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The association between dental malocclusion and gastrointestinal disorders: A scoping review

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Keywords: Dental malocclusion; Gastrointestinal problems; Gastric emptying rate; Gastroesophageal reflux disease, GERD.

Abbreviations: GI: Gastrointestinal; GE: Gastric Emptying; GERD: Gastroesophageal Reflux Disease.

Introduction

Malocclusion refers to the misalignment or incorrect positioning between the teeth of the upper and lower dental arches [1]. The occurrence of dental malocclusion is high in the general population with an estimated prevalence of 56%, with no gender differences [2]. The etiology of malocclusion can be dento-

Abstract

Introduction: Gastrointestinal problems are often reported by patients with moderate to severe dental malocclusion. However, a direct link between dental malocclusion and the health of the gastrointestinal tract has not been established. The aim of this scoping review is to summarize and evaluate the existing evidence regarding the association between dental malocclusion and gastrointestinal problems.

Methods: A thorough review of the literature was conducted. Five databases were searched for peer-reviewed human studies, including information about gastrointestinal problems in patients with dental malocclusion. Article screening was performed independently by two reviewers using predefined eligibility criteria. Data extraction was performed by two independent reviewers using a customized data extraction tool. Information about article type, study design, participants' characteristics, interventions, and outcomes were extracted, summarized, and synthesized. A qualitative appraisal of the included studies was also conducted.

Results: Four prospective cohort studies met the review eligibility criteria. The results of three out of four studies indicated an association between dental malocclusion and reduced gastric emptying rate, as well as a higher incidence of gastroesophageal reflux disease. However, the amount of current evidence is currently limited, and the quality of the studies is moderate.

Conclusion: According to the results of this review, it is likely that there is a positive direct association between dental malocclusion and GI symptoms. More clinical studies are required to address this research question of high clinical significance.

alveolar, skeletal, or a combination of the two elements. In the case of dentoskeletal malocclusion differences in the size, shape and/or position of the maxilla and the mandible contribute to the malocclusion phenotype [3,4]. Dental or dentoskeletal malocclusion can manifest in various forms, such as overbite, under bite, open bite, cross bite, dental crowding, or caused by several missing teeth (Figure 1).

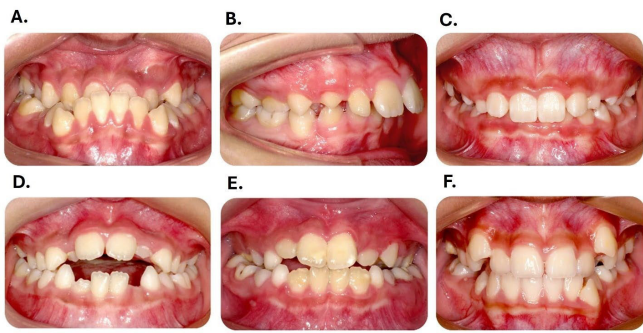


Figure 1: Figure depicting the main types of dental malocclusions. A. Underbite, B. Overbite, C. 488 Deep bite, D. Open bite, E. Posterior crossbite, F. Crowding. (Figure adapted from: Zhou et al. *Int J Oral Sci.* 2024; 16(1): 32).

Previous studies have reported that malocclusion can negatively affect oral function [5]. Specifically, moderate to severe dental malocclusion has been associated with speech impairments, obstructive sleep apnea, compromised periodontal health, increased carries risk, and increased risk for dental trauma [6-13]. Various types of malocclusions have also been directly associated with compromised masticatory efficiency [14,15]. Moreover, previous patient survey studies between orthodontic treatment and/or orthognathic surgery candidates concluded that the improvement of masticatory function is one the main motivating factors to undergo treatment, with percentages varying depending on the subjects' gender, age, and geographic location [16-19].

Mastication is the initial processing of food in the oral cavity. During oral processing, food is comminuted by a mixture of cutting and grinding by the teeth and squeezing by the tongue and cheeks [20]. This process stimulates the production of saliva, which contributes in the creation of a bolus. Saliva also contains enzymes that decompose the nutritional ingredients of the food into simpler forms that the rest of the GI tract elements can more easily absorb [21]. Thus, oral processing is the first part of the digestive tract, consisting of the initial stage of digestion [20,22]. Ineffective mastication can impact the release of digestive enzymes in the mouth, potentially increasing the subsequent digestion me and compromising nutrient absorption in the stomach and intestines, whereas a direct functional relationship between mastication and gastric function has also been reported [23-25]. Additionally, adults with multiple missing teeth or edentulous patients reportedly have more digestive complaints and GI disorders, which have been attributed to compromised masticatory function [26,27].

Therefore, there is a possible association between malocclusion and digestive problems, but it has not yet been established. The aim of this scoping review is to summarize and qualitatively evaluate the existing evidence regarding a possible direct link between malocclusion of variable etiology and GI symptoms.

Methods

Protocol

This scoping review was reported under the Preferred Reporting Items for Systematic Reviews and Meta-analyses extension for scoping reviews checklist (PRISMA-ScR) [28].

Eligibility criteria

Eligible articles included human studies, with participants of any age and gender. Only studies including subjects with a reported history of dental or dentoskeletal malocclusion and gastrointestinal problems were included. Investigations of patients with syndromes or congenital gastrointestinal disorders were excluded. No studies were excluded based on geographic location, racial, or gender-based interests, or details about the specific study setting. This review considered both experimental and quasi-experimental study designs. Analytical observational studies including prospective and retrospective cohort studies, case-control studies, and analytical cross-sectional studies, as well as descriptive cross-sectional studies, case series and case reports. Only primary research studies were included. Text and opinion papers, narrative reviews, conference abstracts were not included in this review, due to the lack of peer review processing. Table 1 provides a more detailed outline of the inclusion/exclusion criteria for article selection.

Information sources and search

To ensure a comprehensive search, the following databases were queried from inception to August 2023: MEDLINE (PubMed), EMBASE (Elsevier), Web of Science Core Collection (Clarivate), Scopus (Elsevier), and Cochrane Library (Wiley) with the use of the Title/Abstract or Topic, English Language and Humans filters, and the Cochrane human studies hedge applied to MEDLINE, Scopus and EMBASE. The complete search strategy for all databases and search engines is provided in Supplementary Table A. Upon selection of the eligible studies, all authors (AD,RK,KA) independently reviewed the references for each article included in the review as well as the references of relevant systematic reviews and/or meta-analyses to identify other potentially relevant studies for inclusion. The most recent search was executed on August 18, 2023.

An example search strategy that was used is the PubMed full electronic strategy:

((gastrointestinal [Title/Abstract]) or (gastro enteric [Title/Abstract])) or

(Reflux [Title/Abstract]) or (digestion [Title/Abstract]) or (indigestion [Title/Abstract]) or

(Dyspepsia [Title/Abstract]) or (heartburn [Title/Abstract]) or (GERD [Title/Abstract]) or

(Constipation [Title/Abstract]) or (irritable bowel syndrome [Title/Abstract]) or

(IBS [Title/Abstract]) or (Nausea [Title/Abstract]) or (gas [Title/Abstract]) or

(Bloating [Title/Abstract]) or (diarrhea [Title/Abstract])) AND ((malocclusion [Title/Abstract]) or

(Dental occlusion [Title/Abstract]) or (crowded teeth [Title/Abstract]) or (dental Crowding [Title/Abstract]) or (retrognathism [Title/Abstract]) or (prognathism [Title/Abstract]) or

(Under bite [Title/Abstract]) or (overbite [Title/Abstract]))

Selection of sources of evidence and data charting process

All authors (AD,RK,KA) screened the titles and abstracts of the identified articles based on the predefined inclusion and exclusion criteria. If eligibility could not be decided by title or abstract, the full text of the article was retrieved to determine eligibility. The individual eligibility decisions from the initial screening process of the articles were compared, and disagreements were resolved through discussion.

Data was extracted from the studies included in the scoping review by two independent reviewers (AD,EK) using a data extraction tool developed for this study, including information about the article, study design, participant characteristics, and pertinent results to the research question. The data extraction entries were reviewed by the third author (KA).

Data items and synthesis of the results

The information that was extracted from the articles included was summarized by two reviewers (AD,RK) and is presented within the Results section of the manuscript as well as in a tabular form. A narrative summary accompanied the tabulated results and described how the results were related to the review objective and questions.

Qualitative appraisal of the included studies

The qualitative assessment of the included studies was conducted with the use of the tool developed by the JBI for quasi-experimental studies [29].

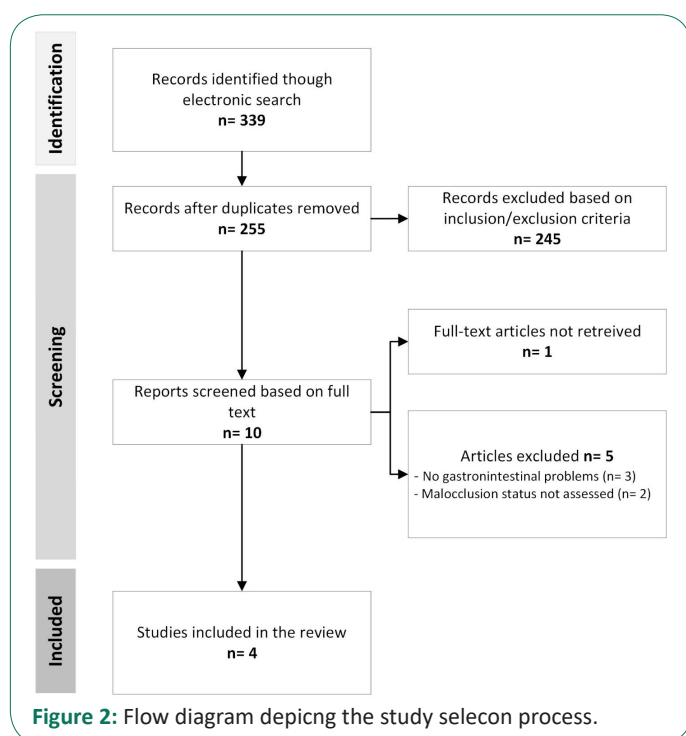


Figure 2: Flow diagram depicting the study selection process.

Results

Selection, characteristics of sources of evidence, and summary results

Of the 399 records identified by the initial search and the removal of 85 duplicates, 255 studies remained for screening. After reviewing their titles and abstracts, 10 studies qualified for retrieval of their full texts and further assessment. 1 full text could not be retrieved, and 5 studies were excluded based on the additional information acquired by the full text. 4 studies were included in the review. No additional studies were de-

tected after a manual search of the references of the studies included as well as reviews with relevant subjects. The study selection process is described with a flow diagram in Figure 2.

Synthesis of the results

Publication dates ranged from 2018 to 2013 and were all conducted in Japan. The parameters examined as well as the assessment modalities varied between studies. Two studies included only female subjects [30,31], whereas one study included only male subjects [32], and the last study a mixed group of subjects [33]. The main characteristics of the studies included in the review are presented in Table 2. Table 3 provides a summary of the participant characteristics and pertinent results for each of the included studies.

The first study by Hattori et al. [32] was a prospective trial with a crossover design. The study cohort included n=13 male participants, with a median age of 20 years and median Body Mass Index (BMI) of 20.4 kg/m². They all had natural dentitions and were in good health. Exclusion criteria included a history of previous abdominal surgery and the use of medication that might affect GI motility. The aim of this study was to investigate the effect of the absence of molar occlusion on the Gastric Emptying (GE) rate. A splint-like mandibular intraoral appliance was used to simulate shortened dental arches. The reason for the use of a crossover design is that there is a wide interindividual variability in the GE rate [34-36]. In this study, each subject functioned as their own control, by measuring their GE with and without the appliance. GE rate was evaluated using a 13C-octanoic acid breath test [36,37]. The experimental protocol involved two sessions on separate days, each including ingestion of a test meal followed by four hours of gastric emptying measurement and masticatory performance assessment. Regarding the GE parameters, no discernible difference was detected between the two occlusal conditions.

The second study by Koike et al. [30] was a prospective cohort study. N=11 female adults with malocclusion seeking orthodontic treatment were randomly selected, alongside n=11 female controls with normal occlusion. Patients had various types of malocclusions, while controls had normal occlusion. The mean age for patients was 25.5±4.8 years, with a mean BMI of 19.6±2.4 kg/m², and for controls had a mean age of 26.5±1.0 years, and a mean BMI of 19.6±1.8 kg/m², with no significant age or BMI differences. Exclusions included orthodontic treatment history, cleft lip or palate and craniofacial syndromes abdominal surgery, medication that could affect the motility of the GI tract, pregnancy, acute illness, alcohol use, heavy smoking, and denture wear. Gastric emptying rate was assessed with a [13C]-labeled acetate breath test and masticatory function with color-changeable gum. Questionnaires evaluated gastroesophageal reflux symptoms and food intake difficulties. According to the results of this study, there was a strong correlation between dental malocclusion and delayed gastric emptying. This delay was detected through the breath test as changes in the amount and rate of [13CO₂] recovery. The hypothesis is that the suppression of the GE of the [13C]-labeled food led to a decrease in the recovery of [13CO₂]. Regarding the results of the questionnaires, the conclusion was that subjects with malocclusion have a higher incidence of digestive complaints and gastrointestinal disorders, but there was a statistically significant difference between the experimental groups.

The third study by Suzuki et al. [31] was a pilot prospective cohort study. The study cohort included n=7 female subjects

seeking orthodontic treatment for malocclusion, and seven female subjects with no malocclusion that served as controls. All subjects were randomly selected. Only healthy females, aged 18-39, with normal BMI, no craniofacial syndromes, no history of craniofacial surgery, no use of medications, diseases, non-smoking, alcohol use, or pregnancy. Gastric emptying and masticatory function were assessed using [13C]-labeled acetate breath test and color-changeable chewing gum, respectively. The Frequency Scale for the Symptoms of GERD (FSSG) questionnaire was used for the evaluation of dyspeptic and dysmotility symptoms [38]. Outcome variables were measured pre-and post- orthodontic treatment for the malocclusion group and at two time points for the controls. Gastric emptying was assessed by measuring the maximum [13CO₂] excretion me. Breath samples were collected using special sampling bags before ingestion of a test meal and at several time points after ingestion. Based on the results of this study, there was a significant difference in the maximum extraction me between malocclusion subjects and controls, that was eliminated after orthodontic treatment. Hence, the correction of malocclusion improved gastric emptying in these patients.

Finally, the last study by Togawa et al. [33] was also a prospective cohort study. Their cohort included n=19 (7 males and 12 females) adult subjects with severe skeletal Class III malocclusion with mandibular prognathism. N=20 control subjects with normal occlusion were also enrolled.

The subjects had to be over 18 years of age, with no underlying diseases, and no current medications. Patients with cleft lip or palate and craniofacial syndromes were excluded. Gastroesophageal Reflux Disease (GERD) symptoms were evaluated using the Carlsson-Dent self-administered Questionnaire (QUEST) and the Frequency Scale For GERD Symptoms (FSSG) [38,39]. The results of this study indicate that patients with severe skeletal Class III malocclusion have a higher incidence of GERD symptoms than subjects with normal occlusion.

Qualitative appraisal of the included studies

The results of the qualitative appraisal of the included studies are included in Table 4. An overall moderate level of evidence was detected.

Table 1: Eligibility criteria used for the selection of the studies.

Category	Inclusion criteria	Exclusion criteria
Participant characteristics	Studies on human patients with dental or dentoskeletal malocclusion and gastrointestinal problems	<ul style="list-style-type: none"> - Animal studies - Syndromic patients - Subjects with congenital gastrointestinal problems - Edentulous patients
Outcome	Studies not providing information on the presence of dental or dentoskeletal malocclusion and/or gastrointestinal problems	<ul style="list-style-type: none"> - Studies not providing information on the presence of dental or dentoskeletal malocclusion and/or gastrointestinal problems - Ongoing studies
Study design	<ul style="list-style-type: none"> - Randomized clinical trials - Prospective clinical trials - Retrospective clinical trials - Case-control observational studies - Cross-sectional surveys - Case series - Case reports 	<ul style="list-style-type: none"> - Narrative reviews - Unsupported opinion of expert - Editor's choices - Replies to the author/editor - Books' abstracts - Conferences' abstracts - In vitro studies - In silico studies - Meta-analyses* - Systematic reviews*

*After checking the reference lists for relevant studies.

Table 2: Characteristics of the included studies.

Author, Year	Title	Country	Study Type	Study Design	Study Aim/Objective
Hatori et al. 2008 [32]	Gastric emptying rate in subjects with experimentally shortened dental arches: a pilot study	Japan	Prospective	Crossover Clinical Trial	To investigate the effect of the experimental loss of molar occlusion on gastric emptying rate.
Koike et al. 2013 [30]	Gastric emptying rate in subjects with malocclusion examined by breath test	Japan	Prospective	Controlled Trial	To investigate the relationship between malocclusion and digestion, specifically gastric emptying rates, masticatory function, and gastrointestinal symptoms.
Suzuki et al. 2016 [31]	Gastric emptying rate before and after orthodontic treatment examined with the [13C] breath test: A pilot study	Japan	Prospective	Controlled Trial	To evaluate the change in gastrointestinal function, using the [13C] breath test with a liquid test meal, before and after orthodontic treatment, in patients with malocclusion and investigate the relationship between occlusion, mastication, and gastrointestinal function.
Togawa et al. 2008 [33]	Gastroesophageal reflux symptoms in adults with skeletal Class III malocclusion examined by questionnaires	Japan	Prospective	Cross-sectional	To examine the symptoms of Gastroesophageal Reflux Disease (GERD), occlusal contact area, maximal voluntary bite force, and salivary flow rate in patients with skeletal Class III malocclusion.

Table 3: Summary of the results of the included studies.

Author, Year	Population studied	Sample size (females, males)	Mean age and/or range (years) or SD	Pertinent results summary
Hatori et al. 2008 [32]	Healthy dentate males with artificially shortened dental arches	0:13	20 (20-21)	Experimental loss of molar occlusion did not lead to significant changes in GE parameters
Koike et al. 2013 [30]	Female adult patients, with and without malocclusion, who sought orthodontic treatment	Malocclusion group: 11:0; Control group: 11:0	Malocclusion group: 25.5±4.8 (19.3-35.9); Control group: 26.5+/-1.0 (25.0-27.8)	Patients with malocclusion had a trend toward higher incidence of delayed GE parameters compared to controls
Suzuki et al. 2016 [31]	Female adult patients, with and without malocclusion, who sought orthodontic treatment	Malocclusion group: 7:0; Control group: 7:0	Malocclusion n group: 26.7±5.5 (18-39); Control group: 25.4±1.0	Patients with malocclusion had delayed GE rates compared to controls. Orthodontic treatment resulted in an improvement in GE rates.
Togawa et al. 2008 [33]	Two groups: adults with severe skeletal Class III malocclusion and a control group of individuals with normal occlusion	Class III group: 12:7; Control group: 12:8	Class III group: 24.4 (18-37) Control group: 25.2 (22-31)	Patients with skeletal Class III malocclusion had significantly higher scores on the GERD symptom questionnaires compared to the control group.

Table 4: Results of the qualitative appraisal of the studies based on the JBI checklist for quasi-experimental studies.

Study	1. Is it clear in the study what is the 'cause' and what is the 'effect' (i.e. there is no confusion about which variable comes first)?	2. Were the participants included in any comparisons similar?	3. Were the participants included in any comparisons receiving similar treatment/care, other than the exposure or intervention of interest?	4. Was there a control group?	5. Were there multiple measurements of the outcome both pre and post the intervention/exposure?
Hattori et al. 2008 [30]	Yes	Yes	Yes	No (crossover)	Yes
Koike et al. 2013 [31]	Yes	Yes	Yes	Yes	Yes
Suzuki et al. 2018 [32]	Yes	Yes	Yes	Yes	Yes
Togawa et al. 2008 [33]	Yes	Yes	Yes	Yes	No

Study	6. Was follow-up complete and if not, were differences between groups in terms of their follow-up adequately described and analyzed?	7. Were the outcomes of participants included in any comparisons measured in the same way?	8. Were outcomes measured in a reliable way?	9. Was appropriate statistical analysis used?
Hattori et al. 2008 [30]	No	Yes	Yes	Yes
Koike et al. 2013 [31]	No	Yes	Yes	Yes
Suzuki et al. 2018 [32]	No	Yes	Yes	Yes
Togawa et al. 2008 [33]	No	Yes	No	Yes

Limitations

One of the main limitations of this review is the small number of included studies, reporting a direct association between malocclusion and GI disorders. These were the results of a thorough search in six large databases. In addition, a manual search of the references of the included studies as well as of previous review studies on similar topics was also conducted. Therefore, every effort has been made to locate relevant studies. Nevertheless, one of the main objectives of a scoping review is to elucidate the need for additional studies on a specific subject, which has been achieved with this study.

Discussion

Scoping reviews are exploratory studies that aim to systematically map the literature available on a topic and identify gaps in the research, that will indicate future lines of inquiry [40]. According to the results of this review, only four studies were detected delineating the direct link between malocclusion and GI tract symptoms. The results of three out of the four studies indicated a direct association between dental malocclusion and GI symptoms.

Specifically, GE rates were slower than age and sex-matched controls in patients with malocclusion. Both studies reporting these results [30,31] were conducted in young adult females, because a significant difference has been previously document-

ed in GE between males and females [41]. Moreover, based on the results of a clinical trial by Klingensmith et al. [42] the range of GE was found to be narrower in females, and, therefore, comparisons can be more consistent. However, the sample size in both studies was small. Moreover, the participants had different types of malocclusions, and the results could not be stratified accordingly due to the small cohort sizes.

In the third study [33], a higher incidence of GERD symptoms was detected in a group of female severe Class III malocclusion patients. The main limitation of this study, apart from the small sample size, was that the diagnosis of GERD was based solely on self-reported questionnaires. The validity of the first Questionnaire (QUEST) has been previously questioned in terms of diagnostic value in several studies [43-47], although a few studies support its diagnostic value [48]. Respectively, the FSSG questionnaire is reportedly more reliable in the diagnosis of GERD [47,49]. However, there are not enough validation studies on this questionnaire. Therefore, the conclusions of this study are relatively subjective.

Finally, in the study by Hatori et al. [32], no correlation was detected between the presence of an artificially shortened arch and GE. In this case, the study design did not allow for the investigation of the chronic effect of malocclusion in the GI system, since all the participants were healthy individuals. In addition, the masticatory function is not necessarily compromised in the case of shortened dental arches. Specifically, results from previous studies support the functional efficiency of the shortened dental arch, indicating that arches with intact premolar regions provide satisfactory chewing ability [50-53]. Regardless, based on the results of this study, masticatory efficiency was negatively affected by the shortening of the arches. However, the interference of the splint could have affected the masticatory performance of the subjects and cannot be directly compared with naturally shortened dental arches.

Based on the results of the above studies, there is a need for additional studies that will contribute in the elucidation of the association of dental malocclusion and GI problems. Specific types of malocclusions could be more likely to cause problems along the GI tract than others.

Therefore, stratification of future cohorts based on malocclusion type should be considered. Moreover, the status of pre- and post-treatment GI symptoms in orthodontic and/or orthognathic surgery treatment studies should also be more consistently monitored and reported. Additional scientific evidence on this matter is required for more informed treatment consultations and improvement in the quality of clinical care of patients with dental malocclusion.

Conclusion

The digestion of food is essential to human nutrition, health, and overall quality of life [54]. Despite the clinical significance of the possible correlation between dental malocclusion and GI problems, the level of evidence on this matter is currently low. Additional clinical studies, with larger cohort sizes and strategic study designs, are required to provide more knowledge on this subject.

Declarations

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