.04 (1.00-1.10)	0.02

All variables entered simultaneously in hierarchical generalized linear models.

* Activity count derived from physiologic monitor (range: 1 - 3,623; SD: 96.7).

KPAS: Kaiser Physical Activity Survey

CESD: Center for Epidemiological Studies Depression Survey

STAI: State-Trait Anxiety Inventory

[#] Morning: Waking-12:00 pm

Afternoon: 12:01-4:00 pm

Evening: 4:01 pm-bed