Influence of Snack Intake on Cardiac Autonomic Nervous System in **Patients with Type 2 Diabetes**

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Abstract

Food intake induces significant changes in heart rate (HR) and heart rate variability (HRV) among healthy individuals. It is known that the cardiac autonomic nervous system (CANS) is impaired in patients with type 2 diabetes. The main goal of this pilot study was to investigate whether snack intake can affect CANS in this patient population. For this purpose, ECG was recorded pre- and post-snack intake in 15 volunteers with type 2 diabetes. Mean HR and short-term HRV measures (e.g., SDNN, RMSSD, percentage of power in LF and HF band, and LF/HF) were extracted and compared using the Wilcoxon signed-rank test between pre- and post-snack intake. From pre- to post-snack intake, mean glucose level increased by 24.1 points (p=0.10) while mean heart rate increased by 1.9 beats per minute (p=0.30). Changes in SDNN, RMSSD, power in LF band (%), power in HF band (%), and LF/HF were not existent (p=0.39, 0.80, 0.45, 0.98, 0.36, respectively). As expected, an increase in glucose count was observed shortly after snack intake. No cardiac neural response, however, was observed in our sample of type 2 diabetics.

1. Introduction

Food intake induces significant changes to heart rate (HR) [1] and heart rate variability (HRV) among healthy individuals [2]. While cardiac neural response is impaired in type 1 diabetics who exhibit inconsistent patterns of heart rate variability (HRV) after eating [3], less is known about the influence of snack intake on cardiac neural response in individuals with type 2 diabetes. Furthermore, it is known that the cardiac autonomic nervous system (CANS) is impaired in patients with type 2 diabetes and low HRV is a risk factor for sudden cardiac death in this group [4, 5]. We hypothesized that snack intake can negatively affect CANS in this population group. Therefore, the objective of this study was to evaluate CANS response in type 2 diabetics after consuming a light snack.

2. Data and Method

This study is a secondary analysis of a retrospective data that was recorded in 2012.

2.1. Data and Study Protocol

Fifteen volunteers with type 2 diabetes (Age: 44.9±7.4, BMI: 27.4±4.6, 14% male) with no history of cardiovascular disease participated in this study. After signing a consent form, blood glucose level and electrocardiography (ECG) data were collected pre- and post-snack intake. ECG was recorded using the Novin s1800 monitoring device (Pooyandegan Rah Saadat Corporation, Tehran-Iran) and blood glucose level was measured using Glucosure Star (ApexBio, Taiwan).

ECG for baseline condition (pre-snack intake) was recorded for 3 minutes and then blood glucose was measured at the end of the baseline measurement. Participants were then asked to consume a light snack and after 15 minutes post-snack intake, ECG was recorded for a duration of 6 minutes. Blood glucose was measured for a second time at the end of the post-snack intake measurement. During the measurements, all participants were seated on a chair. The whole recorded ECG during the pre-snack intake (3 minutes) and the first 3 minutes of ECG recording during the post-snack intake were used for HR and HRV analysis. The study protocol is visualized in Figure 1.



Figure 1. Study protocol

2.2. Method

2.2.1. QRS Detection

Baseline wander was corrected using a moving average filter [6] and then Pan–Tompkins algorithms were used for QRS detection [7]. Detected QRS peaks were verified by manual inspection and were corrected if it was required. Using corrected QRS peaks, inter-beat (RR) interval series were created for HR and HRV analysis.

2.2.2. Heart Rate and HRV

RR intervals were used for estimation of different HR and HRV features for 3 minute recordings pre- and postsnack intake. The Kubios HRV Software was used for HRV analysis [8]. HR (minimum, maximum, and mean) and the following HRV measures were calculated in this study:

- Time-domain measures: standard deviation of RR intervals (SDNN), root mean square of successive differences (RMSSD), the relative number of successive intervals differing more than 50ms (pNN50).

- Frequency-domain measures: percentage of power in low frequency (LF) band and high frequency (HF) band as well as LF/HF. HRV spectrum was estimated with FFT based Welch's periodogram method.

- Poincare plot measures: axis of ellipse fitted to points' distribution in Poincare plot (SD1 and SD2).

- Nonlinear measures: approximate and sample entropy.

Please refer to [8-10] for details on calculation of above-mentioned parameters.

2.2.3. Statistical Analysis

Means and standard deviations for continuous variables are reported. Pre- and post-test parameters were compared and tested using the Wilcoxon signed-rank test [11] at the significance level of 0.05. For comparison of mean differences between pre- and post-snack intake, cohen's d effect size was calculated; values of 0.2, 0.5, and 0.8 were considered as small, moderate, and large effect sizes, respectively [12]. SAS version 9.3 was used for all statistical analysis (SAS Institute, Inc., Cary, NC).

3. Results

Means and standard deviations of HR and HRV measures pre- and post-snack intake are reported in Table 1. The obtained p-values are shown in Table 1 for extracted parameters. From pre- to post-snack intake,

mean glucose level increased by 24.1 points (p=0.10) while mean heart rate increased slightly by 1.9 beats per minute (p=0.30). Changes in SDNN, RMSSD, power in LF band (%), power in HF band (%), and LF/HF were not existent (p=0.39, 0.80, 0.45, 0.98, 0.36, respectively). Approximate entropy and sample entropy also did not change significantly from pre- to post-snack intake (p=0.40 and 0.14, respectively), although increasing trends were noted. Cohen's d effect size for extracted parameters is reported in Table 2. Except for SD2 that had a moderate effect size (d=0.52), all other parameters had small to almost moderate effect sizes.

Table 1. Means and standard deviations of HR and HRV measures pre- and post-snack intake along with p-value for paired comparison. \checkmark and \uparrow shows the decreasing and increasing trend, respectively.

	Pre-snack intake	Post-snack intake	P-value	
Blood Glucose, [mg/dl]	174.07±45.90	198.13±52.06	0.10 个	
Min HR, [bpm]	71.73±9.26	72.87±12.51	0.57 🛧	
Max HR, [bpm]	90.27±15.07	90.00±12.45	0.83 🗸	
Mean HR, [bpm]	77.98±9.26	79.92±10.62	0.30 个	
SDNN, [ms]	32.52±20.31	24.74±11.74	0.39 🗸	
RMSSD, [ms]	22.17±14.32	20.02±14.61	0.80	
pNN50, [%]	3.51±5.70	5.02 ± 9.49	0.96 个	
pLF, [%]	32.53±15.67	36.40±10.20	0.45 🔨	
pHF, [%]	19.93±11.46	19.07 ± 12.87	0.98 🗸	
LF/HF	2.31±1.85	3.12±2.33	0.36 个	
SD1, [ms]	15.71±10.15	14.19±10.35	0.78 🗸	
SD2, [ms]	42.91±27.30	31.58±13.82	0.21 ↓	
Approximate Entropy	0.93±0.16	0.97±0.12	0.40 个	
Sample Entropy	1.56±0.38	1.73±0.35	0.14 个	
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Min: Minimum, Max: Maximum

Table 2. Cohen's d (d) for pairwise comparison of extracted parameters for between group differences.

	Cohen's d effect size
Blood Glucose, [mg/dl]	0.49
Minimum HR, [bpm]	0.10
Maximum HR, [bpm]	0.02
Mean HR, [bpm]	0.19
SDNN, [ms]	0.47
RMSSD, [ms]	0.15
pNN50, [%]	0.19
pLF, [%]	0.29
pHF, [%]	0.07
LF/HF	0.39
SD1, [ms]	0.22
SD2, [ms]	0.52
Approximate Entropy	0.28
Sample Entropy	0.47

4. Conclusion

In this article, blood glucose, heart rate, and heart rate variability for patients with type 2 Diabetes were compared pre- and post-snack intake. As expected, an increase in glucose count was observed shortly after snack intake. No cardiac neural response, however, was observed in our sample of type 2 diabetics. Recognizing the limitation of a small sample size, our findings suggest that HR and HRV in type 2 diabetics are insensitive to eating a snack.

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