

## **Effects of carbon-dioxide lasers on preventing caries: A literature review**

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### *Abstract*

**Purpose:** This study aimed to review the effects of carbon dioxide lasers ( $\lambda = 10,600$  nm) on preventing dental caries.

**Methods:** A systematic search was performed of articles published in English through 2018 and archived in the PubMed, Scopus, and Web of Science databases. The keywords used to identify the relevant publications were “*CO<sub>2</sub> laser*” OR “*carbon dioxide laser*” AND “*dental caries*” OR “*tooth remineralization.*” The titles and abstracts of the initially identified publications were screened. Duplicate records, case reports, reviews, and irrelevant studies were removed. Full texts that focus on the effects of carbon dioxide lasers on preventing dental caries were retrieved from publications.

**Results:** The search identified 543 potentially relevant publications. A total of 285 duplicate records were removed. Twenty-two articles investigating the effect of carbon dioxide lasers on enamel and dentin were included. Results of the studies showed that carbon dioxide lasers could cause chemical and morphological changes to mineralized tissues, including reduction of carbonate content and recrystallization of mineral structure. All studies showed carbon dioxide lasers increased the micro-hardness and decreased the mineral loss of enamel (nine studies) and dentin (three studies). Two out of eight studies showed increased resistance to demineralization of enamel after carbon dioxide laser and fluoride treatment. All studies revealed carbon dioxide laser and fluoride treatment reduced mineral loss of dentin against acid challenge.

**Conclusions:** Carbon dioxide (10,600 nm) lasers can prevent demineralization of enamel and dentin. Furthermore, they have a synergistic effect with fluoride in preventing dentin caries.

**Keywords :** carbon dioxide laser, dental caries, remineralization, caries prevention, review

## **Introduction**

Although dentistry has benefitted from technological advancements in recent years, dental caries remain a major oral health problem in most industrialized and non-industrialized countries affecting more than half of schoolchildren and a large majority of adults [1]. Dental caries are a localized chemical dissolution of dental hard tissues caused by the action of acidic by-products of the metabolic processes of cariogenic biofilm [2]. Conventional noninvasive strategies for caries prevention include oral health education, reduction of sugar consumption, use of fluoride, and application of pit and fissure sealants.

Fluoride principally works by hindering the process of demineralization of enamel and dentin and promoting remineralization of tooth surfaces with early signs of mineral loss. Pit and fissure sealants generate a physical barrier to prevent the access of dental plaque and its acid products from damaging the enamel surface. However, dental sealants are only effective on the pit and fissures and not the smooth surfaces of teeth. New strategies, such as casein phosphopeptide amorphous calcium phosphate [3] and micro- or nano-hydroxyapatite compounds, have been proposed to control the balance between demineralization and remineralization. It is essential to develop new caries-preventive methods to control the disease. A novel, noninvasive approach is the use of laser irradiation on enamel or dentin in preventing caries development.

A number of studies reported the potential of laser irradiation on tooth roots or enamel in inhibiting formation of caries lesions [4-9]. Several types of lasers were studied for caries prevention. The wavelengths of neodymium-doped yttrium aluminum garnet (Nd:YAG) lasers ( $\lambda = 1,064$  nm) and argon lasers ( $\lambda = 488-514$  nm) make them difficult to be absorbed by

enamel. On the other hand, carbon dioxide (CO<sub>2</sub>) lasers ( $\lambda = 9,000-11,000$  nm) are highly absorbed by dental hard tissues and thus have good potential for use in caries prevention [7,9].

The first CO<sub>2</sub> laser adopted a mixture of nitrogen, helium and CO<sub>2</sub>, and CO<sub>2</sub> acted as the active laser medium [10]. Subsequent CO<sub>2</sub> lasers produced various emission laser lines with wavelengths ranging from 9,000 nm to 11,000 nm. The most common laser lines of CO<sub>2</sub> lasers are centered at 9,300 nm, 9,600 nm, 10,300 nm, and 10,600 nm, respectively. CO<sub>2</sub> laser wavelengths have a higher absorption coefficient to hydroxyapatite than water.

A conventional CO<sub>2</sub> laser emitting light at 10,600 nm is well absorbed by minerals, while 9,600 nm has the best absorption coefficient to hydroxyapatite followed by 9,300 nm [1]. However, the 10,600 nm CO<sub>2</sub> laser has been commonly used in medicine and dentistry, and most of the commercially available CO<sub>2</sub> lasers operate only at this wavelength.

The effect of CO<sub>2</sub> lasers in caries prevention has been studied since the 1960s [1], when CO<sub>2</sub> lasers (10,600 nm) were discovered to significantly inhibit enamel caries progression [7]. Significantly less demineralization was also found in CO<sub>2</sub> laser-treated ( $\lambda = 10,600$  nm) dentin than nonlased dentin [9]. Furthermore, promising effects of combined CO<sub>2</sub> laser irradiation and fluoride treatment in preventing enamel and dentine caries were reported [11-13]. However, the mechanism of caries prevention of CO<sub>2</sub> lasers remains to be elucidated.

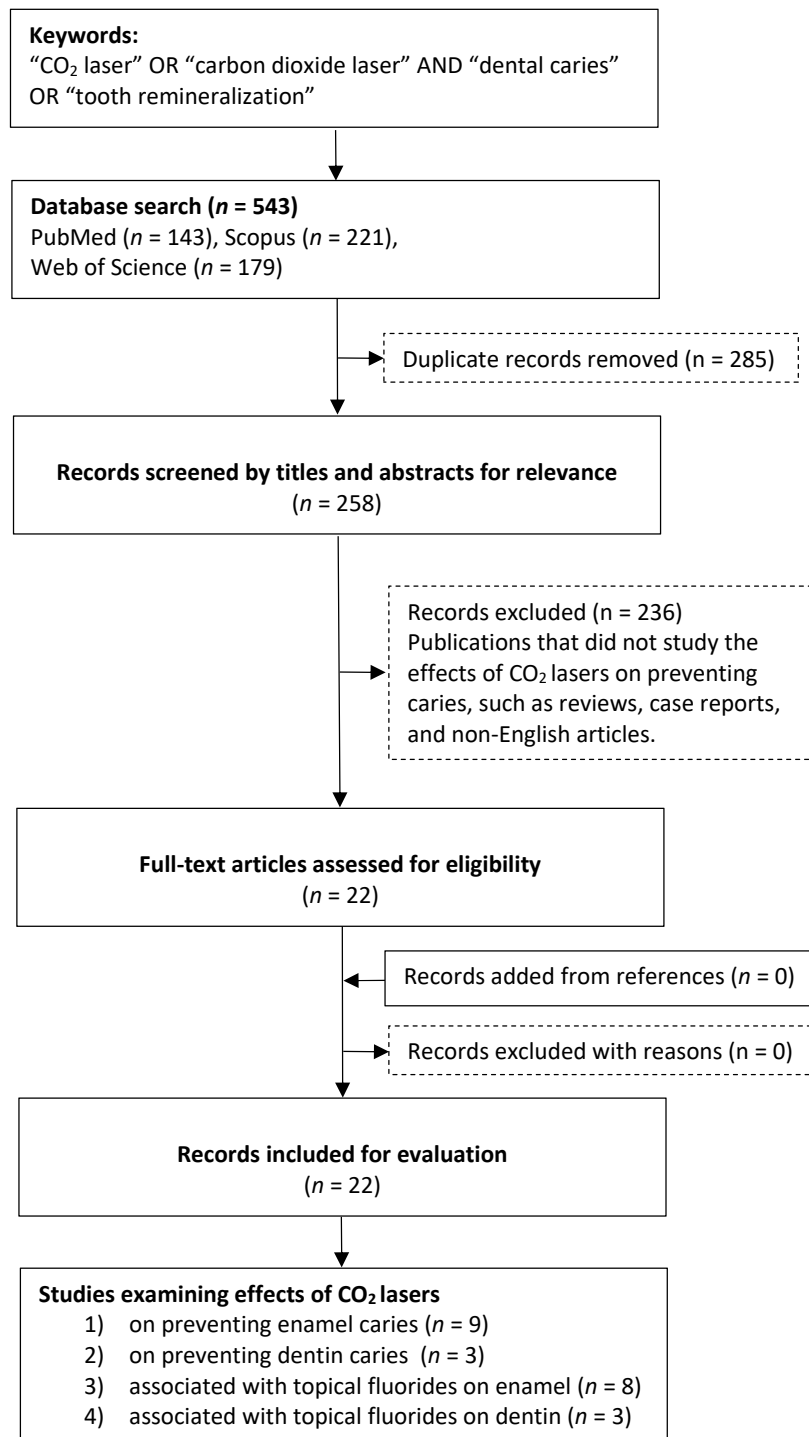
A literature search using the databases PubMed, Scopus and Web of Science revealed no comprehensive review to evaluate studies investigating the effects of actions of CO<sub>2</sub> lasers in

caries prevention. Therefore, the objective of this paper was to systematically review the evidence regarding the effects of CO<sub>2</sub> lasers ( $\lambda = 10,600$  nm) on preventing dental caries.

### **3.2 Materials and methods**

Two investigators (KL and ISZ) performed a systematic search of articles archived in three databases: PubMed, Scopus, and Web of Science. The following keywords were used to identify relevant articles: “CO<sub>2</sub> laser” OR “carbon dioxide laser” AND “dental caries” OR “tooth remineralization.” There was no publication-year limit, and the last search was made on January 31, 2019. Studies published in English through 2018 and archived in the PubMed, Scopus, and Web of Science databases were chosen. A list of potentially eligible articles was compiled of publications featuring the keywords (Figure .1).

**Figure 3.1. Flowchart of the literature search**



### *Study selection and data extraction*

Records identified from database searches were checked for duplication. The titles and abstracts from the potentially eligible list were screened after removing duplicate publications. Articles that did not focus on the effects of CO<sub>2</sub> lasers on preventing dental caries were excluded after screening of titles and abstracts. The remaining articles were retrieved with full texts. The references lists of all the included articles were screened to identify all possibly eligible studies. The inclusion criterion for selecting studies for this review was studies that investigated the effects of CO<sub>2</sub> lasers on preventing enamel and dentin caries, including their combined effect with topical fluorides on caries progression. For the included papers, the following information was recorded: publication details (authors and years), methods, outcome assessments (various criteria for studying the prevention of caries: reduction of carbonate content, lesion depth, microhardness, mineral loss, surface morphology, and bacterial counts), and the main findings. Any disagreements on study inclusion or data extraction were discussed by the two authors with a third author (OYY) until consensus was reached.

### **Results**

The initial literature search identified 543 potentially relevant articles (143 articles in PubMed, 221 in Scopus, and 179 in Web of Science). A total of 285 duplicate records were removed. After screening the titles and abstracts, 258 articles that were classified as literature reviews, case reports, non-English articles, or irrelevant studies were excluded. No additional relevant publications were found from the references of the selected papers. Finally, 22 articles were found to meet the inclusion criteria and were included in this review. Among these articles, nine studies examined the action of CO<sub>2</sub> lasers on preventing enamel caries (Table 1).

**Table 1. Studies of effects of CO<sub>2</sub> (10,600 nm) lasers on enamel**

Authors, Year	Methods	Main findings
Vieira et al., 2015 [14]	Human enamel was irradiated with CO <sub>2</sub> laser before FTIRS and EDX. They were subjected to pH cycling before MHT and SEM.	Irradiated enamel increased microhardness and reduced carbonate content more than negative control. Melting and recrystallization of enamel occurred after irradiation.
Correa-Afonso et al., 2012 [15]	Human enamel irradiated with CO <sub>2</sub> laser was subjected to pH cycling before MHT, PLM, and SEM.	Enamel increased microhardness and reduced demineralization after irradiation. No fusion, melting, or exposure of enamel prisms were found on the irradiated enamel.
Souza-Gabriel et al., 2010 [18]	Human and bovine enamel were irradiated with CO <sub>2</sub> laser before SEM. They were subjected to pH cycling before MHT.	Enamel showed melting and recrystallization after irradiation. It was more resistant to demineralization than the negative control.
Esteves-Oliveira et al., 2009 [17]	Bovine enamel was irradiated with CO <sub>2</sub> laser before SEM. It was subjected to pH cycling before PLM.	Irradiated enamel showed no ablation, melting, or cracks. It was more resistant to demineralization than the negative control.
Steiner-Oliveira et al., 2006 [20]	Human enamel was irradiated with CO <sub>2</sub> laser before FTRS and SEM. It was subjected to pH cycling before MHT.	Irradiated enamel had less carbonate content and mineral loss than the negative control. Enamel showed melting and fusion after irradiation.
Klein et al., 2005 [19]	Human enamel was irradiated with CO <sub>2</sub> laser before SEM. It was subjected to pH cycling before MHT.	Enamel surfaces showed melting and fusion after laser irradiation. They had less demineralization than the negative control.
Hsu et al., 2000 [16]	Human sound enamel with organic matrix removal or not was treated with CO <sub>2</sub> laser and subjected to pH cycling before PLM, microradiography, and SEM.	Irradiated enamel had no surface melting or crater after irradiation. It had less mineral loss and lesion depth than other groups. An interaction effect was found between laser irradiation and enamel organic matrix content.
Featherstone et al., 1998 [7]	Human enamel was irradiated with CO <sub>2</sub> laser and then subjected to pH cycling before MHT.	Irradiated enamel showed less mineral loss than the negative control.
Kantorowitz et al., 1998 [8]	Human enamel irradiated with CO <sub>2</sub> laser was examined by SEM and subjected to pH cycling before MHT.	Irradiated enamel surfaces showed little or no morphological changes. They had less mineral loss than the negative control.

EDX, energy-dispersive X-ray; FTRS, Fourier transformed Raman spectroscopy; MHT, microhardness testing; PLM, polarized-light microscopy; SEM, scanning electron microscopy



Three studies investigated the effect of CO<sub>2</sub> lasers on preventing dentin caries (Table 2), eight studies investigated the effect of CO<sub>2</sub> lasers combined with topical fluorides on enamel (Table 3), and three studies examined the effect of CO<sub>2</sub> lasers combined with topical fluorides on dentin (Table 4).

**Table 2. Summary of studies of effects of CO<sub>2</sub> (10,600 nm) lasers on dentin**

Authors, Year	Methods	Main findings
de Melo et al., 2014 [21]	Human root surfaces irradiated with CO <sub>2</sub> laser were subjected to pH cycling before MHT.	Irradiated root surfaces were more resistant to demineralization than the negative control.
de Souza-Zaroni et al., 2010 [22]	Human root surfaces were irradiated with CO <sub>2</sub> laser before SEM. They were subjected to pH cycling before MHT.	Irradiated root surfaces showed melting and resolidification. They showed less mineral loss than negative control.
Nammour et al., 1992 [9]	Human root surfaces irradiated with CO <sub>2</sub> laser were subjected to cariogenic challenge before SEM, microdensitometry measurements, and microradiography.	Root surfaces appeared smooth after irradiation. Root surfaces showed melting with no dentinal openings, and they showed more resistant demineralization than the negative control.

MHT, microhardness testing; SEM, scanning electron microscopy

**Table 3. Summary of studies of CO<sub>2</sub> (10,600 nm) lasers with fluoride on preventing enamel demineralization**

Authors, Year	Methods	Main findings
Mahmoudzadeh et al, 2018 [29]	Human enamel was irradiated with CO <sub>2</sub> laser associated or not with NaF in different order before SEM. It was subjected to cariogenic biofilm challenge before MHT.	No difference in microhardness was found among enamel treated with laser plus NaF, laser alone, or NaF alone. Irradiated enamel showed melting, cracks, and craters with discontinuities.
Mirhashemi et al., 2016 [24]	Bovine enamel irradiated with CO <sub>2</sub> laser was treated with TiF <sub>4</sub> and subjected to pH cycling before AAS and MHT.	Irradiated enamel treated with TiF <sub>4</sub> was more resistant to demineralization and had an increased microhardness compared to those without TiF <sub>4</sub> treatment.
Esteves-Oliveira et al., 2017 [26]	Bovine enamel was irradiated with CO <sub>2</sub> laser associated or not with NaF. It was subjected to cariogenic biofilm challenge before TMR, CFU, and SEM.	No difference in mineral loss was found between irradiated enamel treated with or without fluoride. The bacterial numbers on enamel were not affected by laser or fluoride.
Seino et al., 2015 [27]	Human enamel was irradiated with CO <sub>2</sub> laser associated or not with APF before SEM. It was subjected to pH cycling before QLF.	Enamel surface showed no melting after laser irradiation. Irradiated enamel treated with fluoride showed similar resistance to demineralization compared to those treated with laser only.
Tagliaferro et al., 2007 [28]	Human enamel was irradiated with CO <sub>2</sub> laser associated or not with APF. It was subjected to pH cycling before MHT.	Irradiated enamel treated with fluoride showed similar resistance to demineralization compared to those treated with laser alone.
Tepper et al., 2004 [25]	Human enamel was irradiated with CO <sub>2</sub> laser associated or not with amine fluoride before SEM. It was subjected to acid challenge before AAS.	Irradiated enamel had higher fluoride uptake. Irradiated enamel treated with fluoride had less morphological change and similar demineralization compared to those treated with laser alone.
Hsu et al., 2001 [12]	Human enamel treated with sodium fluoride was irradiated with CO <sub>2</sub> laser before pH cycling. It was studied by microradiography and SEM.	Irradiated enamel treated with NaF was more resistant to demineralization than the control group. Irradiated enamel showed no evidence of surface melting.
Hsu et al., 1998 [23]	Enamel irradiated with CO <sub>2</sub> laser associated or not with fluoride was exposed to a demineralizing solution before light microscopy and microradiography.	The irradiated and nonirradiated enamel was more resistant to demineralization in the presence of fluoride. There was a synergism between laser irradiation and fluoride in reduction of enamel solubility.

AAS, atomic absorption spectrometry; APF, acidulated phosphate fluoride; CFU, colony-forming unit; NaF, sodium fluoride; QLF, quantitative light-induced fluorescence; TiF<sub>4</sub>, titanium tetrafluoride; TMR, transverse microradiography; MHT, microhardness testing; SEM, scanning electron microscopy

**Table 4 Summary of studies of CO<sub>2</sub> (10,600 nm) lasers with fluoride on preventing dentin demineralization**

Authors, Year	Methods	Main findings
Esteves-Oliveira et al., 2017 [26]	Bovine dentin irradiated with CO <sub>2</sub> laser was treated with NaF and subjected to cariogenic biofilm challenge before TMR, CFU, and SEM.	Irradiated dentin treated with NaF was more resistant to demineralization and yielded lower CFUs than those without NaF treatment.
Esteves-Oliveira et al., 2011 [13]	Bovine dentin was irradiated with CO <sub>2</sub> laser associated or not with APF and then subjected to pH cycling before ICP-OES.	Irradiated dentin treated with fluoride had lower calcium and phosphorus concentration in the demineralization solution than those without fluoride treatment.
Gao et al., 2006 [11]	Human root dentin was irradiated with CO <sub>2</sub> laser associated or not with NaF. They were subjected to pH cycling before PLM. Fluoride uptake was characterized by the ToF-SIMS analysis.	Irradiated dentin treated with NaF was more resistant to demineralization than the control group. Irradiated dentin had a significantly higher fluoride uptake than nonirradiated groups.

APF, acidulated phosphate fluoride; CFU, colony-forming unit; ICP-OES, inductively coupled plasma optical emission spectrometer; NaF, sodium fluoride; PLM, polarized-light microscopy; TMR, transverse microradiography; SEM, scanning electron microscopy; ToF-SIMS, time of flight-secondary ion mass spectrometry.

#### *Effects of CO<sub>2</sub> lasers on preventing caries of enamel and dentin*

Nine studies investigated effects of 10,600 nm CO<sub>2</sub> lasers on morphological and chemical changes of enamel as well as on the reduction of enamel demineralization. Concerning the microhardness analyses, CO<sub>2</sub> laser-treated enamel surfaces showed significantly higher values than that of negative control [14,15]. Polarized light microscopy evaluation revealed that laser-treated enamel groups showed significantly lower lesion depth compared with the nonlased controls [16,17], and the decreased lesions represented up to 87% inhibition of the caries progression [8]. The values of mineral loss of enamel calculated from cross-sectional microhardness analyses were statistically lower in laser-irradiated groups than those in nonlased groups [7,8,16,18-20].

There was an interaction effect between the CO<sub>2</sub> laser irradiation and the organic matrix content of enamel on lesion depth and mineral loss of enamel ( $p < 0.05$ ) [16]. The enamel with the removal of organic content (nonorganic enamel) had a 38% reduction in lesion depth and a 74% reduction in mineral loss after laser irradiation when compared to the nonorganic enamel without laser irradiation. Both reductions were significantly lower than the results for the lased normal enamel but greater than those for the nonlased enamel [16]. CO<sub>2</sub> laser irradiation could induce chemical changes in enamel.

Fourier transform Raman spectroscopy showed a reduced carbonate content for laser-treated enamel when compared with non-treated ones [14,20], while there were no statistical differences for calcium and phosphorus components between irradiated groups and the control by energy-dispersive X-ray fluorescence spectrometry measurements [14]. Observations under scanning electron microscopy revealed evidence of melting and fusion in the enamel samples treated with CO<sub>2</sub> lasers [14,19,20], and melting and fusion were more frequent in the treated groups that underwent more than one laser application [14].

Enamel surfaces exhibiting melted structures were several times bigger than the prismatic structures and fusion across the prism boundaries [14]. It was verified that a laser-modified layer with a coalescence of crystals was presented in human enamel surfaces, forming an irregular solid mass [18]. A homogeneous and smooth recrystallization was also observed on the fused enamel surfaces [14,18]. However, some studies found that the treated enamel areas showed little or even no morphological changes [8,16].

The irradiated enamel surfaces exhibited a similar appearance to the nonlased controls [17], and no signs of surface melting, ablation, crater formation, cracks, or fissures could be observed using the parameters showing caries inhibition effects [16,17]. Three studies investigated the

effects of a 10,600 nm CO<sub>2</sub> laser on morphological features of dentin as well as on the reduction of dentin demineralization [9,21,22].

Root dentin surfaces irradiated with a CO<sub>2</sub> laser with fluences from 4.0 to 6.0 Jcm<sup>-2</sup> showed evidence of melting and resolidification under scanning electron microscopy, and a cracking appearance was more evident on those samples treated with 5.0 and 6.0 Jcm<sup>-2</sup> [22]. Longitudinally fractured samples had a layer of dentin with no tubular structure (20-70 μm thick), whereas the dentinal tubules retained the normal structure below the sealed layer [9]. The values of mineral loss were significantly lower in the laser-irradiated dentin groups than the nonirradiated controls [21,22]. Additionally, laser-induced inhibitory effects on dentin demineralization were observed when fluences reached or exceeded 4.0 Jcm<sup>-2</sup>.

#### *Effects of CO<sub>2</sub> lasers combined with topical fluorides on enamel and dentin*

Eight studies investigated the combined effects of 10,600 nm CO<sub>2</sub> laser irradiation and the application of topical fluoride on enamel. The results of these studies varied. Four studies found that there was a synergistic effect between laser irradiation and fluoride application in reducing enamel demineralization [12,23-25], whereas the other four studies found that the combination of laser and fluoride had no additional impact on decreasing lesion progression when compared with the laser alone [26-29].

Enamel samples treated with laser irradiation in conjunction with the application of titanium tetra fluoride solution showed less calcium loss and increased microhardness than those treated with laser only [24]. Combined sodium fluoride treatment and laser irradiation showed overwhelming effectiveness in preventing enamel demineralization based on the data analysis of mineral loss and lesion depth [12].

When enamel surfaces were treated with amine fluoride solution immediately followed by laser irradiation, they had markedly reduced cracking and fused and melted areas compared to counterparts treated with the laser alone [25]. Moreover, laser irradiation through the amine fluoride solution significantly increased fluoride uptake to the enamel surface layer compared to fluoride treatment alone [25].

Three studies investigated the combined effects of 10,600 nm CO<sub>2</sub> laser irradiation and application of topical fluorides on dentin, and the results of these studies were consistent, showing a significant synergistic effect of laser irradiation combined with fluoride treatment on the prevention of dentin demineralization [11,13,26]. The laser combined with fluoride treatment resulted in an 84.5% decrease in lesion depth [11], significantly less mineral loss [26], and lower calcium and phosphorous losses [13]. In addition, the combination of laser and fluoride yielded significantly lower bacterial numbers on dentin surfaces than laser only.

## **Discussion**

The publication search in this review used three databases: PubMed, Scopus, and Web of Science. They are three of the most commonly used and readily available databases. PubMed, which focuses on medicine and biomedical science, provides open access to the public and includes articles published after 1950.

One advantage of PubMed is that it provides printed literature as well as early versions before printing publication. Scopus and Web of Science belong to commercial companies, and subscription fees are required to use these two databases, which cover most scientific fields. Scopus includes a wider range of journals than PubMed and Web of Science [30], and it includes articles published since 1966. Web of Science is the first comprehensive scientific

citation indexing database covering the oldest publications. The earliest records date back to 1900 [30]. The articles in Web of Science are published in journals with an impact factor. With the use of these three commonly used databases, the results of the retrieval were exhaustive, and they provided comprehensive information on the effects of CO<sub>2</sub> lasers on caries prevention in previous publications.

This systematic review only included 10,600 nm CO<sub>2</sub> wavelength, while studies with 9,300 nm and 9,600 nm wavelengths were excluded. The 10,600 nm wavelength was selected because it has been extensively studied and marketed in the dental field [31]. Moreover, the absorption coefficient of 10,600 nm of enamel and dentin is lower than that of 9,300 nm and 9,600 nm wavelengths. This allows lasers of 10,600 nm wavelength to be six times and ten times deeper in penetration depth in dental hard tissue than those of 9,300 nm and 9,600 nm, respectively. Further, 10,600 nm might be the preferred wavelength for caries prevention studies because of its optical property [24,26].

Enamel and dentin surface irradiated by CO<sub>2</sub> laser either showed little to no surface morphological change [7,8,15-17] or melting, fusion, and recrystallization [14,18-20,22]. Regardless of surface change, irradiated surfaces showed reductions in carbonate content and increases in microhardness and demineralization resistance. Surface changes were thought to be related to demineralization inhibition [14]. Sealing of enamel surface was also suggested to reduce the permeability of enamel surfaces during ion exchange [32,33].

Studies showed that fluences between 5 and 16 Jcm<sup>-2</sup> produced melting, fusion, resolidification, and recrystallization on enamel surface [14,18-20]. Lower fluence is required on dentin than on enamel, as higher surface temperature is achieved due to the lower reflectance loss and lower thermal diffusivity of dentin [21]. Morphological variations on melted surfaces

such as droplets, holes [14], and smooth recrystallined surfaces [18] might foster resistance to demineralization. However, undesirable cracks were also observed [8,22,24].

Although surface melting and fusion reduced permeability, surface morphological change was not necessary for enhancing resistance of enamel to demineralization [8,15-17]. The effect of temperature rise induced by laser in an organic matrix was suggested by Sato [34]. He suggested that the melting and swelling of the organic matrix under heat (300°C to 400°C) blocked the diffusion pathway of calcium, which resulted in calcium loss reduction. Hsu [16] suggested there may be partial blockage of inter and intraprismatic spaces. This affected the ion diffusion in enamel and restricted enamel demineralization. Dentin contains organic and inorganic components similar to enamel, but it is higher in water and organic content by volume. A laser induced thermal effect at 600°C to 900°C was suggested as a mechanism to reduce or eliminate carbonate content and increase crystallinity to reduce tooth [21,22].

Calcium fluoride crystals were present on fluoridated surfaces after topical fluoride application [35,36]. The concentration of fluoride on enamel surfaces was shown to increase with application of acidic fluoride agents [37]. Fluoride agents from neutral to strong acidity (low pH value) were studied in this review [12]. Enamel treated with 4% titanium tetrafluoride (pH1.2) alone had better resistance to demineralization than those treated with laser and titanium tetrafluoride [24].

Laser irradiation before or after fluoride application of 1.23% acidulated fluoride phosphate and 1.23% and 5% sodium fluoride (NaF) at pH between 3.5 and 4.5 improved demineralization resistance and microhardness similar to fluoride alone or laser alone [26-28]. There was a synergistic effect in mineral loss reduction using a laser with 2% NaF neutral solution [12]. It was suggested that laser irradiation in the presence of fluoride resulted in fluorapatite formation as organic matrix was removed [12]. Concentrations of NaF did not



seem to be a factor in synergism with lasers [12,26,28]. It might be deduced from the above studies that acidic fluoride affected the synergism of laser and fluoride, but it appears that neutral fluoride is related to synergism.

Amine fluoride has been demonstrated to superior to sodium fluoride [38]. The amino group binds readily onto enamel enabling fast, even distribution of fluoride over the surface. It was reported that there was a synergistic effect in fluoride uptake and a reduction in enamel solubility using 1% amine fluoride solution (pH not reported), but there was no significant difference in acid resistance between fluoride alone and laser with a fluoride group [25]. However, the same article concluded that laser with fluoride showed beneficial effect in acid resistance.

Synergistic effects of laser and fluoride on dentin did not seem to depend on the sequence of laser and fluoride application [11,13,26]. The use of acidulated phosphate fluoride (pH 3.5 and 1.25% F) and 5% NaF (pH4.5) on dentin gave synergistic effect in reduction in calcium loss and mineral loss [13,26]. However, laser with the same fluoride concentration and acidity did not show synergistic effect on enamel. The use of 2% neutral NaF solution with laser showed synergistic effect on dentin as well as enamel, as discussed earlier. The combination of NaF and laser irradiation resulted in significantly higher uptake of fluoride and lesion depth reduction than fluoride or laser alone [11].

Fluoride interacts with teeth by incorporating into hydroxyapatite crystals. The formation of calcium fluoride-like material on the surface was thought to be the main factor in caries reduction. There were a few postulated mechanisms of combined laser and fluoride. Laser-

induced temperature increase resulted in loss of carbonate in the crystalline structure, which was substituted by fluoride ions, enhancing fluoride uptake [11].

The use of laser to modify dentin surface energy might cause more stable absorption of calcium fluoride [13,26]. The fluoride ions released by calcium fluoride could reduce demineralization and enhance remineralization. The main reason for reduction in lesion depth might be the result of fluoride that firmly bound to root surfaces, acting as a fluoride reservoir against demineralization [11]. Therefore, the effect of CO<sub>2</sub> laser irradiation on the lesion depth of dentin needs further investigation.

The combined effect of laser and fluoride was also likely to be related to effect of on enamel and dentin surfaces. Synergistic effects were achieved with different sets of laser parameters on enamel and dentin with 2% neutral NaF solution [11,12]. There were many variables in laser parameters that could influence the temperature increase and changes on enamel and dentin surfaces [13]. Parameters such as pulse durations, energy densities, irradiation methods, and total pulses irradiated per spot area are not discussed in this review.

## **Conclusion**

In this review, studies found that CO<sub>2</sub> (10,600 nm) lasers could prevent demineralization of enamel and dentin. Although the exact mechanism is not well elucidated, current evidence suggests that CO<sub>2</sub> lasers have a synergistic effect with fluoride in promoting the remineralization of enamel and dentin.

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