

Hydrophobicity of Denture Base Resins: A Systematic Review and Meta-analysis

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ABSTRACT

Objectives: The aim of this article is to review the factors that attract *Candida albicans* to denture base resin (DBR) and to verify the influence of different surface treatments, chemical modification, or structural reinforcements on the properties of DBR. **Materials and Methods:** Searches were carried out in PubMed, Scopus, WOS, Google Scholar, EMBASE, and J-stage databases. The search included articles between 1999 and 2020. This study followed the Preferred Reporting Items for Systematic Reviews and Meta-Analyses statement. The keywords used during the search were “*Candida albicans*,” “Denture base,” “PMMA,” “Acrylic resin,” “Surface properties,” “hydrophobicity/hydrophilicity,” “contact angle,” and “surface free energy.” English full-text articles involving *in-vitro* studies with different acrylic resin modifications were included, whereas abstracts, dissertations, reviews, and articles in languages other than English were excluded. A meta-analysis was performed where appropriate. **Results:** Out of the 287 articles, 21 articles conformed to inclusion criteria. Sixteen articles were subjected to meta-analysis using random-effects model at 95% confidence interval. Results showed that DBR coatings/plasma coatings were effective methods to modify surface properties with estimated contact angle (CA) of 59.37° [95% confidence interval (CI): 53.69, 65.04]/55.87° (95% CI: 50.68, 61.06) and surface roughness (R_a) of 0.55 μm (95% CI: 0.52, 0.58)/0.549 μm (95% CI: 0.5, 0.59), respectively. Antifungal particle incorporation into poly(methylmethacrylate) DBR also produced similar effects with an estimated R_a of 0.16 μm (95% CI: 0.134, 0.187). **Conclusion:** The three properties responsible for *C. albicans* adhesion to DBR were R_a , CA, and surface free energy in terms of hydrophobicity. Therefore, the correlations between the hydrophobicity of DBR and *C. albicans* adhesion should be considered during future investigations for *Candida*-related denture stomatitis.

KEYWORDS: *Candidiasis*, *PMMA denture base*, *surface properties*

INTRODUCTION

Candida-associated denture stomatitis (DS) infection depends mainly on the denture base (DB) properties and the ability of *Candida albicans* (*C. albicans*) (the most common pathogen in DS) to adhere to the denture surface.^[1,2] Reports have confirmed that *Candida* adhesion to acrylic is associated with hydrophobic interactions between the two.^[2,3] Because

C. albicans are hydrophobic, they can easily adhere to the hydrophobic poly(methylmethacrylate) (PMMA) DB.^[4] Therefore, hindering this interaction may help prevent various infections including DS. Achieving

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this would be of extreme benefit for elderly patients with dentures and their caretakers.^[2] The most relevant factors of a DB that influence microbial attachment are surface roughness (R_a), hydrophobicity/hydrophilicity, and surface free energy (SFE), in addition to salivary pellicles and the presence of other microorganisms.^[5,6]

Higher microbial adhesion is linked to R_a and hydrophobicity of the DB material,^[7] in which roughness is capable of providing more surface area and protective hideout spot for microorganisms away from denture cleaning forces.^[7] To limit the microbial colonization, R_a of DB should not exceed 0.2 μm .^[7,8]

The chemical composition of PMMA which includes carboxylate, methyl ester groups, as well as other additives, cross-linking agents, fillers, and colorants affects the hydrophobicity and SFE of the DB.^[9] Studies have reported that SFE and wettability of different denture base resins (DBRs) are related to variations in these additives.^[10] In recent years, several nanoparticles such as ZrO_2 , SiO_2 , TiO_2 , and diamond nanoparticles have been incorporated within the PMMA in an attempt to enhance the physio-mechanical properties of the material. These fillers were also found to increase the resistance of the material to microbial adhesion.^[10,11]

Researchers have used surface coating, chemical modifications, or synthesized and incorporated fillers with antimicrobial properties within PMMA to solve the issue of *Candida* adhesion. However, reviews of the effect of these treatment modifications on PMMA properties with correlation to hydrophobicity are not yet available. The aims of this study were to (1) systematically review literature pertaining to the modifications of DBR and (2) to correlate the variables to *Candida* adhesion/biofilm formation. The null hypothesis of this study was that alteration of the DBR in the form of filler addition, chemical composition modification, or surface coating will not affect the hydrophobicity of the resin surface and therefore will not affect *Candida* adhesion.

MATERIALS AND METHODS

SEARCH STRATEGY

This systematic review was completed according to the Preferred Reporting Items for Systematic Reviews and Meta-analyses (PRISMA). Focus question was generated through the PICO(S) approach and research strategy [Table 1] to systematically review the available literature. Two PICO questions were formulated as follows: first, do the modifications of DB alter the hydrophobicity and *Candida* adhesion thereafter? Secondly, what factors will influence the hydrophobicity of modified DBR? An electronic search of English-language dental literature on PubMed, Scopus, WOS, Google Scholar, EMBASE, and J-stage databases was conducted for articles published between January 1999 and March 2020 [Figure 1]. To identify all relevant articles, a list of keywords was used for the search. These included “Denture base,” “PMMA,” “Surface properties,” “hydrophobicity/hydrophilicity,” “contact angle,” “surface energy,” and “*C. albicans*.” The inclusion criteria included full-text articles in the English language, with *in-vitro* design, investigating heat-polymerized DBR, *C. albicans* adhesion, contact angle (CA), surface wettability, R_a , and/or SFE with different DB modifications (antimicrobial additives, surface coating, chemical composition modification). In contrast, papers in languages other than English, *in-vivo* clinical study, case reports, abstracts, short communication, letters to the editors, reviews, and dissertations and materials other than heat-polymerized acrylic resin or resin not used for DBs were excluded.

ELIGIBILITY CRITERIA

Two investigators (MMAG and RA) reviewed the articles independently according to the same parameters. Studies that (1) measured the effect of incorporated antifungal agents, surface coating, or chemical composition modifications of heat-polymerized PMMA, (2) evaluated the *C. albicans* adhesion and one of the following properties: CA, SFE, R_a , or hydrophobicity/hydrophilicity, (3) reported

Table 1: Systematic search strategy

Focus questions	What are the influencing factors of the hydrophobicity of modified denture base resin?
PICOS	
P: Participant	Modified denture base materials
I: Interventions	Incorporating antifungal agents Surface coatings Chemical composition modification
C: Comparison	Unmodified heat-polymerized/microwave-polymerized acrylic resin Modified heat-polymerized acrylic resin
O: Outcomes	Effectiveness of modifications on surface free energy/hydrophobicity
S: Study design	Networking meta-analysis

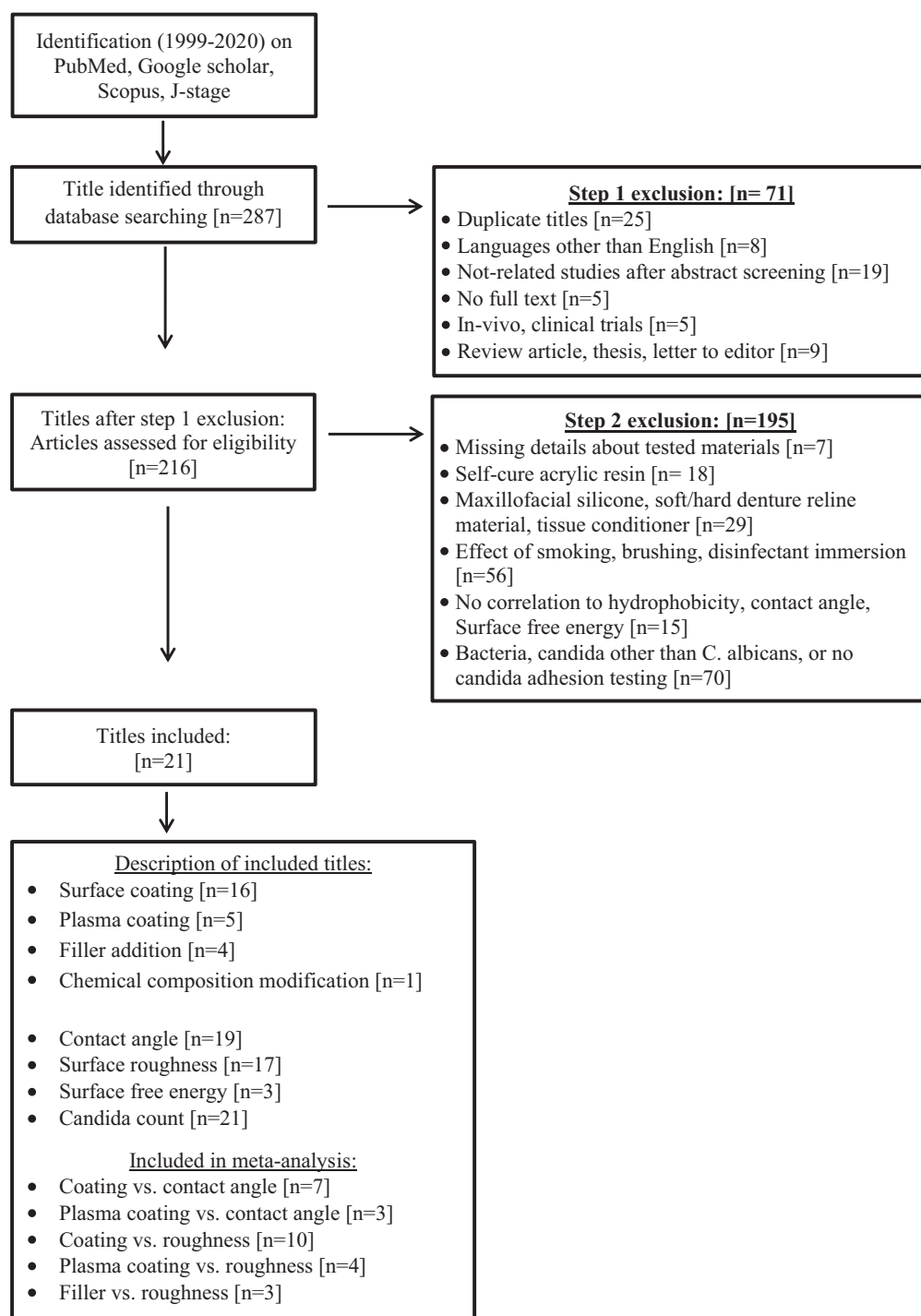


Figure 1: Flowchart of the study design

sample size, mean, and standard deviation values, and (4) included brand names and specifications of tested materials were included in this review. DB modifications were categorized as follows: control (unmodified PMMA), antifungal additive, surface coatings, and chemical composition modifications.

DATA MANAGEMENT, SCREENING, AND SELECTION

Two independent investigators (MMG and RA) used a standardized Excel sheet to extract the data of the

studies. The search was conducted in three steps. First, the titles were reviewed according to the inclusion/exclusion criteria. Secondly, the abstracts of the selected titles were screened to select those of interest for full-text analysis. At the third step, all full-text articles were analyzed. At all stages, any discrepancies between investigators were resolved by discussion. The extracted data included: the authors' names, year of publication, materials of the study, processing method, *Candida* species, tests employed, presence of control group,

number and dimensions of specimens, type of resin modification, results, statistical analysis and significance, and conclusions. Studies with similar methodology were selected to undergo meta-analysis. Among the scopes of this systematic review is to conduct a meta-analysis taking into consideration the diverse designs (resin modifications) of the studies and the various properties tested and to assess their effect qualitatively (surface properties) and quantitatively (number of *Candida* colony-forming units) [Tables 2 and 3].

ASSESSMENT OF RISK OF BIAS

A modification of the method used in previous systematic reviews was used by two authors (MMAG and RA) to independently assess the quality and risk of bias of each study.^[12-14] The characteristics were tabulated ($n=21$) and the parameters were reported as “+ve” if the parameter was described in the text or “-ve” if the information was missing or unclear. The parameters assessed were: sample size calculation, the use of a control group, stating the treatment method, statistical analysis performed, reliable analytical methods, blinding of the evaluators, and correlation of the reported properties with hydrophobicity. The risk of bias was classified according to the sum of “+ve” marks obtained as follows: 1 to 3= high-, 4 to 5= medium-, 6 to 7= low-risk of bias.^[15]

Meta-analysis was performed for each treatment modality separately. Moreover, due to the variability of outcomes and methodology per treatment method, quantitative meta-analysis was done for 16 studies, whereas the rest of the studies were descriptively analyzed.

DATA ANALYSIS

Comprehensive meta-analysis (version 3, NJ, USA) was used for analysis. Visual inspection of forest plots and χ^2 tests were used to evaluate the presence of heterogeneity. Random-effects model was used when the data were found to be heterogenic, whereas the fixed-effects model was used otherwise. Egger's and Begg's tests were used to check for the possibility of publication bias. P -values less than 0.05 were considered statistically significant.

RESULTS

DATA SELECTION

Twenty-one studies met the inclusion criteria [Figure 1] and submitted for data extraction and result analysis. Tables 2 and 3 summarize the studies' details, methods, results, and outcomes.

RISK OF BIAS

Figure 2 presents the risk of bias for the included studies. Out of the 21 studies, 19 showed medium risk of bias and two showed low risk of bias. The risk of

bias was mainly linked to the absence of sample size calculation and non-blinding of investigators.

Applying the inclusion criteria, out of the 21 included articles,^[16-36] 16 used surface coating, 4 added antimicrobial fillers, and 1 modified the chemical composition of PMMA (refer to Tables 2 and 3 for details). In addition to that, several included studies compared between smooth and rough surfaces of the modified specimens.^[17,30] Results revealed that hydrophobicity of DBRs was affected by surface coating, antimicrobial additives, or chemical composition modifications. Therefore, the results of this study were categorized based on the effects of these modifications on the hydrophobicity of DBR and its correlations with CA, R_a , and *C. albicans* adhesion.

META-ANALYSIS

In coating vs. CA (Supplementary Appendix 1), after exclusion of outliers, 74 groups underwent meta-analysis. Due to the considerable heterogeneity found ($I^2 >75\%$, $P < 0.001$), random-effects model was used and the average CA after coating was found to be 59.37° [95% confidence interval (CI): 53.7–65.0]. The trim and fill method suggested inclusion of 33 more groups to remove publication bias after getting significant results of Begg's and Eggers' tests ($P=0.002$ and $P=0.001$, respectively).

In the plasma coating vs. CA (Supplementary Appendix 2) and coating vs. R_a (Supplementary Appendix 3), a total of 38 and 91 observations were, respectively, included in the analysis. Due to significant heterogeneity ($I^2 >70\%$, $P < 0.001$) in both the groups, random-effects model was used. The average CA and R_a were found to be 55.87° (95% CI: 50.68–61.06) and $0.552 \mu\text{m}$ (95% CI: 0.524–0.58), respectively. In plasma coating vs. CA, Begg's and Eggers' tests provided insignificant results; hence, the trim and fill method was not used. However, in coating vs. R_a , the trim and fill method provided insertions of 32 more observations to avoid publication bias.

In plasma coating vs. R_a (Supplementary Appendix 4) and filler vs. R_a (Supplementary Appendix 5), 27 and 13 observations were included in the analysis. Both data sets reflected the presence of heterogeneity ($I^2 >70\%$, $P < 0.001$) and hence the random-effects model was used for both. The estimated average R_a for plasma coating and filler addition were $0.549 \mu\text{m}$ (95% CI: 0.504–0.593) and $0.161 \mu\text{m}$ (95% CI: 0.134–0.187), respectively. Significant P -values for Eggers' and Begg's tests proved the presence of publication bias for both data sets. Hence, the trim and fill method suggested to insert 12 and 7 observations, respectively, to remove the publication bias.

Table 2: Included studies

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
Yildirim et al., 2005 ^[6]	Denture acrylic, Meliodent (Bayer Dental, Newbury Berkshire, UK)	*Heat-cured *102 discs (17×1 mm) *n=60 for wettability *n=30 for <i>Candida</i> adhesion *n=12 for surface analysis	*One surface polished, the other was not (ground with 500 grid sandpaper) *Plasma surface treatment for 15 min at the O ₂ level of 0, 50, or 100 W. n=34 *Saliva contact	*Contact angle * <i>Candida</i> adhesion	*O ₂ surface modification sig. improved wettability (lowered contact angle) compared with control *The reduction in contact angle is directly related to plasma power *Saliva reduced contact angle of control, and increased it for plasma-treated	*O ₂ gas is effective in increasing wettability of PMMA even with salivary pellicle. * <i>Candida</i> adherence increased as hydrophilicity increased
Nevzatoglu et al., 2007 ^[17]	ACRON Shade No. 3, GC	*Heat-cured	*Polishing up to 1000 grit *Buff polished	* <i>Candida</i> adherence	* <i>Candida</i> adhesion to control was less than surface treated → increased surface wettability, increased <i>Candida</i> adhesion *Increase in O/C ratio → more hydrophilic * <i>Candida</i> count was lowest in the coated specimens < buff polished < control	*Straight silicone coating is capable of improving surface properties of denture base material so that it becomes difficult for <i>C. albicans</i> to adhere
Zamperini et al., 2010 ^[8]	Vipi Wave; VIPI	*Dises (20×1 mm) *Control (uncoated) *Microwave-cured	*20% straight silicon coating for 5, 30 min *Processing technique (against glass or against stone)	*Contact angle *Surface roughness	*Contact angle of the coated specimens was larger than control or buff polished *Surface roughness of specimens processed against glass was lower. No difference between all groups regarding surface roughness in each investigation.	*Adherence of <i>C. albicans</i> was sig. reduced by ArO ₂ /70 W and ArSF6/70 W plasma, regardless of the presence or absence of saliva and surface roughness (smooth or rough).

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
	Industria e Comercio Exportacao e Importacao de Produtos Odontologicos Ltda Pirassununga, SP, Brazil <i>C. albicans</i> (ATCC 90028)	*Control *180 discs (13.8×2 mm) in 10 groups *n=18	*Plasma treatment for 5 min:	*Contact angle	*Groups 2 and 4 were not sig. different from each other and showed sig. lower absorbance reading.	*Hydrophobicity was altered by the plasma treatments and water immersion
		*Processed against glass or stone	1: argon atmosphere at 50 W 2: argon/oxygen atmosphere at 70 W 3: atmospheric air at 130 W 4: argon atmosphere, followed by sulfur hexafluoride atmosphere at 70 W *Saliva exposure (30 min unstimulated whole human saliva)	* <i>Candida</i> adhesion	*Contact angle was altered by plasma tx and water immersion for all groups except controls. *For control, contact angle was sig. different b/w rough and smooth. *XPS showed incorporation of fluorine into the surface of group 4	*No sig. effect of surface roughness and saliva on adherence of <i>C. albicans</i>
Zamperini et al., 2010 ^[9]	Vipi Wave; VIPI	*Microwave-cured	*Processing technique (against glass or against stone)	*Surface roughness	*Surface roughness of specimens processed against glass was lower.	*Contact angle was altered by the plasma treatments. However, mean contact angles of treated specimens were similar to those of control specimens, after 48 h of immersion in water.
	Industria e Comercio Exportacao e Importacao de	*Control *180 discs (13.8×2 mm) in five groups	*Plasma treatment for 5 min: 1: argon atmosphere at 50 W	*Contact angle	*Contact angle for all groups changed after water immersion except control. All test groups showed an increase in contact angle after water immersion except group 4 which showed a reduction.	*Adherence of <i>C. albicans</i> was not sig. reduced by plasma treatments, surface roughness, or presence of saliva
	Produtos Odontologicos Ltda Pirassununga, SP, Brazil	*n=18 *Processed against glass or stone	2: argon/oxygen atmosphere at 70 W 3: atmospheric air at 130 W	* <i>Candida</i> adhesion	*No sig. difference between all groups regarding <i>Candida</i> adhesion irrespective of ± saliva, surface roughness, treatment	

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
Wady et al., 2012 ^[20]	<i>C. albicans</i> (ATCC 90028) Vipi Wave; VIPI	*Microwave-cured	4: argon atmosphere, followed by sulfur hexafluoride atmosphere at 70 W *Saliva exposure *AgNPs solution mixed with 75g acrylic powder at concentrations of (1000, 750, 500, 250, 30, 0 ppm), dried, sieved, ball milled	*Surface roughness	*No sig. difference in contact angle between 0 and 7 days or 90- and 180-day storage periods.	*AgNPs had no effect on <i>C. albicans</i> adherence and biofilm formation regardless of concentrations
Lazarin et al., 2013 ^[21]	Industria e Comercio Exportacao e Importacao de Produtos Odontologicos Ltda Pirassununga, SP, Brazil <i>C. albicans</i> (ATCC 90028) Vipi Wave; VIPI	*Control *72 discs (13.8×2 mm) *n=18 *Microwave-cured	*Different storage periods (0, 7, 90, 180 days) (n=18) *Processed against glass (smooth) or against stone (rough)	*Contact angle *Adherence biofilm formation *Surface roughness	*After 90 and 180 days, contact angles were sig. higher than that at 0 and 7 days *Contact angles were lower than control for all experimental groups *No sig. difference b/w 0-7, 90-180 days regarding <i>Candida</i> adhesion and biofilm *Significant absorbance value noted for 90 and 180 days *Sig. increase in surface roughness for all rough specimens.	*Experimental S and HP coatings showed sig. reduction of short-term attachment (90 min) of <i>C. albicans</i> to PMMA
	Industria e Comercio Exportacao e Importacao de Produtos Odontologicos Ltda Pirassununga, SP, Brazil	*468 discs (13.8×2 mm) in 13 groups	*Photopolymerized coatings: 1. 2-hydroxyethyl methacrylate (HE) (HEMA) (cured for 4 min)	*Surface free energy through contact angle measurement	*Total surface free energy was generally higher in all experimental groups compared with controls	

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
	<i>C. albicans</i> (ATCC 90028)	*n=36 *Control	2. hydroxypropyl methacrylate (HP) (HPMA) (cured for 4 min) 3. 2-tri-methyl-ammonium ethyl methacrylate chloride (T) (TMAEMC) (cured for 4 min) 4. sulfobetaine methacrylate (S) (oven at 80°C for 2 h) *Concentrations of coatings at 0%, 25%, 30%, and 35% of the total composition in mmol. Additional components in the coating: MMA, TEGDMA, bis-GMA, 4-methyl benzophenone. Also, amino propyl methacrylate for group 4 *± saliva (non-stimulated) for 30 min at room temp.	* <i>Candida</i> adhesion	*Generally, no sig. difference of surface free energy b/w saliva-coated and uncoated specimens *For smooth specimens, no sig. difference b/w all groups *For rough surfaces, S30, S35, and HP30 had sig. lower absorbance values than control	
Queiroz et al., 2013 ^[27]	Lucitone 550; Dentsply Ind. Com. Ltda, Petropolis, Brazil	*Heat-cured *45 discs (10×5mm) in three groups	*Polishing of both sides to 1200 grit silicon carbide paper	*Surface roughness (optical, non-contact)	*RBS confirms the presence of carbon in groups 2 and 3 and silver in group 3 *DLC thin films significantly diminished <i>C. albicans</i> biofilm formation	

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
Al-Bakri et al., 2014 ^[23]	<i>C. albicans</i> (ATCC 18804)	*n=15 *Control (no surface treatment)	*Surface treatment for 15 min: 1. no coating (Gc) 2. surface coating with DLC fil (Gdlc) 3. surface coating with DLC doped with Ag-Nps (Gag) DLC = diamond-like carbon	*Rutherford backscattering (RBS) and atomic force microscopy (AFM) for film characterization *Anti-microbial activity assessment after 24 h at 37°C b CFU count	*Surface roughness did not affect the number of <i>Candida</i> adhered *Surface treatment reduced <i>Candida</i> adhesion in groups 2 and 3 compared with control *No additional reducing effect was seen with Ag addition *DLC increased hydrophobicity and lowered surface energy	*The films undoped and doped with silver nanoparticles presented similar behavior.
		*Heat-cured *50 discs (10×1.5 mm) in 5 groups	*Silane-coated glass fibers (1.5 µm, with 15% w/w fluoride) were added to PMMA at concentrations of 0.5%, 1.0%, 2.5%, 5.0%, 10% *Polishing of both sides with 400 grit Al ₂ O ₃	*Contact angle	*No sig. difference between all groups regarding contact angle and surface free energy → fluoride did not have an effect	*Increased loading of the fillers produced increased surface roughness
Lazarin et al., 2014 ^[24]	<i>C. albicans</i> (GDH 2346)	*n=10 *Control	*Processed against glass (smooth) or against stone (rough)	*Surface free energy (contact angle cosine value) *Surface roughness (non-contact) *Adherent <i>Candida</i> count using a light microscope *Surface roughness	*10% filler produced sig. rougher surface than control and 1.0% *Fluoride addition sig. reduced <i>Candida</i> adhesion to PMMA *Coating PMMA with saliva sig. reduced <i>Candida</i> adhesion	*No direct correlation between surface roughness and microbial adhesion *Presence of saliva and fluoride glass fillers significantly reduced <i>Candida</i> adhesion
		*Microwave-cured *468 discs (13.8×2 mm) in 13 groups			*No sig. differences in surface roughness among groups within each fabrication method	*Experimental photopolymerized coatings did not alter hydrophobicity but changed chemical composition.

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
Industria e Comercio Exportacao e Importacao de Produtos Odontologicos Ltda Pirassumunga, SP, Brazil <i>C. albicans</i> (ATCC 90028)	*n=36 *Control	*Photopolymerized coatings:	*contact angle	*Samples prepared against stone were sig. rougher than those prepared against glass	* <i>C. albicans</i> adhesion decreased with coatings sulfo betaine, 2-hydroxypropyl methacrylate, and 2-hydroxyethyl methacrylate	
	1. 2-hydroxyethyl methacrylate (HE) (HEMA) (cured for 4 min) 2. hydroxypropyl methacrylate (HP) (HPMA) (cured for 4 min) 3. 2-tri-methyl-ammonium ethyl methacrylate chloride (T) (TMAEMC) (cured for 4 min) 4. sulfo betaine methacrylate (S) (oven at 80°C for 2 h)	* <i>Candida</i> adhesion	*Smooth groups HE30, T25, T30, and T35 had sig. higher contact angle *Contact angles for rough surface were not sig. different *No sig. different in <i>Candida</i> adhesion for saliva-coated specimens *Smooth and non-saliva-coated specimens showed sig. lower <i>Candida</i> with S35, HP35, and HE35 *Rough specimens ± saliva → no sig. difference between groups regarding <i>Candida</i> adhesion *Rough S25, S30, HP35, HE30, HE35, T35 with no saliva showed higher <i>Candida</i> adhesion than same smooth groups *XPS showed increase in C, O, Si after HE, HP, and T coating, and S for S coating			
	*Concentrations of coatings 0%, 25%, 30%, and 35% of the total composition in mmol. Additional components: MMA, TEGDMA, Bis-GMA, 4-methyl benzophenone. Also, amino propyl methacrylate for group 4 *± saliva (non-stimulated) for 30 min at room temp.					

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
Yodmongkol et al., 2014 ^[25]	Rodex (Australia)	*Heat-cured	*Silane-SiO ₂ nanocomposite dip-coating evaporating solvent at 65°C for 20 min and then heating to 110°C for 2 h	* <i>Candida</i> adhesion after 1 h using optical microscope (<i>n</i> =6)	*Sig. higher cell adhesion was seen on uncoated specimens than coated	*Silane-SiO ₂ nanocomposite films can make acrylic resin more hydrophobic, which decreases <i>C. albicans</i> adhesion.
Sawada et al., 2014 ^[26]	<i>C. albicans</i> (ATCC 10231)	*Rectangular specimens (1.5×1.5×1 mm)	*Addition of 5 wt.%; —FAP-TiO ₂ (100 nm) —HAP-TiO ₂ (100 nm) —TiO ₂ (25 nm)	*FTIR (<i>n</i> =3)	*FTIR for coated showed a peak for Si-O-Si	*This film improved surface and physical properties of acrylic
		*Control		*Surface roughness (contact) (<i>n</i> =5)	*Surface roughness was the same for coated and uncoated	
		*Roughness (<i>n</i> =5) *Contact angle and SFE (<i>n</i> =3)		*Contact angle of three liquids were used: deionized water 18 MΩ/cm, diiodomethane, and glycerol (<i>n</i> =3) → SFE *SEM	*Surface energy was slightly reduced on coated (not sig.) *Average thickness of coating was 6.8 ± 1.0 μm	
Compagnoni et al., 2014 ^[27]	Lucitone 550	*Heat-cured	*Polishing up to 2000 grit polishing paper *Modification with PTBAEMA (0% or 10%)	*Viable cells determination after incubation for 2 h at 37°C and UVA irradiation	* <i>Candida</i> binding was sig. lower in the FAP-TiO ₂ - and TiO ₂ -containing discs than in the HAP-TiO ₂ -containing discs	*Influence of surface characteristics on the adhesion of <i>C. albicans</i> to various denture lining mats
		* <i>n</i> =12		*Contact angle measurement using 1.0 μL deionized water drop	* <i>Candida</i> binding was sig. decreased in all test groups compared with controls	
		*Control (pure PMMA)		*Surface roughness increased with PTBAEMA addition		*PTBAEMA slightly increases wettability and roughness of acrylic resin

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
Pan et al., 2015 ^[28]	Dentsply International Inc., York, PA, USA	*Control (unmodified)	PTBAEMA = (2-tert-butylaminoethyl)	*Atomic force microscopy observations of 100 and 400 μm^2	*Contact angles of PTBAEMA-modified acrylic is lower than controls	*PTBAEMA into acrylic resins did not have an effect against <i>C. albicans</i> at 10%
	<i>C. albicans</i> (ATCC 90028)	*Disks (15×3mm) in two groups	Methacrylate	*Adherence assay using CFU counts after 90 min at 37°C	*Contact angle decreased as roughness increased	
Pan et al., 2015 ^[28]	Vertex Rapid Simplified, Vertex-Dental, Zeist, The Netherlands	*Heat-cured	*Polishing to 600 or to 2000 grit silicon carbide	*Contact angle of 2 μL ultrapure water drop ($n=18$)	*No sig. difference b/w controls and PTBAEMA-modified acrylic regarding <i>Candida</i> count.	
	<i>C. albicans</i> (ATCC 10231)	*36 discs (12×1 mm) in two groups	* \pm saliva (non-stimulated whole saliva)	*Fungal adherence test after 90 min at 37°C ($n=18$) using gradient dilution method.	*Contact angle sig. reduced after plasma treatment \rightarrow more hydrophilic	*Ar/O ₂ plasma treatment improved surface wettability of PMMA without degrading physical properties
Qian et al., 2016 ^[29]	Vertex Rapid Simplified, Vertex-Dental	* $n=18$	*Cold plasma treated or not (98% argon, 2% oxygen, at atmospheric pressure). Discs were treated for 90 s, and rectangular discs were treated for 8.5 min	*Surface roughness (contact)	*Sig. reduction in early <i>Candida</i> adhesion for plasma treated	*Ar/O ₂ plasma treatment sig. reduced early <i>C. albicans</i> adhesion
		*Rectangular (64×10×3.3 mm)		*SEM	*No sig. difference in surface roughness	
		*Control		*X-ray photoelectron spectroscopy analysis (XPS)	*XPS revealed fluorine on the surface of plasma treated and reduction of C/O	
		*Heat-cured	*Polished to silicon carbide grit 1000	*Optical emission spectroscopy (OES)	*OES revealed abundance of O and OH as active components	
				*Contact angle (after plasma TX, 48 h, 15 days, 30 days)	*Contact angle decreased after plasma treatment	*Cold plasma treatment resulted in increased hydrophilicity and reduced <i>Candida</i> adhesion

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
	BV, Zeist, The Netherlands	*45 discs (12×1 mm) in five groups	*Plasma surface treatment with argon 98%/oxygen 2% for 0, 30, 60, 90, and 120 s	*Surface roughness non-contact (<i>n</i> =9)	*No difference between plasma-treated groups (immediately)	*Prolonged plasma treatment did not improve wettability but affected durability.
	<i>C. albicans</i> (ATCC 10231)	*Control * <i>n</i> =9		* <i>Candida</i> adhesion by CFU analysis (<i>n</i> =9)	*Contact angle increased after water immersion, 48 h, 15 days	*Reduction in the ratio of C/O, direct relation with treatment time
				*XPS	*No difference b/w all groups after 30 days	*No relation between surface roughness and <i>Candida</i> adhesion
					*Surface roughness not significantly different b/w all groups	
					*Lower <i>Candida</i> adhesion for all test groups (90S lowest)	
Liu et al., 2017 ^[30]	Lucitone 199; Dentsply Intl Inc. <i>C. albicans</i> (ATCC 18804)	*Heat-cured *60 discs (10×2 mm) in four groups * <i>n</i> =15 *Control	*Smooth and rough surfaces *Coated with TMS or not coated (trimethylsilane)	*Contact angle of sessile drop of distilled water *Absorbance of OD (optical density) for <i>Candida</i> *MTT assay	*Contact angle of coated specimens was higher *Absorbance intensity of coated specimens is less than that of controls	*Hydrophobicity increased by TMS coating * <i>Candida</i> adhesion was decreased by TMS coating
Türkcan et al., 2018 ^[31]	Meliocent	*Heat-cured	*Polished with silicon carbide paper 600 grit	*SEM and EDS *Contact angle of 2 µL sessile drop of pure water (<i>n</i> =3)	*Significant decrease in contact angle for 0.25 and 0.75 mol/L MPC → increased wettability	*Surface roughness alone did not affect <i>Candida</i> adhesion *Surface modification with MPC coating decreased contact angle in 0.25 and 0.75 mol/L MPC groups.
	Heat Cure, Heraeus Kulzer, Germany	*Disc (6×1.5 mm) in four groups	*Surface coating with MPC (2-methacryloyloxyethyl phosphorylcholine) dissolved in degassed pure water at concentrations of 0.25, 0.5, 0.75 mol/L	*Surface roughness (contact) (<i>n</i> =3)	*MPC coating increased surface roughness, no difference between groups	*Graft polymerization of MPC does not cause a significant change in surface roughness.
	<i>C. albicans</i> (ATCC 90028)	*Contact angle and roughness (<i>n</i> =3)		*FTIR spectroscopy with attenuated total reflection (ATR) equipment (<i>n</i> =2)	*MCP increased hydrophilicity (increased water absorption)	*Graft polymerization of MPC decreased <i>C. albicans</i> adhesion onto PMMA surface.

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
Hirasawa et al., 2018 ^[32]	Natural resin, Nissin Co., Kyoto, Japan <i>C. albicans</i> (JCM2085)	* <i>Candida</i> adhesion (n=10) *Control *Heat-cured *250 discs (12×2 mm)	*Polished on both sides to 8000 grit *In laboratory-made co-polymer coating plasma cleaning → primer → drying → immersion (10 s) in prepared polymer at concentrations SM0%, SM15%, SM30%, and SM50% → UV (27s)	* <i>Candida</i> adhesion assay using CFU (n=10) *SEM (n=2) *XTT reduction assay (n=10) *CFU (n=10)	*Reduction in <i>Candida</i> adhesion as concentration increased, no difference between 0.5 and 0.75 mol/mL *Significant difference among all groups for XTT and CFU *Significant reduction in biofilm for all test groups compared with controls	*Coating with cross-linkable co-polymers containing SBMAm significantly reduced the initial adhesion of <i>C. albicans</i>
Darwish et al., 2019 ^[33]	Lucitone 199 (Dentsply Intl, York, PA, USA)	*Heat-cured *Rectangular specimens (20×20×1 mm)	*Polished to 4000 grit silicon carbide paper *TiO ₂ coating at 65°C for 3 h to form 30 nm film	*SEM (n=10) *Surface roughness (non-contact) *Film thickness (spectroscopic ellipsometer) *Contact angle of 1 mL purified water drop *Surface roughness (non-contact) (n=10) *Contact angle using sessile drop of 5 µL deionized water (n=10)	*Surface roughness was less than 0.005 µm for all groups (no difference) *Film thickness was less than 5 µm for all groups—thicker for SM30% than SM0% and SM50% *All coated groups had lower contact angle than control, SM15% had lowest contact angle and highest hydrophilicity *Surface roughness of coated specimens was less than that of non-coated *Contact angle of coated was lower than that of non-coated	*Titanium oxide coating improved wettability, surface smoothness, and increased resistance to microbial adherence.

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
Acosta et al., 2019 ^[34]	Lucitone 199 (Dentsply Sirona) and ProBase Hot (Ivoclar Vivadent AG)	*Roughness and contact angle (n=10)	Acrylic acid or itaconic acid coatings	* <i>Candida</i> adhesion after 12 h at 37°C	*Sig. reduction in viable attached <i>Candida</i> cells to coated surfaces	PMMA acrylic resin base material was superficially modified through the incorporation of carboxylic acid groups by using PAA and PIA coatings that reduced the adherence of <i>C. albicans</i> biofilm by 90%.
		* <i>Candida</i> adhesion (n=5)		*Biofilm formation (n=5)	*Sig. reduction in viable <i>Candida</i> biofilm on coated surfaces	
		*Control		*Heat-cured	Affected surface roughness	
Fouda et al., 2019 ^[35]	Major.Base.20 Resin	*Disks (13–14×4–5 mm)	Nano-diamond at 0.5%, 1.0%, and 1.5%	*Contact angle using sessile drop of 5 µL deionized water (n=30) <i>Candida</i> biofilm adhesion	Increased surface wettability	PMMA disks modified with PIA or PAA showed a 90% reduction of <i>C. albicans</i>
		*n=30		*Surface roughness (non-contact) (n=30)	Decreased surface roughness at 1% NDs and 0.5% NDs	
		*Control		*Heat-cured	Decreased <i>C. albicans</i> adhesion	
	<i>C. albicans</i>	*Square (10×10×3 mm)		*Contact angle using sessile drop of 5 µL deionized water (n=30) <i>C. albicans</i> adhesion	No significant effect was observed on the contact angle.	PMMA/ND composites could be valuable in the prevention of denture stomatitis, which is considered one of the most common clinical problems among removable denture wearers.
	(ATCC 10231)	*n=30				
		*Control				

Table 2: Continued

Author, year	Acrylic brand, composition, <i>Candida</i> species	Processing method/sample dimensions	Modification	Tested properties	Results	Conclusions
AlBin-Ameer et al., 2020 ^[36]	Major: Base 20, Major Prodotti Dentari SPA, Moncalieri, Italy <i>C. albicans</i> (ATCC 10231)	*Heat-cured *Rectangular specimens (12×10×2.5 mm) *n=14	Nanocoat Optiglaze Nano-silica Cyanoacrylate	*Surface roughness (non-contact) *Contact angle using sessile drop of 5 µL deionized water * <i>C. albicans</i> adhesion	Nano-coat, Optiglaze, Nano-silica decrease R_a whereas cyanoacrylate increased Nano-coat, Optiglaze, Nano-silica decrease contact angle, whereas cyanoacrylate increased Nano-coat, Optiglaze, Nano-silica decrease <i>C. albicans</i> adhesion, whereas cyanoacrylate increased	Coating of removable prosthesis with nano-coat, Optiglaze, or nanosilica is an effective method to reduce <i>C. albicans</i> adhesion

CA = contact angle, SFE = surface free energy, SR = surface roughness, H = hydrophobicity, GS = Google scholar, S = Scopus

Concerning these factors and their direct and indirect relations to *C. albicans* adhesion, almost all treatment modalities decreased *C. albicans* adhesion to modified DB in comparison to unmodified DB.

DISCUSSION

The results of this review revealed that the different treatment modalities (filler incorporation, surface coating, and chemical composition modification) affected the R_a , CA, and hydrophobicity of the DBR resulting in *Candida* adhesion modification, and therefore, the null hypothesis was rejected.

WHY COATING? AND WHAT IS THE OUTCOME?

In recent years, the DB surface has been modified with various coatings in an attempt to increase its hydrophilicity and to reduce *C. albicans* adhesion.^[3,21] These coatings can be in the form of plasma-based treatment, photopolymerized coatings, and hydrophilic polymer coatings, among others. In plasma-based treatment, partial ionization of the gas is brought up by electrical discharge which creates an environment that contains reactive species such as electrons, ions, and free radicals. Plasma treatment helps clean debris, generates reactive groups on the surface, and makes the surface more attractive to specific cells depending on the treatment atmosphere.^[16] The newly formed surface has higher SFE, improved wettability, and diminished CA, which reduces the adherence of *C. albicans*.^[18,19,28,29]

Plasma treatment of PMMA in the presence of O₂ gas improved the wettability of the surface even in the presence of salivary pellicle.^[16] Similarly, plasma coating in argon, argon-oxygen, and atmospheric air resulted in lower CAs.^[18] Conversely, TMS coating increased the hydrophobicity, lowered the wettability of the DB surface, and significantly reduced *C. albicans* adhesion.^[30] Silane-SiO₂ nanocomposite films were found to improve the surface, augment the physical properties of PMMA, and increase surface hydrophobicity which decreases *C. albicans* adhesion.^[25]

Coating with TiO₂ created smoother surfaces that are more resistant to wear and less porous, which prevent microorganisms from diffusing into the acrylic resin and colonizing on the surface.^[33] UV irradiation of TiO₂ activates oxidative species that produce irreversible damage to the cells.^[33] Additionally, TiO₂ coating creates a super-hydrophilic surface with “water sheathing” effect. The ability of TiO₂ to improve surface wettability is essential to reduce or inhibit *Candida* attachment on DBR.^[33]

Surface modification with photopolymerized coating of poly(acrylic acid) (PAA) or poly(itaconic acid)

Table 3: Type of tests applied in the included studies

No.	Year	Author	Tests undertaken in the study				
			Contact angle	Surface free energy	<i>Candida albicans</i>	<i>Candida</i> adherence	Surface roughness
1	2005	Yildirim <i>et al.</i>	×		ATCC 10321	×	
2	2007	Tokita <i>et al.</i>	×		JCM 1542	×	
3	2010	Zamperini <i>et al.</i>	×		ATCC 90028	×	×
4	2010	Zamperini <i>et al.</i>	×		ATCC 90028	×	×
5	2012	Wady <i>et al.</i>	×		ATCC 90028	×	×
6	2013	Lazarin <i>et al.</i>	×	×	ATCC 90028	×	×
7	2013	Queiroz <i>et al.</i>			ATCC 18804	×	×
8	2014	Al-Bakri <i>et al.</i>	×	×	GDH 2346	×	×
9	2014	Lazarin <i>et al.</i>	×		ATCC 90028	×	×
10	2014	Yodmongkol <i>et al.</i>	×	×	ATCC 10231	×	×
11	2014	Sawada <i>et al.</i>			ATCC 1002	×	×
12	2014	Compagnoni <i>et al.</i>	×		ATCC 90028	×	
13	2015	Pan <i>et al.</i>	×		ATCC 10231	×	×
14	2016	Qian <i>et al.</i>	×		ATCC 10231	×	×
15	2017	Liu <i>et al.</i>	×		ATCC 18804	×	
16	2018	Turkan <i>et al.</i>	×		ATCC 90028	×	×
17	2018	Hirasawa <i>et al.</i>	×		JMC 2085	×	×
18	2019	Darwish <i>et al.</i>	×		ATCC 90028	×	×
19	2019	Acosta <i>et al.</i>	×		ATCC 90028	×	×
20	2019	Fouda <i>et al.</i>	×		ATCC 10231	×	×
21	2020	AlBin-Ameer <i>et al.</i>	×		ATCC 10231	×	×

(PIA) followed by UV irradiation has been achieved. The coatings decreased the CA and increased the SFE, which may have resulted from changes in the surface polar groups after coating^[21] and the acidic environment in the presence of (-OH) groups.^[34] In a similar manner, surface modification by polymerization of 2-methacryloyloxyethyl phosphorylcholine (MPC) polymer provided a statistical decrease in CA and *C. albicans* adhesion.^[31]

Other hydrophilic coatings like 3-hydroxypropyl methacrylate (HPMA) and polymers containing sulfobetaine methacrylate (SBMA) were found to enhance the wettability of the DB surface and to reduce *C. albicans* adhesion as a result of limited hydrophobic interactions.^[2,21,22] The CA and *C. albicans* adhesion were significantly reduced after coating the DBR with nanocoat or Optiglaze. This effect was brought up by changes in the carbon and oxygen content and different types of interactions.^[36] Conversely, cyanoacrylate coating increased the *C. albicans* adhesion with no effect on CA.^[36]

The main advantage of coating is that it allows surface alteration at a relatively low cost and preserves the properties of the original material.^[18,19] The low viscosity can produce thin films (~50 nm^[22] and <5 µm^[32]) on the surface that do not interfere with the fit of the denture.^[22,36] Coating PMMA with

ceramic materials improves its resistance to abrasion^[33] and protects the surface from attacks of different solutions.^[22] Cold plasma treatment is performed at room temperature avoiding possible damage or warpage of acrylic resin with thermal treatments.^[30] As most of the aforementioned coatings produce hydrophilic surfaces and improve the wettability of the PMMA, their application on the fitting surface of a denture could enhance the retention of the DBs by increasing affinity to saliva/liquid molecules that would create a denture seal.^[28] In contrast, some of the coating materials require a certain preservation temperature and consumption within a short duration after preparation.^[25] Also, the durability of different coatings needs further investigation.

ROUGHNESS (R_a), HYDROPHOBICITY, RESIN SURFACE CHEMISTRY, AND *candida* ADHESION

High R_a may enhance microbial retention because a rougher surface provides more area for microbial adhesion and promotes fungal adhesion and colonization.^[7,8,37-39] Hahnel *et al.*^[40] did not find a linear relationship between R_a and *C. albicans* adhesion. However, many other studies reported that greater *C. albicans* adhesion is associated with higher R_a .^[7,38,39] Studies indicated that R_a was not altered following plasma treatment or film deposition process.^[29] Thus, these opposing results suggest that the

reduced *C. albicans* biofilm was due to the chemical modification of the PMMA surface represented by increased hydrophilicity and SFE that was promoted by film coating.^[22] Hirasawa *et al.*^[32] reported that roughness of different coated specimens was not the main determining factor in *Candida* reduction, rather it was surface hydrophilicity that played the major role.

REINFORCEMENT/R_a/CA

Incorporation of antifungal agents within DBR affected *C. albicans* adhesion and the development of DS.^[41] The antimicrobial efficiency of the added AgNPs is associated with ingress of water molecules into the material and the outward movement of the silver ions to the aqueous solution.^[20] Others suggested that the

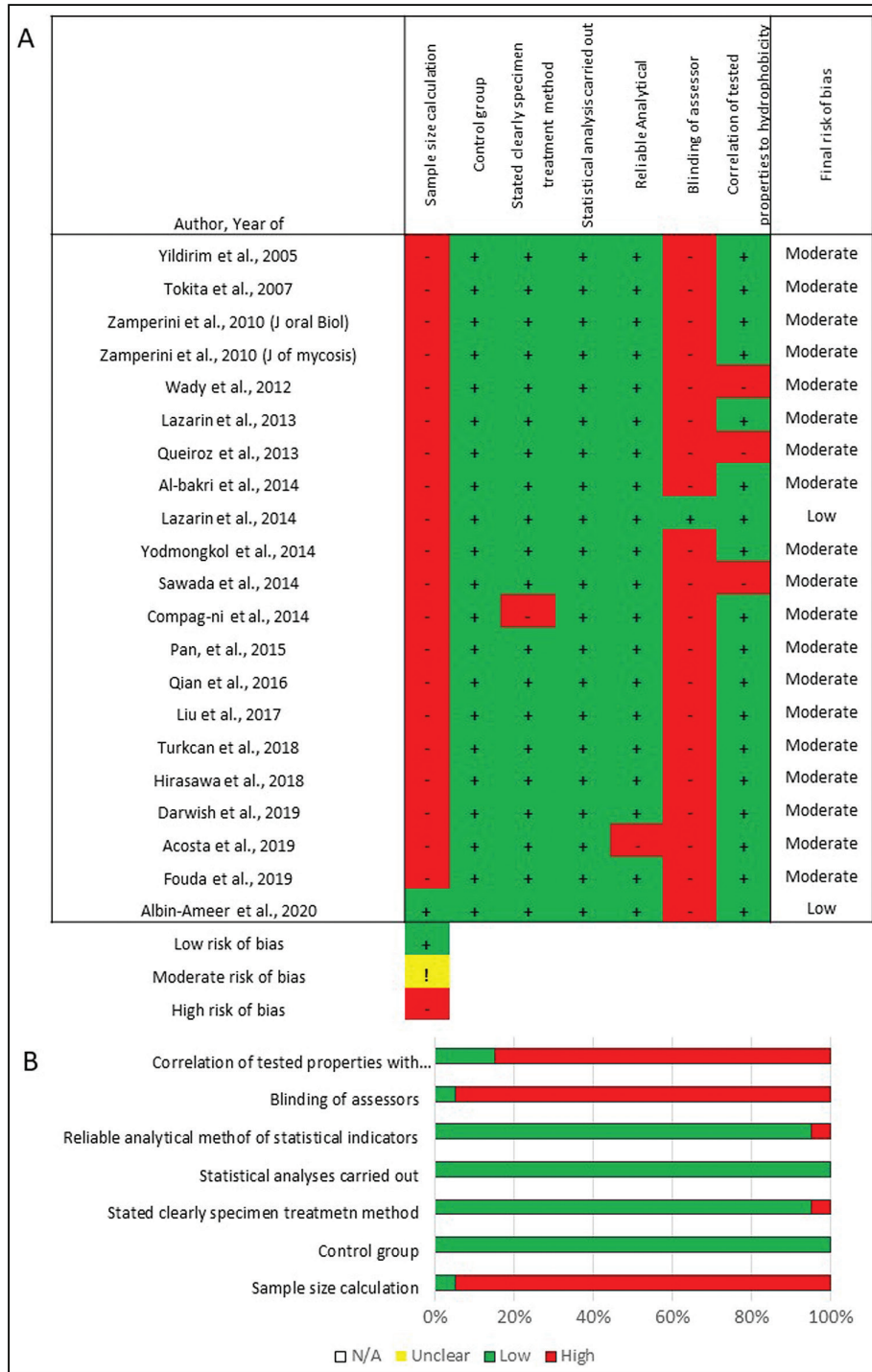


Figure 2: Risk of bias for the included studies

inhibitory effect was due to the greater antimicrobial effect of the smaller particles which provides more surface area in direct contact with the nanoparticles.^[20] PMMA containing FAp-TiO₂ exhibited strong photocatalytic activity following irradiation through the production of reactive oxygen species such as (-OH) and (H₂O₂) which inhibit *C. albicans* attachment.^[26] This filler has clinical advantages especially for elderly patients through maintenance of proper denture hygiene.^[26]

The addition of nano-diamonds showed an improvement of the specimen surface, which may contribute to the significant reduction in *C. albicans* adhesion. Regardless of the increase in R_a at a high concentration, a reduction in *Candida* adhesion was detected. Moreover; the inclusion of nano-diamonds within PMMA did not alter the CAs of the modified specimens in comparison to the unmodified specimen.^[35] However, the mechanism of antifungal activity of ND was not described clearly and requires further investigations.

CHEMICAL COMPOSITION MODIFICATION

The addition of phosphate into DBR by monomer substitution was reported to improve the surface hydrophilicity.^[28,42] The quantity of adherent *C. albicans* was associated with the wettability properties of the DB, emphasizing the role of acrylic resin chemistry on the initial attachment of *C. albicans*.^[16]

CLINICAL SIGNIFICANCE

The literature reported that hydrophobicity and R_a of DBs influence the attachment and colonization of *C. albicans*. Therefore, to reduce *Candida* adhesion, the surface of the DB must be smooth, hydrophilic, and has no porosities.^[5] Improving the hydrophilicity of the DB allows contact with more liquid molecules which helps in forming the seal that keeps the denture tight to air leakage.^[28] Additionally, it has been reported that hydrophilic surfaces have fewer adherent *C. albicans*.^[2] Therefore, increasing the surface hydrophilicity would hinder *Candida* attachment.^[36]

Additionally, the intaglio surface provides the best environment for *C. albicans* adhesion, as it cannot be finished or polished to preserve its accuracy and fit. Therefore, surface coatings can be of great use in such situations in which the coating films are extremely thin and less likely to induce any misfit between the DB and oral tissues, affect the occlusion, or affect the texture of the resin.^[24,25,43] The different coating modalities mentioned earlier can reduce *C. albicans* adhesion and biofilm formation.^[25]

The limitations of this review could be attributed to a wide range of different treatments in each section, such as different coating materials, fillers, and minimal studies on chemical modification, which made the comparison more difficult as a result of the wide range of properties of each material and its effect on the studied properties.

CONCLUSION

Based on this review, it could be concluded that the hydrophobicity of DBRs and *C. albicans* adhesion were affected by the interrelated following factors: wettability (CA), SFE, and surface structure of DBR. Incorporation of antifungal agents or surface coating of DBR affected its hydrophobicity. Future studies evaluating the long-term biocompatibility and antifungal efficacy of different modifications are required to correlate between factors affecting the hydrophobicity and *C. albicans* adhesion.

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CONFLICTS OF INTEREST

There are no conflicts of interest.

AUTHORS' CONTRIBUTIONS

Not applicable.

ETHICAL POLICY AND INSTITUTIONAL REVIEW BOARD STATEMENT

Not applicable.

PATIENT DECLARATION OF CONSENT

Not applicable.

DATA AVAILABILITY STATEMENT

Not applicable.

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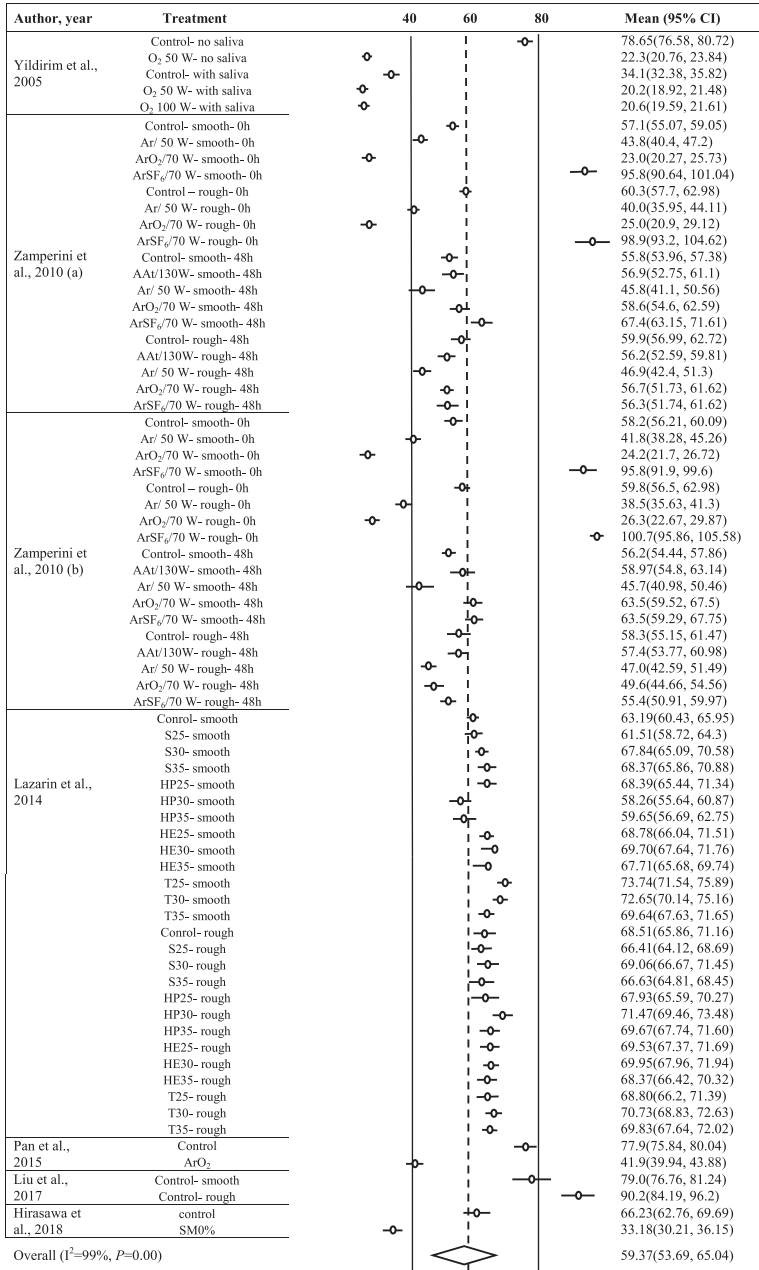
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SUPPLEMENTARY MATERIAL

SUPPLEMENTARY APPENDIX 1: FOREST PLOT FOR COATING VS CONTACT ANGLE (°)



0h; immediate reading after plasma treatment

48h; reading after 48 hours in water

Ar/50W; Argon atmosphere at 50 W

ArO₂/70W; Argon/oxygen atmosphere at 70 W

AAT/130W; Atmosphere air at 130 W

Ar/SF₆/70W; Argon atmosphere then plasma treatment in Sulphur hexafluoride atmosphere at 70W

S; zwitterionic monomer (sulfbetaine methacrylate)

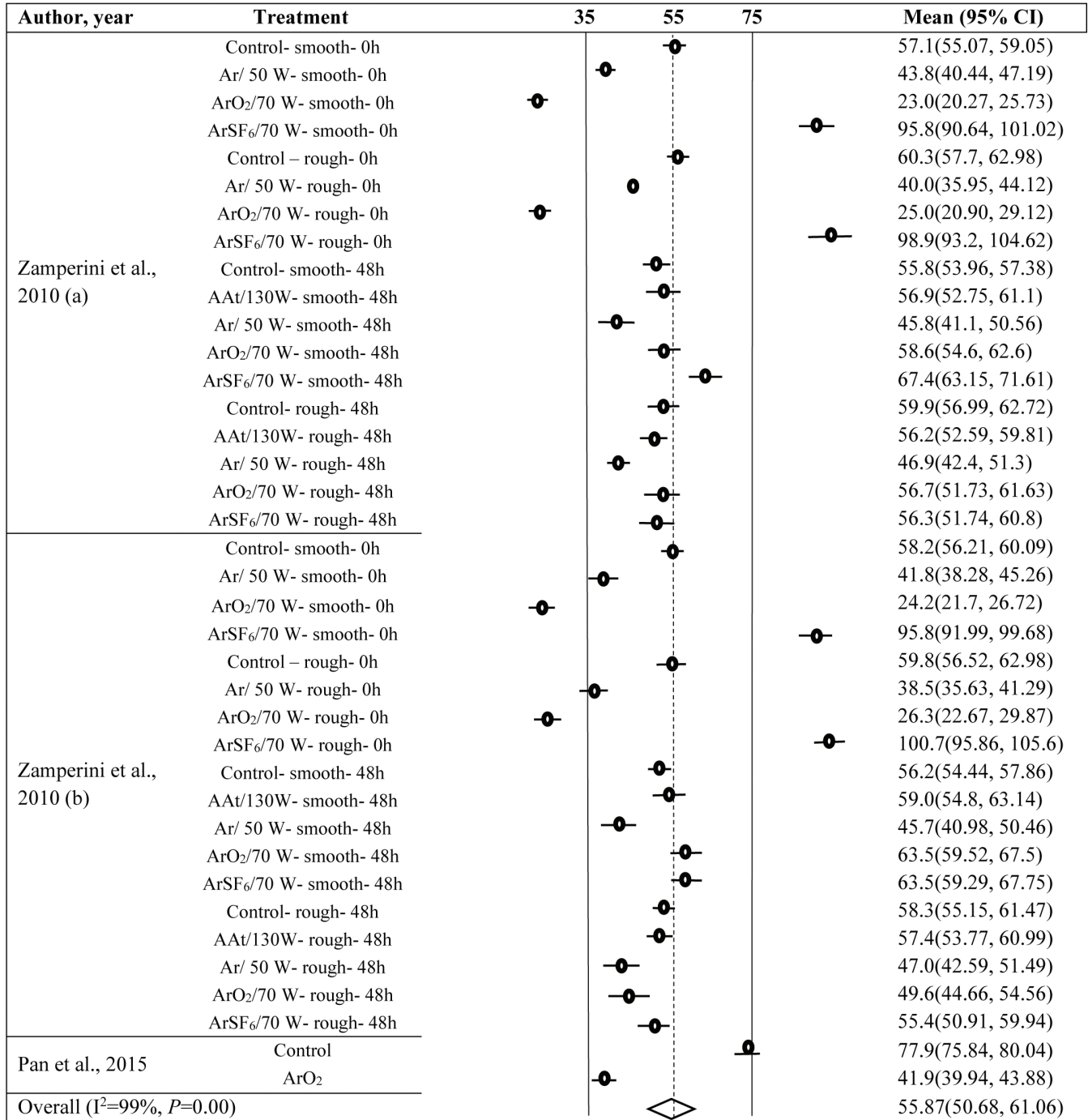
HP; 2-hydroxypropyl- methacrylate (HPMA)

HE; 2-hydroxyethyl methacrylate (HEMA)

T; 2-trimethylammonium ethyl methacrylate chloride (TMAEMC)

(SM0%); sulfbetaine methacrylamide:N,N'-(4,7,10-trioxa-1,13-tridecanediamine) diacrylamide in ratio 0:100

SUPPLEMENTARY APPENDIX 2: FOREST PLOT FOR PLASMA SURFACE TREATMENT VS CONTACT ANGLE (°)



0h; immediate reading after plasma treatment

48h; reading after 48 hours in water

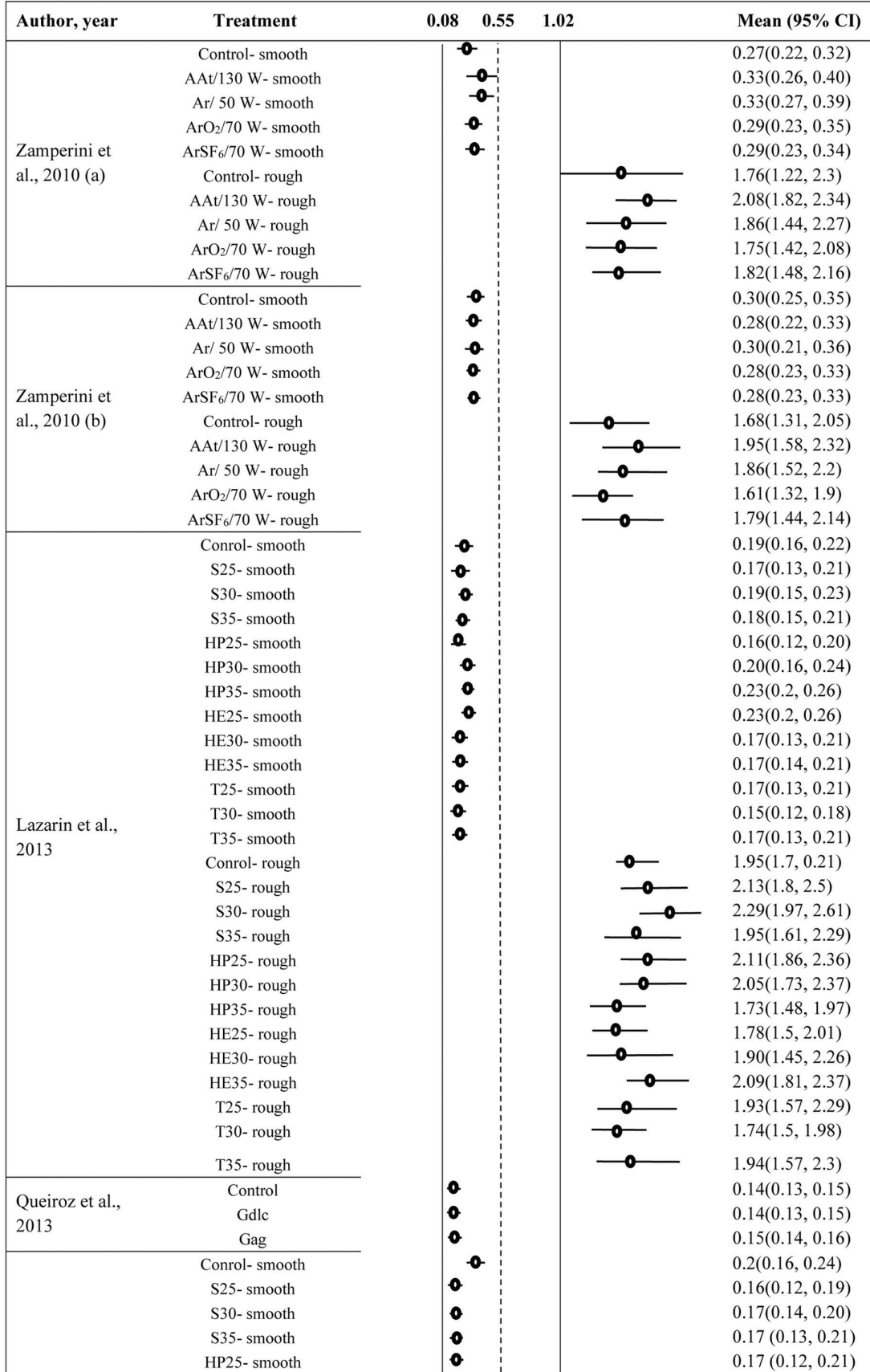
(Ar/50W)- Argon atmosphere at 50 W

(ArO₂/70W)- Argon/oxygen atmosphere at 70 W

(AAT/130W)- Atmosphere air at 130 W

As/SF₆/70W)- Argon atmosphere then plasma treatment in Sulphur hexafluoride atmosphere at 70W

SUPPLEMENTARY APPENDIX 3: FOREST PLOT FOR SURFACE COATING VS. SURFACE ROUGHNESS (MM)



SUPPLEMENTARY APPENDIX 3: CONTINUED

	HP30- smooth		0.17 (0.13, 0.21)
	HP35- smooth		0.21(0.18, 0.24)
	HE25- smooth		0.18(0.14, 0.22)
	HE30- smooth		0.16(0.12, 0.19)
	HE35- smooth		0.18(0.14, 0.22)
	T25- smooth		0.20(0.16, 0.24)
	T30- smooth		0.17(0.13, 0.21)
Lazarin et al., 2014	T35- smooth		0.16(0.13, 0.19)
	Control- rough		2.17(1.8, 2.4)
	S25- rough		2.17(1.8, 2.4)
	S30- rough		1.96(1.68, 2.2)
	S35- rough		1.94(1.64, 2.25)
	HP25- rough		1.85(1.58, 2.1)
	HP30- rough		1.68(1.43, 1.9)
	HP35- rough		1.82(1.5, 2.1)
	HE25- rough		1.71(1.5, 1.9)
	HE30- rough		1.74(1.6, 1.93)
	HE35- rough		1.76(1.4, 2.1)
	T25- rough		1.85(1.66, 2.03)
	T30- rough		1.65(1.46, 1.84)
	T35- rough		1.81(1.64, 1.98)
Yodmongkol et al., 2014	Control		0.62(0.45, 0.79)
	Silane-SiO ₂		0.54(0.43, 0.65)
Pan et al., 2015	Control		0.07(0.05, 0.09)
	Ar/O ₂		0.07(0.05, 0.09)
Qian et al., 2016	Control		0.22(0.21, 0.23)
	Cold plasma/ Atmos./ 30 S		0.23(0.21, 0.24)
	Cold plasma/ Atmos./ 60 S		0.23(0.22, 0.24)
	Cold plasma/ Atmos./ 90 S		0.21(0.19, 0.23)
	Cold plasma/ Atmos./ 120 S		0.22(0.21, 0.23)
Darwish et al., 2019	Control		0.12(0.11, 0.13)
	TiO ₂		0.08(0.06, 0.099)
Albin-Ameer et al., 2020	Control		0.21(0.17, 0.25)
	Nano-coat		0.15(0.12, 0.18)
	Optiglaze		0.12(0.08, 0.16)
	Nano-silica		0.19(0.10, 0.28)
	Cyanoacrylate		0.29(0.23, 0.35)
Overall (I ² =99%, P=0.00)			0.55(0.52, 0.58)

Ar/50W; Argon atmosphere at 50 W

ArO₂/70W; Argon/oxygen atmosphere at 70 W

AAT/130W; Atmosphere air at 130 W

Ar/SF₆/70W; Argon atmosphere then plasma treatment in Sulphur hexafluoride atmosphere at 70W

S; zwitterionic monomer (sulfobetaine methacrylate)

HP; 2-hydroxypropyl- methacrylate (HPMA)

HE; 2-hydroxyethyl methacrylate (HEMA)

T; 2-trimethylammonium ethyl methacrylate

Gdlc; diamond-like carbon

Gag; diamond-like carbon doped with silver nanoparticles

Silane-SiO₂; silane silicaAr/O₂; Argon/Oxygen

Atmos.; Atmospheric pressure

S; seconds

TiO₂; Titanium dioxide

SUPPLEMENTARY APPENDIX 4: FOREST PLOT FOR PLASMA TREATMENT VS ROUGHNESS (MM)

Author, year	Treatment	0	0.55	1.1	Mean (95% CI)
Zamperini et al., 2010 (a)	Control- smooth	●			0.27(0.23, 0.31)
	AAAt/130 W- smooth	●			0.33(0.28, 0.38)
	Ar/ 50 W- smooth	●			0.33(0.29, 0.37)
	ArO ₂ /70 W- smooth	●			0.29(0.25, 0.33)
	ArSF ₆ /70 W- smooth	●			0.29(0.25, 0.33)
	Control- rough			—●—	1.76(1.38, 2.1)
	AAAt/130 W- rough			—●—	2.08(1.89, 2.26)
	Ar/ 50 W- rough			—●—	1.86(1.57, 2.15)
	ArO ₂ /70 W- rough			—●—	1.75(1.5, 2.0)
	ArSF ₆ /70 W- rough			—●—	1.82(1.6, 2.06)
Zamperini et al., 2010 (b)	Control- smooth	●			0.30(0.27, 0.33)
	AAAt/130 W- smooth	●			0.28(0.24, 0.32)
	Ar/ 50 W- smooth	●			0.30(0.26, 0.34)
	ArO ₂ /70 W- smooth	●			0.28(0.24, 0.32)
	ArSF ₆ /70 W- smooth	●			0.28(0.24, 0.32)
	Control- rough			—●—	1.68(1.4, 1.9)
	AAAt/130 W- rough			—●—	1.95(1.69, 2.21)
	Ar/ 50 W- rough			—●—	1.86(1.62, 2.1)
	ArO ₂ /70 W- rough			—●—	1.61(1.4, 1.8)
	ArSF ₆ /70 W- rough			—●—	1.79(1.54, 2.0)
Pan et al., 2015	Control	●			0.07(0.05, 0.09)
	Ar/O ₂	●			0.07(0.05, 0.09)
Qian et al., 2016	Control	●			0.22(0.21, 0.23)
	Cold plasma/ Atmos./ 30 S	●			0.23(0.22, 0.24)
	Cold plasma/ Atmos./ 60 S	●			0.23(0.22, 0.24)
	Cold plasma/ Atmos./ 90 S			—●—	0.21(0.19, 0.23)
	Cold plasma/ Atmos./ 120 S	●			0.22(0.21, 0.23)
Overall (I ² =99%, P=0.00)		◇			0.549(0.5, 0.59)

(Ar/50W)- Argon atmosphere at 50 W

(ArO₂/70W)- Argon/oxygen atmosphere at 70 W

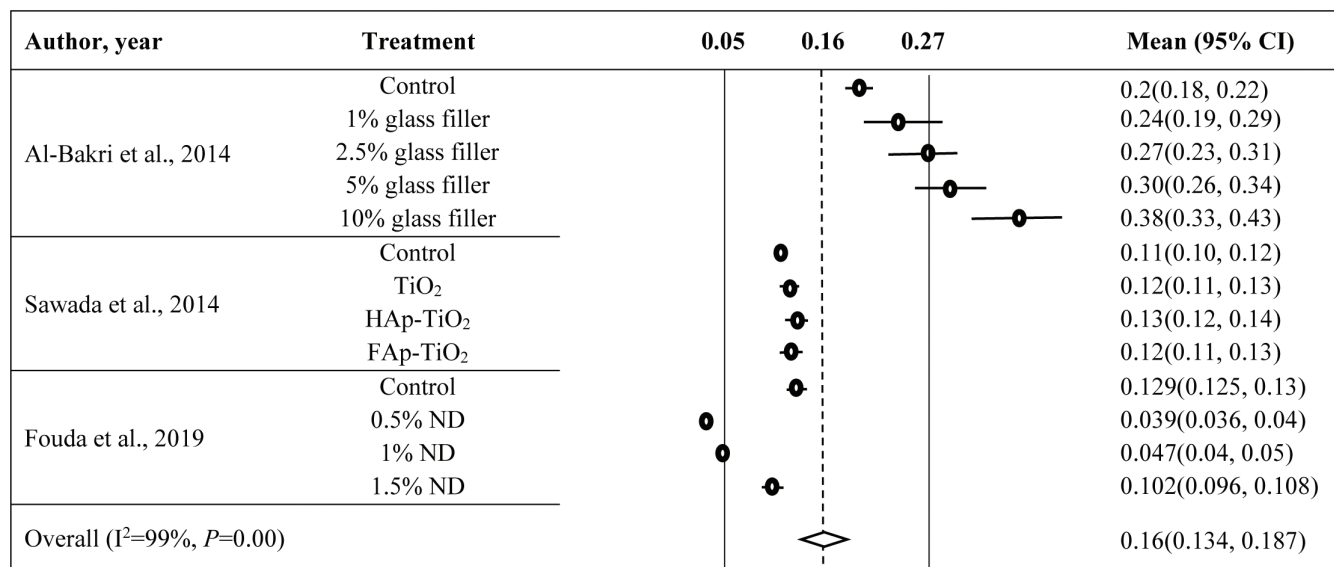
(AAT/130W)- Atmosphere air at 130 W

(As/SF₆/70W)- Argon atmosphere then plasma treatment in Sulphur hexafluoride atmosphere at 70W

Atmos.- Atmospheric pressure

S- seconds

SUPPLEMENTARY APPENDIX 5: FOREST PLOT FOR FILLERS VS ROUGHNESS (MM)



Glass filler: silane coated glass filler with 15% w/w fluoride

TiO₂: Titanium dioxide

HAp-TiO₂: Hydroxyapatite-coated titanium dioxide

Fap- TiO₂: Fluoroapatite-coated titanium dioxide

ND: Nano-diamond