Minimizing the aerosol-generating procedures in orthodontics in the era of a pandemic: Current evidence on the reduction of hazardous effects for the treatment team and patients

Theodore Eliades and Despina Koletsi
Zurich, Switzerland

The purpose of this critical review is to list the sources of aerosol production during orthodontic standard procedure, analyze the constituent components of aerosol and their dependency on modes of grinding, the presence of water and type of bur, and suggest a method to minimize the quantity and detrimental characteristics of the particles comprising the solid matter of aerosol.

Minimization of water-spray syringe utilization for rinsing is suggested on bonding related procedures, while temporal conditions as represented by seasonal epidemics should be considered for the decision of intervention scheme provided as a preprocedural mouth rinse, in an attempt to reduce the load of aerosolized pathogens. In normal conditions, chlorhexidine 0.2%, preferably under elevated temperature state should be prioritized for reducing bacterial counts. In the presence of oxidation sensitive viruses within the community, substitute strategies might be represented by the use of povidone iodine 0.2%-1%, or hydrogen peroxide 1%. After debonding, extensive material grinding, as well as aligner related attachment clean-up, should involve the use of carbide tungsten burs under water cooling conditions for cutting efficiency enhancement, duration restriction of the procedure, as well as reduction of aerosolized nanoparticles. In this respect, selection strategies of malocclusions eligible for aligner treatment should be reconsidered and future perspectives may entail careful and more restricted utilization of attachment grips. For more limited clean-up procedures, such as grinding of minimal amounts of adhesive remnants, or individualized bracket debonding in the course of treatment, hand-instruments for remnant removal might well represent an effective strategy. Efforts to minimize the use of rotary instrumentation in orthodontic settings might also lead the way for future solutions.

Measures of self-protection for the treatment team should never be neglected. Dressing gowns and facemasks with filter protection layers, appropriate ventilation and fresh air flow within the operating room comprise significant links to the overall picture of practice management. Risk management considerations should be constant, but also updated as new material applications come into play, while being grounded on the best available evidence. (Am J Orthod Dentofacial Orthop 2020; - - - -)

The pandemic outbreak of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) has had a large impact on the frontline of health care workers, and among those, on dentists and orthodontists. Besides the public health and economic burdens of the coronavirus disease 2019, it is now evident that its massive spread around the world has imposed great occupational challenges, with the implementation of routine dental services being at stake. The nature of the virus’ infectious route, with direct implication of airborne droplets in the form of aerosol, has revealed certain potential hazards underlying conventional and standard oral health care procedures. Orthodontic practices are not to be left aside. An aerosol is defined as a suspension system of solid or liquid particles in a gas. The term was introduced by Frederick G. Donnan to describe an aero-solution—
clouds of microscopic particles in air. The various types of aerosol, classified according to physical form and how they were generated, include dust, fume, mist, smoke, and fog. Aerosol should be differentiated from solid particles staying airborne for some time in the air and the splatter of relatively large sized droplets of water generated by splashes in a dental setting, such as those produced by using the water syringe.

Aerosol-producing dental procedures, along with upcoming concerns, are not new to the dental discipline and at most, these concerns should not be selectively twisted, hampered, or emphasized under the light of the present pandemic or potential future endemics. They are effectively there since more than 20 years, and protective measures for dentists and clinic personnel should be prioritized in practice, irrespective of the presence of a pandemic, epidemic, normalized conditions, or otherwise. Furthermore, these concerns and protective measures should effectively be carried forward through advancements in technologies as well as evidence directed by new knowledge over the years. The current pandemic situation has boosted our thinking and endorsements on how to efficiently manage and minimize aerosol production in contemporary practice.

Evidently, common categories and burdens of orthodontics-related applications producing aerosol and/or airborne particulates are focusing on bonding and debonding strategies. The former involve application of water-spray practices in connection to enamel etching, before conditioning with bonding agents and bracket bonding; the latter pertain to enamel clean-up practices after removal of fixed appliances on completion of orthodontic treatment. Of late, in the line of debonding strategies, an additional procedure liable to aerosol generation has emerged in the clinical field; composite attachment removal after aligner therapy or possible attachment replacement and/or removal cycles during treatment with aligners should not be neglected. This is particularly striking if one considers that most orthodontists and/or other clinicians utilizing aligner methods to straighten teeth and treat malocclusions have adopted wide application of these adjuncts in everyday practice.

With regard to bonding strategies, the conventional acid-etching stage may be employed with the use of a gel etchant of very thick consistency, a gel of lower viscosity, or a liquid etchant (Fig 1). Implications for the first alternative are rather straightforward, as it might require a considerably higher pressure of water flow to be rinsed off, as well as a longer rinsing period; but practically, there is more. Very thick consistencies of gel essentially negate the action of acid for the amount of material not in contact with the enamel surface owing to limited wetting. Thus, the 2 other alternatives are often selected. However, high water pressure used generates splatter, which does not belong to the aerosol classification, but may too contribute to the contamination of the operatory. Water pressure is normally set at 40 PSI in the dental units, with existing air pressure at 80 PSI. The American Dental Association (ADA) has suggested testing of water squirt of more than 1.3 m (~4 ft), as a practical measure of raised water pressure.

Regarding debonding strategies of fixed appliances, implication of rotary instruments used to remove remnants of composite compounds after fixed appliance removal, as well as utilization of water as cooling agent during handpiece usage form priority factors that should be considered. Cutting efficiency and aerosolized dust formation are also discussed. This narrative article aims to discuss the hazards arising from routine orthodontic practices implicated to aerosol generation, sometimes on par with and following examples from standard dental procedures, and also to elucidate potential interventions or alterations of conventional orthodontic applications as an attempt to minimize substantial hazards or adverse effects. The narrative is built on 2 basic pillars regarding aerosol generation; the microbiologic on one side, and particulate production and toxicity related implications on the other.
MICROBIOLOGIC CONSIDERATIONS AND BIOAEROSOLS

The pathogenic pervasiveness of dental aerosol rests in its dependence on the concentration of bacterium or virus load in compressed air, or water-spray spatter mixed-up with tooth material, plaque, blood, calculus, and saliva debris that are theoretically and practically produced during routine dental practice, which makes use of an intraoral service handpiece. As such, orthodontic practices fall within the range of these procedures, seemingly within a more limited extent, but it is important that they are not neglected. The presence of dental unit waterlines (DUWLs) microbiota has also been considered an additional intriguing factor, especially because pathogens get carried forward through the water supply system directly to the handpiece in use. When coolants are used during service, the interaction of the cooling agent with fluids and debris produced within the oral environment as a result of composite or tooth grinding practices or use of ultrasonic scaling is present, and inductively, it may be detected in air-suspended particles and aerosol. The Centers for Disease Control and Prevention has established a safety maximum level of colony forming units (CFUs) emitted and detected in the air at the threshold of 500 CFUs per mL as a result of dental handpiece and water and/or air supply instrumentation usage, excluding coliform bacteria for nonsurgical procedures. These levels are liable to reduction when immunocompromised patients are in chair, and are lowered to 200 CFUs per mL. Evaluation of pathogen levels may be done through simply available test strips or kits. In addition, air and/or water related dental instrumentation (handpiece, spray syringe, and/or ultrasonic scaler) in direct usage to patients’ oral cavity should be flushed and pseudotested for 2 minutes at the start of each day, as well as for 30 seconds between patients.

A recent systematic review on bioaerosols in dental environment has pinpointed the presence of 38 types of micro-organisms, including 19 bacteria and 23 fungal genera, indicating a high variety of a range of species, whereas it was interesting that none of the included articles reported on the presence of viruses or parasites; seemingly, this is not linked to their absence from air-suspended droplets, but rather to the line of focus of the primary studies, partially in favor of the abundance and commonness of the former pathogens and their easier and nonspecific detection through wide air sampling techniques. A mean bacterial load range of 1 to 3.9 CFUs in logarithmic scale has been reported after procedural produced aerosol, while the most eminent load has been reported in the range of 1.5 meters from the oral cavity, even higher compared with closer distance measures such as that of 1 meter from the patient. Fusobacterium family pathogens have been identified in aerosols produced after ultrasonic scaling in practice through checkerboard DNA–DNA hybridization techniques. Of the family, Fusobacterium nucleatum has been identified as a bacterium related to pathologic, ophthalmic, and respiratory implications, while also inductive of cellular apoptosis in vivo. However, the results of checkerboard hybridization techniques should be interpreted with caution as per the exact bacteria species eligible for identification, because such practices are close ended, checked in preselected DNA-probe panels and other pathogens not presupposed might be present within droplet spatters as well. Nonetheless, studies assessing mostly periodontal pathogens have identified an increased prevalence of species belonging to the so-called orange complex in aerosols generated during usage of ultrasonic scaler. These mostly pertained to Campylobacter rectus, Prevotella intermedia, and others, including F. periodonticum in addition to F. nucleatum. Apart from directly exposed aerosolized bacteria, another potential contamination source within dental offices or in hospital based dental units has been identified and special attention has been placed to the presence of Legionella pneumophila as well as Pseudomonas spp in DUWLs. These might well serve as routes of infection for patients and/or dental personnel indirectly and via droplet suspension after aerosol-generating handpiece or water and/or spray syringe usage. Other sources of L. pneumophila constitute air-conditioning systems or cooling towers within dental settings. Interestingly, the novel SARS-CoV-2 has also been lately reported to demonstrate capacity of emanation via the airflow of air-conditioning units in business environments.

An array of clinical studies, since more than 25 years and until recently, have attempted to identify effective methods of reducing pathogen load stemming from aerosol forming procedures in dental settings (Fig 2). The vast majority have studied in-service utilization of ultrasonic scaling, whereas some have reported on orthodontic related strategies of debonding procedures, or other dental prophylaxis or restorative procedures. Largescale efforts have been lately endorsed to collectively appraise all available evidence and provide justifiable ranking of the efficiency of these methods. The most prevalent recorded approaches were preprocedural mouth rinse using a wide
variety of potentially antimicrobial agents, such as, chlorhexidine (CHX) 0.12%, CHX 0.2% or tempered CHX 0.2%, cetylpiridinium chloride 0.05%, povidone iodine (PI) 1%, chlorine dioxide, herbal-based agents, or others pertaining to ozone irrigation, use of high volume evacuators and/or dental isolation systems, or agents added to DUWLs to reduce the load. Evidence from a study on bacterial load during orthodontic procedures comparing bracket debonding followed by enamel clean-up with high-speed handpiece and water cooling versus standard orthodontic care involving archwire and/or ligation change, and replacing procedures, highlighted the increased pathogenic state of aerosols produced by the former, with a mean difference of 49.2 (95% CI, 19.4–79.0) in total CFUs. This highlights the exposure hazards of orthodontists related to certain orthodontic procedures in practice and draws attention to additional prophylactic measures to be selectively taken within the dental operating office. Effectively, bacterial load in aerosol in the dental and/or orthodontic cabinet has shown to be significantly raised immediately within 5 min of service for an aerosol-generating procedure, including enamel clean-up.

Further evidence on microbiologic assessment of aerosol produced after debonding of fixed orthodontic appliances and during composite clean-up has elucidated the increased potential of aerosolized particles, particularly those with aerodynamic diameters of 50 µm or less, to surpass the respiratory barriers and invade deep into the lungs, along with pathogen contaminants. Bioaerosol infiltration has been detected in simulation studies all the way to the respiratory tree from the pharynx to the bronchial alveoli of the lungs. Although decreased particulate size seems to exhibit increased potential to penetrate deep into the lungs, the viability of pathogens has been shown to simultaneously decrease, also impacting biodiversity at the deep respiratory levels. Use of preprocedural mouthrinse with CHX of either 0.12% or 0.2% concentration has been identified by individual studies as an important decontaminating agent contributing to identification of decreased bacterial amounts of infected aerosol; latest data coming from an endorsement to compare all direct and indirect evidence from examined interventions (mouth rinses, evacuators, decontamination of DUWLs, and others) across studies and within dental settings, has revealed this supremacy of preprocedural chlorhexidine mouthrinse over other measures for 30 s to 1 min, but also with documented prevailing of tempered (47°C) CHX 0.2%. Tempered CHX solution at 47°C, has been reported to offer increased anti-microbiologic action against bacteria of the human dental plaque, while also preserving adverse effects on tooth and pulp vitality to the minimum. The increase in bacterial kill rate has been determined to reach as high as 25% surplus, while to avoid storage contamination with toxic compounds such as p-chloroaniline, freshly made CHX solutions should undergo heating. As this measure might be potentially considered impractical for the routine management of clinical practice, it might still be the treatment of choice for highly prone to aerosol induction procedures, with water cooling involvement; other solutions could also be considered for more conservative procedures. Among the priority treatments of choice and apart from CHX solutions (either tempered or nontempered), PI 1% has also been considered a viable alternative.

Aforementioned documented evidence originates, as discussed, primarily from ultrasonic scaling clinical studies, randomized in most cases, while total bacterial count in generated aerosol has been the outcome of interest, leaving virus load aside. Extrapolation to other potentially producing aerosolized compounds procedures, however, seems reasonable within a dental cabinet setting and certain orthodontic procedures, such as fixed appliance debonding, may benefit from such measures.

At present and in the middle of SARS-CoV-2 pandemic mid-2020, there is no evidence from clinical
trials on the effectiveness of interventions taken prepro-
cedurally in dental offices against viral load in air-
suspended droplets or aerosols. However, it would be
reasonable to assume that mouth rinses or irrigates
with proven capacity to interact with viral molecules
and its cellular membranes might prove beneficial. On
the basis of the oxidative action of such agents against
the lipid membrane of coronaviruses, latest reports as
well as primary guidelines of the National Health Com-
mision by the People’s Republic of China on measures
against SARS-CoV-2, have indicated a decreased effec-
tiveness of chlorhexidine as a measure of choice, mostly
because of the lack of oxidative action, while use of
hydrogen peroxide 1%, or PI 0.2% to 1% appear more
realistic as effective alternatives.44-46 Oxidative agents
act directly on the lipid shell membrane of the virus
and destroy cellular components. In particular, PI
action is enhanced by the slow and gradual release of
iodine as carried by the povidone vehicle, while any
adverse effects of iodine are reduced, allowing for a
toxicity-free simultaneous interaction.47 Based on the
absence of clinical trials in the field of virus load of aero-
sol, latest calls have emerged and suggest the use of
flavonoids or cyclodextrine agents to fight or attenuate
SARS-CoV-2 infection through saliva expectorations or
sputters secretions.48 However, their effectiveness re-
mains to be tested.

COMPOSITE GRINDING AND PARTICULATE
PRODUCTION

Cutting instrumentation

Composite grinding and particulate production dur-
ing handpiece instrumentation usage in routine dental
practice has been considered an additional source of
potentially hazardous concern for dentists and ortho-
dontists in general, but also in particular in the middle
of a pandemic of a novel SARS-CoV-2, with unprece-
dented impact worldwide.1

An initial notion before any consideration of pro-
duced aerosolized dust is cutting efficiency and types
of dental rotary instruments that might effectively
reduce grinding duration. Knowledge on the topic may
largely be attributed to the extensive research and
work on this field by A.J. von Fraunhofer et al.49-53

Type of cutting bur and mode of action

First, discrimination between commonly used burs in
terms of cutting mechanism is discussed, roughly be-
tween 2 of the most prevalent cutting instruments in
use, tungsten carbide and diamond burs. The tungsten
carbide burs differ from diamond burs, as they are
considered to achieve material removal through a
flow-dependent fracture process (plastic flow), occurr-
ing as a result of elevated shear forces between the car-
bide blades and the material surface; this makes them
rotary instruments of choice for cutting ductile sub-
strates including composites, dentin, or metals. Dissim-
ilarly, diamond cutting burs induce brittle fracture of
substrates, functioning by creating grooves and making
use of dislocation motion and subsequent radial flow of
the material, ultimately leading to propagation of cracks
by the generated tensile stresses produced and chip for-
mation. Evidently, diamond burs are mostly efficient for
ceramics or enamel surface.49 Latest innovations for ad-
hesive removal after completion of orthodontic treat-
ment, entail the use of fiber-glass or fiber-reinforced
composite burs, which have been reported to exhibit a
potential for reduced enamel surface roughness on
enamel clean-up, compared with standard carbides.54,55

In reality, no data is currently available with respect to
the effect of these cutting burs on particulate composite
dust dynamic.

Moreover, water supplementation and spray patterns
of the handpiece during tooth or material grinding, apart
from the straightforward effect on preservation of tem-
perature within tooth and pulpal tolerable standards,
have also been implicated as a medium for achieving ef-
ciciency during the cutting procedure.36 Water spray
during tooth preparation within a proximal value of
40 mL/min room temperature has been considered
reasonable for avoiding pulp interactions.36 In reality,
water or other lubrication medium has been considered
to play a significant role in cutting efficiency following
Reynold’s hydrodynamic lubrication theory.

In particular, across dental setting environments
where standard and known length and material cutting
instruments are used for commonly used 400,000 rpm
bur rotation speed, it appears unlikely that effects of dy-
namic viscosity of coolant media may be significant.
Testing across water coolant, alcohol (1%) as well as
glycerol (2%) solution has revealed comparable effects.59

Further, water application as coolant usage during
material grinding in practice, including enamel clean-
up from bonding remnants after orthodontic treatment,
offer a thin line layer of interproximal matter between
the carbide and material interface. This is considered
to induce surface adsorption alterations in the substrate
material after reduction of the surface-free energy, pro-
duced by changes in the strength of association of the
interatomic bonds between interactive entities, thus re-
sulting to surface hardness changes.59 To this respect,
and as discussed above, cutting with carbide burs in
ductile substrates such as resin remnants after debond-
ing of fixed appliances or bonded attachment removal
after or during aligner therapy, shall be advantaged, in
terms of cutting efficiency, by water supplementation targeted directly to the carbide–composite interface, in the following manner: initial groove formation after bur application is generated, followed by lateral displacement of the substrate, pilling-up material dislocation, and crack propagation, resulting in chip formation. The described procedure broadly follows the original work of Rehbinder et al back in 1940s, who suggested that chemically-induced surface hardness changes bear the potential to increase drilling efficiency of the cutting tool in mining settings with aqueous surfactant solutions, within a range of 30%-50%. Gain is 2-fold, with subsequent extrapolation to orthodontic and dental practice: faster advancement of the bur into the substrate and decreased demand for heavy load application in practice, thus reduction in operating time and total amount of aerosol production.

**Material substrate, composite dust, and aerosol**

Resin composites are known to possess a wide range of applications in dentistry, with orthodontics usage in bonding procedures of both fixed appliances as well as treatment with aligners and attachment adjuncts being in the spotlight. Normal composite composition comprises of the resin matrix (usually represented by bisphenol-A [BPA] diglycidyl dimethacrylate, triethylene glycol dimethacrylate, and ethoxylated BPA glycol dimethacrylate), the inorganic filler compounds as well as a coupling agent to guarantee bonding between the two. Filler compounds usually fall below 0.4 μm and may serve in a wide range of particulate sizes and even fall within the nano-range. Orthodontic adhesives have also been considered to acquire quartz-type filler particles as well. Heavy metal oxides are preferred, namely barium, strontium, zinc, aluminum, or zirconium, while their primary service remains to offer enhanced physical and mechanical properties to the material, including polymerization shrinkage water sorption and solubility, radiopacity, and reduction of biodegradation in-service.

During debonding strategies, but also lately increasingly during attachment removal in the course of and/or after the end of aligner treatment with thermoplastic devices, breakdown of the bulk of composites takes place, with material micro- and/or nano-fragments being aerosolized. These particulates bear the aerodynamic potential to surpass the respiratory fraction barriers and natural defense mechanisms of the clinician, patient and office personnel and find their way deep into the lungs.

A foremost effort to provide evidence in the field of aerosolized composite compounds in dental settings, has been mainly initiated and driven by 2 separately working groups in Leuven, Belgium, and Bristol in the United Kingdom, in essence after simulation in clinical conditions. Evidently, aerosols comprising of particles lower than 10 μm or 2.5 μm (PM_{10} or PM_{2.5}, particulate matter) are gaining attention due to their potential to enter the respiratory tract; interestingly, even smaller particulates within the range of dozens of nanometers (<100 nm) have been associated with an increased dynamic to surpass the primary boundaries of the respiratory system and reach deepest levels of the terminal epithelial bronchioles of the lungs because of their increased surface to volume ratio, offering an amplified reactive potential when in interaction with cellular interfaces.

Several studies have investigated the content compounds of composite dust produced in aerosols in dental and orthodontic setting, and it has been claimed that percentage and concentration of nano-sized identified filler particles in the aerosols might be related to the original filler content of the composites. However, this is far from the case, because all types of composites, irrespective of filler size, have been reported to exhibit significant amounts of nanoparticles within the range of 38–70 nm during grinding and clean-up. In particular, surface friction and heating shock during composite grinding results to matrix decomposition of the substrate, aging, C=C conversion of bonds on surface, and ultimately production of respirable composite dust.

**Wet or dry conditions**

Apart from water supplementation contribution to the cutting efficiency of grinding tools on the composite substrate during debonding, thus offering minimization of (bio-)aerosol production duration, the effect of water as per emanation, and generation of airborne dust has been disputed, however, with scarce evidence from few research efforts, across variable settings. In essence, a recent study inspected the effect of water cooling in slow-handpiece usage on bulk composite sticks containing an array of filler sizes under simulated conditions of dry and wet grinding. Their work suggested consistent findings for all types of composites, which demonstrated a significant reduction in the number of detected nanoparticles being released when water spray was in-service (5.6 \times 10^5 - 13.7 \times 10^7 numbers per cubic centimeter), denoting a half-pace reduction, compared with dry settings. Interestingly though, both dry and wet grinding alternatives produced high numbers of nano-sized particulates being aerosolized overall. The highest amounts have been detected during the last minute of grinding, reaching levels of approximately 33 \times 10^5 numbers.
per cubic centimeter. Particulate agglomeration has been considered to occur across time, thus contributing in increasing average particulate diameter, overall. To this respect, under water usage conditions, airborne generated nanoparticles have been considered particularly prone to being trapped within water droplets, resulting in increased matter sizes, which are less likely to achieve penetration of the epithelial bronchial barriers and find their way to the lungs.

The aforementioned conditions and settings could be considered as vastly resembling to the bulk attachment material removal during orthodontic treatment with aligners. As previously discussed, aligner usage for treatment of malocclusions currently involves increasingly frequent adoption of composite grips bonded to tooth enamel, sometimes more than 1 per tooth, as attachments of various sizes and shapes, with nonnegligible dimensions, varying within the range of 2–5 mm and also width or thickness that may exceed 1 mm. These adjuncts target to the achievement of modes of tooth movement, either rotational or translational, within all 3 planes of space, which would otherwise be non-manageable with the early phase plain thermo-plastic aligner usage, that do not necessitate enamel involvement. This compares to the thin layer of composites used as a layer of “sandwich-type” pattern between the bracket base and the enamel surface in a conventional case fixed appliance treatment, with an average estimated thickness of 150 to 250 μm; one may evidently cognize that the bulk and thickness of the attachment grips in aligner therapy is implicated in 2 conditions: first, the occurrence of an excessive amount of composite polymerized material within the oral cavity, allowing for the potential risk of BPA release or monomer leaching, depending on the number and shape or size; second, grinding procedures for attachment removal may prove extremely exhaustive and timely, bearing an increasing risk of excessive production of aerosolized composite dust.

Handpiece role

Furthermore, an earlier report on human extracted teeth and subsequent simulated bracket removal and enamel clean-up, has examined the effect of handpiece, water coolant, and high volume evacuator as well as surgical facemask, on the amount of particulate production and particle concentration during composite grinding after debonding; however, the baseline effect of handpiece was variable, because slow-speed handpiece was used in absence of water coolant, whereas high-speed handpiece only under water-spray emission. Findings structured on nonparametric data revealed a significantly higher concentration of airborne particulates under wet conditions and the use of high-speed instrumentation. In addition, use of facemask appeared considerably effective, contributing to the reduction of the detected concentration, while high volume evacuator was not identified as a critical parameter in this respect. To date, there is no further evidence on the direct crude effect of handpiece variation and rotary instrumentation speed with regard to airborne particulate generation, under otherwise comparable conditions.

Cytotoxicity and Estrogenicity of aerosolized particulates

Following research about cytotoxicity and xenogenic effects of BPA and/or monomer release of adhesive compounds within the oral cavity, airborne particulates produced during grinding of composites after fixed appliance removal or aligner’s attachment elimination, are seemingly a potential source of similar concerns. A mild but gradual reduction of human bronchial epithelial cell viability in laboratory conditions has been documented, giving rise to speculations on the reactive dynamic of such particulates. Composite filler particles and matrix composition of restorative adhesives did not appear to play a role. Interestingly, the latest report encompassing orthodontic adhesive material evaluation at grinding stages after simulated conventional orthodontic treatment, pinpointed the aptitude of aerosolized particles of adhesives comprising of quartz-type fillers to demonstrate disrupting effects on interacting cell membrane integrity and cellular viability, while also to intervene with cellular growth potential of epithelial bronchial populations at an early stage. These effects are probably related to the size and shape of such fillers’ configuration, following the increased surface to volume ratio they present.

Related evidence on orthodontic adhesives comes also from the assessment of in vitro estrogenicity of orthodontic composites ground under simulated bonding–debonding settings. Estrogenic effects appear as a result of residual monomer release (BPA), which follows action as an endocrine disruptor because of the very similar structure with beta-estradiol. Under the use of highspeed handpiece without water-spray, eluents containing airborne particulates, after grinding different types of adhesives (ie, chemically or light-cured), have shown an increased proliferating capacity on MCF-7 breast cancer cells in vitro.

Such findings are of particular interest and raise considerable awareness when it comes to the large-scale removal of attachment grips implicated in aligner therapy. The bulkiness and volume of these adjuncts
evidently requires a great amount of grinding efforts and intraoral cutting instrumentation service. It is therefore likely that a significant amount of heat influx occurs first at the surface of the composite substrate if not substantially cooled, resulting in heat shock of the matrix.\textsuperscript{8} Resultant effects on chemical decomposition of the produced aerosolized dust with further implications on monomer release and BPA diglycidyl dimethacrylate

| Procedure          | Aerosol-liable actions (conventional)                                                                 | Safety measures                               | Future perspectives                          |
|--------------------|---------------------------------------------------------------------------------------------------------|-----------------------------------------------|---------------------------------------------|
| Etching            | High thickness and/or viscosity gel                                                                     | Liquid gel and/or low viscosity               | Nonetching mediated bonding                  |
|                    | Self-etching primer and/or no rinsing                                                                    | Glass-ionomer cement and/or no rinsing        |                                             |
| Bonding            | Conventional resin-based adhesive                                                                     | Glass-ionomer cement                          | Biomimetic based bonding with use of L-DOPA primers |
|                    | BPA-free adhesives                                                                                        |                                               |                                             |
| Debonding          | Standard debonding with considerable amounts of adhesive remnants on enamel surface                     | Alteration of adhesive-bracket base interface | Command-debond adhesives                     |
|                    |                                                                                                        |                                               | (thermally expandable particles and ferrous micro-particles) |
|                    |                                                                                                        | Identify bracket base mesh and/or shape and/or size and adhesive combination for cohesive resin fracture | Irradiation of specific wavelength to reverse polymerization |
|                    | Standard rotary grinding to clean-up enamel                                                              | Removal of significant amounts of resin remnants with hand-instruments—avoid rotary instrumentation as much as possible | Temperature control and variation of adhesives (heat and/or freezing) plasticization and/or brittleness |
|                    | Use of tungsten burs*, w/o water cooling for limited trace composite remnants (ie, individually debonded brackets during treatment) |                                               |                                             |
|                    | Use of tungsten burs*, under water cooling for enamel clean-up after debonding and/or attachment removal |                                               |                                             |
| Attachment grips   | Careful selection of patients and/or malocclusions for treatment with aligners; abandon company preset distribution of arrays of attachments |                                               |                                             |
| Attachment-free    | Use of BPA-free composite to eliminate estrogenic activity (ie, PCDMA)                                    |                                               |                                             |
| treatment          |                                                                                                        |                                               |                                             |
| Preprocedural       | Mouthrinse with (47°C) CHX 0.12%-0.2% for bacterial pathogens (0.5-1 min)                               |                                               |                                             |
| measures            |                                                                                                        |                                               |                                             |
| Personnel equipment | Mouthrinse with 0.2%-1% PI or 1% H$_2$O$_2$ for oxidation vulnerable viruses (0.5-1 min)                 |                                               |                                             |
| and/or settings     | Facemask, shield, gown, apparel for all clinic personnel, and fresh air and surgical suction             |                                               |                                             |

L-DOPA, L-3,4-dihydroxyphenylalanine; w/o, without; PCDMA, phenylcarbamoyloxy-propane dimethacrylate; H$_2$O$_2$, hydrogen peroxide.

*Smaller number of flutes in the beginning of removal, advancing to 20-fluted for polishing.
compounds might be alarming. Thus, broad and time-consuming composite removal, as required in extensive removal of attachments, with no water cooling in-service, should largely be avoided, while further research in the field is critical to detect specific effects of water supplementation to the emanation of monomer and other potentially estrogenic compounds.

**IMPLICATIONS AND RECOMMENDATIONS FOR CLINICAL PRACTICE**

Direction of measures taken to minimize effects of aerosol production in orthodontic practice should target in 2 basic routes: bonding and debonding procedures, in essence those are interconnected (Table).

**Bonding**

The former basically comprises procedures that take place before bracket placement on tooth surface and involve rinsing actions for enamel preparation agents and use of certain types of bonding materials. As previously stated, very thick consistencies and substantial amounts of etchant acid gels applied on tooth surface, apart from presenting compromised action per se, evidently require higher water and/or spray pressure to be rinsed off, thus increasing the likelihood for spatter emanation and droplet formation, but also resulting in prolonged working times. Conventional acid-etching agents entailing low viscosity or even liquid gels should be prioritized. Self-etching primer alternatives have also been proposed, although these may require careful pumicing to ensure a precipitations-free enamel substrate. In the same line and to avoid rinsing application and aerosol production, glass-ionomer cements as compared with conventional light-cured counterparts may be preferred. These material alternatives present a chemical interaction and adherence with enamel surface, do not involve prior conventional enamel conditioning, or involve a thin layer of polyacrylic acid agent in contact with enamel, with an induced shallow depth of penetration of approximately 5 to 7 μm. They are also less susceptible to moisturized oral cavity conditions, thus offering a viable alternative to classic adhesives bringing the aforementioned advantages, but also bearing a reduced risk for iatrogenic damage to the enamel surface. However, all currently and widely adopted bonding alternatives do not target on the desirable minimization of adhesive remnants covering the enamel surface after debonding.

Starting from the necessity of an enamel-friendly bonding agent, there has been an endorsement and inspiration, following nature and wildlife environment, to design new material structures on par with living creatures’ observations. These form the so-called biomimetic materials. For example, gekkonidae lizards (geckos) acquire a unique adhesion ability attributed to their foot pad, the “contact splitting.” In particular, geckos’ foot pad contains densely packed ultrafine hair, split in the endings, thus offering increased number of contact points per unit area, contributing to greater adhesion forces generated. As such, geckos are capable of sustaining their weight upside-down, with a gravity-defying ability, without mediation of any chemical agent, relying only to physical forces, otherwise being impossible to achieve. This type of strong gecko-feet grip has inspired the design of...
medical adhesives and might attain applicability in orthodontic bonding agents for dry environments. Moreover, to overcome failures of geckos’ inspired materials, in wet conditions, scientists have studied the use of mussel adhesion as a combination approach, with a resulting new material named “geckel,” which might exhibit enhanced adhesion potential both in dry and wet conditions. Mussel biomimetic polymers are based on L-3,4-dihydroxyphenylalanine (DOPA), offering “sticky” and “glue” resembling properties in the materials. In essence, biomimetic based bonding offers a combination approach, with a resulting new material named “geckel,” which might exhibit enhanced adhesion potential both in dry and wet conditions. Mussel biomimetic polymers are based on L-3,4-dihydroxyphenylalanine (DOPA), offering “sticky” and “glue” resembling properties in the materials.101 In essence, biomimetic based bonding primers such as L-DOPA might offer clinicians a significant tool against oral environment conditions. In combination with geckos’ related properties and applicability to bracket bases, sufficient bond strength to enamel surface might be achieved, without necessitation for prior enamel conditioning, also making debonding practices and enamel clean-up at the end of treatment, effortless.

Debonding

Pertaining to debonding procedures, calls and endorsements for aerosol containment, in general, should be focused first on preventive measures to minimize composite remnants after bracket removal in conventional orthodontics, and second on effective grinding patterns to reduce dust, particulate generation, and operating time, with further speculations on bio-aerosol formation and microbiologic perspectives, as well as xenoestrogenic action of the produced particulate matter. The composite-bracket base interface may play a significant role in achieving a desirable limited amount of adhesive remnant for grinding. Alterations in the adhesive-base interlocking characteristics may take place by induced modifications in the resin filler content and also in the adhesive retention patterns within the bracket base.106 Targeting an efficient combination of bracket base mesh, size, and shape with adhesive composition that may result in a cohesive composite fracture on debonding, would allow for minimal enamel clean-up (Fig 3).

In this respect, applications from high technology and automotive industries might offer reformatory solutions in orthodontic procedures in the near future. Lately, adhesives that debond on command have been used in interlocking joint positions in technology adjuncts to allow for a temperature-controlled initiation of the debonding process.106,107 This is achieved mostly through the embedding of thermally expandable particles (TEMs) into the adhesive matrix.107 The idea about TEMs dates many decades back and resides in the transformation of the particles through heat shock, occurring by softening of the cell particulate matter jointly with gasification of the inner liquid phase hydrocarbon.103,104 In the same line, ferrous microparticles within the micron range, have been introduced as fillers and act by being preferentially distributed after external magnet polarity reversal, thus inducing destabilization of the polymer structure and initiating crack states within the resin matrix that may easily be diffused. Other initiatives might also entail application of irradiation to reverse polymerization and produce a highly viscous adhesive state easily to be removed.96

Wide adoption of BPA-free adhesives has been suggested for a range of dentistry applications including orthodontic bracket or fixed retainer bonding. To this line, advantages of such alternatives which miss BPA monomer derivatives, have been directed towards the elimination of the reactive oxygen species produced after BPA leaching in the oral cavity, after incomplete polymerization of the adhesives and being able to incite an estrogenic potential. The majority of such alternatives make use of aliphatic co-monomers based on triethylene-glycol dimethacrylate, urethane dimethacrylate, and cycloaliphatic dimethacrylates or are effectively represented by a single aromatic dimethacrylate derivative. These efforts might prove beneficial also with regard to elimination of BPA release in aerosolized dust at the debonding stage.96,105

CONCLUSION

In all, wide and consistent adoption of occupational measures to control generation of aerosol in orthodontic practice should be universal, with microbiologic considerations, particulate matter production as well as toxicity related perspectives being on the spot, even more within the course of a pandemic. Realistic management in practice, should focus on bonding and debonding strategies, while careful selection of procedures and application of safety measures depending on individualized patient needs is fundamental.

In particular, minimization of water-spray syringe utilization for rinsing is anticipated on bonding related procedures, while temporal conditions as represented by seasonal epidemics should be considered for the decision of intervention scheme provided as a procedural mouthrinse, in an attempt to reduce the load of aerosolized pathogens. In normal conditions, CHX 0.2%, preferably under elevated temperature state should be selected for minimization of bacterial load. In the presence and spread of oxidation vulnerable viruses within the community, substitute strategies should be opted, effectively represented by the use of PI 0.2%-1%, or hydrogen peroxide 1%.

After debonding, largescale enamel clean-up strategies should entail the use of carbide tungsten burs under
water cooling conditions, to augment cutting efficiency, timely fulfillment of the procedure, as well as reduction of aerosolized nanoparticles. Attachment clean-up at the end of aligner therapy falls into this category; however, selection strategies of malocclusions eligible for aligner treatment should be reconsidered, and a more confined use of attachment grips might also be a viable future perspective. For more limited clean-up procedures, with traces of adhesive remnants left on enamel substrate or individual “re-bracketings” or grinding after bracket breakage in the course of treatment, water cooling rotary instrumentation might not be the treatment of choice, whereas hand-instruments for remnant removal might represent better an effective strategy.

Furthermore, in-office measures of self-protection should never be neglected. Dressing gowns and face-masks with filter protection layers and face shields for all clinic personnel, appropriate ventilation, and fresh air flow within the operating room are of paramount importance. Risk management considerations should be constant but also updated as new material applications come into practice and/or epidemiologic equilibrium of the community is disrupted.

REFERENCES

1. World Health Organization. Coronavirus situation report. Available at: https://www.who.int/emergencies/diseases/novel-coronavirus-2019. Accessed April 22, 2020.
2. Meng L, Hua F, Bian Z. Coronavirus Disease 2019 (COVID-19): emerging and future challenges for dental and oral medicine. J Dent Res 2020;99:481-7.
3. Tang D, Comish P, Kang R. The hallmarks of COVID-19 disease. PLoS Pathog 2020;16:e1008536.
4. Hinds WC. Aerosol technology: properties, behavior, and measurement of airborne particles. 2nd ed. Hoboken: Wiley-Interscience; 1999.
5. Porter SR. Infection control in dentistry. Curr Opin Dent 1991;1:429-35.
6. Eliades T, Papageorgiou S, Ireland A. The use of attachments in aligner treatment: analyzing the “innovation” of expanding the use of acid etching-mediated bonding of composites to enamel and its consequences. Am J Orthod Dentofacial Orthop; July 2, 2020 [Epub ahead of print].
7. Iliadi A, Koletsi D, Papageorgiou SN, Eliades T. Safety considerations for thermoplastic-type appliances used as orthodontic aligners or retainers. A systematic review and meta-analysis of clinical and in-vitro research. Materials (Basel) 2020;13:1843.
8. Weckmann J, Scharf S, Graf I, Schwarze J, Keilig L, Bourauel C, et al. Influence of attachment bonding protocol on precision of the attachment in aligner treatments. J Orofac Orthop 2020;81:30-40.
9. Dai FF, Xu TM, Shu G. Comparison of achieved and predicted tooth movement of maxillary first molars and central incisors: first premolar extraction treatment with Invisalign. Angle Orthod 2019;89:679-87.
10. Infection control recommendations for the dental office and the dental laboratory. ADA council on scientific affairs and ADA council on dental practice. J Am Dent Assoc 1996;127:672-80.
11. Tuvo B, Totaro M, Cristina ML, Spagnolo AM, Di Cave D, Profeti S, et al. Prevention and control of Legionella and Pseudomonas spp. Colonization in dental units. Pathogens 2020;9:305.
12. Centers for Disease Control and Prevention. Summary of infection prevention practices in dental settings: basic expectations for safe care. Atlanta, GA: Centers for Disease Control and Prevention, US Dept of Health and Human Services; 2016.
13. Australian Dental Association. ADA’s guidelines for infection control. 3rd ed. St Leonards, Australia: Australian Dental Association; 2015.
14. Zemouri C, de Soet H, Crielaard W, Laheij A. A scoping review on bio-aerosols in healthcare and the dental environment. PLoS One 2017;12:e0178007.
15. Rautemaa R, Nordberg A, Wuolijoki-Saaristo K, Meurman JH. Bacterial aerosols in dental practice – a potential hospital infection problem? J Hosp Infect 2006;64:76-81.
16. Feres M, Figueiredo LC, Faveri M, Stewart B, de Vizio W. The effectiveness of a preprocedural mouthwash containing cetylpyridinium chloride in reducing bacteria in the dental office. J Am Dent Assoc 2010;141:415-22.
17. Retamal-Valdes B, Soares GM, Stewart B, Figueiredo LC, Faveri M, Miller S, et al. Effectiveness of a pre-procedural mouthwash in reducing bacteria in dental aerosols: randomized clinical trial. Braz Oral Res 2017;31:e21.
18. Bhattacharya S, Livesy SA, Wiselka M, Bukhari SS. Fusobacteriosis presenting as community acquired pneumonia. J Infect 2005;50:236-9.
19. Kang W, Jia Z, Tang D, Zhang Z, Gao H, He K, et al. Fusobacterium nucleatum facilitates apoptosis, ROS generation, and inflammatory cytokine production by activating AKT/MAPK and NF-κB signaling pathways in human gingival fibroblasts. Oxid Med Cell Longev 2019;2019:1681972.
20. Kang W, Ji X, Zhang X, Tang D, Feng Q. Persistent exposure to Fusobacterium nucleatum triggers chemokine/cytokine release and inhibits the proliferation and osteogenic differentiation capabilities of human gingiva-derived mesenchymal stem cells. Front Cell Infect Microbiol 2019;9:429.
21. Laheij AMG, Kistler JO, Belibasakis GN, Välimaa H, de Soet JJ, European Oral Microbiology Workshop (EOMW) 2011. Healthcare-associated viral and bacterial infections in dentistry. J Oral Microbiol 2012;4.
22. Paluśnińska-Szysz M, Cendrowska-Pinkosz M. Pathogenicity of the family Legionellaceae. Arch Immunol Ther Exp (Warsz) 2009;57:279-90.
23. Lu J, Gu J, Li K, Xu C, Su W, Lai Z, et al. COVID-19 outbreak associated with air conditioning in restaurant, Guangzhou, China, 2020. Emerg Infect Dis 2020;26:Epub 2020 Apr 2.
24. Gupta G, Mitra D, Ashok KP, Gupta A, Soni S, Ahmed S, et al. Efficacy of preprocedural mouth rinsing in reducing aerosol contamination produced by ultrasonic scaler: a pilot study. J Periodontol 2014;85:562-8.
25. Holloman JL, Mauriello SM, Pimenta L, Arnold RR. Comparison of suction device with saliva ejector for aerosol and spatter reduction during ultrasonic scaling. J Am Dent Assoc 2015;146:27-33.
26. Jawade R, Bhandari V, Ugale G, Taru S, Khaparde S, Sukarni A, et al. Comparative evaluation of two different ultrasonic liquid coolants on dental aerosols. J Clin Diagn Res 2016;10:ZC53-7.
27. Joshi AA, Padhye AM, Gupta HS. Efficacy of two pre-procedural rinses at two different temperatures in reducing aerosol contamination produced during ultrasonic scaling in a dental set-up - a microbiological study. J Int Acad Periodontol 2017;19:138-44.
28. Paul B, Baiju RMP, Raseena NB, Godfrey PS, Shanmoogle PL. Effect of aloe vera as a preprocedural rinse in reducing aerosol contamination during ultrasonic scaling. J Indian Soc Periodontol 2020;24:37-41.

29. Waghmare SV, Kini VV, Srivastava S. Comparative evaluation of colony forming unit count on aerobic culture of aerosol collected following pro-procedural rinses of either 0.2% chlorhexidine gluconate or 1% stabilized chlorine dioxide during ultrasonic scaling: a clinical and microbiological study. J Contemp Dent Pract 2018;8:70-6.

30. Swaminathan Y, Thomas JT, Muralidharan NP. The efficacy of preprocedural mouth rinse of 0.2% chlorhexidine and commercially available herbal mouth containing Salvadora pensca in reducing the bacterial load in saliva and aerosol produced during scaling. Asian J Pharm Clin Res 2014;7:21-4.

31. Toroglu MS, Haytaç MC, Köksal F. Evaluation of aerosol contamination during debonding procedures. Angle Orthod 2001;71:299-306.

32. Dawson M, Soro V, Dymock D, Price R, Griffiths H, Dudding T, et al. Microbiological assessment of aerosol generated during debond of fixed orthodontic appliances. Am J Orthod Dentofac Orthop 2016;150:831-8.

33. Logothetis DD, Martinez-Welles JM. Reducing bacterial aerosol contamination with a chlorhexidine gluconate pre-rinse. J Am Dent Assoc 1995;126:1634-9.

34. Parohit B, Priya H, Achariya S, Bhat M, Ballal M. Efficacy of pre-procedural rinsing in reducing aerosol contamination during dental procedures. J Infect Prevent 2009;10:190-2.

35. Koletsi D, Belibasakis GN, Eliades T. Interventions to reduce aerosolized pathogens in dental practice. A protocol for a systematic review and meta-analysis. Available at: https://osf.io/ewph9/. Accessed May 2, 2020.

36. Koletsi D, Belibasakis GN, Eliades T. Interventions to reduce aerosolized pathogens in dental practice. A systematic review with network meta-analysis of randomized controlled trials. J Dent Res 2020 [Submitted manuscript].

37. Kaur S, White S, Bartold PM. Periodontal disease and rheumatoid arthritis: a systematic review. J Dent Res 2013;92:399-408.

38. Mamajiwala AS, Sethi KS, Raut CP, Karde PA, Khedkar SU. Comparative evaluation of chlorhexidine and cinnamon extract used in dental unit waterlines to reduce bacterial load in aerosols during ultrasonic scaling. Indian J Dent Res 2018;29:749-54.

39. Micik RE, Miller RL, Mazzarella MA, Ryge G. Studies on dental microbiology. I. Bacterial aerosols generated during dental procedures. J Dent Res 1969;48:49-56.

40. Tham KW, Zuraimi MS. Size relationship between airborne viable bacteria and particles in a controlled indoor environment study. Indoor Air 2005;15(Suppl 9):48-57.

41. Shetty SK, Sharath K, Shenoy S, Sreekumar C, Shetty RN, Biju T. Compare the efficacy of two commercially available mouth rinses in reducing viable bacterial count in dental aerosol produced during ultrasonic scaling when used as a preprocedural rinse. J Contemp Dent Pract 2013;14:848-51.

42. Reddy S, Prasad MGS, Kaul S, Satish K, Kakarala S, Bhowmik N. Efficacy of 0.2% tempered chlorhexidine as a pre-procedural mouth rinse: a clinical study. J Indian Soc Periodontol 2012;16:213-7.

43. König J, Storcks V, Kocher T, Bößmann K, Plagmann HC. Antiplaque effect of tempered 0.2% chlorhexidine rinse: an in vivo study. J Clin Periodontol 2002;29:207-10.

44. Peng X, Xu X, Li Y, Cheng L, Zhou X, Ren B. Transmission routes of 2019-nCoV and controls in dental practice. Int J Oral Sci 2020;12:9.

45. Izzetti R, Nisi M, Gabriele M, Graziani F. COVID-19 transmission in dental practice: brief review of preventive measures in Italy. J Dent Res 2020;22:0345:209:20580. Epub 2020 Apr 17.

46. National Health Commission PRC. Guidance for corona virus disease 2019. Prevention, control, diagnosis and management. 5th ed. China: National Health Commission by the People’s Republic of China; 2020.

47. Yoo JH. Review of disinfection and sterilization - back to the basics. Infect Chemother 2018;50:101-9.

48. Carrouel F, Conte MP, Fisher J, Gonçalves LS, Dussart C, Llodra JC, et al. COVID-19: a recommendation to examine the effect of mouthrinses with β-Cyclodextrin combined with Citrox in preventing infection and progression. J Clin Med 2020;9:1126.

49. von Fraunhofer JA, Siegel SC. Enhanced dental cutting through chemomechanical effects. J Am Dent Assoc 2000;131:1465-9.

50. von Fraunhofer JA, Siegel SC, Feldman S. Handpiece coolant flow rates and dental cutting. Oper Dent 2000;25:544-8.

51. Siegel SC, von Fraunhofer JA. The effect of handpiece spray patterns on cutting efficiency. J Am Dent Assoc 2002;133:184-8.

52. Siegel SC, von Fraunhofer JA. Comparison of sectioning rates among carbide and diamond burs using three casting alloys. J Prosthodont 1999;8:240-4.

53. Siegel SC, von Fraunhofer JA. Dental burs—what bur for which application? A survey of dental schools. J Prosthodont 1999;8:258-63.

54. Shah P, Sharma P, Goje SK, Kanazariya N, Parikh M. Comparative evaluation of enamel surface roughness after debonding using four finishing and polishing systems for residual resin removal in an in vitro study. Prog Orthod 2019;20:18.

55. Garg R, Dixit P, Khosla T, Gupta P, Kalra H, Kumar P. Enamel surface roughness after debonding: a comparative study using three different burs. J Contemp Dent Pract 2018;19:521-6.

56. Ercoli C, Rotella M, Funkenbusch PD, Russell S, Feng C. In vitro comparison of the cutting efficiency and temperature production of 10 different rotary cutting instruments. Part I: turbine. J Prosthodont 2009;101:248-61.

57. Rehländer PA, Wenström EK. The effect of medium and adsorption layers on plastic flow of metals. Bull Acad Sci USSR ser PhD 1983:4-531-8.

58. Darwell BW. Materials science for dentistry. 9th ed. Cambridge: Woodhead Publishing Limited; 2009.

59. Eliades T, Bisphosphon A and orthodontics: an update of evidence-based measures to minimize exposure for the orthodontic team and patients. Am J Orthod Dentofacial Orthop 2017;152:435-41.

60. Chen MH. Update on dental nanocomposites. J Dent Res 2010;89:549-60.

61. Villarroel M, Fahl N, De Sousa AM, De Oliveira OB Jr. Direct esthetic restorations based on translucency and opacity of composite resins. J Esthet Restor Dent 2011;23:73-87.

62. Illiad A, Koletsi D, Eliades T, Eliades G. Particulate production and composite dust during routine dental procedures. A systematic review. Available at: https://osf.io/so9m4/. Accessed May 1, 2020.

63. Illiad A, Koletsi D, Eliades T, Eliades G. Particulate production and composite dust during routine dental procedures. A systematic review with meta-analyses. Materials (Basel) 2020;13:E2513.

64. Cokic SM, Ghosh M, Hoet P, Godderis L, Van Meerbeek B, Van Larebeke N. Cytotoxic and genotoxic potential of respirable fraction of composite dust on human bronchial cells. Dent Mater 2020;36:270-83.

65. Ilie N, Hickel R. Investigations on mechanical behaviour of dental materials. 4th ed. China: National Health Commission by the People’s Republic of China; 2020.

66. Meyer GR, Ernst CP, Willershausen B. Determination of polymerization stress of conventional and new “Clustered” Microfilled Composites in comparison with Hybrid Composites. J Dent Res 2003;81:921-35.
67. Anusavice KJ, Shen C, Rawls RH. Phillips’ science of dental materials. 12th ed. St Louis: Elsevier; 2013.
68. Eliades T. Dental materials in orthodontics. In: Graber LW, Vanarsdall RL Jr., Vig KWL, Huang GJ. editors. Orthodontics: current principles and techniques. 5th ed. Philadelphia: Elsevier; 2012. p. 187-200.
69. Ireland AJ, Moreno T, Price R. Airborne particles produced during enamel cleanup after removal of orthodontic appliances. Am J Orthod Dentofacial Orthop 2003;124:683-6.
70. Van Landuyt KL, Yoshihara K, Geebelen B, Peumans M, Godderis L, Hoet PH. The effect of orthodontic adhesive particulates produced by simulated debonding of dental composite resins. J Dent Res 1982;61:791-5.
71. Cokic SM, Hoet P, Godderis L, Wiemann M, et al. Nanoparticle release from dental composites. Acta Biomater 2014;10:365-74.
72. Cokic SM, Hoet P, Godderis L, Wiemann M, Asbach C, Reichl FX, et al. Cytotoxic effects of composite dust on human bronchial epithelial cells. Dent Mater 2016;32:1482-91.
73. Cokic SM, Duca RC, Godderis L, Hoet PH, Seo JW, Van Meerbeek B, et al. Release of monomers from composite dust. J Dent 2017;60:56-62.
74. Cokic SM, Asbach C, De Munck J, Van Meerbeek B, Hoet P, Seo JW, et al. The effect of water spray on the release of composite dust. Clin Oral Investig 2019: Epub 2019 Dec 6.
75. Johnston NJ, Price R, Day CJ, Sandy JR, Ireland AJ. Quantitative and qualitative analysis of particulate production during simulated clinical orthodontic debonds. Dent Mater 2009;25:1155-62.
76. Hext PM, Rogers KO, Paddle GM. The health effects of PM2.5 (including ultrafine particles). Brussels: CONCAWE; 1999.
77. Möller W, Häussinger K, Winkler-Heil R, Stahlhofen W, Meyer T, Hofmann W, et al. Mucociliary and long-term particle clearance in the airways of healthy nonsmoker subjects. J Appl Physiol 1985;59:475.
78. Oberdörster G. Pulmonary effects of inhaled ultrafine particles. Brussels: CONCAWE; 1999.
79. Borm PJA, Robbins D, Haubold S, Kuhlbusch T, Fissan H, Horß S, Panzer J, et al. The strong-yet-gentle grip, inspired by mussels and geckos. Nature 2007;448:338-41.
80. Napier D, Thomassen LCI, Lison D, Martens JA, Hoet PH. The nanosilica hazard: another variable entity. Part Fibre Toxicol 2006;3:111.
81. Klæsens CD, Casarett and Doulil’s toxicology: the basic science of poisons. 7th ed. NY: McGraw-Hill; 2008.
82. Schmalz G, Hickel R, van Landuyt KL, Reichl FX. Scientific update on nanoparticles in dentistry. Int Dent J 2018;68:299-305.
83. Vankerckhoven H, Lambrechts P, van Beylen M, Davidson CL, Vanheule G. Unreacted methylacrylate groups on the surfaces of composite resins. J Dent Res 1982;61:791-5.
84. Góka I, Eliades T, Zinelis S, Pratsinis H, Athanasiou AE, Eliades G, et al. Characterization and in vitro estrogenicity of orthodontic adhesive particulates produced by simulated debonding. Dent Mater 2009;25:376-82.
85. Bradna P, Ondrácková L, Zdímal V, Navrátil T, Pecelova D. Detection of nanoparticles released at finishing of dental composite materials. Monatsh Chem 2017;148:531-7.
86. Dasy H, Dasy A, Asatryan G, Rózsa N, Lee HF, Kwak JH. Effects of variable attachment shapes and aligner material on aligner retention. Angle Orthod 2015;85:934-40.
87. Kechagia A, Zinelis S, Pandis N, Athanasiou AE, Eliades T. The effect of orthodontic adhesive and bracket-base design in adhesive remnant index on enamel. J World Fed Orthod 2015;4:18-22.
88. Eliades T, Voutsa D, Sifakakis I, Makou M, Katsaro C. Release of bisphenol-A from a light-cured adhesive bonded to lingual fixed retainers. Am J Orthod Dentofacial Orthop 2011;139:192-5.
89. Kloukos D, Sifakakis I, Voutsa D, Doulis I, Eliades G, Katsaro C, et al. BPA qualitative and quantitative assessment associated with orthodontic bonding in vivo. Dent Mater 2015;31:887-94.
90. Eliades T, Hiskia A, Eliades G, Athanasiou AE. Assessment of bisphenol-A release from orthodontic adhesives. Am J Orthod Dentofacial Orthop 2007;131:72-5.
91. Bettencourt AF, Neves CB, de Almeida MS, Pinheiro LM, Oliveira SA, Lopes LP, et al. Biodegradation of acrylic based resins: a review. Dent Mater 2010;26:e171-80.
92. Atabek D, Aydinmiit I, Alacaşan A, Berkkan A. The effect of temperature on bisphenol: an elution from dental resins. J Contemp Dent Pract 2014;15:576-80.
93. Pandis N, Polychronopoulos A, Eliades T. Failure rate of self-ligating and edgewise brackets bonded with conventional acid etching and a self-etching primer: a prospective in vivo study. Angle Orthod 2006;76:119-22.
94. Fleming PS, Johal A, Pandis N. Self-etch primers and conventional acid-etch technique for orthodontic bonding: a systematic review and meta-analysis. Am J Orthod Dentofacial Orthop 2012;142:83-94.
95. Iliaidi A, Baumgartner S, Athanasiou AE, Eliades T, Eliades G. Effect of intraoral aging on the setting status of resin composite and glass ionomer orthodontic adhesives. Am J Orthod Dentofacial Orthop 2014;145:425-33.
96. Eliades T. Future of bonding. In: Eliades T, Brantley WA, editors. Orthodontic applications of biomaterials. A clinical guide. Cambridge: Woodhead Publishing; 2017. p. 267-71.
97. McComb D. Luting in orthodontic practice. In: Davidson C, Mjör IA, editors. Advances in glass-ionomer cements. Carol Stream, Quintessence Publishing; 1999. p. 149-70.
98. Eliades T. Orthodontic materials research and applications: part 1. Current status and projected future developments in bonding and adhesives. Am J Orthod Dentofacial Orthop 2006;130:445-51.
99. Berengueres J, Saito S, Tadakuma K. Structural properties of a scaled gecko foot-hair. Bioinspir Biomim 2007;2:1-8.
100. National Institute of General Medical Sciences. Gecko foot hairs inspired the design of medical adhesives. Available at: https://www.nigms.nih.gov/education/life-magnIfied/Pages/9_kunkegeckofoot hairs.aspx. Accessed May, 24, 2020.
101. Lee H, Lee BP, Messersmith PB. A reversible wet/dry adhesive agent and method of foaming the same. Available at:https://www.google.com/patent/US3615972A/en. Accessed May 25, 2020.
102. Banea MD, da Silva LFM, Carbas RJC. Debonding on command of adhesive joints for the automotive industry. Int J Adhes Adhes 2020;59:14-20.
103. Banea M, da Silva F, Campbell D. An overview of the technologies for adhesive debonding on command. The Annals of “Dunarea de Jos” University of Galati 2013;24:11-4.
104. Morehouse DS Jr, Mich M, Tetreault RJ, Mass S. Expandable thermoplastic polymer particles containing volatile fluid foaming agent and method of foaming the same. Available at: https://patents.google.com/patent/US8856972A/en. Accessed May 25, 2020.
105. Iliaidi A, Eliades T, Silikas N, Eliades G. Development and testing of novel bisphenol A-free adhesives for lingual fixed retainer bonding. Eur J Orthod 2017;39:1-8.