

Article

Not peer-reviewed version

Clinical Assessment of Plaque Removal using a Novel Dentifrice containing Cellulose Microfibrils

[Mohamed E. Labib](#)^{*}, Antonio Perazzo , James Manganaro , Yacoub Tabani , Kimberly R. Milleman , Jeffery L. Milleman , [Laurence James Walsh](#)

Posted Date: 6 October 2023

doi: 10.20944/preprints202310.0248.v1

Keywords: oral hygiene; dentifrices; cellulose; nanoparticles; dental plaque



Preprints.org is a free multidiscipline platform providing preprint service that is dedicated to making early versions of research outputs permanently available and citable. Preprints posted at Preprints.org appear in Web of Science, Crossref, Google Scholar, Scilit, Europe PMC.

Copyright: This is an open access article distributed under the Creative Commons Attribution License which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Article

Clinical Assessment of Plaque Removal Using a Novel Dentifrice Containing Cellulose Microfibrils

Mohamed E. Labib ^{1,*}, Antonio Perazzo ¹, James Manganaro ¹, Yacoob Tabani ¹,
Kimberly R. Milleman ², Jeffery L. Milleman ² and Laurence J Walsh ³

¹ NovaFlux, Inc, Princeton, NJ, USA; labib@novaflux.com

² Salus Research Inc, Fort Wayne, IN, USA; Milleman@SalusResearch.us

³ The University of Queensland School of Dentistry, Brisbane, Australia; l.walsh@uq.edu.au

* Correspondence: labib@novaflux.com; Tel.: +1 609 683 0215

Abstract: Dentifrice performance in the removal of dental plaque is influenced by the interactions of dentifrice components with tooth surfaces. This randomized controlled clinical study assessed the effectiveness and safety of a novel fluoride dentifrice formulation that included a micro-fibrillated cellulose network with entangled microcrystalline cellulose and silica particles (Protegera™), and compared this to a positive control fluoride dentifrice (Crest Cavity Protection™). Whole mouth dental plaque levels in 82 healthy adults were measured after the first supervised use, and following a week of twice daily use at home. Overall, the test dentifrice was at least three times more effective in whole mouth plaque reductions than the reference dentifrice ($p < 0.001$), with a 38.6% reduction on first use, a 30.9% reduction at day 7, and a 41.6% reduction from day 1 to day 7, compared to reductions of 12.0%, 9.6%, and 11.6% respectively for the positive control. No safety issues arose from the use of the test dentifrice. These results indicate that including micro-fibrillated cellulose enhances plaque removal effectiveness, without causing adverse changes to oral soft tissues.

Keywords: oral hygiene; dentifrices; cellulose; nanoparticles; dental plaque

1. Introduction

Twice daily mechanical toothbrushing using a fluoride dentifrice and interdental cleaning are considered key self-performed oral health behaviors [1,2]. While toothbrushing has value for preventing dental plaque-induced oral diseases [3–5], many patients struggle with effectively performing mechanical tooth cleaning [6], and this has prompted the development of powered brush technologies, and enhancements to the formulation of fluoride dentifrices. Despite such improvements, it is common for patients to have dental plaque deposits remaining in most regions of the dentition at the completion of a toothbrushing episode [7–13].

A contributing factor to the less-than-ideal performance of modern toothbrushes and dentifrices is that the collective mechanical action of a toothbrush with a dentifrice may be less than ideal, in terms of the forces that are generated on the tooth surfaces where plaque deposits are present [14,15]. A specific issue is the interaction of tooth brush bristles with the tooth surface. Viscosity modifiers (including glycerol and polymeric thickeners) and detergents may act as lubricants between dental plaque deposits and toothbrush bristles, thus limiting the mechanical removal of dental plaque [14–17]. This lubricating effect is enhanced by stimulated saliva that dilutes the dentifrice within the oral cavity, creating a slurry that does not have suitable mechanical properties to remove plaque.

Given these considerations, it is not surprising that numerous studies have demonstrated that dentifrices are ineffective in improving the action of the toothbrush in mechanically removing plaque biofilm [7–13]. The 2016 systematic review undertaken by Valkenburg et al. [13] included 20 comparisons of toothbrushing with a dentifrice compared to brushing without a dentifrice. The extent of plaque removal from brushing alone was on average 50.3%, while brushing with a dentifrice achieved a reduction of just 49.2%. Several meta-analyses confirmed that there is no significant difference when toothbrushing with and without a dentifrice.

To enhance the plaque removal capabilities of a dentifrice, one approach is to include a material made of fibrils that when formulated as a dentifrice forms a microstructural network with tailored mechanical properties [18,19] which can create the shear stress needed to remove dental plaque as it moves across the surface of dental plaque deposits. This allows the dentifrice to remove plaque even in dental areas not reached by the brush bristles, and makes it less prone to interference from lubricant effects. Such materials can be of natural origin, e.g., such as micro-fibrillated cellulose (MFC) derived from wood pulp, and can include (micro)fibrils of different thicknesses and lengths. Trapped within these microscale three-dimensionally entangled networks of fibers can be included abrasive particles such as silica [18,19], or microcrystalline cellulose and polymeric thickeners as needed. Due to the interconnected entangled microstructure, the clusters of microfibrils do not fully breakdown into a slurry upon saliva-induced dilution, and maintain sufficient mechanical properties that enhance biofilm removal [20,21]. Any abrasive particles that are trapped within the fiber network will be dragged along the surface of the tooth as fluid flow occurs, causing some of these to drag against the surfaces being cleaned. This allows the fiber networks to make direct contact with the plaque biofilm under normal brushing conditions. At the same time, fibers can become entangled with stacks of the biofilm, and cause these to become detached. This technology has been deployed for cleaning channels within medical devices [18,19,22,23].

A fluoride dentifrice based on this same concept is Protegera™ (Protegera, Inc., Madison, WI, USA), where the cellulose fibers are derived from the wood pulp of Norwegian spruce trees using a mechanical process [20,21]. MFC is a "Generally Recognized As Safe" (GRAS) ingredient by the US Food and Drug Administration and is approved as a food additive. In this dentifrice, MFC fibers form an extended toothpaste-containing matrix that is designed to dislodge deposits of dental plaque, including from regions beyond the reach of toothbrush bristles. The water-insoluble MFC fiber matrix protects the incorporated toothpaste ingredients from significant saliva-induced material microstructural breakdown known to occur in traditional toothpaste made with soluble polymers such as carboxymethyl cellulose (CMC), xanthan gum and carrageenan. Brushing provides the shear stress needed to force the MFC fiber matrix to dislodge the plaque biofilm from tooth surfaces. MFC has a low relative dentin abrasivity (RDA) score (RDA = 5), and hence the fiber complexes clean tooth surfaces without damaging the enamel, dentin, or oral mucosal tissues. MFC does not exert any direct antimicrobial actions.

Since no past studies have explored the clinical performance of a dentifrice containing cellulose microfibrils, the aim of the present study was to assess the plaque removal efficacy and safety of Protegera™ dentifrice immediately after the first, supervised use and following a week of twice, daily use at home, and compare this to a commonly-used fluoride dentifrice (Crest Cavity Protection™). The null hypothesis was that there would be no difference between the two dentifrices.

2. Materials and Methods

2.1. Study design

The study used a randomized, single-center, examiner-blinded, parallel design. Subjects were randomized 1:1 between the test product (Protegera™ dentifrice, Protegera, Inc., Madison, WI, USA) and the control (Crest Cavity Protection™, Procter & Gamble, Cincinnati, OH, USA) based on a computer-generated randomization schedule prepared in advance of the study. The staff responsible for random assignments were blinded to the treatment protocol.

The control fluoride dentifrice contained 0.243% sodium fluoride (1500 ppm fluoride), with the following additional ingredients: sorbitol, water, hydrated silica, sodium lauryl sulfate, trisodium phosphate, flavor, cellulose gum filler, sodium phosphate, carbomer, sodium saccharin, titanium dioxide, and blue 1 color. The test dentifrice also contained 0.243% sodium fluoride (1500 ppm fluoride), with the following additional ingredients: water, glycerol, hydrated silica, microcrystalline cellulose, microfibrillated cellulose, peppermint flavor, cocamidopropyl betaine, sodium gluconate, carbomer, sodium benzoate, Gantrez™ S-97, Poloxamer 407, xanthan gum, and sodium hydroxide.

The test dentifrice had an RDA value of 88. The flavor was peppermint (provided by DSM-Firmenich), to match the most common flavor used in commercial dentifrices.

Plaque levels were assessed at baseline, and then following a one-minute single supervised brushing period with the assigned dentifrice, and then finally after 7 days of twice daily unsupervised use at home. For the post-brushing plaque evaluations, the clinical examiner was blinded to the product codes/identification and product assignments. Subjects were queried regarding adverse events both after the initial brushing episode and at the 7-day review.

2.2. Recruitment and selection

A total of 82 subjects, aged between 19 and 69 years of age, were enrolled to ensure that approximately 80 subjects would complete the study. All subjects resided in Fort Wayne, IN, USA. The sample size was based on a power analysis. With 40 subjects per group, there is 80% power to detect a 0.634 effect size (mean difference between treatment groups divided by the pooled SD for the mean change from baseline) for a two-sided t-test with 5% type 1 error. This corresponds to detecting a large effect size, corresponding to a meaningful superiority in plaque reduction.

Informed consent was obtained from each subject prior to participation, in line with US Food and Drug Administration Good Clinical Practice guidelines for clinical studies. Information was given in both oral and written form, and subjects were given ample opportunity to inquire about details of the study prior to signing and dating the consent form. A copy of the signed consent form was given to each subject. This study was conducted according to the guidelines of the declaration of Helsinki, and received ethics approval from U.S. Investigational Review Board Inc, which is a US Department of Health and Human Services (DHHS) registered independent ethics committee (#IRB00007024).

The inclusion criteria included having good general health (based on a medical history review), having a minimum of 18 natural teeth (excluding all those with gross caries, cervical abrasion and/or enamel abrasion, orthodontic bands, crowns, veneers, third molars, and implants). Subjects were required to have, at the baseline visit, a mean full-mouth pre-brushing plaque score of ≥ 2.00 at the baseline visit (based on the Soparkar modification of the Turesky modification of the Quigley and Hein plaque index) [26–28].

Subjects were required to not undergo dental prophylaxis or any other elective, non-emergency dental procedures (other than those provided for the study) any time during the 7-day period of the investigation. Subjects also agreed to refrain from using any oral hygiene products (i.e. floss, mouthwash, etc.), other than the toothbrush and toothpaste supplied, for the 7-day period of the study. Subjects refrained from all oral hygiene procedures (including chewing gum) for 24 hrs, and from eating, drinking and smoking for 2 hours prior to the two clinical visits of the study.

The exclusion criteria included: having undergone a periodontal treatment or having received a dental prophylaxis within 30-days prior to the baseline visit; pregnancy; breastfeeding; known adverse reactions, allergies or sensitivities to oral hygiene products or to any ingredient in the test materials; any physical limitations or restrictions that might preclude normal tooth brushing; the presence of fixed or removable orthodontic appliances, bridges, peri/oral piercings, or removable partial dentures; the use of antibiotics or any medications that could influence salivary flow or oral soft tissues within 30 days prior to the study; the presence of severe periodontal disease, undergoing periodontal treatment; the presence of oral soft tissue pathology; rampant caries; and the presence of extrinsic stain or calculus deposits that could interfere with plaque assessments.

2.3. Plaque scoring

At the initial visit, each subject rinsed their mouth with an erythrosine disclosing solution. Plaque was scored on the facial and lingual tooth surfaces using the Soparkar modification of the Turesky modification of the Quigley and Hein plaque index. As mentioned earlier, all subjects were required to have a mean full-mouth pre-brushing plaque score of ≥ 2.00 at baseline to be eligible for the study. Plaque was scored in six areas per tooth (mesio-facial, midfacial, disto-facial, mesio-lingual, mid-lingual, and disto-lingual) from 0 to 5, giving a maximum possible score of 30 per tooth.

After randomization, subjects were provided with a toothbrush (Oral-B 35) [29] and then performed the first supervised brushing intervention using a known (weighed) amount of the designated dentifrice for one minute, after which a second soft and hard tissue examination was completed. The purpose of this second examination was to identify any changes in color or texture, soft tissue abrasion, and any irregularities. After a second application of disclosing solution, a post-brushing assessment of plaque levels was undertaken. The plaque score at baseline was used to calculate the extent of plaque removal from the first brushing episode.

At the end of the initial visit, subjects were reminded to brush twice daily for the next week, with the toothbrush and dentifrice that they received. They were reminded to refrain from all oral hygiene (including chewing gum) for 24 hours prior to their next visit and to refrain from eating, drinking, and smoking for 2 hrs. prior to their next visit. These instructions were repeated by telephone prior to the second visit. During the one-week period, subjects completed a toothbrushing diary.

The second visit occurred 7 days later, and followed a similar workflow, with an initial soft and hard tissue examination, followed by disclosing and scoring plaque levels. These data would then later be compared to the baseline plaque levels at the start of first visit. During the second visit, a second supervised 1-minute brushing intervention occurred using the assigned dentifrice, after which there was a further soft and hard tissue examination, followed by disclosing and scoring plaque levels. All examinations in the study were performed by the same dentist, who was blinded to the treatment allocations.

At both visits, the examining dentist assessed subjects for adverse changes, and also queried subjects regarding any adverse events.

2.4. Data analysis

A plaque index score for each subject was calculated by adding all the individual plaque scores (six per tooth), and dividing this sum by the total number of measurements (number of teeth scored multiplied by six). Only those subjects who completed the 1-week test period and had no protocol violations were included in the efficacy analysis.

The primary efficacy parameter was reduction in subject-wise mean plaque index scores from pre-brushing to post-brushing for all surfaces of the whole mouth. Within-treatment comparisons of the pre-brushing plaque scores to the post-brushing plaque scores were performed using paired t-tests. An unpaired t-test was used to compare treatment groups for the difference in the mean changes from baseline.

Additionally, secondary efficacy parameters were assessed, namely the corresponding reduction in subject-wise mean plaque scores based on evaluations made on various subsets, including proximal, facial, lingual, gingival, lingual proximal, lingual gingival, posterior proximal, posterior facial proximal, posterior lingual proximal, posterior gingival, posterior facial gingival and posterior lingual gingival.

Continuous variables for demographic data were also evaluated using an unpaired (two-sample) t-test, while data for categorical variables were assessed using Fisher's exact test. All statistical tests were two-sided and conducted at the 0.05 significance level.

3. Results

3.1. Participants

After applying the selection criteria, a total of 82 subjects (63 females, 19 males) were enrolled, and all subjects completed the study, with 41 in each group. The mean age was 48.7 (range 19-69 years). There were no statistically significant differences between the two groups for any demographic variables (age, gender, race, and ethnicity), and likewise no significant differences in baseline whole mouth plaque scores (Crest 3.13 + 0.31 (SD), Protegera 3.02 + 0.32).

3.2. Plaque removal

Data for the primary outcome of whole mouth plaque reduction are shown in Table 1. Both dentifrices gave statistically significant reductions from the baseline levels ($p < 0.001$ for both), however across all three assessments, the test dentifrice was three times more effective than the control ($p < 0.001$ for each). Thus, the null hypothesis was rejected.

Due to the accumulative effect of the test dentifrice lowering the pre-brushing plaque scores at day 7, plaque scores differed between the two treatments at the start of the Day 7 visit. An analysis of covariance model was used for the between-treatment analysis to adjust for this difference, using the Day 7 pre-brushing plaque score as the covariate, to improve the power and estimates of the treatment effects.

3.3. Site-specific effects

When secondary efficacy parameters were assessed, plaque removal for the test dentifrice was superior to the control at each of the three-time intervals ($p < 0.001$ for each) for all 14 subsets, including proximal, facial, lingual, and gingival (Table 1), as well as for lingual proximal, lingual gingival, posterior proximal, posterior facial proximal, posterior lingual proximal, posterior gingival, posterior facial gingival and posterior lingual gingival (Figure 1). Of note, the test dentifrice was superior for reducing dental plaque levels in the gingival aspect of teeth, with a 62.5% reduction on first use, a 69.6% reduction at day 7, and a 37.4% reduction from day 1 to day 7, all of which were superior to the positive control ($p < 0.001$). Data for the mean difference in plaque score between treatments (i.e. test dentifrice plaque score versus control dentifrice plaque score for the Day 1 baseline to the Day 7 post-brushing score) are summarized in Table 2.

3.4. Adverse outcomes

No safety issues or adverse responses arose from the use of either dentifrice.

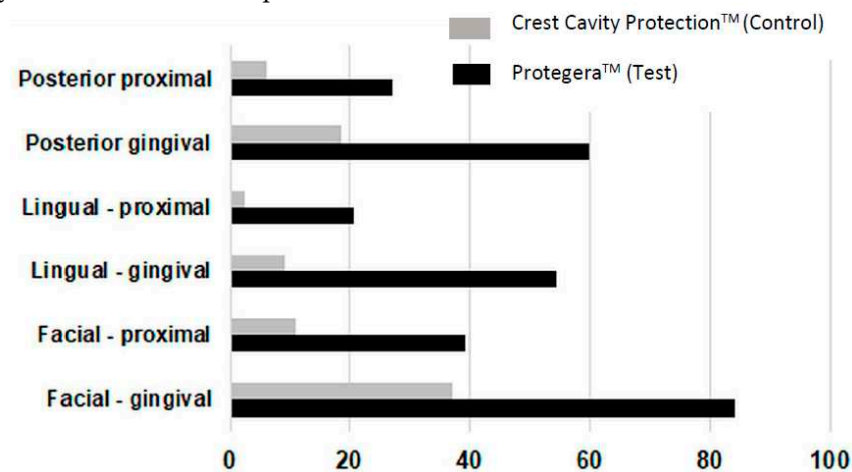


Figure 1. Plaque reduction (in percent) for one week of brushing. In all cases, the test (Protegera™) dentifrice was superior to the control (Crest Cavity Protection™) dentifrice ($p < 0.001$).

Table 1. Site-specific reductions in dental plaque levels.

Parameter	Product	Day 1	Day 7	One week use
Whole mouth	Test	38.6%	30.9%	41.6%
	Control	12.0%	9.6%	11.6%
Gingival	Test	62.5%	56.3%	68.7%
	Control	21.5%	20.3%	23.0%
Proximal	Test	28.6%	22.4%	30.1%
	Control	7.9%	5.0%	6.7%
Facial	Test	49.5%	40.5%	52.2%

Lingual	Control	19.5%	16.8%	18.6%
	Test	27.7%	22.3%	31.0%
	Control	4.4%	2.2%	4.5%

* Data show mean reductions in plaques scores in percent for the three relevant time points (N = 41 subjects per group). For all analyses, the test dentifrice was superior to the control ($p < 0.001$). Day 1 = Day 1 baseline to post-brushing; Day 7 = Day 7 baseline to post-brushing; One week = Day 1 baseline to Day 7 post-brushing.

Table 2. Plaque index difference between treatments at one week.

Subset	Difference
Whole Mouth	0.89 (0.77-1.01)
Gingival	1.19 (1.02-1.36)
Proximal	0.74 (0.62-0.87)
Facial	0.98 (0.80-1.17)
Lingual	0.80 (0.68-0.93)

¹ Data show the mean difference in plaque score between the test and control groups for the Day 1 baseline to the Day 7 post-brushing score, and the 95% confidence limits. All differences between treatments were statistically significant ($p < 0.001$).

4. Discussion

The results of this clinical study show that the test dentifrice based on MFC and tailored mechanical properties, was at least three times more effective in whole-mouth plaque reductions than the positive control dentifrice. As a result of superiority in all comparisons that were made, including reductions in whole mouth scores and all 14 site subsets, the null hypothesis of equivalence was rejected. Both dentifrices had the same levels of sodium fluoride, and both used sodium lauryl sulfate as the surfactant. It is thus unlikely that antimicrobial actions of either of these components contributed to differences in performance.

The results of the present study also show that the superiority of the cleaning action extends to sites of different types across the dentition, including areas involving the interproximal space, a region within the dentition that has limited accessibility to the bristles of a toothbrush. It is likely that the novel dentifrice can effectively transfer the forces of brushing to the dental plaque biofilm that is being removed without direct contact between the toothbrush bristles and the biofilm. This can occur since, under the forces of brushing, MFC fibers, and fibrils are forced to flow into the interproximal space where they can then dislodge plaque deposits. Such a fluid-based cleaning action is an essential extension to concepts of dental plaque removal, given the well-established principle that physical plaque removal by a toothbrush, with or without a traditional dentifrice, is due to direct contact between the bristles and the dental plaque biofilm [8,9,13].

An important consideration with a novel cleaning approach is that the method is safe for teeth and oral soft tissues [28–30]. The test toothpaste had an RDA of 88, and was designed to give a similar mouth feel and taste as regular dentifrices. It did not cause any abrasions of the oral soft or hard tissues, and no concerns or adverse effects were reported by the subjects. Given the short time scale of the present trial, longer clinical studies would be informative regarding safety aspects. Since the fiber complexes themselves have low abrasivity, issues such as abrasion of root surfaces are unlikely to arise.

A further note is that the same toothbrush (Oral-B 35) was used with both products. It is feasible that different designs of toothbrushes could be explored to maximize the ability of the MFC to remove dental plaque deposits. Moreover, changes in the formulation of the dentifrice could be explored to provide some degree of chemical plaque control such as calculus control, or others. This may give even more outstanding performance when the dentifrice is used over extended periods of time.

5. Conclusions

The present clinical study shows that a dentifrice designed to maximize the cleaning action of microfibrillated cellulose (MFC) outperforms a traditional dentifrice when used with the same toothbrush. The overall gain in performance was approximately threefold when considering whole-mouth plaque scores. No safety issues arose from the use of the novel dentifrice. These results indicate that including micro-fibrillated cellulose enhances plaque removal effectiveness, without causing adverse changes to oral soft tissues.

6. Patents

Authors M.E.L., J.M. and Y.T. are named inventors on a patent (U.S. patent 10,266,793), M.E.L. and A.P. are named inventors on patent application US 2021/0121386 A1, and M.E.L., A.P., J.M. and Y.T. are named inventors on patent application US 2021/0330557 A1 that are relevant to this clinical trial.

Author Contributions: Conceptualization: M.E.L. and A.P, Methodology: M.E.L., K.R.M. and J.L.M.; Oral Assessment: K.R.M.; analysis: L.J.W., writing the original draft: L.J.W.; revisions to the draft: M.E.L., A.P., J.M., Y.T. and J.L.M.; project administration, M.E.L. All authors have read and agreed to the published version of the manuscript.

Funding: This research was funded by Protegera Inc, and received no external funding.

Institutional Review Board Statement: This study received ethics approval from U.S. Investigational Review Board Inc, which is a US Department of Health and Human Services (DHHS) registered independent ethics committee (#IRB00007024).

Informed Consent Statement: Informed consent was obtained from all subjects involved in the study.

Data Availability Statement: Data are available upon reasonable request to the corresponding author.

Acknowledgments: We thank Anthony Winston for assistance and advice with product development. We thank Kaylie S. Wills for coordinating the clinical study. We thank Reinhard Schuller for assistance with statistical analysis.

Conflicts of Interest: Authors M.E.L., A.P., J.M. and Y.T. are employees of NovaFlux, Inc. and consultants to Protegera, Inc. The authors declare no other conflicts of interest.

References

1. Forward, G.C. Role of toothpastes in the cleaning of teeth. *Int. Dent. J.* **1991**; *41*: 164-170.
2. Forward, G.C.; James, A.H.; Bartnett, P.; Jackson, R.J. Gum health product formulations: what is in them and why? *Periodontol.* **2000** *1997*; *15*:32-39. doi: 10.1111/j.1600-0757.1997.tb00102.x.
3. Anopa, Y.; McMahon, A.D.; Conway, D.I.; Ball, G.E.; McIntosh, E.; Macpherson, L.M.D. Improving child oral health: cost analysis of a national nursery toothbrushing programme. *PLoS One* **2015**; *10*(8):e0136211. doi: 10.1371/journal.pone.0136211.
4. Soares, R.C.; da Rosa, S.V.; Moysés, S.T.; Rocha, J.S.; Bettega, P.V.C.; Werneck, R.I.; Moysés, S.J. Methods for prevention of early childhood caries: Overview of systematic reviews. *Int. J. Paediatr. Dent.* **2021**; *31*(3):394-421. doi: 10.1111/ipd.12766.
5. Gray-Burrows, K.; Day, P.; El-Yousfi, S.; Lloyd, E.; Hudson, K.; Marshman, Z. A national survey of supervised toothbrushing programmes in England. *Br. Dent. J.* **2023**. doi: 10.1038/s41415-023-6182-1.
6. Ashley, P. Toothbrushing: why, when and how? *Dent. Update.* **2001**; *28*(1):36-40. doi: 10.12968/denu.2001.28.1.36.
7. Parizotto, S.P.; Rodrigues, C.R.M.D.; Singer, J.da M; Sef, H.C. Effectiveness of low cost toothbrushes with or without dentifrice, in the removal of plaque in deciduous teeth. *Pesqui Odont. Bras.* **1993**; *17*(1): 17-23. doi: 10.1590/s1517-74912003000100004.
8. Paraskevas, S.; Timmerman, M.F.; van der Velden, U.; van der Weijden, G.A. The additional effect of dentifrices on the instant efficacy of toothbrushing. *J. Periodontol.* **2006**; *77*:1522-1527. doi: 10.1902/jop.2006.050188.
9. Paraskevas S, Rosema NA, Versteeg P, Timmerman MF, van der Velden U, van der Weijden GA. The additional effect of a dentifrice on the instant efficacy of toothbrushing: a crossover study. *J. Periodontol.* **2007**; *78*(6):1011-1016. doi: 10.1902/jop.2007.060339.

10. Creeth, J.; Gallagher, A.; Sowinski, J.; Bowman, J.; Barrett, K.; Lowe, S.; Patel, K.; Bosma, M.L. Effects of brushing time and dentifrice on dental plaque removal in vivo. *J. Dent. Hyg.* **2009**; *83*(3):111-116.
11. Jayakumar, A.; Padmini, H.; Haritha, A.; Pathakota Reddy, K. Role of dentifrice in plaque removal: A clinical trial. *Indian J. Dent. Res.* **2010**; *21*(2):213-217. doi: 10.4103/0970-9290.66629.
12. Zanatta, F.B.; Antoniazzi, R.P.; Pinto, T.M.; Rösing, C.K. Supragingival plaque removal with and without dentifrice: A randomized controlled clinical trial. *Braz. Dent. J.* **2012**; *23*(3):235-240. doi: 10.1590/s0103-64402012000300009.
13. Valkenburg, C.; Slot, D.E.; Bakker, E.W.P.; Van der Weijden, F.A. Does dentifrice use help to remove plaque? A systematic review. *J. Clin. Periodontol.* **2016**; *43*(12): 1050-1058. doi: 10.1111/jcpe.12615.
14. Lewis, R.; Dwyer-Joyce, R.S.; Pickles, M.J. Interaction between toothbrushes and toothpaste abrasive particles in simulated tooth cleaning. *Wear* **2004**; *257* (3-4):368-376. doi: 10.1016/j.wear.2004.01.015.
15. Lewis, R.; Dwyer-Joyce, R.S. Interactions between toothbrush and toothpaste particles during simulated abrasive cleaning. *Proc. Inst. Mech. Eng. Part J: J. Engin. Tribol.* **2006**; *220*(8): 755-765. doi: 10.1243/13506501JET96.
16. Bongaerts, J.H.H.; Rossetti, D.; Stokes, J.R. The lubricating properties of human whole saliva. *Tribology Lett.* **2007**; *27*:277-287. doi: 10.1007/s11249-007-9232-y.
17. Lewis, R.; Barber, S.C.; Dwyer-Joyce, R.S. Particle motion and stain removal during simulated abrasive tooth cleaning. *Wear* **2007**; *263*(1-6): 188-197. doi: 10.1016/j.wear.2006.12.023.
18. Labib, M.E.; Duhkin, S.S.; Tabani, Y.; Lai, C.Y.; Manganaro, J.L.; Materna, P.; Roberston, J.C. Compositions for cleaning and decontamination. U.S. patent 10,266,793. 2019.
19. Labib, M.E.; Duhkin, S.S.; Tabani, Y.; Lai, C.Y.; Manganaro, J.L.; Materna, P.; Roberston, J.C.; Sohn, S.Y. Cleaning composition with superabsorbent polymer. U.S. patent application publication US 20200270551. 2020.
20. Labib, M.E.; Perazzo, A. Oral cavity cleaning composition method and apparatus. U.S. patent application publication US 20210121386. 2021.
21. Labib, M.E.; Perazzo, A.; Winston, A.E.; Tabani, Y.; Manganaro, J.L.; Franz, L.L.; Sohn, S.Y.; Kuchar, C. Oral cavity cleaning composition, method, and apparatus. U.S. patent application publication US 202100330557. 2021.
22. Tabani Y, Labib ME. Method for cleaning hollow tubing and fibers. US patent No. 6,945,257. 2005.
23. Sohn, S.Y.; Alfa, M.J.; Lai, R.; Tabani, Y.; Labib, M.E. Turbulent fluid flow is a novel closed-system sample extraction method for flexible endoscope channels of various inner diameters. *J. Microbiol. Methods* **2020**; *168*:105782. doi: 10.1016/j.mimet.2019.105782.
24. Lobene, R.R.; Soparkar, P.M.; Newman, M.B. Use of dental floss – Effect on plaque and gingivitis. *Clin. Prev. Dent.* **1982**; *4*(1):5-8.
25. Turesky, S.; Gilmore, N.D.; Glickman, I. Reduced plaque formation by the chloromethyl analogue Victamine C. *J. Periodontol.* **1970**; *41*(1):41-43. doi: 10.1902/jop.1970.41.41.41.
26. Quigley, G.A.; Hein, J.W. Comparative cleansing efficiency of manual and power brushing. *J. Am. Dent. Assoc.* **1962**; *65*:26-29. doi: 10.14219/jada.archive.1962.0184.
27. Heasman, P.A.; Stacey, F.; Heasman, L.; Sellers, P.; Macgregor, I.D.M.; Kelly, P.J. A comparative study of the Philips HP 735, Braun/Oral B D7 and the Oral B 35 Advantage toothbrushes. *J. Clin. Periodontol.* **1999**; *26*: 85-90. doi: 10.1034/j.1600-051x.1999.260204.x.
28. Hunter, M.L.; Addy, M.; Pickles, M.J.; Joiner, A. The role of toothpastes and toothbrushes in the aetiology of tooth wear. *Int. Dent. J.* **2002**; *52* (Suppl. 5): 399-405. doi: 10.1111/j.1875-595X.2002.tb00729.x.
29. Lewis, R.; Dwyer-Joyce, R.S. Wear of human teeth: a tribological perspective. *Proc. Inst. Mech. Engin. Part J: J. Engin. Tribol.* **2005**; *219*(1): 1-18. doi:10.1243/1350650053295394.
30. Addy, M.; Hunter, M.L. Can tooth brushing damage your health? Effects on oral and dental tissues. *Int. Dent. J.* **2003**; *53* (Suppl. 3):177-186. doi: 10.1111/j.1875-595x.2003.tb00768.x.

Disclaimer/Publisher's Note: The statements, opinions and data contained in all publications are solely those of the individual author(s) and contributor(s) and not of MDPI and/or the editor(s). MDPI and/or the editor(s) disclaim responsibility for any injury to people or property resulting from any ideas, methods, instructions or products referred to in the content.