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Oral microbiota in children with type 1 diabetes mellitus

Microbiota jamy ustnej u dzieci z cukrzycą typu 1

¹Marek Pachoński, ²Aleksandra Koczor-Rozmus, ²Katarzyna Mocny-Pachońska, ²Patrycja Łanowy, ³Anna Mertas, ⁴Przemysława Jarosz-Chobot

¹NZOZ Pachońscy Dental Clinic Marek Pachoński, Tarnowskie Gory, Poland

²Department of Conservative Dentistry with Endodontics, Faculty of Medical Science, Medical University of Silesia, Katowice, Poland

³Department of Microbiology and Immunology, Faculty of Medical Science, Medical University of Silesia, Katowice, Poland

⁴Department of Children's Diabetology, Medical University of Silesia, Katowice, Poland

Abstract

Introduction: The oral cavity is a unique ecosystem in which the microbiome is formed by the colonization of billions of bacteria, viruses, and fungi.

Aim of the study was to assess the quality and quantity of bacterial microbiota of the oral cavity in children with type 1 diabetes in comparison to healthy children.

Material and methods: The study group consisted of 50 randomly selected type 1 diabetic children aged 10–18 years divided into 2 groups (25 people each) according to the level of diabetes control: well-controlled group (WC) – $HbA_{1c} \le 7.5\%$) and poorly-controlled group (PC) – $HbA_{1c} \ge 7.5\%$. The control group (GC) consisted of 25 randomly selected children with no systemic diseases. The material for microbiological tests was collected as two swabs: one from the bottom of the oral cavity and the other one from the posterior part of the dorsum of the tongue).

Results: The statistical analysis revealed statistically significant differences in the total number of isolated microorganisms between PC and GC (*post hoc* test p = 0.003) and WC and GC (*post hoc* test p < 0.001). There were no statistically significant differences in the number of isolated microorganisms between PC and WC (p = 0.195).

Conclusions: Oral microbiome in type 1 diabetic children is significantly quantitatively different in comparison to healthy children. There are also visible qualitative differences in the profile of oral microbiota in type 1 diabetic children and healthy children.

Key words:

periodontitis, microbiota, dental caries, Candida albicans, type 1 diabetes.

Streszczenie

Wprowadzenie: Jama ustna stanowi unikalny ekosystem, który kolonizują biliony bakterii, wirusów i grzybów.

Cel pracy obejmował ocenę ilościową i jakościową mikroflory jamy ustnej u dzieci z cukrzycą typu 1 w porównaniu z dziećmi zdrowymi. **Materiał i metody:** Materiał mikrobiologiczny pobierano w postaci dwóch wymazów: z dna jamy ustnej oraz z grzbietu języka u 50 losowo wybranych dzieci z cukrzycą typu 1 w wieku 10–18 lat. Uczestników badania zakwalifikowano do dwóch grup (po 25 osób każda) w zależności od stopnia wyrównania cukrzycy: z cukrzycą wyrównaną (WC, HBA_{1c} < 7,5%) i z cukrzycą niewyrównaną (PC, HBA_{1c} \geq 7,5). Grupa kontrolna (GC) obejmowała 25 losowo wybranych zdrowych dzieci w wieku 10–18 lat.

Wyniki: Analiza statystyczna wykazała istotną statystycznie różnicę w całkowitej ilości wyizolowanych szczepów drobnoustrojów pomiędzy grupami PC i GC (*post hoc* test p = 0,003) and WC i GC (*post hoc* test p < 0,001). Różnicy tej natomiast nie wykazano pomiędzy grupami PC i WC (p = 0,195).

Wnioski: Mikrobiota jamy ustnej dzieci z cukrzycą typu 1 różni się istotnie pod względem ilościowym w stosunku do dzieci zdrowych. Zaobserwowano również wyraźne różnice jakościowe w składzie mikroflory jamy ustnej pomiędzy badanymi dziećmi z cukrzycą typu 1 a zdrowymi.

Słowa kluczowe:

mikrobiota, cukrzyca typu 1, próchnica zębów, Candida albicans, choroby przyzębia.

Introduction

The oral cavity is a unique ecosystem in which teeth, gingival groove, tongue, cheeks, soft and hard palate, and tonsils coexist to create a special environment. It is covered with mucosa, which consists of keratinized or nonkeratinized squamous stratified epithelium, depending on the localization and function. The oral cavity, being an extension of the surrounding external environment, is colonized by billions of bacteria, viruses, and fungi that form the microbiome of the oral cavity. This commensal microbiota coexists in symbiosis with its host, but when an imbalance occurs, the disease develops. This imbalance is called dysbiosis. Dental caries and periodontal diseases, the main dental conditions, are caused by dysbiosis. In addition to the commensal flora in the oral cavity, there also occurs a transient flora, e.g. the intestinal flora, which resides in the oral cavity for a short period of time due to difficulty in acquiring nutrients [1, 2].

The microbiota of the oral cavity is also a risk factor for many systemic diseases such as tumours [3], diabetes [4], cardio-vascular diseases [5], sepsis [6], preterm births, and low birth weight of infants [7]. More than 700 species of bacteria were identified with culture-independent approaches, and more than 250 were isolated, cultured, and named [8]. Undoubtedly, new species are still being identified [9, 10].

Salivary microbial

The greatest number and diversity of microbiota are observed in dental plaque, then in saliva, and on the oral mucosa. The dominant fraction of microbiota in the saliva is like that found in the throat and tonsils. In addition, saliva also contains bacteria from supra and subgingival niches. The main types of bacteria noticed in saliva include *Actinobacteria*, *Bacteroidetes*, *Firmicutes*, *Fusobacteria*, and *Proteobacteria* [11]. It has been proven that the salivary microbial configuration is more dependent on the influence of the environment than on genetic conditions [12].

Dental plaque

Dental plaque is a type of biofilm located on the surface of the teeth. There are 2 plaque classifications: supragingival plaque (placed above the gingival margin) and subgingival plaque (below the gingival margin). The composition of the subgingival plaque depends on the balance between health and disease and is associated with periodontal destruction. Microbiological research of this niche focuses mainly on changes in the microbiota that occur during the development of periodontal diseases. In gingivitis, which is an inflammatory process confined to the gum and most often associated with the accumulation of plague, the dominant microorganisms are: Streptococcus, Actinomycetes, Capnocytophaga, Campylobacter, Eikenella, Fusobacterium, and Prevotella. On the other hand, in the case of periodontitis, which is a chronic inflammatory process involving not only the destruction of the gingiva, but also tooth-supporting tissues (periodontal membrane and alveolar bone), the subgingival microbiota in the course of periodontitis is very complex. Over 400 phenotypes have been isolated from periodontal pockets [13].

Oral mucosa

Compared with the other parts of the oral cavity, microorganisms existing on the mucosa are few. The dorsum of the tongue is the most colonized, by bacteria associated with halitosis. Among the bacteria on its surface in healthy populations, we can identify *Streptococcus salivarius*, *Rothia mucilaginosa*, and unidentified, cultivable species of *Eubacterium* strain FTB41 [13].

Diabetes mellitus

Various research studies on the composition of the subgingival bacterial microbiota in diabetics compared to healthy individuals have been performed. However, no clear influence of diabetes on bacterial flora has been demonstrated.

Certain species of bacteria are more common in diabetic patients. However, it is difficult to confirm whether their occurrence is directly related to a change in the subgingival environment or indirectly due to a change in the host's immune response. Diabetic individuals are more prone to chronic periodontitis as a result of hyperglycaemia that alters the subgingival environment such that pathogenic bacteria become dominant. Alternatively, diabetes can alter the host response to plaque, causing more tissue destruction. Considering the elevated glucose level in the subgingival environment and the altered or inhibited host immune response, real differences can be found in the subgingival microbiome in diabetic patients in comparison to healthy individuals [14].

Aim of the study

The aim of the study was to analyse and compare the composition of bacterial oral microbiota in children with type 1 diabetes and healthy children.

Material and methods

Characteristics of the research groups

The study group consisted of 50 randomly selected children aged 10–18 years, who had had type 1 diabetes for at least 5 years. The subjects were patients of the regional diabetic clinic at the Upper John Paul II Silesian Children's Health Centre University Hospital in Katowice, Poland.

Patients from the study group were divided into 2 groups (25 people each) according to their level of diabetes control. The allocation of patients to the groups was based on the qualification criterion, which was the level of glycated haemoglobin (HbA_{1c}). According to the recommendations of the American Diabetes Association (ADA) and the International Society for Paediatric and Adolescent Diabetes (ISPAD 2014), the threshold value of HbA_{1c} was set at 7.5% [15]. Patients with good glycaemic control were assigned to the Well Controlled group (WC, HbA_{1c} \leq 7.5%). The mean age of the subjects in the WC group was 14.40 ±2.0 years. The group consisted of

10 girls and 15 boys. Twenty-three patients in this group were using a personal insulin pumps (continuous subcutaneous insulin infusion-CSII) and 2 patients were using insulin pens (multiple daily insulin injection-MDII). The average level of gly-cated haemoglobin was 6.70 \pm 0.40%. The Poorly Controlled (PC, HbA_{1c} > 7.5%) group consisted of patients with poor gly-caemic control. The mean age of the subjects in the PC group was 14.92 \pm 1.87 years. The group consisted of 14 girls and 11 boys. Eighteen patients in this group were using personal insulin pumps (CSII) and 7 were using insulin pens (MDII). The mean glycated haemoglobin level in this group was 8.23 \pm 0.64%.

The control group (GC) consisted of 25 randomly selected healthy children who were treated at the NZOZ Pachońscy Dental Clinic in Tarnowskie Góry, Poland. The mean age of the subjects was 14.52 \pm 1.29 years. The group consisted of 13 girls and 12 boys.

The study groups were homogeneous in terms of sex and age (χ^2 Pearson test: p = 0.50, p = 0.41). The study was conducted in the period 2014–2018.

Exclusion criteria from the research

The following groups were excluded: Patients and/or legal guardians, parents not agreeing to participate in the study, patients not willing to cooperate in the study, and those additionally burdened with other diseases (thyroid diseases, celiac disease, arterial hypertension).

In the control group, the exclusion criteria were diabetes and all systemic disorders, as well as refusal to participate in the study.

The tests were performed anonymously, and each patient was given a code number. The parents or legal guardians of all participants gave their written consent to participate in the study. The research project was approved by the Bioethical Commission of the Silesian Medical University in Katowice (No. KNW / 0022 / KB1 / 26 / I / 14 of 22/04/2014) and conducted in accordance with the latest version of the Helsinki declaration

Collection of the samples to microbiological tests

The material for microbiological tests was collected from each of the subjects as 2 swabs: one from the bottom of the oral cavity (BOC bacteria) and the other from the posterior part of the dorsum of the tongue (DT bacteria). The swabs were collected using sterile swabs, which were then placed in the transport medium and delivered to the Department of Microbiology and Immunology of the Medical University of Silesia in Zabrze, where the laboratory microbiological tests were carried out. The material was collected in the morning between 08:00 and 10:00 a.m. The participants were asked to attend the examination with an empty stomach and before completing any hygienic procedures in the oral cavity.

Laboratory microbiological tests

Microbiological tests were carried out using classical methods routinely used in microbiological diagnostics. The material collected from the examined patients was cultured on appropriate culture media to multiply and then isolate pure microbial cultures. Aerobic bacteria were multiplied on solid medium Columbia agar with a 5% addition of sheep blood at 37°C. Anaerobic bacteria were multiplied on Schaedler K3 solid medium with a 5% addition of sheep blood at 37°C in an-aerobic conditions obtained using Biomerieux Genbag Anaer kits (Marcy l'Etoile, France). Candida fungi were multiplied and initially identified using chromogenic medium ChromID Candida from Biomerieux (Marcy l'Etoile, France).

After isolation and multiplication of the cultured strains of microorganisms, their species identification was carried out using the following sets of reagents: ENTEROtest 24 N, NEFER-Mtest 24 N, STREPTOtest 24, STAPHYtest 24, ANAEROtest 23, OXItest, PYRAtest, and the computer program TNW_lite 6.5 for species identification of Erba-Lachema microorganisms (Brno, Czech Republic). The following biochemical tests from Biomerieux (Marcy l'Etoile, France) were also used: Catalysis, Slidex Staph Kit, and API Candida. Execution, reading, and interpretation of the test results were performed in accordance with the recommendations of manufacturers of diagnostic reagent kits.

Statistical analysis

All statistical analyses were performed using the STATIS-TICA Version 13 program package (StatSoft, Tulsa, OK, USA), the SciPy module, and the Statsmodels Phyton module. All charts were prepared using Excel (Microsoft Office, Redmond, Washington, USA). Continuous variables between the groups were compared using the Kruskal-Wallis and Mann-Whitney U tests because the analysed variables were not normally distributed.

All statistical analyses were supplemented with the results of 2 post-hoc tests: the Dunn-Bonferroni and Conover tests. To test the differences of nominal variables between the groups, the Pearson chi-square test was used. P values <0.05 were considered statistically significant.

Results

The results are presented in the form of box plots. The upper and lower boundaries of the box determine the first (Q1) and third quartiles (Q3) of the variable distribution, respectively. The line in the centre of the box is the median. Lower and upper whisker indicate the minimum and maximum values. The cultures were assessed according to the number of microbial species. The number of strains of a given species of microorganisms was determined. The authors counted the number of different microorganism strains that were isolated from mucosa of each particular subject per participant for the 3 mentioned groups of children: PC, WC, and GC.

Figure 1 presents the results of the Kruskal-Wallis test used to assess the presence of any significant differences in the number of all bacteria isolated from the bottom of the oral cavity (Bacteria BOC) between all study groups. These differences were shown to be statistically significant (p < 0.001), and as such the results of the statistical analysis were supported by 2 of the post-hoc tests: the Dunn-Bonferroni test and

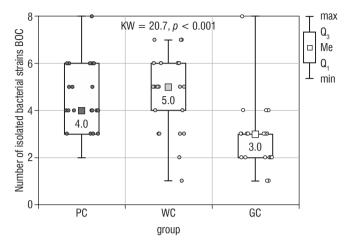


Figure 1. The number of bacterial strains isolated from the bottom of the oral cavity in the study groups of children (Kruskal-Wallis test)

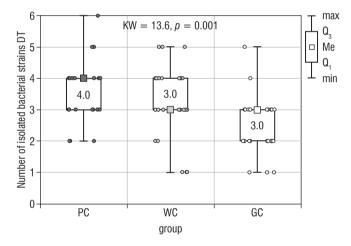


Figure 2. The number of bacterial strains isolated from the dorsum of the tongue in the study groups of children (Kruskal-Wallis test)

the Conover test. *Post-hoc* analysis revealed that the median number of different strains isolated by BOC swab was significantly lower in the GC group compared to the PC (p = 0.001) and WC groups (p < 0.001) (Table I). The obtained median in PC group was 4.0, 5.0 in WC group, and 3.0 in GC group. The highest number of isolated strains of bacteria [8] in one participant was found in the PC and GC groups. In the WC group this value was 7.

The number of all isolated bacteria from the tongue's dorsum (Bacteria DT) also differed significantly between the examined groups of children (the Kruskal-Wallis test; p = 0.001; Fig. 2). The results of the Dunn-Bonferroni test showed statistically significant differences between the WC (median 3.0) and GC (median 3.0) groups as well as between the PC (me**Table I.** Results of post hoc tests – number of isolated bacterial strains (BOC)

	Post hoc	Dunn-Bonferroni		
	р	PC	WC	GC
Conover	PC		0.623	0.001
	WC	0.222		< 0.001
	GC	0.028	0.019	

 $\mathsf{PC}-\mathsf{poorly}$ controlled group; $\mathsf{WC}-\mathsf{well}$ controlled group; $\mathsf{GC}-\mathsf{control}$ group

 Table II. Results of post hoc tests – number of isolated bacterial strains (DT)

	Post hoc	Dunn-Bonferroni		
	р	PC	WC	GC
Conover	PC		0.368	0.001
	WC	0.166		0.021
	GC	0.029	0.058	

 $\mathsf{PC}-\mathsf{poorly}$ controlled group; $\mathsf{WC}-\mathsf{well}$ controlled group; $\mathsf{GC}-\mathsf{control}$ group

dian 4.0) and GC groups. The second post-hoc test revealed the statistically significant difference only between PC and GC groups (Table II). There were no statistical differences between the studied groups of children with diabetes.

The conducted statistical analysis revealed statistically significant differences in the total number of all isolated microorganisms (Bacteria BOC & DT) between PC and GC groups (Kruskal Wallis test; p < 0.001; Fig. 3). The highest number of isolated bacterial strains [10] was obtained in the PC group. In the WC and GC groups the value was 9. The medians obtained in the PC, WC, and GC groups were: 6, 7, and 4, respectively. There were no statistically significant differences in the number of isolated microorganisms between the PC and WC groups (Table III).

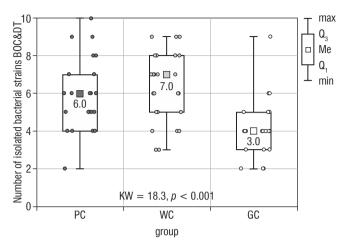


Figure 3. The number of bacterial strains isolated from the bottom of the oral cavity and the dorsum of the tongue in the study groups of children (Kruskal-Wallis test)

As a result of the microbiological examination, the number of total_microorganisms isolated in the PC group was 160 species. In the WC group, 158 microorganisms were cultured, and in the control group 103.

The microorganisms cultured were assigned to the following groups: gram-positive streptococci, gram-negative streptococci, gram-positive coccobacillus, and gram-negative coccobacillus (Table IV).

A separate group isolated during the microbiological examination was the yeast *Candida albicans*. In the PC group, *Candida albicans* was isolated in 8 children, in the WC group in 4 diabetic children, and in the GC group in 5 healthy children. The obtained results were not statistically significant (p = 0.21; Table IV).

The largest group of isolated microorganisms comprised bacteria from the Streptococcus genus. As a result of the study, the following streptococci were cultured: *Streptococcus acidominimus*, *S. cristatus*, *S. mitis*, *S. oralis*, *S. pneumoniae*, *S. pyogenes*, *S. sanguinis*, and *S. salivarius*. Statistical calculations demonstrated that the total number of *Streptococcus* differed significantly between the studied groups (p = 0.018; Fig. 4). Both of the post hoc tests showed a significant difference between the WC and GC groups (Table V).

Statistical analysis was also performed using the Mann-Whitney test between the combined groups of children with diabetes (PC & WC = GD) and the control group. This analysis showed a statistically significantly higher number of bacterial strains isolated from the bottom of the oral cavity and the dorsum of the tongue in the group of children with diabetes (GD) compared to the control group (Mann-Whitney U test; p < 0.001). Similar data were obtained with the *Streptococcus* strain. The GD group was characterized by a significantly higher number of these bacteria as compared to the control group (Mann-Whitney U test; p = 0.014; Figs. 5 and 6).

 Table III. Results of post hoc tests – number of isolated bacterial strains (BOC&DT)

	Post hoc	Dunn-Bonferroni		
	р	PC	WC	GC
Conover	PC		0.507	0.003
	WC	0.195		< 0.001
	GC	0.035	0.022	

 $\mathsf{PC}-\mathsf{poorly}$ controlled group; $\mathsf{WC}-\mathsf{well}$ controlled group; $\mathsf{GC}-\mathsf{control}$ group

Discussion

Caries is the most common disease of the oral cavity to which humans are susceptible during their lifetime. This disease not only results in tooth destruction but also affects the pulp and periapical tissues. The most pathogenic bacteria related to the development of caries is Streptococcus mutans. Nevertheless, other bacteria including Streptococcus, Veillonella, Actinomyces, Granulicatella, Leptotrichia, Thiomonas, Bifidobacterium, and Prevotella species were detected in cases of S-ECC [16, 17]. A significantly higher number of bacteria from the Streptococcus genus were found in the group of children with well-controlled diabetes mellitus compared to healthy children. The samples that were taken from the floor of the oral cavity also revealed differences in the number of described bacteria between both examined WC and PC groups of diabetics. Streptococcus mitis, S. salivarius, S. sanguinis, and S. acidominimus were the most abundantly represented. The study of Janem et al. showed that the microbiological salivary profile of diabetics did not differ statistically significantly from healthy people [18]. Nevertheless, Kampoo et al. noticed distinctly higher levels of Streptococcus and Lactobacillus in diabetics with caries [19].

Our own research also noted an increased amount of *Strep-tococcus mitis* in the group of children with type 1 diabetes in comparison to healthy children. This bacterium was isolated in 88% of children in the PC group, 92% in the WC group, and 64% in the control group. *Streptococcus mitis* is classified as "low pH streptococcus" – this microorganism can reduce the pH of glucose broth to below 4.4. Species with similar abilities include *S. oralis, S. anginosus, and S. gordonii.* The organisms mentioned above are considered to have low carious potential and are listed amongst the primary colonizers of the pellicle. However, their capability of decreasing the pH level and therefore the occurrence of caries is confirmed by numerous studies [20, 21].

Table IV. Species of oral microbiota isolated from studied groups of children

solated microorganisms	Number of microorganisms		
	PC	WC	GC
Gram-positive cocci			
Anaerococcus prevotti	_	_	1
Gemella spp.	5	9	1
Globicatella sanguinis	1	_	_
Microccus spp.	_	1	_
Peptococcus niger	_	1	1
Streptococcus acidominimus	7	14	8
Streptococcus cristatus	_	_	1
Streptococcus mitis	22	23	16
Streptococcus oralis	4	4	_
Streptococcus pneumoniae	_	2	6
Streptococcus pyogenes	_	_	1
Streptococcus sanguinis	14	12	9
Streptococcus salivarius	16	16	11
Staphylococcus aureus MSSA	6	4	1
Staphylococcus epidermidis MSCNS	6	1	_
Staphylococcus lentus	_	1	_
Streptococcus warneri	1	_	_
Sarcina spp.	1	_	_
Gram-negative cocci			
Nesiseria subflava	7	8	1
Neisseria spp.	16	13	24
Veillonella parvula	7	12	2
Gram-positive rods			
Atopobium parvulum	_	_	1
Bifidobacterium adolescentis	6	_	_
Bifidobacterium breve	1	2	3
Bifidobacterium dentium	1	_	_
Bifidobacterium longum	_	4	_
Eubacterium tortuosum	_	1	_
Lactobacillus acidophilus	_	1	_
Lactobacillus catenaformis	_	1	1

Table IV. Species of oral microbiota isolated from studied groups of children (cont.)

solated microorganisms	Number of microorganisms		
	PC	WC	GC
Lactobacillus fermentum	_	1	_
Propionibacterium acnes	1	1	_
Propionibacterium propionicum	1	2	_
Pseudoflavonifractor capilosus	_	1	1
Actinomyces israeli	4	_	_
Actinomyces naeslundi	4	6	1
Actinomyces odontolyticus	2	1	_
Actinomyces viscosus	_	1	_
Blauta producta	-	1	-
Clostridium barati	_	_	2
Clostridium chauvoei	2	4	-
Clostridium clostridioforme	3	_	-
Clostridium histoliticum	1	_	1
Clostridium novyi	_	1	_
Clostridium perfringens	4	_	-
Clostridium ramosum	2	_	-
Clostridium tertium	_	_	1
Gram-negative rods			
Bacteroides ovatus	_	1	_
Campylobacter gracilis	_	_	1
Campylobacter ureolyticus	_	1	2
Capnocytophaga ochracea	5	_	_
Citrobacter freundii	_	1	_
Enterobacter cloacae	_	1	_
Klebsiella oxytoca	_	1	_
Klebsiella pneumoniae	_	1	_
Mitsuokella multiacida	1	_	_
Proteus mirabilis	_	_	1
Serratia marcescens	1	_	_
Fungi			
Candida albicans	8	4	5
he total number	160	158	103

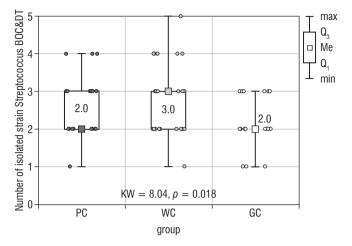


Figure 4. The number of *Streptococcus* strain isolated from the bottom of the oral cavity and the dorsum of the tongue in the study groups of children (Kruskal-Wallis test)

Candida spp. are commensals detected in the oral cavity among healthy individuals. However, their occurrence and virulence are significantly increased in immunocompromised patients. They may become pathogenic in the case of physiological changes in the host and cause oral candidiasis or invasive systemic infection [22]. Numerous studies identified an increased incidence of Candida spp. in type 1 diabetes, which was closely related to diabetes duration and level of decompensation [23]. Soysa et al. demonstrated that the incidence of that fungus on mucosa may concern 80% of subjects [24]. The study conducted by the authors did not reveal statistically significant differences in the number of isolated Candida spp. (p = 0.21). In the PC group Candida spp. were detected in 8 cases, in the WC group in 6 cases, and in the GC group in 5 cases. It is certainly related to the fact that the study group consisted of children aged 10-18 years with no periodontitis [25]. Candida albicans strains are often isolated in cases of co-existing diabetes and periodontitis. Sardi et al. revealed that the incidence of Candida albicans reaches 63.63% among patients with diabetes mellitus and chronic periodontitis [22].

Hintao *et al.* showed an increased incidence of *Trepone-ma denticola*, *Streptococcus sanguinis*, *Prevotella nigrescens*, *Staphylococcus intermedius*, and *Streptococcus oralis* in the subgingival plaque of people with type 2 diabetes compared to healthy people, despite the lack of statistically significant differences in the examined subgingival plaque samples [26].

Table V. Results of post hoc tests – number of Streptococcusstrain (BOC&DT)

	Post hoc	Dunn-Bonferroni		
	р	PC	WC	GC
Conover	PC		0.242	0.228
	WC	0.140		0.007
	GC	0.137	0.049	

 $\mathsf{PC}-\mathsf{poorly}$ controlled group; $\mathsf{WC}-\mathsf{well}$ controlled group; $\mathsf{GC}-\mathsf{control}$ group

A similar composition of bacterial flora was observed among patients with type 1 DM and in the follow-up of nondiabetic diabetics with periodontitis [27].

Limitations of the study

The performed research relates to an interesting issue. However, we may be limited by an insufficient number of examined patients. On the other hand, the research was carried out on patients of the largest Diabetology Clinic in Upper Silesia, and applying the inclusion and exclusion criteria also significantly reduced the number of children who could take part in the research. Conducting the study in other regions of Poland to examine the possible influence of environmental factors on oral microbiota would be informative.

Conclusions

The oral microbiome of children with type 1 diabetes in quantitative terms is significantly different in comparison to healthy children. There are also visible qualitative differences in the profile of oral microbiota in children with type 1 diabetes versus healthy children.

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