

RESEARCH ARTICLE

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Continuous or intermittent walking, the effect on hemoglobin in sedentary employees during 10-week intervention

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Abstract: To examine long term changes on glycated hemoglobin in sedentary employees exposed to two different walking programs during a 10-week intervention. A total of 68 sedentary employees participated in a 10week walking intervention and were randomly assigned to one of three groups: intermittent walking, continuous walking or control group. Hemoglobin A1cNOW+ device tested glycated hemoglobin and accelerometry assessed physical activity. Results showed glycated hemoglobin significantly decreased over the ten weeks (5.82±0.49, 5.66±0.44) F(1,64) =4.229, p=.044) in the continuous walking group. Post-Hoc test showed the continuous walking group was significantly affected, F=8.463, p=.009, with a large size effect n2=.297. There were no changes within the intermittent group $(5.69\pm0.63, 5.63\pm0.6)$ or control group $(5.59\pm0.6, 5.6\pm0.54)$ (p>0.05). Accelerometry showed a main effect of time by group interaction F(4,124) =4.688, p=0.001). Post-Hoc indicated that the continuous walking group took significantly longer bouts of moderate to vigorous intensity walking at week-6 compared to pre-test (p=0.006) at this could have influenced the results, there were no changes in the length of bouts in the intermittent or control groups (p>0.05). Sedentary employees who perform a moderate intensity continuous walking program show benefits in controlling blood glucose, thereby, reducing the risk of developing type 2 diabetes.

Keywords: Physical activity, Sedentary behavior, Blood glucose, Glycated hemoglobin



Mynor Rodriguez-Dr. Hernandez lives in San Ramon Costa Rica, He is a full Professor at the University of Costa Rica, Western Campus. His research focuses on projects that would impact community needs in terms of physical activity, health,

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Dr. McDonald has been a professor clinical at Auburn University since 2012. His TigerFit laboratory conducts health and fitness testing for the Auburn community. program allows students to get "hands-on" experience in all

aspect of health and fitness testing and prescription working with a cross section of the local community. Dr. McDonald came to academia later in life following a 30-year career in the U.S. Air Force where he served as a fighter pilot and military diplomat, retiring at the rank of Colonel. After a few years working in industry, he came to Auburn in pursuit of a PhD in exercise physiology and after completing the degree, he was hired as clinical faculty. He has done research in basic muscle oxygen on-kinetics, lactate metabolism, and role of exercise in weight loss and health across various human populations.





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adherence and translating these findings to effective evidence-based interventions, primarily for women and children. Dr. Wadsworth has served as a co-investigator on National Institutes of Health and Robert Wood Johnson Foundation grants aimed at examining the impact of physical activity policies on children's physical activity levels at school and during physical education. Over the past year, she has concentrated her efforts on identifying factors to long-term exercise adherence in young children. Dr. Wadsworth lives with her husband and three children in Auburn, Alabama.

1. Introduction

Sedentary behavior and physical activity are two inversely related behavioral scenarios that have significant effects on health outcomes. While physical activity produces positive effects on overall health [1] chronic sedentary behavior is linked to health impairments such as increased obesity, diabetes, cardiovascular disease, cancer, and early mortality [2-With sedentary behavior leading development of several health complications, it is important to consider the positive effect that small amounts of exercise may produce on overall well-being [1]. Walking behavior is linked to reductions in type 2 diabetes (T2D) risk. A previous review that included 300,000 participants, showed that T2D was 0.69 (95% CI 0.58-0.83) for regular participation in physical activity of moderate intensity as compared with being sedentary. Also, the relative risk was 0.70 (0.58–0.84) for moderate walking approximately 2.5 hours/week the risk was 0.69 (95% CI 0.58-0.83) compared to inactive [5]. An additional prospective study that examined approximately 70,000 healthy female nurses over 8 years, determined that a higher walking activity level was associated with significant reductions in T2D risk [6].

High levels of blood glucose are related to risk of developing cardiovascular disease (CVD) and increase the risk of all-cause mortality. To control glycemic levels in long term, patients with T2D should stablish the goal of keeping glycated hemoglobin (HbA1c) below 7% [7-10], which can reduce the long term complications in about 76% [11]. According to the experts, HbA1c indicates the average blood glucose

levels over several months and it is not a measure that changes over the course of one day when the blood glucose can variate depending on the content and frequency of the meals and physical activity achieved [12-14].

Previous epidemiological studies have shown that small reductions (1%) on HbA1c, is associated with 14% to 21% of reduction on cardiovascular and diabetes complications and 37% microvascular detriments [15]. Some studies indicate that the intensity level of the exercise is related to improvements of blood glucose control as reflected in reductions in HbA1c [16].

Looking at studies that targeted participants with T2D with twelve or more weeks interventions, and focused on different models of exercise training, some have found significant differences post intervention in favor of reductions of HbA1c [8, 17, 18]. Meanwhile, results obtained from a meta-analysis that included exercise interventions with ≥8 weeks [19], show that when exercise groups were compared to control groups, the weighted mean difference (WMD) HbA1c was significantly lower in the exercise groups, pooling the post-intervention results, the exercise groups had a lower HbA1c compared with the control groups (7.65% vs 8.31%; WMD, −0.66%; *P*<.001). However, others have not found significant changes. For example, in a 16 week aerobic training conducted by Tessier and colleagues, no significant changes regarding HbA1c were found [20]. Although physical activity has constantly been linked to reductions in T2D and results seem promising [21, 22], the effect of walking on HbA1c needs to be further clarified [23], particularly in terms of which type of walking activity has the greatest effective on HbA1c and also, it is important to clarify the timeline to see positive changes on this variable. Furthermore, most of the interventions have included participants with type 2 diabetes, focusing on physical activity and its effect on T2D versus a prevention model.

Intermittent walking activity to disrupt sedentary behavior may be a modality of exercise to reduce the risk of developing T2D [24]. In acute interventions, some studies have compared intermittent physical activity versus continuous physical activity. In the short term, intermittent physical activity appears to attenuate, to a larger extent, glucose excursions compared to 1-h morning exercise time and intensity matched [25]. In T2D patients and in a counterbalanced acute intervention, intermittent



walking improved glycemic control compared to a time duration-matched continuous walking activity [26]. Both models, independently of continuous or intermittent physical activity, have been determined to provide similar glucose outcomes when time and intensity are matched, meaning that physical activity is beneficial to improve blood glucose levels [27].

However, to our knowledge, most interventions have focused on acute effects of physical activity on blood glucose levels when comparing intermittent versus continuous physical activity. Thus, our experimental study investigated the long-term effect of two different walking activity protocols in free-living conditions on HbA1c in sedentary office workers.

2. Materials and Methods

The methodology for this study has been published before [28, 29]. For instance, this section of the study adds the particular effect of walking activity on HbA1c. A brief procedure is defined below.

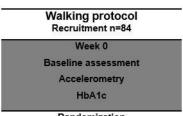
2.1 Participants and study protocol

This study involved initially 84 sedentary office employees, males and females who participated in a 10-week walking- free living conditions- intervention.

The sample size was determined following Cohen's recommendations [30]. Then, for a conservative calculation a_1 =.05, r=.30, and power =.80 the desire sample size was 68. Certainly, a total of 68 subjects completed the trial as explained before [28].

The walking intervention included a 10-week program consisting of two walking protocols; multiple micro-bout and continuous walking. For both experimental groups, time and intensity were matched, and with intensity and duration increasing incrementally over the 10- week program (Figure 1).

All procedures described herein were approved by the Auburn University Institutional Review Board and conformed to the standards set by the latest revision of the Declaration of Helsinki. Prior to participation, all subjects were asked to sign an informed consent and complete the Physical Activity Readiness Questionnaire (PAR-Q) followed by baseline assessments for height, weight, HbA1c and physical activity. Height, weight and HbA1c measures were taken again at week 11. Physical activity was measured with an ActiGraph GT3X (ActiGraph Corp., Pensacola, FL), accelerometer worn on the waist at baseline, week 6 and week 11.



Randomization

Gender

• DIVII										
Intermittent walking group n=28	Continuous walking group n=28	Control group n=28								
Multiple bouts/day Walking program + Advise 10,000 steps per day	1 single bout /day Walking program + Advise 10,000 steps per day	No PA prescription + Advise 10,000 steps per day								
Week 1 4 break/5-min/3 day/week 30-60% HR/3-6 RPE Week 2-3 6 break/5-min/3 day/week 40-60% HR/4-6 RPE Week 4-5 6 break/5-min/4 day/week 40-60% HR/4-6 RPE	Week 1 20 min/3 day/week 30-60% HR/3-6 RPE Week 2-3 30 min/3 day/week 40-60% HR/4-6 RPE Week 4-5 30 min/4 day/week 40-60% HR/4-6 RPE									
	Week 6 accelerometry assessmen	nt								
Week 6-7 6 break/5-min/5 day/week 40-60% HR/4-6 RPE Week 8-10 8 Break/5- min/5 day/week 40-60% HR/4-6 RPE	Week 6-7 30 min/5 day/week 40-60% HR/4-6 RPE Week 8-10 40 min/5 day/week 40-60% HR/4-6 RPE									
Post-test week 11 n=23 Accelerometry HbA1c	Post-Test week 11 n=23 Accelerometry HbA1c	Post-Test week 11 n=22 Accelerometry HbA1c								

Figure 1 study design in a 10-week intervention. BMI (Body Mass Index), HR (heart rate), RPE (rated perceived exertion).



At baseline groups were randomized to one of three groups, with equal randomization between males and females and BMI status. The two experimental groups were given walking exercise prescriptions based on group assignments in written and electronic format. A weekly email and three text messages a week were sent to remind and encourage walking participation. The control group was told that "10,000 steps a day is a minimum for good health". Figure 1 details the intervention design.

2.2 Measurements

2.2.1 Body Mass Index

Height and weight were assessed using a stadiometer (SECA Model 769, Seca, Hamburg, Germany) and Body Mass Index (BMI) was determined by weight (kg) / [height (m)]².

2.2.2 Hemoglobin A1c test

A1cNOW⁺ system was used to measure hemoglobin HbA1c Polymer Technology Systems, Inc. (Indianapolis, IN). The HbA1c test is a blood test to analyze the average levels of blood glucose over a period of three months to determine diabetes development. Outcomes of the test are interrupted as: normal range below 5.7%, pre-diabetes 5.7-6.4%, and diabetes 6.5% or above.

When comparing the A1CNOW+ system to the National Glycohemoglobin Standardization Program (NGSP) certified method [31], the accuracy of the A1cNOW on average is 99%, meaning that a true 7.0 %A1c could be approximately 6.9 %A1c or 7.1%A1c [32]. A small amount of blood sample (5 microliters (μ L)) was collected from a finger prick using a 28-gauge lancet (Unistick 3 comfort, Owen Mumford, Marietta, GA). The sample was placed in a portable A1CNOW device with a test cartridge and the results were recorded after 5 minutes.

2.2.3 Physical Activity measures

To measure Physical Activity behavior an Actigraph accelerometer GT3X (ActiGraph GT3X; ActiGraph Corp., Pensacola, FL) was attached on the right hip of each participant for seven days at baseline, week 6 and week 11. The device is a small trial-axial device weighing 27g and measuring 3.8 cm x 3.7 cm. x 1.8 cm. The GT3X records accelerations ranging from 0.05 to 2 g at a rate of 30 Hz in three different axes: vertical, antero-posterior, and medio-lateral [33]. Based on previous studies and best practice guidelines [34, 35], an epoch length of 1-minute was chosen as

the standard for the current study with a sampling rate of 30 Hz. Additional criteria for analysis include a minimum of 10 hours daily wear time and 3-5 days of monitoring. There is relative consensus of a minimum of 10 hours per day of wear time needed for sampling wake-time behavior with 3-5 days of monitoring required to achieve 80% reliability for total and moderate-to-vigorous intensity physical activity [36-38]. Non-wear time was identified by participants completing a daily log of wearing time and non-wear time was removed from the analysis. Previously validated cut points were used to classify accelerometer data as sedentary (<100 counts/minute), moderate (<5,999 counts/minute) and vigorous (>5,999), [39]. Light activity was defined as 500-2019 counts per minute [40]. After meeting the above criteria, the data were analyzed to obtain on each assessment a percentage of time per hour spent in sedentary behavior, light, and moderate to vigorous physical activity per group. Also, to clarify the results, a single regular week day of accelerometer data was used to describe participants' physical activity behavior as an example, specifically MVPA.

2.3 Statistical analysis

All analyses were performed with SPSS 23.0. To answer the research questions, a mixed design ANOVA approach was performed to examine the main effect over time and the main effect of time and group interaction in HbA1c and physical activity (sedentary behavior, light, and moderate to vigorous physical activity (MVPA). Between factors examined differences between the three groups, whereas, within factors assessed change over time. The Bonferroni Post-Hoc testing was used if a significant interaction occurred.

3. Results and Analysis

A total of sixty-eight sedentary office employees completed the intervention. Initial analysis showed that groups did not differ by BMI p=0.272. The adherence and attrition rates of the study were 80.95% and 19.05% respectively, which is considered positive rates [41, 42].

3.1 Hemoglobin A1c results

The analysis show that HbA1c was not significantly different between groups at the onset of the study. The results from the mixed-design ANOVA showed that HbA1c decreased significantly from pretest to post-test F(1,64) = 5.709, p=.020) with a medium size effect of $n^2=082$.



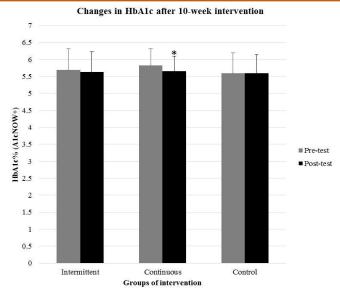


Figure 2 compares the A1cNOW+ results for the three intervention groups at pre-test and post-test.

* significantly different from pre-test (p<0.05)

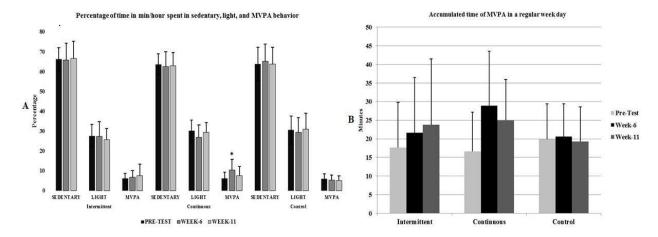


Figure 3A Time percentage per hour spent in sedentary, light, and MVPA, and **3B** Example of time accumulated in a regular week day shown as MVPA.

*significantly different from pre-test (p<0.05)

		PRE- TEST	WEEK-6	WEEK-11	Main effect of group			Main effect of time			Time by group interaction			Post- Hoc
					F	p	\mathbf{n}^2	F	P	\mathbf{n}^2	F	p	\mathbf{n}^2	р
Overall mixed	Sedentary				1.318	0.275		0.019	0.982		0.440	0.780		
ANOVA	Light				2.067	0.135		2.435	0.092		1.605	0.177		
	MVPA				3.963	0.024*	.115	4.895	0.009*	.074	4.688	0.001*	.133	
Intermittent	Sedentary	66.2±5.9	65.9±8.4	66.6±8.7										
	Light	27.6±5.6	27.3±7.5	25.7±5.6										
	MVPA	6.2±2.6	6.8±3.4	7.7±5.7										
Continuous	Sedentary	63.6±5.4	62.7±7.4	62.97±6.5										
	Light	30.2±5.4	26.9±6.3	29.4±4.98										
	MVPA	6.21±3	10.4±5.6	7.62±4.6										**
Control	Sedentary	63.8±8.5	65.3±8.7	63.9±8.4										
	Light	30.6±7.1	29.3±7.6	30.9±8.1										
	MVPA	5.6±2.6	5.34±2.5	5.18±2.24										

Figure 4 Results of the mixed ANOVA in terms of percentage of time spent in sedentary, light, and MVPA behavior per group of intervention.

^{**} Continuous group MVPA was significantly higher on week-6 compared to pre-test (p<0.05).



^{*}Significantly different (p<0.05)

There was a significant effect of time by group interaction F(2,64)=3.158, p=.049. The Bonferroni *Post-Hoc* test shows a significant reduction on HbA1c from pre-test to post-test within the continuous group (p=.005). There were no significant changes within the multiple break group (p=.266) or control group (p=.661). When analyzing the results by delta change, the continuous walking group showed a significant reduction of HbA1c by 0.16% (40 to 38 mmol/mol) with a total mean change of 2.75% from pre-test to post-test.

The multiple breaks group reduced HbA1c by 0.06% (39 to 38 mmol/mol) with a total mean change of 1.1% but it was not significant, and the control group increased HbA1c levels by 0.02% (38 mmol/mol) with a mean increment of 0.18%.

3.2 Physical activity results

Accelerometer data were obtained and analyzed as time percentage per hour spent in sedentary, light intensity, MVPA bouts. Figure 3A shows the percentage of time per hour spent in each of those behaviors per intervention group. Figure 3B represents daily behavior in terms of MVPA; it reflects the MPVA bouts accumulation for all groups in a regular week day. Continuous group shows higher level of accumulated MVPA during waking time.

Figure 4 shows the Mixed ANOVA results as percentage of minutes per hour in sedentary, light, and MVPA in each intervention group. Participants in the continuous group significantly increased MVPA at week 6 and they were able to maintain higher levels of MVPA at the end of the intervention.

4. Discussion

The present study examined changes in HbA1c for sedentary employees exposed to two different walking programs during a 10-week intervention. The results show that participants in the continuous group (20-to-40 minutes 3-to-5 days/week, 10-week intervention) reached moderate intensity walking within the continuous bout and showed significant reductions on glycated hemoglobin or HbA1c, potentially reducing the risk of developing type 2 diabetes and subsequent health disorders.

Our findings are in concordance with results from two meta-analyses [43, 10] showing programs that included 12 or more weeks of structured physical activity with more than 150 minutes per week were associated with significant HbA1c reductions compared

to control participants. For our study, participants were not asked to participate in 150 minutes/week of walking until week 6. In our free-living trial, experimental groups were asked to incrementally increase walking time from week 1 to the end of the program and were not asked to walk 150 minutes per week until week 6 (figure 1). The accelerometer analysis shows that both experimental groups spent on average less than 12 percent of the time in MVPA per hour. The continuous group spent in average almost 29 minutes in MVPA per bout on week 6 and 25 minutes on week 11, approximately 10 percent (week-6) and 8 percent (week-11) which may have stimulated physiologic pathways to significantly reduce HbA1c after almost three months. Exercise is a physiological stressor that affects the glucose homeostasis and energy balance through multiple hormonal activators and regulators, nervous system, regulators within the muscle and liver, which positively reduces glucose content [44]. Whereas, HbA1c in the intermittent group was reduced, changes were not significant. Based on these results changes in HbA1c might occur with longer bouts of MVPA and specific accumulation of MVPA during the day needs to be explored as intermittent physical activity seems to be important to break sedentary time and improve metabolic markers. Exercise prescription characteristics such us time, intensity, and frequency may play a fundamental role on glucose changes throughout walking interventions to produce significant changes on HbA1c [16, 45, 46]. For example a program with only 3 workouts per week was not enough to change glycated hemoglobin in a nonrandomized trial even though they accumulated more than 150 minutes per week [47]. In addition, the intensity of walking is an important determinant of HbA1c changes [16, 48]. A group of scientists found exercise intensity as a predictor of HbA1c reductions [16], and other researchers concluded that moderate intensity walking significantly decreased glycated hemoglobin compared to the control participation [48], which may explain the lack of significance for the intermittent group due to the lower amount of moderate intensity accumulated throughout the day. The accelerometer data grouped in percentage of sedentary behavior, light intensity, and MVPA per hour, as well as the accumulated time of MVPA in a regular day, showed that the continuous group reached longer bouts of MVPA which could explain changes in HbA1c in this group. These results are supported by the study of Gay, Buchner, and Schmidt [49] who showed that the relationship between physical activity and HbA1c was associated with short bouts of MVPA (~10minutes)



for individuals at risk of developing T2D, the length of the MVPA bouts positively associates to HbA1c reductions. Like those findings, in our study we were able to see reduction on HbA1c on those who achieved longer bouts of MVPA. Putting the results of both studies together [48, 49], there may be a specific doseresponse relationship that needs to be studied in further detail to determine the amount of MVPA that people need to perform to reduce the risk of developing T2D.

Early acute trials where intermittent physical activity was applied in short time bouts (2 min) to see effects on glucose have shown that short bouts compared to continuous bouts of PA are a similar or better approach to control insulin and glucose excursions [25, 50]. In our 10-week study, intermittent physical activity did not reduce HbA1c significantly, possible explained by the lower levels of MVPA compared to the continuous trial. More research is required to explore the possible benefits of short versus long bouts of physical activity and specifically, accumulated MVPA effects on HbA1c sedentary adults at risk for developing TD2. Although the intermittent group was prescribed with the same dose of physical activity, this group participated in less MVPA and did not have significant changes in HbA1c. Intermittent activity throughout the day may be harder to regulate and reach a moderate pace. Short walking breaks with no supervision may be even harder to achieve the desired intensity when comparing to longer periods of walking, in which people have more time to improve exertion compared to the intermittent walking. In time percentage per hour, at week 6, the continuous walking group achieved significantly longer bouts of moderate intensity during their walking activity (10.4±5.55%) compared to the intermittent walking group (6.79±3.44%) and control group (5.34±2.5%). However, at week 11, the continuous achieved shorter MVPA bouts (7.62±4.59%) compared to week 6, intermittent walking group slightly increased MVPA (not significant) about 1% from week 6 (7.7±5.68%), and the control group decreased MVPA (5.18±2.24%). Further research is necessary to determine the effects of these two-walking program on long-term walking behavior and how long-term walking behavior relates to HbA1c.

We faced some limitations, we did not test for hemoglobin levels to compare with A1cNOW results, but previous validations have been performed [51-53]. We did not test participants for any other blood related disease (e.g. hemolytic anemia or other hemolytic

disease, blood loss) that may change HbA1c levels and produce inaccurate readings.

The walking program was meant to replicate free living conditions to mimic how exercise prescriptions are giving in a clinical setting, thus, physical activity intensity and frequency were not recorded every day to ensure participants met the program prescription; however, participants indicated they following the instruction as asked, and participants were followed closely to ensure they were following the instructions and meeting recommendations. This is a major concern in this study; however, people's behavior is a real scenario that accounts when interventions are based on free living condition prescriptions.

5. Conclusion

Based on our results and previous reports, we conclude that long term continuous walking activity at moderate to vigorous intensity seems to be enough to reduce glycated hemoglobin percentages in adult sedentary employees. Since breaking sedentary time is a major concern to reduce health impairment risks, intermittent walking activity may have some positive impact on reducing HbA1c percentages, however, further research is necessary to identify the dose of intermittent physical activity necessary to accumulate enough MVPA during the day to decrease HbA1c levels and reduce the risk of developing T2D on sedentary population.

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Ethics approval

This study was approved by Institutional ethics committee.

Authors Contribution

Mynor Rodriguez-Hernandez - Conceived the study design, methodology and analysis Collected the data, Performed the analysis, Wrote the paper. James R. McDonald - Contributed to the study design, Proved and edited the paper. Danielle W. Wadsworth -Contribute to the study design and analysis, contributed to the data analysis, Edited and proved the manuscript.

Does this article screened for similarity?

Conflict of interest

The authors have no conflicts of interest to declare that they are relevant to the content of this article.

Informed consent

Written consent was obtained from the participants

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