

Correlation Between Hand Grip Strength and Glycemic Control Among Saudi Children with Chronic Type 1 Diabetes Mellitus (T1DM)

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Abstract

Objectives: We aimed to study the correlation between the hand grip strength as an indicator of the musculoskeletal affection and the degree of glycemic control in Type 1 Diabetic pediatric patients. **Methods:** Cross-sectional interventional study conducted among children having chronic T1DM recruited from the pediatric diabetes clinic at King Fahd hospital of the University, Saudi Arabia, they were divided into 3 groups according to their HbA1c level to well controlled, fairly controlled and poorly controlled. Anthropometrics measure taken then handgrip strength for both dominant and nondominant hands were measured using valid and reliable digital JAMAR PLUS hand dynamometer, data collection

was performed according to the American Society of Hand Therapists (ASHT) guidelines. **Results:** Total of 150 patients, aged 5-18 years, with 56% females and 44% males, two third of them were poorly controlled. Well controlled group showed better hand grip strength than the other 2 groups however it was not statistically significant. **Conclusions:** The handgrip strength in Type 1 DM children is affected by the degree of the glycemic control and it might give a clue of early musculoskeletal functional derangement by the effect of chronic hyperglycemia in these affected children.

Keywords: handgrip strength · glycemic control · diabetes mellitus

1. Introduction

Diabetes Mellitus (DM) consists of a heterogenous group of disorders characterized by the presence of hyperglycemia due to the inability of the human body to metabolize glucose properly [1]. Type 1 Diabetes Mellitus (T1DM) occurs as a result of chronic insulin deficiency due to the destruction of pancreatic beta cells by an aberrant autoimmune response [2]. The onset of T1DM is influenced by multiple genetic and environmental factors [3]. However, the varying viral and nutritional factors across the globe may influence the clinical appearance of T1DM [4]. Type 2 Diabetes Mellitus, on the other hand, results from the development of insulin resistance [1]. DM is considered a global health burden with a rapidly rising incidence and prevalence. As per the 8th edition of Diabetes Atlas, 35,000 children and adolescents in Saudi Arabia suffer from T1DM, which ranks Saudi Arabia as the 8th country in terms of prevalence of T1DM and the

4th country in terms of incidence with a rate of 33.5 per 100,000 individuals [5], whereas the World Health Organization ranks Saudi Arabia the 7th in prevalence and the 5th in incidence of T1DM [6].

Over the past few decades, vast advancements in the management of T1DM were achieved. Glycosylated hemoglobin A1c (HbA1c) remains an important guide for the initial diagnosis, management, control, and prediction of complications in diabetic subjects [6]. Nevertheless, individuals with uncontrolled T1DM will invariably develop long-term consequences due to microvascular and macrovascular complications such as blindness, renal failure, and cardiovascular disease. Though often overlooked, musculoskeletal system is also affected by T1DM, and adverse outcomes can be anticipated with suboptimal control of the disease [7].

Diabetic skeletal muscle disease is a common clinical condition observed among individuals with T1DM [8]. It is characterized by lower muscle mass,

generalized weakness, functional weakness, in addition to an overall reduction in physical capacity [9]. Muscle weakness contributes to the increased risk of physical disability associated with diabetes in children [10]. Impaired muscle strength has been reported in subjects with diabetes as a late complication of severe diabetic peripheral neuropathy (DPN) with motor nerve involvement [11]. However, other studies have indicated that reduced muscle strength, involving the upper body, may occur earlier in the course of diabetes independent of diabetic peripheral neuropathy [12].

According to American Society of Hand Therapists (ASHT), the evaluation of hand function, by measuring grip and pinch strength, reflects the overall strength of the upper limb [13,14]. Hand grip strength (HGS) is the sum of the strength of the flexor muscles against the palm, and used to evaluate the maximum static force a hand can handle [15,16]. Handheld dynamometry (HHD) provides a simple, inexpensive and versatile alternative for assessing muscle strength [17]. Moderate to good reliability was found across all patient groups with reliability coefficients ranging from 0.80 to 0.99 [18]. In addition to its use to assess disease severity and evaluate the effectiveness of certain interventions, HGS can reflect the general health and level of physical activity of the individual, predicting the overall strength [19-21]. An accumulating body of literature has documented a decline in HGS among individuals with DM, compared with healthy individuals with normal glucose tolerance [22-24], reflecting the link between metabolic and mechanical functions of the muscle [24,25].

Despite the high prevalence of T1DM and the importance of hand function in the activities of daily living of individuals, there is insufficient data concerning the association between glycemic control and hand strength among diabetic children. Therefore, the present study aimed to investigate the correlation between strength measurement through HGS and glycemic control amongst Saudi children diagnosed with T1DM aged between 5 and 18 years with good glycemic control in comparison to children with poor glycemic control following in a pediatric diabetes clinic at King Fahad Hospital of the University in Khobar, Saudi Arabia.

2. Materials and methods

2.1 Design and subjects

A cross-sectional study was conducted among children diagnosed with T1DM recruited from the pediatric diabetes clinic at King Fahd hospital of the University, Imam Abdulrahman Bin Faisal University in Khobar, Saudi Arabia. 150 children with T1DM were initially screened and assessed to determine age, diagnosis, and inclusion and exclusion criteria. Children aging between 5 and 18 years with a confirmed diagnosis of chronic T1DM for at least 2 years, not known to have another medical condition such as hypertension, anemia, neurological, cardiopulmonary,

Abbreviations:

DM	Diabetes Mellitus
T1DM	Type 1 diabetes Mellitus
DNP	Diabetic Neuropathy
ASHT	American Society of Hand Therapists
HGS	Hand Grip Strength
HHD	Handheld dynamometry
HbA1c	Glycosylated Hemoglobin A1c
BMI	Body Mass Index

or renal diseases, and not taking regular medications beside insulin were included in the study. Patients with any other disease affecting their physical activity level, diagnosed with other types of DM, upper limb pain, trauma or fracture around the hand within the past year, severe social deprivation and any history of known psychiatric disease or treatment and mental impairment, that might interfere with their response based on parent report were excluded from the study. All participants were apparently healthy, cognitively competent and able to understand and follow instructions with no evidence of musculoskeletal injuries.

2.2 Ethical approval

Ethical approval for this study was granted by the Institutional Review Board at Imam Abdulrahman bin Faisal University and issued the approval number (IRB-2022-01-184). Parents signed the consent form authorizing the child's participation. The study was conducted in accordance with the Helsinki Declaration of 1975, as revised in 1996 (World Medical Association, 1996).

2.3 Biochemical measurements

The degree of control of diabetes was evaluated by measuring the average of the last 3 readings of glycosylated haemoglobin (HbA1c, HPLC method). Blood samples were collected by a trained paediatric nurse.

2.4 Anthropometric parameters

Participants' weight and height were measured, and the body mass index (BMI) was calculated as $\text{Weight (W) in kg} / [\text{Height (H) in meter}]^2$. Medical conditions of the participants, such as the T1DM chronicity, appropriate medication doses and HbA1c, were assessed by a pediatric endocrinologist in the clinic. Participants were assessed physically and clinically by a pediatrician, and according to their HbA1c they were categorized into three groups: group (A) with poor glycemic control with an HbA1c above 8.5%, group (B) with fairly controlled DM with an HbA1c between 7% and 8.5%, and group (C) with well controlled DM with an HbA1c of 7%. All groups were matched by

average age, weight, height, and BMI. All participants have received the same verbal cues and directions for outcome measurement.

2.5 Handgrip strength assessment

Under direct supervision of an expert physical therapy faculty member, handgrip strength for both dominant and nondominant hands were measured using valid and reliable digital JAMAR PLUS hand dynamometer (Patterson Medical, Sammons Preston, Jackson, MI). Data collection was performed according to the American Society of Hand Therapists (ASHT) guidelines [26]. The participants were seated on a chair without armrests with their feet flat on the floor, shoulder was abducted and neutrally rotated, elbow flexed at 90°, forearm in a neutral position, and wrist between 0 and 30° extension and between 0 and 15° ulnar deviation. Testing positions demonstrations and verbal instructions were provided. The participants received verbal encouragement to squeeze the gauge as hard as possible to exert their maximal force during each trial and hold for 5 seconds. Each test was performed three times to collect HGS data, and the average was calculated and expressed in kilograms. If a measurement displayed a difference of over 10% from the previously achieved measurements, it would lead to performing a fourth trial [26,27]. The measurements of HGS was done in alternating order between the dominant and nondominant hands, with a 1-min rest between them to minimize fatigue effect [28, 16]. The calibration of Jamar hand dynamometer was tested periodically during the study [16].

2.6 Statistical analysis

The data analysis was performed using SPSS version 26(Armonk, NY: IBM Corp. USA). The normality of the data distribution was assessed using Kolmogorov-Smirnov test which showed normal data distribution if p

> 0.05 . Continuous variables were expressed as mean \pm SD (for variables with normal distributions) or medians (interquartile range) (for variables with nonnormal distributions, and categorical variables were expressed as frequency and percentages (%). The demographic and baseline characteristics, between levels of glycemic control were compared using the Chi square test or Fisher's exact test for categorical variables, ANOVA/Kruskal-Wallis tests for continuous variables, as appropriate for more than two groups. Student's t-test or Mann-Whitney U test was used to compare the dexterity and the various characteristics whichever was appropriate. Spearman's rank correlation coefficient was used to analyse the relationship between the hand grip strength with duration of diabetes and also with BMI. Two-way ANOVA was used to test for the differences between the levels of glycemic control and hand dexterity and also its interaction effect in the assessment of hand grip strength. A p-value of < 0.05 was considered significant for the analysis.

3. Results

Details of the characteristics of the study participants are presented in Table 1. A total of 150 children with age range, 5 to 18 years participated in the study. Among the 150 (66 males, 84 females) participants included in the study, duration of diabetes was 5.0 years (interquartile range 3.0– 7.0 years). With poorly controlled glycemic level was found in 103 (68.7%) participants, fairly controlled in 31 (20.7%) participants, and well controlled in 16 (10.7%) of the participants. The mean \pm SD for BMI was 20.20 \pm 4.5 kg/m². The right- and left-hand dominance was found in 139 (92.7%) and 11 (7.3%) of the participants, respectively. No significant difference was found in the proportion of participants with different level of glycemic control in all the demographic and clinical characteristics, except for weight and those variables related to glucose level.

Table 1. Baseline characteristics of participant

Variables	Total (n = 150)	Glycemic control			P value
		Poorly-Controlled (n = 103)	Fairly-Controlled (n = 31)	Well controlled (n = 16)	
Sex					
Male	66 (44.0)	46 (44.7)	11 (35.5)	9 (56.3)	0.386
Female	84 (56.0)	57 (55.3)	20 (64.5)	7 (43.8)	
Age	12.00 (10.0,14.3)	13.00 (11.0,14.0)	11.00 (10.0,15.0)	10.00 (8.0,12.8)	0.064
BMI mean \pm SD	20.20 \pm 4.5	20.64 \pm 4.7	19.73 \pm 4.3	18.27 \pm 2.5	0.119
Dominant side					
Right	139 (92.7)	96 (93.2)	29 (93.5)	14 (87.5)	0.623*
Left	11 (7.3)	7 (6.8)	2 (6.5)	2 (12.5)	
Duration of DM	5.00 (3.0,7.0)	5.00 (2.0,7.0)	5.00 (3.0,8.0)	2.00 (1.0,4.0)	0.012
HBA1C level	10.20 (9.0,12.0)	11.00 (10.1,12.5)	8.80 (8.5,9.1)	7.48 (6.9,7.9)	0.000
Systolic mean \pm SD	113.53 \pm 2.9	114.46 \pm 11.6	112.19 \pm 11.0	110.19 \pm 10.8	0.294
Diastolic	73.25 \pm 8.6	73.00 (68.0,80.0)	72.00 (68.0,77.0)	73.00 (68.0,80.0)	0.124
Heart rate	92.00 (85.8,104.0)	92.00 (85.8,104.0)	92.00 (85.8,104.0)	71.00 (65.8,73.5)	0.893

Respiratory rate	17.00 (14.0,20.0)	15.00 (14.0,20.0)	18.00 (14.0,20.0)	15.50 (14.0,18.0)	0.286
Dominant side 1st trial	11.00 (8.0,16.0)	11.00 (8.0,16.0)	11.00 (7.0,17.0)	9.00 (7.3,11.8)	0.364
Dominant side 2nd trial	10.00 (8.0,17.0)	10.00 (8.0,17.0)	10.00 (8.0,17.0)	9.50 (6.3,12.8)	0.608
Dominant side 3rd trial	11.00 (8.0,15.0)	10.00 (8.0,15.0)	10.00 (8.0,16.0)	9.00 (5.3,13.5)	0.577
Mean Dominant side	10.60 (8.0,15.39)	10.60 (8.0,15.7)	11.0 (7.6,16.6)	9.15 (6.6,12.7)	0.465
Non-dominant side 1st trial	10.00 (8.0,14.3)	10.00 (8.0,15.0)	10.00 (7.0,17.0)	9.00 (7.0,12.0)	0.656
Non-dominant side 2nd trial	9.00 (6.8,14.0)	9.00 (7.0,14.0)	10.00 (6.0,16.0)	9.00 (6.0,11.5)	0.581
Non-dominant side 3rd trial	9.00 (6.8,14.0)	9.00 (7.0,14.0)	10.00 (5.0,15.0)	8.00 (6.0,12.0)	0.703
Mean Non-dominant side	9.60 (6.9,13.1)	9.60 (7.3,13.0)	9.60 (5.3,15.3)	8.60 (6.2,12.3)	0.594

Categorical variables are expressed as number (%), continuous Data are expressed as the mean ± standard deviation or median (inter quartile range); * - Fisher's exact test

Table 2 shows the median (interquartile range) of the HGS for the different levels of glycemic control for the dominant and non-dominant hands. The results revealed that individuals with well controlled glycemic level had the best performance in both hands, but the difference was not significantly different from the participants with other levels of glycemic control. The proportion of males were significantly higher in left hand dominance with 81.8% males compared to 41.0% males in the right-hand dominant group. All the other variables were not significantly different between right- and left-hand dominance. (See Table 3)

Table 4 presents the HGS of the participants

measured on both hands are compared between the dominant and non-dominant hands. For the right hand, a median of 10.60 (inter quartile range 8.0,16.3) was measured for the right dominant hand while the same was 11.30 (9.3,16.6) in the non-dominant hand (p=0.686). Similarly, for the left hand, a median of 10.00 (inter quartile range 7.3,15.3) was measured for the left dominant hand while the same was 9.30 (6.6,13.0) in the non-dominant right-hand group (p=0.708) (Table 5).

The results revealed that the participants did not show a statistically significant difference in HGS measured in the right hand for glycemic control,

Table 2. Relationship between handgrip strength with glycemic control for the dominant and the non-dominant hand of the participants (n=150)

Variable	Glycemic control			KW H [#]	P value
	Poorly-Controlled (n = 103)	Fairly-Controlled (n = 31)	Well controlled (n = 16)		
Dominant hand	10.30 (8.0,16.6)	10.60 (6.6,15.3)	9.15 (6.6,12.7)	1.640	0.441
Non-dominant hand	9.60 (7.3,12.7)	10.00 (5.3,16.6)	8.60 (6.2,12.3)	0.998	0.607

#- Kruskal Wallis test statistic

Table 3A. Distribution of baseline and clinical characteristics of the participants

Variables	Right hand (n = 139)	Left hand (n = 11)	P value
Sex			
Male	57 (41.0)	9 (81.8)	0.011*
Female	82 (59.0)	2 (18.2)	
Age	12.00 (10.0,14.0)	14.00 (11.0,15.0)	0.543
BMI mean ± SD	20.37±4.6	18.09±2.5	0.107
Glycemic control			
Poorly-Controlled	96 (69.1)	7 (63.6)	0.623*
Fairly-Controlled	29 (93.5)	2 (18.2)	
Well controlled	14 (10.1)	2 (18.2)	
Duration of DM	5.00 (2.0,7.0)	4.00 (2.0,8.0)	0.811
HBA1C level	10.3 (9.1,12.0)	10.00 (8.5,13.0)	0.945
Systolic mean ± SD	113.22±11.5	117.45±10.4	0.239
Diastolic mean ± SD	73.35±8.7	72.0±6.2	0.616
Heart rate	93.00 (85.0,105.0)	87.00 (86.0,95.0)	0.090

Table 3B. Distribution of baseline and clinical characteristics of the participants

Variables	Right hand (n = 139)	Left hand (n = 11)	P value
Respiratory rate	18.00 (14.0,20.0)	14.00 (14.0,18.0)	0.146
HG RT side 1st trial	10.00 (8.0,17.0)	12.00 (10.0,13.0)	0.400
HG RT side 2nd trial	10.00 (8.0,17.0)	11.00 (10.0,13.0)	0.758
HG RT side 3rd trial	10.00 (8.0,15.0)	10.00 (9.0,14.0)	0.957
Mean HG RT side	10.6 (8.0,16.3)	11.30 (9.3,12.3)	0.686
HG LT side 1st trial	10.00 (7.0,15.0)	11.00 (9.0,14.0)	0.188
HG LT side 2nd trial	9.00 (6.0,14.0)	10.00 (8.0,16.0)	0.625
HG LT side 3rd trial	9.00 (6.0,14.0)	11.00 (10.0,15.0)	0.105
Mean HG LT side	9.30 (6.6,13.0)	10.00 (7.3,15.3)	0.708

Table 4. Comparison of handgrip strength measured on each hand

Variable	Grip strength measured on	
	Right hand	Left hand
Dominant hand	10.60 (8.0,16.3)	10.00 (7.3,15.3)
Non-dominant hand	11.30 (9.3,16.6)	9.30 (6.6,13.0)
P value	0.686	0.708

Table 5. Handgrip strength as a function of glycemic control

Hand	Effect	df	F	P	η ²
Right	Glycemic control	2,144	0.924	0.399	0.013
	Dominant side	1,144	0.003	0.953	0.000
	Glycemic control*Dominant side	2,144	0.687	0.505	0.009
Left	Glycemic control	2,144	0.983	0.377	0.013
	Dominant side	1,144	0.233	0.630	0.002
	Glycemic control*Dominant side	2,144	0.624	0.537	0.009

dominance as well as for the interaction of the two [F(2,144) = 0.924, p = 0.399, η² = 0.013], [F(1,144) = 0.003, p = 0.953, η² = 0.000] and [F(2,144) = 0.687, p = 0.505, η² = 0.009], respectively. Similarly, for the left hand, HGS measured for glycemic control, dominance as well as for the interaction of the two was not statistically significant [F(2,144) = 0.983, p = 0.377, η² = 0.013], [F(1,144) = 0.233, p = 0.630, η² = 0.002] and [F(2,144) = 0.6245, p = 0.537, η² = 0.009]. Also see figure 1 and 2.

Table 6 shows the correlation between age and HGS with each glycemic control group. The correlations showed a positive linear correlation, and the strength of the correlation coefficient was moderate and was statistically significant except for the correlation of HGS of the left hand in the well-controlled glycemic group. From table 7, we see that from age 11 and above, both the sexes were not significantly different from the normative hand grip strength taken from the reference study.

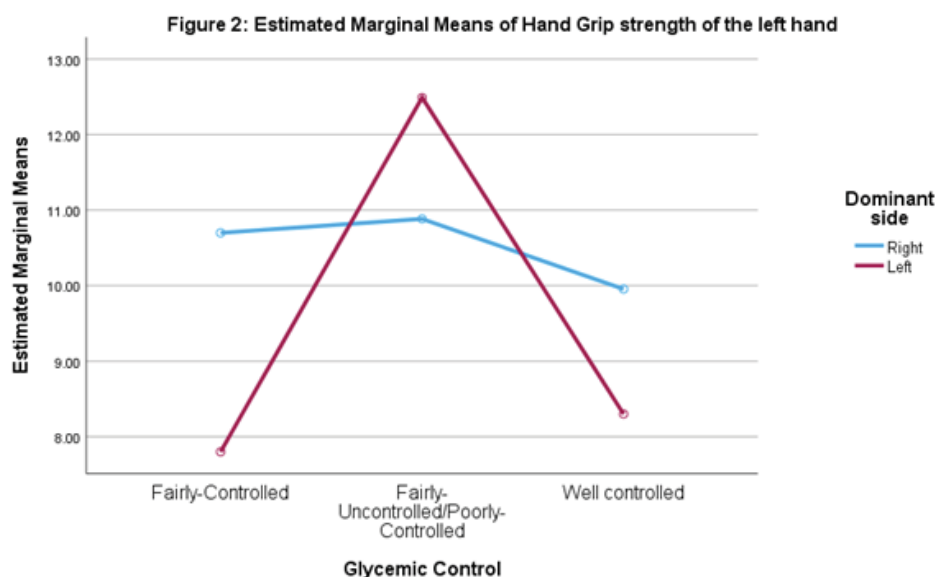
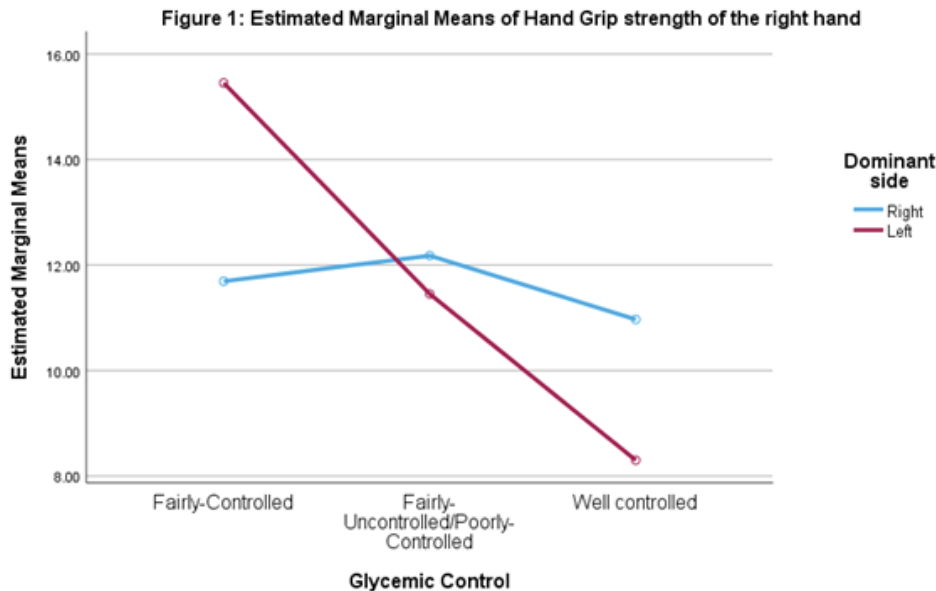


Table 6. Correlation between age and handgrip strength on each hand for each glycemic control groups

Variable	Age correlated with Grip strength measured on			
	Right hand		Left hand	
	r/rho	P value	r/rho	P value
Glycemic control				
Poorly-Controlled	0.481	0.000	0.414	0.000
Fairly-Controlled	0.571	0.001	0.560	0.001*
Well controlled	0.571	0.021*	0.473	0.063*

*- Pearson's correlation

Table 7. Comparison of handgrip strength of both sexes with normative levels from a reference study

Age	Sex	Hand grip strength				Pvalue
		Reference study		Current study		
		N	Mean±SD	N	Mean±SD	
5	Male			2	8.0±2.8	-
	Female					-
6	Male	14	8.5±6.1			-
	Female	14	8.5±5	1	6.0	-
7	Male	61	18.6±10.5	4	8.4±1.5	0.000
	Female	37	13.3±5.8	5	6.1±1.2	0.000
8	Male	22	15.8±9.6	7	7.6±3.1	0.000
	Female	23	13.5±6.5	6	6.8±1.0	0.000
9	Male	31	18.7±9.7	3	14.3±7.8	0.438
	Female	31	12.5±6.3	1	6.0	-
10	Male	15	15.1±9.8	4	7.2±3.0	0.016
	Female	19	12.7±5.1	11	8.4±2.9	0.006
11	Male	27	12.4±4.8	10	11.2±2.7	0.348
	Female	26	12.9±5.7	5	10.0±3.3	0.15
12	Male	31	17.4±6.6	5	12.0±4.9	0.069
	Female	22	12.0±6.5	14	13.1±5.4	0.586
13	Male	29	17.8±9.5	10	15.5±4.9	0.335
	Female	20	13.4±6.4	6	12.0±3.6	0.505
14	Male	23	16.6±8.0	4	15.0±5.5	0.638
	Female	26	12.5±6.7	15	12.0±4.1	0.769
15	Male	23	14.5±8.2	9	16.6±7.0	0.478
	Female	28	13.2±5.1	11	15.5±5.3	0.234
16	Male	19	14.1±8.8	5	17.0±5.7	0.394
	Female	25	12.4±6.2	5	9.1±2.2	0.051
17	Male	16	16.6±8.1	3	24.4±6.9	0.175
	Female	15	12.8±4.6	3	11.2±6.9	0.732
18	Male	7	14.5±6.8			-
	Female	12	14.5±6.8	1	10.60	-

The relationship between the duration of diabetes with hand grip strength of the right hand showed the spearman's rank correlation coefficient was low with $\rho = 0.204$ $p=0.012$. similarly, the relationship between the duration of diabetes with hand grip strength of the left hand showed the Spearman's rank correlation coefficient was low with $\rho = 0.164$ $p=0.045$. Also see figure 3 and 4.

The relationship between the BMI with hand grip strength of the right hand showed the Spearman's rank correlation coefficient was low with $\rho = 0.395$ $p=0.000$. similarly, the relationship between the duration of diabetes with hand grip strength of the left hand showed the Spearman's rank correlation coefficient was low with $\rho = 0.350$ $p=0.000$. Also see figure 5 and 6.

Figure 3: Relationship between duration of diabetes and HGS-Right hand

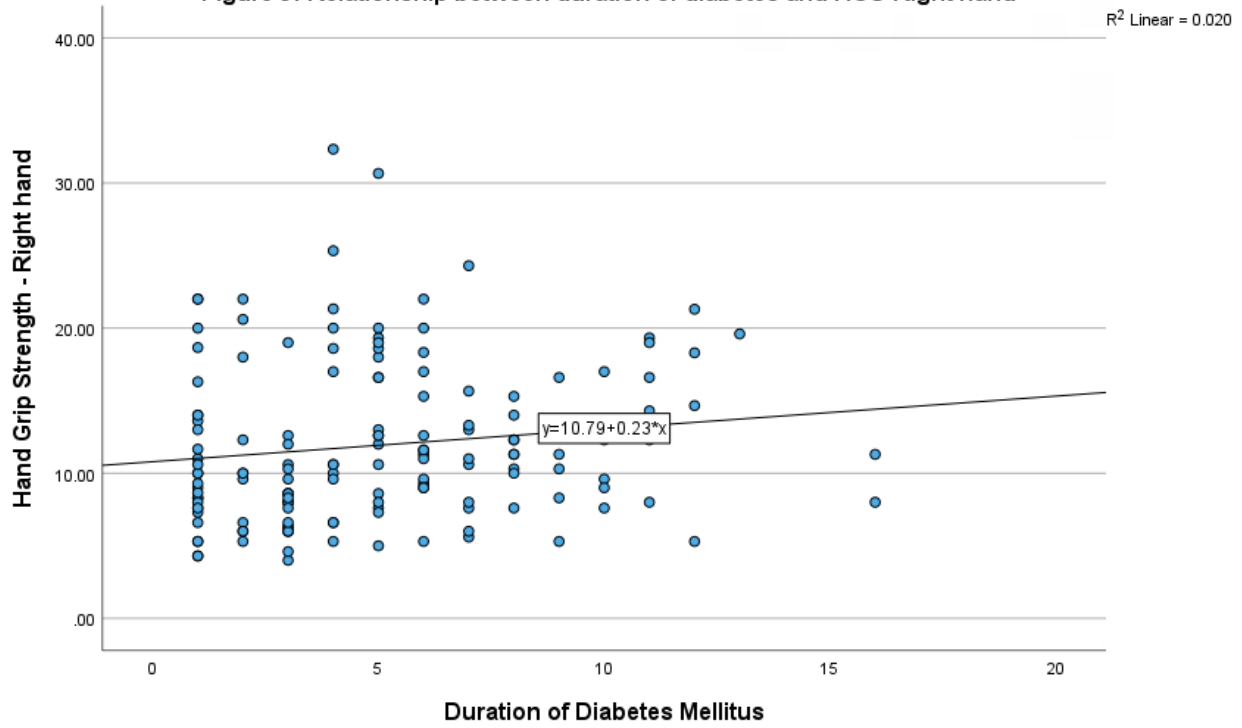
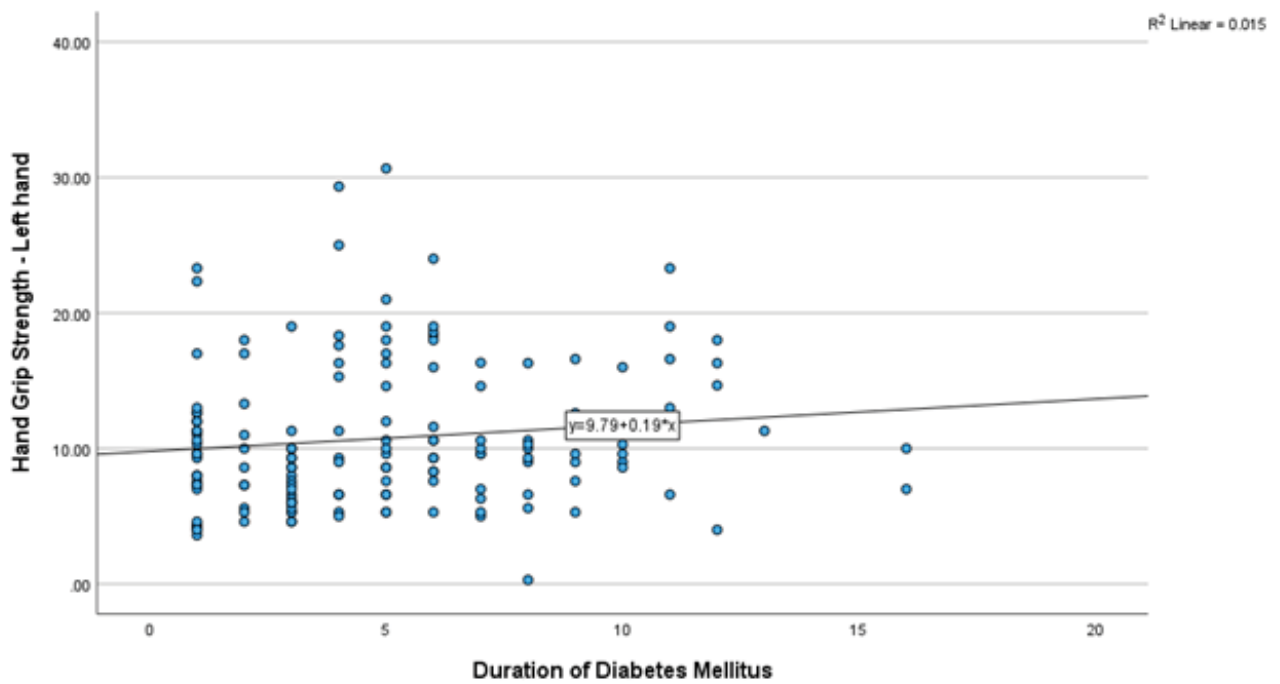
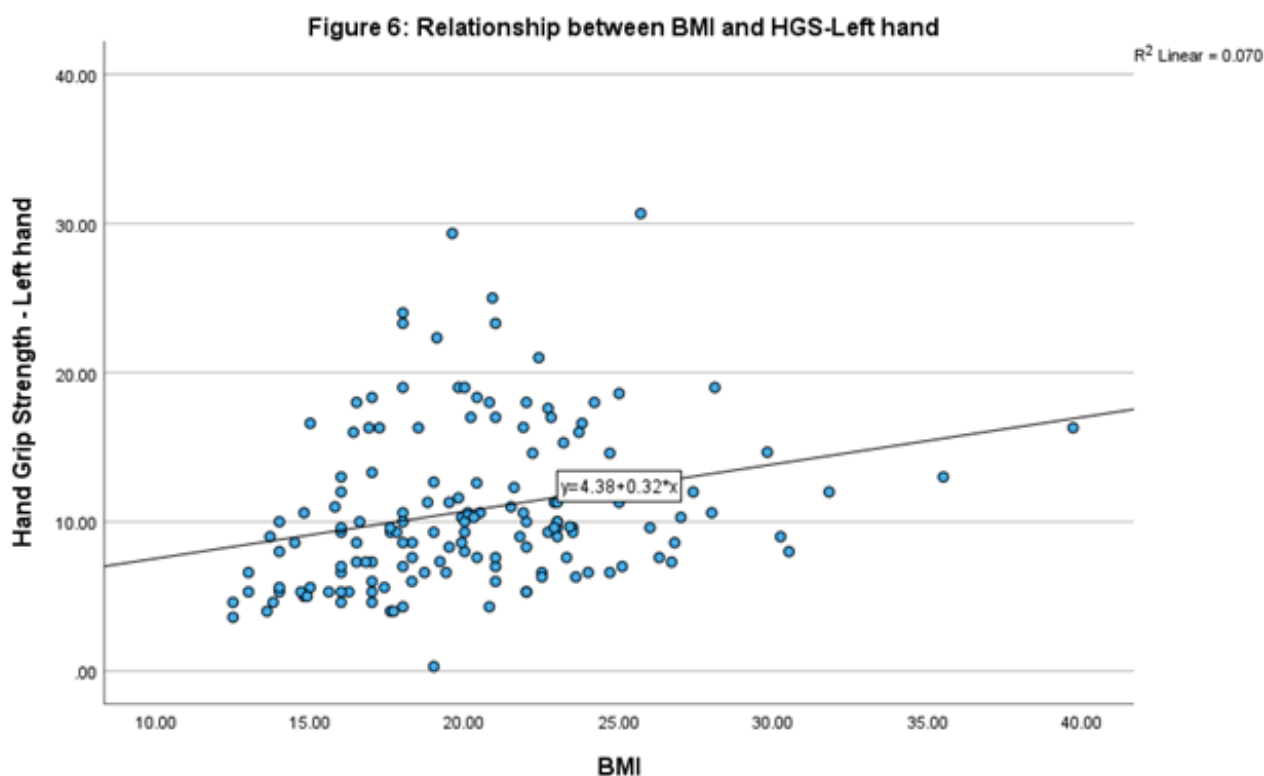
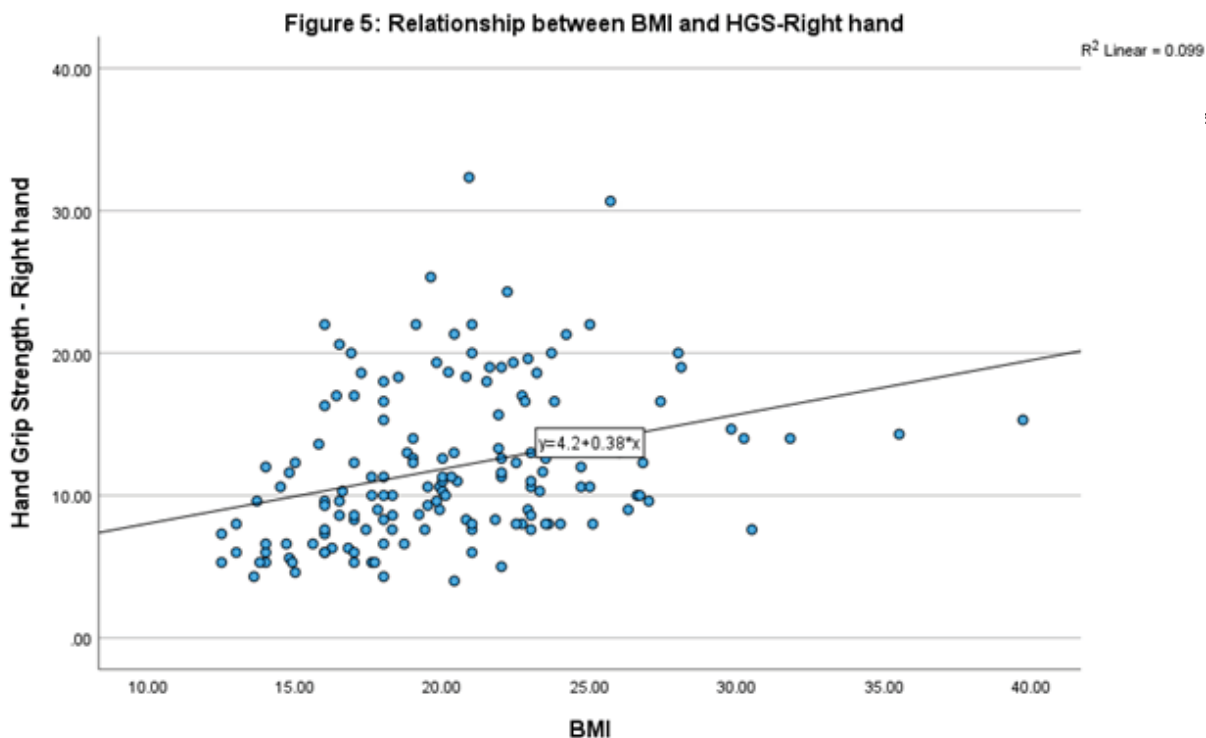


Figure 4: Relationship between duration of diabetes and HGS-Left hand





4. Discussion

The present study has investigated the correlation between T1DM and HGS among Saudi children aged between 5 and 18 years with different levels of glycemic control. No significant differences in HGS were observed between patients with poorly controlled disease in comparison to those with fairly controlled

and well controlled T1DM. Nonetheless, although it was statistically insignificant, participants with well controlled T1DM have shown the best performance in both dominant and non-dominant hands compared to children with other levels of glycemic control. Consistent with our findings, Dongare-Bhor et al. reported a lack of correlation between HGS and HbA1C [29]. In adults

with T1DM, Wallymahmed *et al.* reported a significant negative correlation between HGS and HbA1C which opposes the results of the current study [30]. Fricke *et al.*, in contrast, reported a significantly higher HGS in children with an HbA1C above 8.5% compared to those with lower values of HbA1C [31].

In addition, we report a significantly low correlation between HGS and disease duration. In line with our findings, Dongare-Bhor *et al.* reported the lack of correlation between HGS and disease duration across diabetic children who had the disease for more than 5 years compared to those who had it for shorter durations [29]. Klara *et al.*, on the contrary, evaluated the dynamic muscle function using jumping mechanography and reported a decline in muscle function among adolescents with T1DM which was particularly evident in patients who had T1DM longer than 9 years [32]. The previous study has anticipated a further deterioration in muscle function in adulthood with increased T1DM duration [32]. Reduced HGS and impaired muscle function among adults was reported by several studies to be a complication of T1DM and T2DM especially in patients with DPN and carpal tunnel syndrome [32-34].

A recently published community-based study conducted by Alqahtani *et al.* measured the HGS among 616 Saudi children aged between 6 and 18 years old across different cities in Saudi Arabia [35]. The reported normative values of HGS have been used as a reference for comparison in the present study. Children with T1DM aged 7 to 10 years from both sexes were found to have a significantly lower HGS compared to the reference population except for males aging 9 years. However, no significant difference in HGS was found between the reference population and diabetic children aged 11 years and above.

Fricke *et al.* found a significantly lower maximal isometric force among children with T1DM in all age groups compared to age-matched reference counterparts [31], which is partially congruent with the findings of the current study. Similarly, Dongare-Bhor *et al.* reported that children with T1DM had a significantly lower HGS compared to the controls [29]. Bechtold *et al.*, in contrast, have investigated the impact of T1DM on the development of bone and muscle and reported a significantly higher HGS among diabetic children in comparison to controls which was attributed to intensive motivation of the participants during the assessment [36].

Although knowledge is currently limited and the existing evidence remains equivocal, multiple postulations describing the association between decreased muscle function and T1DM have been proposed. The effect of T1DM on skeletal muscle function is thought to be multifactorial with altered levels of hormones and hyperglycemia being the major contributors [8,32]. Insulin is a potent anabolic hormone that promote protein synthesis and inhibits the degradation of protein in the skeletal muscle [8,37]. A deficiency in insulin leads to a protein catabolic state that results in the loss of muscular tissue [32], which

could be a possible underlying mechanism that leads to impaired muscle function in diabetic individuals. In addition, several studies found that Insulin-like Growth Factor-1 (IGF-1), an essential growth factor of the skeletal muscle, was reduced in adolescents and adults with T1DM, raising the suggestion that it may have a role in muscle growth impairment in T1DM [8].

Furthermore, in a long-standing hyperglycemic state, protein glycation occurs in skeletal muscle, a process at which proteins undergo chemical modification as a result of sugars reduction [8,32]. In early stages of this reaction, myosin motility is found to be reduced [32]. Further oxidation reactions result in advanced glycation end-product (AGE), the contribution of which in T1DM-related complications has been clearly established [8]. In skeletal muscle, the accumulation of AGE was suggested by several studies to be associated with a decline in muscle function in adult patients with T1DM, T2DM and elderly individuals [38].

In the present study, a significant positive linear correlation was observed between age and HGS among diabetic children. Similarly, Bechtold *et al.* found that HGS has increased significantly with age in children. In accordance with previous findings, an overall trend of increasing HGS with age among healthy children was reported by several studies [35,39]. This increase is probably attributed to the physiological changes and development in muscle strength of the upper limbs in both genders with age [35].

Although the correlation between HGS and T1DM remains uncertain with heterogenous findings [29,30,31,36], chronic diseases are often associated with deterioration in muscle function [31]. Several studies have reported a decrease in HGS among pediatric and adult patients suffering from chronic conditions [31,40,41]. Rauch *et al.* reported reduced maximal isometric grip force among children and adolescents with cystic fibrosis and kidney transplantation [40]. In adults, HGS was found to approximately 25-50% lower in patients on hemodialysis compared to general population [41]. In addition, HGS weakness was found to be associated with multimorbidity including chronic diseases such as anemia, stage 3 of chronic kidney disease or above, stroke, and kyphosis [42]. Hence, further research is imperative to assess the impact of T1DM on muscle growth in children and adolescents to facilitate early intervention and prevent long-term complications affecting the quality of life of individuals.

4.1 Limitations

We have some limitations to be considered in interpreting the results of our study; first, this study is cross-sectional with relatively small sample sizes, dealing with single center of the country. However, this study provided a baseline assessment based on grip strength in Type 1 diabetic children, given the scarcity of data on national as well international paediatric population, therefore, comprehensive studies and innovative research are strongly needed and the clinical trials must be well designed, adequately powered, carefully controlled, cautiously conducted,

and multicenter approaches in order to prevent or delay the development of the devastating musculoskeletal complications of the T1DM in pediatric age group.

Second, the study compared the controlled type 1 diabetic children, determined by average A1c level compared to poorly controlled diabetic children, not able to have normal matching children as a baseline comparison since it is difficult to recruit normal children from the pediatric clinics in hospital population, however we compared them to the single Saudi published reference standard.

4.2 Conclusion

Chronic standing uncontrolled type 1 diabetes mellitus has significant impact on all the body systems including the musculoskeletal system, which might be

detected early by performing some standardized motor ability tests, which will help in prevention and early management of such complications.

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References

1. **Baronio F, Mazzanti L, Girtler Y, Tamburrino F, Lupi F, Longhi S, Fanolla A, Radetti G.** The influence of GH treatment on glucose homeostasis in girls with Turner Syndrome: a 7 years study. *J Clin Endocrinol Metab* 2017, 102(3):878-83.
2. **Maahs DM, West NA, Lawrence JM, Mayer-Davis EJ.** Epidemiology of type 1 diabetes. *Endocrinol Metab Clin North Am* 2010, 39(3):481-97.
3. **Robertson CC, Rich SS.** Genetics of type 1 diabetes. 2018, 50:7-16.
4. **Nagesh VS, Kalra S.** Type 1 diabetes: syndromes in resource-challenged settings. *J Pak Med Assoc* 2015, 65(6):681-5.
5. **Robert AA, Al-Dawish A, Mujammami M, AlDawish MA.** Type 1 Diabetes Mellitus in Saudi Arabia: A Soaring Epidemic. *Int J Pediatr* 2018, 2018:9408370.
6. **Alaqeel AA.** Pediatric diabetes in Saudi Arabia: Challenges and potential solutions. A review article. *Int J Pediatr Adolesc Med* 2019, 6(4):125-30.
7. **Coleman SK, Rebalka I, D'Souza DM, Hawke TJ.** Skeletal muscle as a therapeutic target for delaying type 1 diabetic complications. *World J Diabetes* 2015, 6(17):1323-36.
8. **Krause MP, Riddell MC, Hawke TJ.** Effects of type 1 diabetes mellitus on skeletal muscle: clinical observations and physiological mechanisms. *Pediatr Diabetes* 2011, 12(1):345-64.
9. **Atay C, Mutlu EK, Taskiran H, Ozgen IT.** Comparison of Hand Function Between Children with Type 1 Diabetes Mellitus and Children Without Type 1 Diabetes Mellitus. *Pediatr Phys Ther* 2018, 30(1):58-65.
10. **Balducci S, Sacchetti M, Orlando G.** Correlates of muscle strength in diabetes: the study on the assessment of determinants of muscle and bone strength abnormalities in diabetes (SAMBA). *Nutr Metab Cardiovasc Dis* 2014, 24(1):18-26.
11. **Andreassen CS, Jakobsen J, Andersen H.** Muscle weakness: a progressive late complication in diabetic distal symmetric polyneuropathy. *Diabetes* 2006, 55(3):806-12.
12. **Stenholm S, Harkanen T, Sainio P, Heliovaara M, Koskinen S.** Long- term changes in handgrip strength in men and women—accounting the effect of right censoring due to death. *J Gerontol A Biol Sci Med Sci* 2012, 67(10):1068-74.
13. **Martins JC, Teixeira-Salmela LF, Castro e Souza LA, Aguiar LT, Lara EM, Moura JB.** Reliability and validity of the modified sphygmomanometer test for the assessment of strength of upper limb muscles after stroke. *J Rehabil Med* 2015, 47(8):697-705.
14. **Shetty M, Balasundaran S, Mullerpatan R.** Grip and pinch strength: reference values for children and adolescents from India. *J Pediatr Rehabil Med* 2019, 12(3):255-62.
15. **Kim CR, Jeon Y-J, Kim MC, Jeong T, Koo WR.** Reference values for hand grip strength in the South Korean population. Mogi M, editor. *PLoS One* 2018, 13(4):e0195485.
16. **Mathiowetz V, Kashman N, Volland G, Weber K, Dowe M, Rogers S.** Grip and pinch strength: normative data for adults. *Arch Phys Med Rehabil* 1985, 66(2):69-74.
17. **Vermeulen HM, De Bock GH, Van Houwelingen HC, Van der Meer RL, Mol MC, Plus BT.** A comparison of two portable dynamometers in the assessment of shoulder and elbow strength. *Physiotherapy* 2005, 91:101-12.
18. **Dowman L, McDonald CF, Hill CJ, Lee A, Barker K, Boote C, Glaspole I, Goh N, Southcott A, Burge A, et al.** Reliability of the hand-held dynamometer in measuring muscle strength in people with interstitial lung disease. *Physiotherapy* 2016, 102(3):249-55.
19. **Hansen AW, Beyer N, Flensburg-Madsen T, Grønbæk M, Helge JW.** Muscle strength and physical activity are associated with self-rated health in an adult Danish population. *Prev Med* 2013, 57(6):792-8.
20. **Leong DP, Teo KK, Rangarajan S, Lopez-Jaramillo P, Avezum A, Orlandini A, Seron P, Ahmed SH, Rosengren A, Kelishadi R.** Prognostic value of grip strength: findings from the Prospective Urban Rural Epidemiology (PURE) study. *Lancet* 2015, 386(9990):266-73.
21. **Rantanen T, Parkatti T, Heikkinen E.** Muscle strength according to level of physical exercise and educational background in middle-aged women in Finland. *Eur J Appl Physiol Occup Physiol* 1992, 65(6):507-512.
22. **Cetinus E, Buyukbese MA, Uzel M, Ekerbicer H, Karaoguz A.** Hand grip strength in patients with type 2 diabetes mellitus. *Diab Res Clin Prac* 2005, 70(3):278-86.
23. **Mainous AG, Tanner RJ, Anton SD, Jo A.** Grip strength as a marker of hypertension and diabetes in healthy weight

- adults. *Am J Prev Med* 2015, 49(6):850-8.
24. **Sayer AA, Dennison EM, Syddall HE, Gilbody HJ, Phillips DI, Cooper C.** Type 2 diabetes, muscle strength, and impaired physical function: The tip of the iceberg? *Diabetes Care* 2005, 28(10):2541-2.
 25. **Yeh HC, Punjabi NM, Wang NY, Pankow JS, Duncan BB, Cox CE, Brancati FL.** Cross-sectional and prospective study of lung function in adults with type 2 diabetes: The Atherosclerosis Risk in Communities (ARIC) study. *Diabetes Care* 2008, 31(4):741-6.
 26. **MacDermid J, Solomon G, Valdes K.** Clinical assessment recommendations. *Am Soc Hand Therap* 2015.
 27. **Lam NW, Goh HT, Kamaruzzaman SB, Chin AV, Poi PJ, Tan MP.** Normative data for hand grip strength and key pinch strength, stratified by age and gender for a multiethnic Asian population. *Singapore Med J* 2016, 57(10):578-84.
 28. **Mitsionis G, Pakos EE, Papakostas T, Beris AE.** Normative data on hand grip strength in a Greek adult population. *Int Orthop* 2009, 2009:713-7.
 29. **Dongare-Bhor S, Lohiya N, Maheshwari A, Ekbote V, Chiplonkar S, Padidela R, Mughal Z, Khadilkar V, Khadilkar A.** Muscle and bone parameters in underprivileged Indian children and adolescents with T1DM. *Bone* 2020, 130:115074.
 30. **Wallymahmed ME, Morgan C, Gill GV, MacFarlane IA.** Aerobic fitness and hand grip strength in Type 1 diabetes: relationship to glycaemic control and body composition. *Diabet Med* 2007, 24(11):1296-9.
 31. **Fricke O, Seewi O, Semler O, Tutlewski B, Stabrey A, Schoenau E.** The influence of auxology and long-term glycemic control on muscle function in children and adolescents with type 1 diabetes mellitus. *J Musculoskelet Neuronal Interact* 2008, 8(2):188-95.
 32. **Maratova K, Soucek O, Matyskova J, Hlavka Z, Petruzekova L, Obermannova B, Pruhova S, Kolouskova S, Sumnik Z.** Muscle functions and bone strength are impaired in adolescents with type 1 diabetes. *Bone* 2018, 106:22-7.
 33. **Liang X, Jiang CQ, Zhang WS, Zhu F, Jin YL, Cheng KK, Lam TH, Xu L.** Glycaemia and hand grip strength in aging people: Guangzhou biobank cohort study. *BMC Geriatr* 2020, 20(1):1-0.
 34. **Zhang Y, Liu X, Jia J, Zhang Q, Lin Y, Zhang L, Lu Q, Lv H, Zheng X.** Diabetic polyneuropathy and carpal tunnel syndrome together affect hand strength, tactile sensation and dexterity in diabetes patients. *J Diabetes Investig* 2021, 12(11):2010-8.
 35. **Alqahtani BA, Alenazi AM, Elnaggar RK, Alshehri MM, Alhowimel A, Najmi AA, Alasraj M, Alghadeir M.** Normative values for hand grip and pinch strength for 6 to 18-year-olds in Saudi Arabia. *BMC Musculoskelet Disord* 2023, 24(1):1-8.
 36. **Bechtold S, Dirlenbach I, Raile K, Noelle V, Bonfig W, Schwarz HP.** Early manifestation of type 1 diabetes in children is a risk factor for changed bone geometry: data using peripheral quantitative computed tomography. *Pediatrics* 2006, 118(3):e627-34.
 37. **Fujita S, Rasmussen BB, Cadenas JG, Grady JJ, Volpi E.** Effect of insulin on human skeletal muscle protein synthesis is modulated by insulin-induced changes in muscle blood flow and amino acid availability. *Am J Physiol Endocrinol Metab* 2006, 291(4):E745-54.
 38. **Mori H, Kuroda A, Araki M, Suzuki R, Taniguchi S, Tamaki M, Akehi Y, Matsuhisa M.** Advanced glycation end-products are a risk for muscle weakness in Japanese patients with type 1 diabetes. *J Diabetes Investig* 2017, 8(3):377-82.
 39. **Martínez-Torres J, Gallo-Villegas JA, Aguirre-Acevedo DC.** Normative values for handgrip strength in Colombian children and adolescents from 6 to 17 years of age: estimation using quantile regression. *J Pediatr* 2022, 98(6):590-8.
 40. **Rauch F, Neu CM, Wassmer G, Beck B, Rieger-Wettengl G, Rietschel E, Manz F, Schoenau E.** Muscle analysis by measurement of maximal isometric grip force: new reference data and clinical applications in pediatrics. *Pediatr Res* 2002, 51(4):505-10.
 41. **Ran-hui C, Sil LG, Yeon YJ, Bog RO, Duk JY.** Hand Grip and Leg Muscle Strength in Hemodialysis Patients and Its Determinants. *J Korean Med Sci* 2021, 36(11):1-3.
 42. **Cheung C-L, Nguyen U-SDT, Au E, Tan KCB, Kung AWC.** Association of handgrip strength with chronic diseases and multimorbidity. *Age* 2013, 35(3):929-41.