

1 Morning (Fasting) Vs. Afternoon Resistance Exercise in Individuals with Type 1 Diabetes: A  
2 Randomized Cross-Over Study

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31

32 ABSTRACT

33

34 *Objective* – To determine the effect of morning exercise in fasting condition, versus afternoon  
35 exercise on blood glucose responses to resistance exercise (RE).

36 *Research Design and Methods* – Using a randomized crossover design, 12 participants with type  
37 1 diabetes [9 females, aged  $31 \pm 8.9$  years, diabetes duration  $19.1 \pm 8.3$  years, HbA1c =  $7.4 \pm$   
38  $0.8\%$  ( $57.4 \pm 8.5$  mmol/mol)] performed ~40 minutes of RE (three sets of eight repetitions, seven  
39 exercises, at the individual's pre-determined eight repetition maximum) either at 7 am (fasting)  
40 or 5 pm. Sessions were performed at least 48 hours apart. Venous blood samples were collected  
41 immediately pre-, immediately post-, and 60-minutes post-exercise. Interstitial glucose was  
42 monitored overnight post-exercise by continuous glucose monitoring (CGM).

43 *Results* – Data are presented as mean  $\pm$  SD. Blood glucose rose during fasting morning exercise  
44 ( $9.5 \pm 3.0$  to  $10.4 \pm 3.0$  mmol/L) while it declined with afternoon exercise ( $8.2 \pm 2.5$  to  $7.4 \pm 2.6$   
45 mmol/L;  $p=0.031$  for time by treatment interaction). Sixty minutes post-exercise, blood glucose  
46 concentration was significantly higher after fasting morning exercise compared to afternoon  
47 exercise ( $10.9 \pm 3.2$  vs.  $7.9 \pm 2.9$ ;  $p=0.019$ ). CGM data indicated more glucose variability ( $2.7 \pm$   
48  $1.1$  vs.  $2.0 \pm 0.7$  mmol/L;  $p=0.019$ ) and more frequent hyperglycemia (12 events vs. 5 events;  
49  $p=0.025$ ) after morning RE compared to afternoon RE. There were two hypoglycemic events  
50 after morning RE compared to four after afternoon RE (NS).

51 *Conclusions* – Morning (fasting) RE is associated with distinctly different blood glucose  
52 responses and post-exercise profiles from afternoon RE.

53 PRECIS

54 Morning resistance exercise performed while fasting increases blood glucose while resistance  
55 exercise performed later in the day decreases blood glucose in individuals with type 1 diabetes.

56 INTRODUCTION

57

58 Resistance exercise provides the human body with a multitude of benefits including  
59 maintaining and/or building muscle mass, strength and metabolism (1), increasing resting energy  
60 expenditure (2), improving bone mineral density (3), decreasing the risk of type 2 diabetes by  
61 increasing insulin sensitivity (4) and improving cardiovascular health (5,6). In individuals with  
62 type 1 diabetes, resistance exercise is also associated with smaller changes in blood glucose  
63 during activity than aerobic exercise (7), seems to provide more stable blood glucose levels in  
64 the hour post-exercise (7), and may offer a protective effect on blood glucose levels when  
65 performed immediately prior to aerobic exercise (8).

66 While they are few and have small sample sizes, some studies examining the acute  
67 effects of resistance exercise on blood glucose have had divergent outcomes. Studies of  
68 afternoon resistance exercise by Yardley et al. [consisting of three sets of eight repetitions, at the  
69 participants' eight repetition maximum (8RM) of seven different exercises] have been associated  
70 with average declines in blood glucose of approximately 1.5 mmol/L (7,8) during exercise and  
71 an increased risk of nocturnal hypoglycemia (7). Conversely, studies performed by Turner et al.,  
72 (9) where participants performed a very similar protocol while in a fasting state in the morning,  
73 have resulted in either a mean increase in blood glucose levels of 1.5 mmol/L when two sets of  
74 10 repetitions were performed at 60% of the participants' 1RM (a similar intensity to the Yardley  
75 et al. studies) (7,8), or no significant change in blood glucose in either direction when three sets  
76 of eight repetitions (also at 60% of the 1 RM) were performed (10). As CGM data were not  
77 reported for the morning exercise studies, the risk of post-exercise nocturnal hypoglycemia  
78 remains unquantified. Whether the divergent responses are due to differences in participant  
79 characteristics, or differences in metabolism in the fasted versus fed state is uncertain.

80           The present study sought to determine whether or not individuals with type 1 diabetes  
81 would have different blood glucose responses to a standardized resistance exercise protocol if  
82 exercise was performed in the morning while fasting, or in the afternoon. By using a randomized,  
83 cross-over design, participant characteristics are removed as a potential confounder. We  
84 hypothesized that fasting exercise would be associated with smaller declines (if any) in blood  
85 glucose and a lower risk of post-exercise nocturnal hypoglycemia, than when the same resistance  
86 exercise protocol is performed in the late afternoon.

87

## 88 METHODS

89           The study was approved with the University of Alberta research ethics board and all  
90 participants provided informed consent prior to their participation. Twelve non-smoking, non-  
91 obese, complication-free, recreationally active individuals with type 1 diabetes were recruited for  
92 the study. The study was carried out in accordance with the principles of the Declaration of  
93 Helsinki. Participants were required to be between the ages of 18 and 50, to have been diagnosed  
94 with type 1 diabetes for at least one year, to have HbA1c levels below 9.9% (84.7 mmol/mol)  
95 and to be habitually active, performing both aerobic and resistance exercise. Potential  
96 participants were excluded if they had any condition that would render exercise or physical  
97 activity contraindicated (e.g. severe hypoglycemia unawareness, autonomic neuropathy, severe  
98 proliferative, retinopathy, joint or limb injuries preventing weight-bearing activity/requiring limb  
99 immobilization, etc.), if they were performing shift work, or if they were using any medication  
100 (other than insulin) that would impact blood glucose levels. There were no exclusions based on  
101 insulin regimen or mode of insulin administration.

102

103 Experimental Design

104           Research took place in the Physical Activity and Diabetes Laboratory at the Alberta  
105 Diabetes Institute. Participants attended a baseline session where written consent was obtained.  
106 During the same session the participants' aerobic fitness was estimated using a modified  
107 Austrand-Rhyming Submaximal Cycle Ergometer test (11) on an electronically braked cycle  
108 ergometer (Monark Ergomedic 894E; Monark, Varberg, Sweden). A test of muscular strength  
109 (8RM) was performed in order to determine the maximum weight that participants could lift  
110 eight times with good form for the exercises included in the protocol (chest press, leg press,  
111 seated row, leg curl, shoulder press and lat pulldown) all of which were performed on weight-  
112 lifting machines to ensure consistent form and participant safety. At the end of the baseline  
113 sessions the order of exercise sessions was determined by flipping a fair coin before sessions  
114 were scheduled.

115           The study involved a randomized, open-label, cross-over design with two rounds of  
116 testing. One testing session took place at 7 am (as per Turner et al.) (10) with participants in the  
117 fasting state, and the other took place at 5 pm (as per Yardley et al.) (7), with at least 48 hours  
118 between sessions. At least 24 hours prior to the first session participants came to the lab for the  
119 insertion of an Enlite™ sensor with an iPro2® (Medtronic, Northridge, CA) blinded continuous  
120 glucose monitoring (CGM) system. Participants were provided with a log and asked to record  
121 their food intake and insulin dosage over the six days of CGM wear. While wearing the CGM,  
122 participants were asked to match their day to day food intake and insulin dosage (including  
123 adjustments for exercise) as closely as possible, while avoiding alcohol and strenuous exercise.  
124 The study also provided participants with a pedometer (Yamax DigiWalker 200, Yamax  
125 Corporation, Tokyo, Japan) to determine their background physical activity (daily step count),

126 and a OneTouch<sup>®</sup> Ultra<sup>®</sup>2 glucose meter and test strips (LifeScan Milpitas, CA, USA) to record  
127 four daily capillary glucose tests for the purpose of calibrating the CGM. For female participants,  
128 both tests occurred in the same phase of the menstrual cycle.

### 129 Experimental Sessions

130 Participants arrived at the laboratory at 6 am in a fasting state for the morning exercise  
131 session (as per Turner et al.) (10) and at 4 pm for the afternoon session. Upon arriving at the lab  
132 for the afternoon session participants consumed a standardized snack (Glucerna Snack Bar,  
133 Abbott Laboratories, Abbott Park, IL), as per Yardley et al. (7). Exercise started at 7 am and 5  
134 pm respectively for the morning and afternoon exercise conditions. As in previous studies,  
135 resistance exercise was performed with a 2-second count for both the eccentric and concentric  
136 phases of the lift. Participants performed three sets of eight repetitions (8RM) of all exercises  
137 with 90 seconds rest in between. The exercises ensured that all major muscles groups were  
138 targeted, and included leg press, bench press, leg curl, lat pulldown, abdominal crunches,  
139 shoulder press and seated row. Exercise lasted approximately 43 minutes, and was followed by a  
140 60-minute period of seated recovery in the lab. In line with exercise guidelines of both Diabetes  
141 Canada (12) and the American Diabetes Association (13), participants using insulin pumps were  
142 asked to decrease their basal rate by 50% starting one hour before exercise and maintained until  
143 the end of exercise. Individuals using multiple daily injections were asked to reduce their long-  
144 acting insulin dose by 10% the day prior to exercise. The importance of repeating the same  
145 adjustments for both sessions was emphasized.

146 Prior to exercise, intravenous catheters were inserted in the antecubital vein for the  
147 purpose of blood sampling. Blood samples were drawn into 10-mL EDTA vacutainer tubes  
148 immediately before exercise, at the end of exercise and 60 minutes post-exercise. Tubes were

149 immediately centrifuged at 1500 x g for 10 minutes at 4°C to extract plasma. Samples were then  
150 stored in a -80°C freezer until batch analysis could be completed. Plasma glucose values were  
151 determined using the hexokinase timed end point method on a Siemens ADVIA 1800 system  
152 with Siemens ADVIA chemistry glucose hexokinase\_3 (GLUH-c) concentrated reagent.

153 Participants were asked to continue wearing their CGM for a full 24 hours after the lab  
154 testing sessions. CGM units were collected from participants and uploaded to the Medtronic  
155 Carelink web-based platform. Data were exported as Excel spreadsheets. EasyGV<sup>®</sup> Version  
156 9.0.R2 ([www.easygv.co.uk](http://www.easygv.co.uk)) was used to assess interstitial glucose means, and glucose variability  
157 expressed as mean absolute glucose change (MAG). Percentage of time spent in hypoglycemia  
158 ( $\leq 3.9$  mmol/L) and hyperglycemia ( $\geq 10$  mmol/L) were calculated (7) as well as the frequency of  
159 hypoglycemic and hyperglycemic excursions.

#### 160 Statistical Analysis

161 Plasma glucose data were analysed using a repeated measures ANOVA in order to  
162 examine the whether there were significant changes in blood glucose, and if these differed  
163 depending on the timing of the exercise. CGM data that were normally distributed (mean,  
164 standard deviation, MAG) were compared using paired t-tests and are expressed as mean  $\pm$  SD.  
165 Non-normally distributed data (percent of time in hypo/hyperglycemia, frequency of  
166 hypoglycemic events were assessed using Wilcoxon Signed Rank tests, and are expressed as  
167 median  $\pm$  IQR. Statistical analyses were performed using SPSS 25.0 software (IBM, Amonk,  
168 New York, USA). We chose to examine the 6-hour window immediately post-exercise, as this is  
169 when the impact of exercise on blood glucose in individuals with type 1 diabetes is most clearly  
170 seen (7,14). We also chose to assess the overnight period (between 11 pm and 6 am) as nocturnal  
171 hypoglycemia is one of the greatest post-exercise risks for individuals with type 1 diabetes.



172

## 173 RESULTS

174 Participant characteristics can be found in Table 1. Participants were habitually  
175 physically active and were not taking any medications (other than insulin) that would alter  
176 glucose metabolism. Eight of the 12 participants were using insulin pumps, while four were  
177 using multiple daily insulin injections. Of the participants using multiple daily injections, three  
178 were using glargine as basal insulin (administering their dose at bedtime), and one was using  
179 levemir (late morning administration). These basal insulins were combined with either insulin  
180 lispro (n=2) or insulin aspart (n=2). Participant food logs indicate that all participants had  
181 consumed their evening meal at least 8 hours prior to the morning exercise session. Similarly, on  
182 afternoon exercise days, lunch was consumed at least three (n=4) or four (n=8) hours before the  
183 exercise session.

184 Eleven of the 12 participants had identical insulin adjustments prior to both morning and  
185 afternoon exercise, and one participant decreased the basal rate on their insulin pump by 50% for  
186 the afternoon exercise, but made no adjustment for morning exercise. Background physical  
187 activity, measured as pedometer step counts, was not significantly different between conditions  
188 on either the testing day, or the day after testing. None of the participants experienced dangerous  
189 declines in blood glucose, and it was not necessary to provide glucose supplements to  
190 participants at any point during the testing.

191 There was no difference in blood glucose levels at the start of exercise between the  
192 morning ( $9.5 \pm 3.0$  mmol/l) and afternoon ( $8.2 \pm 2.5$  mmol/l;  $p = 0.289$ ) conditions. Changes in  
193 blood glucose were not significant during either the morning or the afternoon exercise sessions  
194 (effect of time  $p = 0.405$ ). Blood glucose levels were higher throughout exercise and recovery

195 after morning exercise (effect of treatment  $p = 0.041$ ). A significant time by treatment interaction  
196 ( $p = 0.031$ ) indicated that blood glucose levels were following different trajectories throughout  
197 the testing session, with morning exercise producing a consistent increase (from  $9.5 \pm 3.0$  to  $10.4$   
198  $\pm 3.0$  mmol/L) in blood glucose (Figure 1), while afternoon exercise caused an initial decline in  
199 blood glucose (from  $8.2 \pm 2.5$  to  $7.4 \pm 2.6$  mmol/L) during exercise, with a return almost to  
200 baseline during the 60-minute recovery. At the end of 60 minutes of recovery, blood glucose  
201 levels were significantly higher after morning exercise ( $10.9 \pm 3.2$  vs.  $7.9 \pm 2.9$ ;  $p=0.019$ ).

### 202 Continuous Glucose Monitoring

203 Complete CGM data sets were available from 10 participants. Mean CGM glucose in the  
204 6 hours post-exercise (Figure 2a) was not significantly different between the morning and  
205 afternoon exercise sessions (am:  $11.5 \pm 3.3$ , pm:  $9.0 \pm 3.9$  mmol/L;  $p = 0.473$ ). Mean absolute  
206 glucose change (MAG) for 6 hours post-exercise was greater ( $p = 0.019$ ) after morning exercise  
207 ( $2.7 \pm 1.1$  mmol/L) compared to afternoon exercise ( $2.0 \pm 0.7$ ) indicating more glycemetic  
208 variability after the fasting exercise session. Where the nocturnal period (midnight until 6 am)  
209 was concerned (Figure 2b), there were no significant differences between morning and afternoon  
210 exercise with respect to mean CGM glucose ( $p = 0.635$ ) or glucose variability as expressed by  
211 MAG ( $p = 0.280$ ).

212 There were no differences between morning and afternoon resistance exercise with  
213 respect to the frequency of hypoglycemia in the 6 hours post-exercise or in the nocturnal period  
214 after exercise (Table 2). Hyperglycemia was more common following morning exercise ( $p =$   
215  $0.011$ ). The percentage of time spent in hypoglycemia, according to CGM, was similar ( $p =$   
216  $0.109$ ) in both groups in the 6 hours immediately post-exercise, as well as during the nocturnal ( $p$   
217  $= 0.345$ ) post-exercise period (Table 2). Time spent in hyperglycemia, on the other hand, was

218 higher in the first six hours after morning exercise [62.0% (36.0-82.6%)] compared to the first  
219 six hours post-afternoon exercise [11.6% (0-36.0%);  $p = 0.037$ ]. The difference between the  
220 amount of time spent in range following morning exercise compared to afternoon exercise  
221 approached statistical significance ( $p = 0.064$ ) with afternoon exercise showing more favourable  
222 blood glucose profiles (less hyperglycemia) in the six hours following exercise. There were no  
223 differences between morning and afternoon exercise with respect to the percent of time spent in  
224 range during the overnight period.

225

## 226 DISCUSSION

227 As hypothesized, fasting resistance exercise in the morning and afternoon resistance  
228 exercise have distinctly different effects on blood glucose levels with the former favouring an  
229 increase in blood glucose, and the latter a decrease. Due to a large amount of variability in the  
230 responses, the overall changes in blood glucose for both of these sessions were not statistically  
231 significant themselves. The time-by-treatment interaction, however, indicates that blood glucose  
232 trajectories over time were in fact significantly different between the morning (an increasing  
233 trend) and afternoon (a decreasing trend) exercise sessions.

234 These outcomes are consistent with what has been observed in previous exercise studies  
235 in type 1 diabetes. While they are not numerous, studies where resistance exercise was  
236 performed in the afternoon resulted in a decline in blood glucose levels (7,8), while studies  
237 where exercise was performed while fasting in the morning (after evening administration of  
238 glargine) either showed no change (10) or a mean increase (9) in blood glucose in response to  
239 resistance exercise. Aerobic exercise studies examining the impact of time of day have found  
240 similar results with repeated measures designs. A study by Ruegamer et al., (15) found distinctly

241 different patterns of blood glucose response to 30 minutes of stationary cycling at 60% of the  
242 participant's aerobic capacity performed while fasting in the morning, or fed in the late  
243 afternoon. Morning exercise produced an increase in blood glucose from  $6.7 \pm 0.4$  mmol/L to  $9.1$   
244  $\pm 0.4$  mmol/L ( $p < 0.01$ ), while a small non-significant decline was found in the afternoon. More  
245 importantly, the inclusion of a non-exercise control day showed that blood glucose levels in  
246 these participants were otherwise stable at this time of day. Meals, snacks and insulin injections  
247 (including basal ultralente in the evening) were also kept consistent across sessions by the  
248 research team. Recently, a study observing both moderate aerobic exercise (65%  $VO_{2peak}$  for 30  
249 minutes) and high intensity interval exercise (6 X 1 minute at 100%  $VO_{2peak}$ , with 1 minute of  
250 recovery in between, for a total of 17 minutes of exercise), performed while fasting did not find  
251 any declines in blood glucose for either type of exercise (16). These results are in contrast to  
252 several studies of aerobic (7,14,17-22) and high intensity interval (17-20,22) exercise in fed  
253 participants where declines in blood glucose were observed, in spite of starting blood glucose  
254 levels being similar to those seen in the fasted exercise studies.

255         A strong potential cause of the divergent blood glucose trends in the present study would  
256 be a difference in circulating insulin. While similar adjustments for exercise were made by all  
257 but one participant, if exercise is taking place 8 to 10 hours after the last meal, it is likely that  
258 only basal insulin will be present. Conversely, when afternoon exercise is performed, it is  
259 generally taking place within 4 to 5 hours of a meal, at which point there may still be some of the  
260 previous meal's bolus remaining in circulation. Unfortunately we were unable to measure insulin  
261 levels as part of this study, as this would have provided more concrete evidence as to insulin's  
262 involvement. It should be noted, however, that when performing afternoon exercise the  
263 participants were provided with a standardized pre-exercise snack (Glucerna snack bar – 19g of

264 carbohydrate) one hour before exercise similar to previous studies by Yardley et al. (7,8), and  
265 consistent with current guidelines for exercise in individuals with type 1 diabetes (13,23). Eight  
266 out of twelve participants did not bolus for this snack, while the final four opted for a reduced  
267 (50%) bolus in order to prevent hyperglycemia. This snack should have, to some extent,  
268 attenuated the impact of potentially higher insulin levels during the afternoon exercise session.

269         Individuals with type 1 diabetes often experience a period of high blood glucose in the  
270 early morning, which is generally referred to as the “dawn phenomenon” (24). This early  
271 morning rise in blood glucose has often been attributed to an increase in growth hormone (25-  
272 27). While starting blood glucose levels in the present study were not significantly higher during  
273 the fasting session compared to the afternoon session, the presence of a higher level of growth  
274 hormone could play a role in the divergent blood glucose outcomes observed in this study.  
275 Previous studies of exercise in type 1 diabetes have suggested that higher growth hormone levels  
276 may have a glucose sparing (8,28) effect by stimulating lipolysis (29). As we did not have the  
277 means to measure growth hormone for the present study, this possibility remains speculative.

278         In addition to higher levels of growth hormone, there is also evidence to suggest that  
279 insulin sensitivity is lower in the morning compared to the afternoon in individuals with type 1  
280 diabetes (30,31) and that there is less suppression of endogenous glucose production (30). A  
281 recent meta-analysis of metabolic responses to fed and fasted exercise also found that fasting  
282 exercise is associated with a higher level of circulating free fatty acids post-exercise than fed  
283 exercise (32). As elevated free fatty acids could lead to an even greater degree of insulin  
284 resistance (33), this may explain not only the change in glucose during exercise, but also the  
285 resulting post-exercise hyperglycemia.

286

287 Besides the fasting versus non-fasting state being a potential driver for the divergent  
288 blood glucose outcomes, it is also possible that, in previous resistance exercise studies, small  
289 samples sizes (varying from  $n=8$  (9,10) to  $n=12$  (7,8)), and differences in the sample composition  
290 had an effect on blood glucose outcomes. Participants in the studies of afternoon exercise by  
291 Yardley et al. had lower mean HbA1c [ $7.1\pm 1.1\%$  ( $54\pm 10$  mmol/mol) (7,8) versus  $8.7\pm 1.1\%$   
292 ( $72\pm 10$  mmol/mol) (9,10)] and also started exercise with blood glucose levels on average  $\sim 2.0$   
293 mmol/L lower than those who took part in the studies of fasting resistance exercise. The  
294 participants in all studies, however, consisted mostly of males (no more than 2 females in any  
295 one studies), who were physically active, with mean ages for the samples falling between  $32 \pm$   
296  $15.3$  (8) and  $38 \pm 6$  yrs (10). While the sample in the present study was similar to previous studies  
297 in terms of sample size ( $n=12$ ), age ( $31.3 \pm 8.9$  years), and HbA1c [ $7.4 \pm 0.8$  ( $57.4\pm 8.5$   
298 mmol/mol)] it had a greater proportion of female participants (9 out of 12). In spite of this major  
299 difference, trajectories of blood glucose change were similar to previous studies with respect to  
300 fasting versus fed exercise, and the randomized repeated measures design removes the potential  
301 confounding effect of physiological factors such as age, sex, and fitness level.

302 Strengths of this study include its randomized repeated measures design, the use of  
303 blinded CGM to assess post-exercise blood glucose levels, and the strict timing/structure of the  
304 exercise sessions. The interpretation of the data are limited, however, due to the lack of hormone  
305 measurements necessary to determine the cause of the divergent blood glucose outcomes during  
306 morning and afternoon resistance exercise. Thus it cannot be determined if the respective blood  
307 glucose trajectories are due to a) natural increases in blood glucose in the morning due to the  
308 “dawn phenomenon”, b) differences in circulating insulin due to administration in the hours  
309 before exercise, or c) potential differences in insulin sensitivity throughout the day. It should be

310 noted, however, that both scenarios tested in this study are ones that individuals with type 1  
311 diabetes who exercise are likely to encounter, thus making the study design ecologically valid.  
312 An additional limitation is that the sample is small (n=12), consisting of relatively young  
313 healthy, habitually active individuals with type 1 diabetes, and may therefore not be applicable to  
314 those who are sedentary or of advancing age. It is also possible that those using multiple daily  
315 injections could have slightly different responses than those using insulin pumps, however this  
316 could not be determined from such a small sample. Further studies are required to elucidate  
317 potential differences in these insulin delivery methods, as well as the potential impact of the  
318 timing of basal insulin injections in those using multiple daily injections.

319         The results of this study help expand the existing evidence related to type 1 diabetes and  
320 exercise, and offer information on two different options related to exercise timing. The  
321 magnitude of counter-regulatory (especially catecholamine) responses to resistance exercise may  
322 vary greatly by age and level of fitness (young fit individuals are likely to have the most  
323 pronounced response) (34). As such some individuals will struggle with hyperglycemia during  
324 and after resistance exercise. These individuals may want to consider performing resistance  
325 exercise later in the day where the activity seems to result in less hyperglycemia. Conversely, for  
326 those who struggle with hypoglycemia during exercise, resistance exercise performed in the  
327 morning may be a better option. It should be noted, however, that the present study only  
328 examined one intensity of resistance exercise (3 sets of 8 repetitions at the participants' 8 RM),  
329 and results may vary with different durations and intensities of resistance exercise. Further  
330 studies to examine the impact of time of fasting on blood glucose responses to a variety of  
331 resistance exercise programs in individuals with type 1 diabetes are warranted. Finally, it should  
332 be noted that several of the current exercise guidelines recommend a decrease in basal insulin for

333 physical activity and exercise. The results of the present study and of those discussed above,  
334 would indicate that this recommendation may not be appropriate for activities performed in a  
335 fasting state. However, patients should always be advised to monitor their glucose closely upon  
336 starting or changing an exercise routine, to ensure that they understand their individual  
337 responses.

338

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343

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346

#### 347 CONTRIBUTION STATEMENT

348 JEY contributed to the conception and design of the project, as well as data collection and  
349 manuscript drafting and editing. SRT contributed to data collection and analysis in addition to  
350 drafting and editing of the manuscript.

351

352 JEY is the guarantor of this work. Data related to the study were presented at the Diabetes  
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461 TABLE AND FIGURE LEGENDS

462

463 *Table 1 - Participant characteristics*

464

465 *Table 2.* Continuous glucose monitoring data the 6-hours post-exercise, as well as overnight  
466 (midnight to 6 am) after exercise

467

468

469 *Figure 1 - Change in blood glucose during exercise and recovery. Gray boxes represent morning*  
470 *exercise, black boxes represent evening exercise. Data are presented as mean  $\pm$  SEM*

471

472

473 *Figure 2 - CGM glucose. Panel A represents CGM glucose in the 6 hours following exercise.*  
474 *Panel B represents overnight blood glucose values. Data are presented as mean  $\pm$  SEM.*

*Table 1 - Participant characteristics*

Characteristic	N or mean $\pm$ SD
Male/Female	3/9
Age (yrs)	31.3 $\pm$ 8.9
Height (m)	1.7 $\pm$ 0.1
Weight (kg)	78.1 $\pm$ 15.0
BMI (kg·m <sup>-2</sup> )	26.6 $\pm$ 3.8
Predicted VO <sub>2</sub> max (mL O <sub>2</sub> · kg <sup>-1</sup> )	39.1 $\pm$ 8.2
HbA1c (%)	7.4 $\pm$ 0.8
HbA1c (mmol/mol)	57.4 $\pm$ 8.5
Diabetes Duration (years)	19 $\pm$ 8
MDI/CSII	4/8
Systolic Blood Pressure (mmHg)	117 $\pm$ 12
Diastolic Blood Pressure (mmHg)	81 $\pm$ 9
Resting Heart Rate (bpm)	74 $\pm$ 13

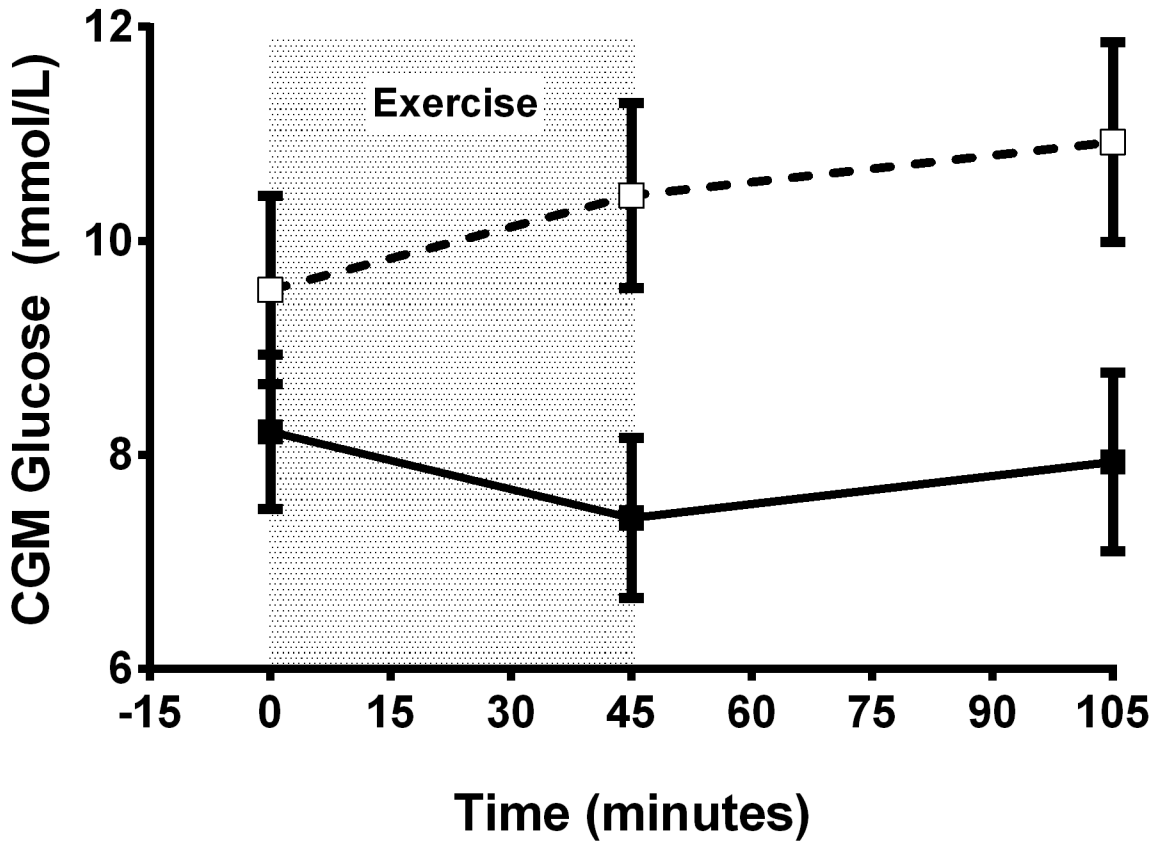
MDI = multiple daily insulin injections; CSII = continuous subcutaneous insulin infusion

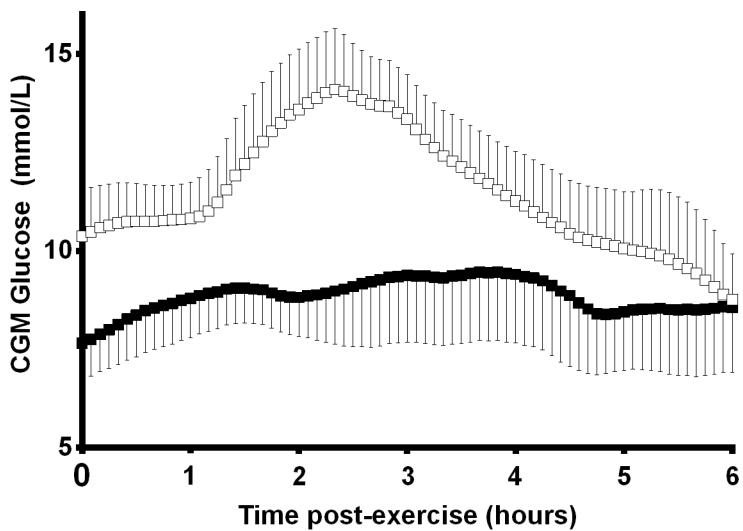
(insulin pump)

Table 2. Continuous glucose monitoring data the 6-hours post-exercise, as well as overnight (midnight to 6 am) after exercise

	Morning	Afternoon	p-values
6-hr mean glucose (mmol/L)	11.5 ± 3.3	9.1 ± 3.9	0.126
6-hr MAG (mmol/L)	2.7 ± 1.1	2.0 ± 0.7	0.076
Nocturnal mean glucose (mmol/L)	7.4 ± 1.7	8.6 ± 4.1	0.428
Nocturnal MAG	1.2 ± 0.8	1.0 ± 0.1	0.105
% high (6 hr)	61.64 [36.30-82.53]	11.64 [0.00-36.03]	0.037
% low (6 hr)	0.00 [0.00-0.00]	0.00 [0.00-4.17]	0.181
% in range (6 hr)	33.56 [27.37-62.84]	82.88 [55.01-100.00]	0.064
# of events ≤ 3.9 mmol/l (6hr)	1	4	0.081
# of events ≥ 10 mmol/l (6hr)	12	5	0.025
% high (nocturnal)	0.00 [0.00-4.41]	15.29 [0.00-47.94]	0.205
% low (nocturnal)	0.00 [0.00-0.00]	0.00 [0.00-6.48]	0.418
% in range (nocturnal)	100.00 [94.41-100.00]	65.17 [45.59-97.93]	0.107
# of events ≤ 3.9 mmol/l (nocturnal)	2	4	0.343
# of events ≥ 10 mmol/l (nocturnal)	4	7	0.434

Data are mean±SD for mean glucose and MAG, otherwise data are presented as median ± [IQR]. SD=standard deviation, MAG = mean absolute glucose change, % high = % of time spent ≥10.0 mmol/L; %low = % of time spent ≤3.9 mmol/L; % in range = % of time spent between 3.9 and 10.0 mmol/L



**A****B**