Better cardiorespiratory fitness associated with favourable metabolic control and health-related quality of life in youths with type 1 diabetes mellitus

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Abstract

Objective: to explore the relationship among health-related quality of life (HRQoL), clinical variables, anthropometric measures, physical activity and cardiorespiratory fitness in children and adolescents with type 1 diabetes. Furthermore, we aimed to find predictors of HRQoL and metabolic control. Methods: A total of 106 patients (sex ratio: 1:1) with mean HbA1c of 8.55 (± 1.44) % and diabetes duration of 5.15 (± 3.24) years were assessed. The average age was 13.22 (± 3.08) years. Results: We observed statistically significant negative medium correlation between HbA₁c and VO₂max (r = –0.343; p < 0.01). There was statistically significant small positive correlation between the HRQoL and the maximal oxygen consumption (r = 0.208; p = 0.032). We found no significant correlation between the HbA₁c and the patients’ HRQoL. In the multiple linear regression analysis both the better metabolic control and the HRQoL was predicted by the VO₂max, other variables had no effect. Physical activity level did not explain the HRQoL. Boys had significantly better HRQoL and less skinfold thickness than girls. Conclusion: Better cardiorespiratory fitness associated with both favourable metabolic control and better HRQoL of diabetic youths. Regular aerobic exercise improves the young patients’ physical fitness and overall health status, and perception of health-related quality of life, respectively.

Keywords: type 1 diabetes, metabolic control, health-related quality of life, cardiorespiratory fitness, physical activity
Type 1 diabetes mellitus (T1DM) is one of the commonest chronic diseases in childhood affecting more and more children worldwide (6). The diabetes care is complex and it takes continuous management from the healthcare experts, caregivers and from the patients. Treatment for T1DM involves the continual use of insulin therapy with appropriate diet and regular physical exercise. The main goal of the diabetes management is to maintain the blood glucose level in the near-normal range in order to prevent the long-term complications. Attention has shifted from treatment to improving overall health condition. The health-related quality of life (HRQoL) is the person’s own perceived state of health. It has been recognised as an important paediatric outcome measurement. We observed controversial results regarding the HRQoL and metabolic control in the literature. Some studies found positive correlation between better HRQoL and good glycemic control (12, 14, 28), others had no evidence about it (7, 9, 17). Physical activity is essential to health in people of all ages, but for children plays a key role for their physical and mental health. The regular exercise in diabetes management is part of the treatment. It is recommended primary for patients with type 2 diabetes, but it can be safe and rewarding experience for the great majority of children and adolescents with T1DM (1). It enhances insulin sensitivity, reduces cardiovascular risk, increases cardiorespiratory fitness, improves glycemic control. There are some evidences that cardiorespiratory fitness is reduced in youths with T1DM compared to healthy peers (21, 29), but limited data exist regarding physical fitness in type 1 diabetic children and adolescents. Majority of diabetics avoid physical activity mainly because of the fear of exercise-induced hypoglycaemia and its severe consequences including loss of consciousness (5), but it can be controlled with careful monitoring of blood glucose level and supervising by healthcare providers (8). The regular exercise especially aerobic exercise is important for reducing the cardiovascular morbidity and mortality (2–4). The best quantitative measure of cardiorespiratory fitness is maximal oxygen consumption, expressed as VO$_2$\text{max}, the amount of oxygen the body uses when a person reaches maximum ability to supply oxygen during exercise.

The purpose of our cross-sectional study was to explore the relationship among the diabetes-specific HRQoL, clinical variables, anthropometric measures, level of physical activity and cardiorespiratory fitness (expressed by VO$_2$\text{max}) in children and adolescents with T1DM. Furthermore, we assessed the predictors of HRQoL and metabolic control from the variables of age, gender, diabetes duration, body composition, physical activity and cardiorespiratory fitness.

**Patients and Methods**

**Study participants**

The sample consisted of 106 patients with type 1 diabetes (53 girls and 53 boys). Their age ranged from 8 to 18 with an average of 13.22 years (SD = 3.08), 13.15 years (SD = 3.10) for females and 13.29 years (SD = 3.09) for males. Glycated haemoglobin in the sample was, in average, 8.55% (SD = 1.44) specifically 8.72% (SD = 1.31) for girls and 8.38% (SD = 1.56) for boys, respectively. Average diabetes duration was 5.15 (SD = 3.24) years with girls being diagnosed for 5.29 (SD = 2.80) years and boys for 5.02 (SD = 3.65) years (Table I).

The diabetic participants were from the Pediatric Diabetes Center of the Borsod-Abaúj-Zemplén County University Hospital, Miskolc representing the northern-east region of the country. The patients had been diagnosed with type 1 diabetes at least 1 year; and had no diabetes complications by regular assessments for retinopathy (fundal photography), nephropathy (microalbuminuria) and neuropathy (nerve conduction velocity and cardiovascular reflex tests). Table 1 presents the characteristics of the study participants by gender.
The parents and the youths were informed about the purpose and the method of the survey verbally and in written form. The parents and youths over 18 gave written consent; the children and adolescents gave their assent before the completion of the study.

The study was approved by the Borsod-Abaúj-Zemplén County Regional Scientific and Research Ethics Committee.

**Pediatric Quality of Life Inventory 3.0 Diabetes Module**
The young patients completed the Pediatric Quality of Life Inventory™ 3.0 Diabetes Module. This instrument is a disease-specific module and includes 28 items with 5 subscales /diabetes symptoms (11 items), treatment barriers (4 items), treatment adherence (7 items), worry (3 items) and communication (3 items)/. Participants had to rate how much of a problem of each item of the questionnaire have been in the previous month on a five-point Likert scale. Five categories were used, ranging from never a problem (0) to almost always a problem (4). All the items were transformed to a 0–100 scale. The higher score represented better quality of life (27). The 3.0 Diabetes Module has been validated for Hungarian usage (20).

**Physical activity**
The Physical Activity Questionnaire for Older Children (PAQ-C) and the Physical Activity Questionnaire for Adolescents (PAQ-A) were used to assess physical activity levels of the patients (15, 16). These tests are self-administered, 7-day recall questionnaires that provide general measure of physical activity for youths from ages 8–19. They classify children and adolescents into five different activity levels. A score of 1 indicates low physical activity, whereas a score of 5 indicates high physical activity.

**Assessment of cardiorespiratory fitness**
The maximal oxygen consumption was estimated from the 20 metre shuttle run test. The test involves continuous running between two lines 20 m apart in time to recorded beeps. The initial running speed is 8.5 km/hr, which increases by 0.5 km/hr each minute. The participant has to keep pace with the beep sound as long as possible. When the runner could no longer follow the pace and reach the 20 meter line consecutively twice with the beep, the last stage announced by the tape recorder was used to predict the VO$_{2\text{max}}$ using Léger et al. regression equation (18). The validity of the test in prediction of maximal oxygen consumption has been previously established (19, 24). The shuttle run test was carried out 08.00-10.00 AM and if morning home blood glucose result of the patient was out of the target range (5–10 mmol/l), a new appointment was given to perform the test on other day.

**Metabolic control**
Latest HbA$_1\text{c}$ values extracted from medical records were used to estimate metabolic control of the patients.

**Anthropometric assessment**
Skinfold thickness was measured at four sites (triceps, biceps, subscapular and suprailliac) on the right side of the body using Harpenden skinfold calliper (HSB-BI, British Indicators Ltd., 80 UK) according to the International Society for the Advancement Kinanthropometry (22). Two non-consecutive measurements for all sites were performed and mean values were used for data analysing. As there is no gold standard to evaluate the body fat content in percentage for children and adolescents the skinfold thickness was summed in mm. Height (to the nearest 0.5 cm) and weight (accurate to 0.1 kg) in light clothing without shoes were measured to calculate the body mass index (kg/m$^2$). Each BMI value was standardized by conversion to a
z-score (BMI z-score) defined by age and gender using the Hungarian National Institute of Child Health Chart (13).

Statistical analysis
SPSS 19.0 software was used for data analysis. Data are presented as mean and ± standard deviation (SD), p-value at ≤ 0.05 was considered significant. Pearson and Spearman correlations were used to assess associations as appropriate, and multiple regression analysis was used to determine significant independent predictors of the HRQoL and the metabolic control from the variables of age, gender, diabetes duration, skinfold thickness, BMZ z-score, physical activity level and maximal oxygen consumption. Gender differences in these variables were explored with independent t-test and Chi-square test.

Results
In our sample diabetic participants had HRQoL scores between 37.04 and 99.07. The mean score was 70.20 ± 12.61. There was a significant effect for gender (p = 0.046), with boys having higher scores than girls. (Fig. 1)

We observed significant gender differences in maximal oxygen consumption (boys: 41.96 ± 5.57 vs. girls 38.30 ± 6.73; p = 0.003), and in skinfold thickness (boys: 57.28 ± 25.57 vs. girls: 77.01 ± 29.56; p < 0.001); but it was not the case in age, diabetes duration, BMI z-score, physical activity and metabolic control.

There was statistically significant negative medium correlation between the metabolic control and the maximal oxygen consumption (r = –0.343; p < 0.001) (Fig. 2) and weak positive correlation between the HRQoL and the maximal oxygen consumption (r = 0.208; p = 0.032). We found no significant correlation between the HbA1c and the patients’ HRQoL.

In the multiple linear regression analysis the better metabolic control was predicted by the VO2max (B = -0.077, SE(B) = 0.021, β = –0.343, t = –3.726, p < 0.001, R = 0.343, R² = 0.118) other variables had no effect. When we examined the independent contributions of age, gender, diabetes duration, BMI z-score, skinfold thickness, metabolic control, physical activity level and maximal oxygen consumption on diabetes-specific HRQoL, significant predictor was likewise the maximal oxygen consumption (B = 0.522, SE(B) = 0.194, β = –0.266, t = –2.700, p = 0.008, R = 0.286, R² = 0.082).

Discussion
In the last decades, the management of type 1 diabetes has improved, with better metabolic control and reduction of long-term complications. It seems that children can live happy life as their healthy peers; however quality of life data are controversial. While there is a growing interest in quality of life issues in diabetes, it is important to identify which variables predict better outcomes. For this reason we examined the relationship among the HRQoL, clinical parameters (age, glycated haemoglobin and duration of disease), body composition (skinfold thickness and BMI z-score), physical activity, cardiorespiratory fitness (expressed by VO2max) in children and adolescents with type 1 diabetes.

HRQoL and body compositions had no relationship in our study similar to Naughton’s result (23). It is known that the overweight is associated with decreased HRQoL perception (25). Our diabetic patients were in normal BMI range, although there was notable gender difference in skinfold thickness. Females have greater skinfold thickness than males, that is typical for the normal pediatric population as well (26). When we compared the youths’ HRQoL by gender, we found that boys had better perception of quality of life than girls. This gender difference was confirmed in Graue and Kalyva’ researches (9, 14). We did not find significant relationship between the HRQoL and the metabolic control. The quality of life is the person’s subjective opinion about his or her well-being and health status; and sometimes
there is no close association between HRQoL and the severity of the disease. Our finding is congruent with Grey’s and Hoare’s study where no relationship was found between glycemic control, as measured by HbA1c, and diabetes-specific quality of life (10, 11). We measured both the physical activity and the maximal oxygen consumption. Most studies found significant association between the physical activity and the HRQoL. In our study the physical activity had no effect on HRQoL. There might be several reasons. We measured the physical activity using questionnaire that might be subjective; and it did not give information on the intensity of exercise. Physical activity generally refers to movement that enhances health condition, but not the type and intensity of exercise. Physical fitness is considered one of the most important health markers. VO_{2max} is the best indicator of the person’s actual fitness status. The higher maximal oxygen consumption correlated with better metabolic control and HRQoL. The better cardiorespiratory fitness explained both the favourable metabolic control and the better quality of life in children and adolescents with type 1 diabetes mellitus in the regression model.

Strength of our study is that we approached the HRQoL in addition to the physical activity level and cardiorespiratory fitness. Findings are important because they suggest that the physical fitness is essential to maintain good metabolic control; assessment of physical activity is not necessarily powerful marker of health status.

Limitation of our study is that we assessed youths from one pediatric centre with relatively small sample size. Self-report physical activity is also a limitation; the questionnaire has been validated in the original population, but not in Hungarian youths. The 20 m shuttle run test is indirect measure of the maximal oxygen consumption. These limitations reduce the possibility to generalize the results.

**Conclusion**

Metabolic control and HRQoL are two important outcomes of diabetes management. We suggest healthcare experts to take into consideration in the treatment and care of diabetes that regular aerobic exercise improves the young diabetic patients’ physical fitness and overall health status, and perception of HRQoL, respectively.

**Acknowledgements**

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**REFERENCES**

Table 1. Characteristics of the diabetic participants by gender

<table>
<thead>
<tr>
<th>Mean, standard deviation (±)</th>
<th>Girls</th>
<th>Boys</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sample size</td>
<td>53</td>
<td>53</td>
<td></td>
</tr>
<tr>
<td>Age (years old)</td>
<td>13.15 (± 3.10)</td>
<td>13.29 (± 3.09)</td>
<td>0.810</td>
</tr>
<tr>
<td>Diabetes duration (years)</td>
<td>5.29 (± 2.80)</td>
<td>5.02 (± 3.65)</td>
<td>0.669</td>
</tr>
<tr>
<td>HbA1c (%)</td>
<td>8.72 (± 1.31)</td>
<td>8.38 (± 1.56)</td>
<td>0.227</td>
</tr>
<tr>
<td>BMI z-score</td>
<td>0.66 (± 0.78)</td>
<td>0.38 (± 1.02)</td>
<td>0.118</td>
</tr>
<tr>
<td>Skinfold thickness (mm)</td>
<td>77.01 (± 22.99)</td>
<td>57.28 (± 25.57)</td>
<td>0.000*</td>
</tr>
</tbody>
</table>

*p ≤ 0.001

Fig. 1. Diabetic participants’ HRQoL by gender (according to the Pediatric Quality of Life Inventory 3.0 Diabetes Module child self-report).

Boys: 72.64 (± 12.64) vs. Girls: 67.77 (± 12.02), p ≤ 0.046

Fig. 2. Scatter plot of the correlation between HbA1c and VO2max (r = -0.343; p = 0.000)