

## Evidence Summary: The relationship between oral health and dementia

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## **In brief**

- The evidence with regard to an association between oral health and dementia is weak. This is due to the lack of well-designed cohort and case-control studies and variation in how dementia and oral health are defined and measured. Increasing evidence, albeit weak, of an *association* between dementia and oral health is sufficient to suggest that this is a very important area for future research.
- Dementia and cognitive decline are risk factors for poor oral health. Those in later stages of the disease tend to have more plaque accumulation, gingivitis, attachment loss, dental caries, poor denture hygiene and denture stomatitis.
- Patients with suboptimal oral health (gingivitis, caries, partial tooth loss) appear to have an associated increased risk of cognitive impairment, but more evidence from different settings is required.
- Securing and maintaining oral health remains important for patients with dementia and those at risk of cognitive impairment.

## **Abstract**

This is the fourth and final paper of a suite of reviews undertaken to explore the relationships between oral health and general medical conditions, in order to support teams within Public Health England, health practitioners and policy makers. This review aimed to explore the most contemporary evidence on whether poor oral health and dementia occurs in the same individuals or populations, to outline the nature of the relationship between these two health outcomes and to discuss the implication of any findings for health services and future research. The review was undertaken by a working group comprising consultant clinicians from medicine and dentistry, trainees, public health and academic staff. Whilst other rapid reviews in the current series limited their search to systematic reviews, this review focused on primary research involving cohort and case-control studies because of the lack of high level evidence in this new and important field. The results suggest that poor oral hygiene is associated with dementia and more so in those in advanced stages. Suboptimal oral health (gingivitis, dental caries, tooth loss, edentulousness) appears to be associated with increased risk of developing cognitive impairment and dementia. The findings are discussed in relation to health and future research.

## Background

Increasing dementia incidence and prevalence in England has led to a commitment to provide the best support for people with dementia and other neurodegenerative diseases, and further research (1). This is timely as dementia, together with cognitive decline, present significant challenges for patients and their carers, practitioners and health systems (2).

Dementia (F00-F03) is described in the International Classification of Disease version 10 (ICD -10) as

*“a syndrome due to disease of the brain, usually of a chronic or progressive nature, in which there is disturbance of multiple higher cortical functions, including memory, thinking, orientation, comprehension, calculation, learning capacity, language, and judgement. Consciousness is not clouded. The impairments of cognitive function are commonly accompanied, and occasionally preceded, by deterioration in emotional control, social behaviour, or motivation. This syndrome occurs in Alzheimer's disease, in cerebrovascular disease, and in other conditions primarily or secondarily affecting the brain.” (3)*

In this paper, we shall use the umbrella term ‘dementia’ to include cognitive impairment, as the majority of the literature used cognitive impairment scores as a diagnostic tool for dementia. It is important to note that dementia is not a specific disease, rather it is a set of symptoms that are the manifestations of different pathology. In this review we shall use the term dementia, though we acknowledge there is more than one type of dementia (3-5).

The common causes of dementia are Alzheimer's disease (AD) and vascular dementia (4, 5). AD is a physical disease where proteins build up to form structures called 'plaques' and 'tangles' in the brain. This leads to the loss of connections between nerve cells, and eventually to the death of nerve cells and loss of brain tissue. There is also a shortage of some important chemicals in the brain. Memory difficulties are usually the earliest symptoms of AD, other symptoms will involve problems with aspects of thinking, reasoning, perception or communication (4). There is currently no cure for dementia, although a number of risk factors have been identified, and modification of these may impact on the presentation and progression of the condition.

The risk factors for dementia are multifactorial and apply throughout the life course. At a population level, educational attainment in early life, such as the number of years spent in education, was found to be protective against dementia (6). A higher level of education appears to delay the onset of dementia by several years (6). Factors such as hypertension, type II diabetes, hyperlipidaemia, cognitive activity, social activity, exercise, alcohol use, diet and smoking are also proposed to play a role in the development of dementia (4, 6).

Oral diseases and cancers affect 3.9 billion people globally, and untreated caries in the permanent dentition is the most prevalent health condition in the world (7). In the UK, the latest national survey suggests that 31% of dentate adults had obvious caries and 6% were edentulous; almost half of dentate adults (45%) had periodontal pocketing  $\geq 4$ mm, although for those affected (37%) the disease level was moderate; the prevalence of pocketing  $\leq 6$ mm increased with age (8). Caries and periodontal disease are thus more common than other chronic health conditions and increase in older age.

Good oral health is an important aspect of general health and wellbeing contributing to self-esteem, dignity, social integration and nutrition. Oral diseases share common risk factors with other non-communicable

diseases (9), and affect quality of life (8). For example, poor oral health in older people has been shown to be associated with pain and discomfort (10), and reduced appetite (11). Those people with moderate and advanced levels of dementia have greater functional dependency, and are often reliant on others for their daily oral care. Some individuals present with behaviour and communication difficulties, and may resist assistance (e.g. not opening mouth, refusing oral care). Furthermore, people with dementia may lose the capacity to clean their teeth regularly resulting in more dental plaque accumulation; thus, increasing their risk of developing periodontal disease and dental caries.

Biological plausibility for the association between poor oral health and development of dementia is suggested by an increase in systemic inflammatory markers in the presence of both conditions. For example, periodontitis has been shown to increase the level of those inflammatory markers which are also implicated in the development of dementia (12, 13). There is emerging evidence from studies that people with dementia have poor oral health but the relationship between the two diseases is not clear. This review aims to synthesise the contemporary evidence exploring the relationship between oral health and dementia.

## **Methods**

A rapid review of articles reporting primary research published between 2005 and October 2015 was undertaken to investigate the relationship between dementia and oral health in line with Khangura et al, (2012) methodology (14). Whilst other reviews in this series limited their scope to systematic reviews, this present review was expanded to focus on primary research, because of the lack of high level evidence in this evolving field. The findings present a review of primary evidence from cohort and case-control studies with separate data tables for each oral health outcome. Given the paucity of primary research and to ensure completeness, the cross-sectional studies were also sourced as background but did not contribute to the results. During data synthesis, two systematic reviews exploring the relationship between cognitive decline and oral health were published and are included with this review to provide a contemporary body of evidence.

Search syntax was developed based on subject knowledge, MESH terms and task group agreements (Figure 1). This was followed by systematic title and abstract searches conducted by two independent researchers (JS and YR) on four electronic databases: Cochrane, Embase, MEDLINE (R), and PsycINFO. The search identified 1,431 potentially relevant abstracts which were screened in duplicate for relevance. This was followed by an assessment of 91 full text articles using explicit inclusion and exclusion criteria planned a priori. The inclusion criteria for papers were as follows: a defined diagnosis of dementia or cognitive impairment using a validated measure, a defined oral health outcome using a validated measure, and for the study to be a systematic review, randomised controlled trial, cohort or case-controlled study. Papers were excluded if dementia was not the primary focus or they contained only a proposed research protocol or were published before September 2005. The following information was extracted from each paper: author, year, title, journal, population studied, oral disease/intervention, definitions used, methods, comparison/intervention and controls, outcomes, results, authors' conclusions, quality and quality justification as shown in the data extraction Tables 1-2.

## Box 1: Search terms

Databases: Embase <1974 to 2015 November 04>, Ovid MEDLINE(R) in-process & other non-indexed citations and Ovid MEDLINE(R) <1946 to Present>, PsycINFO <2002 to November Week 1 2015>

Search Strategy:

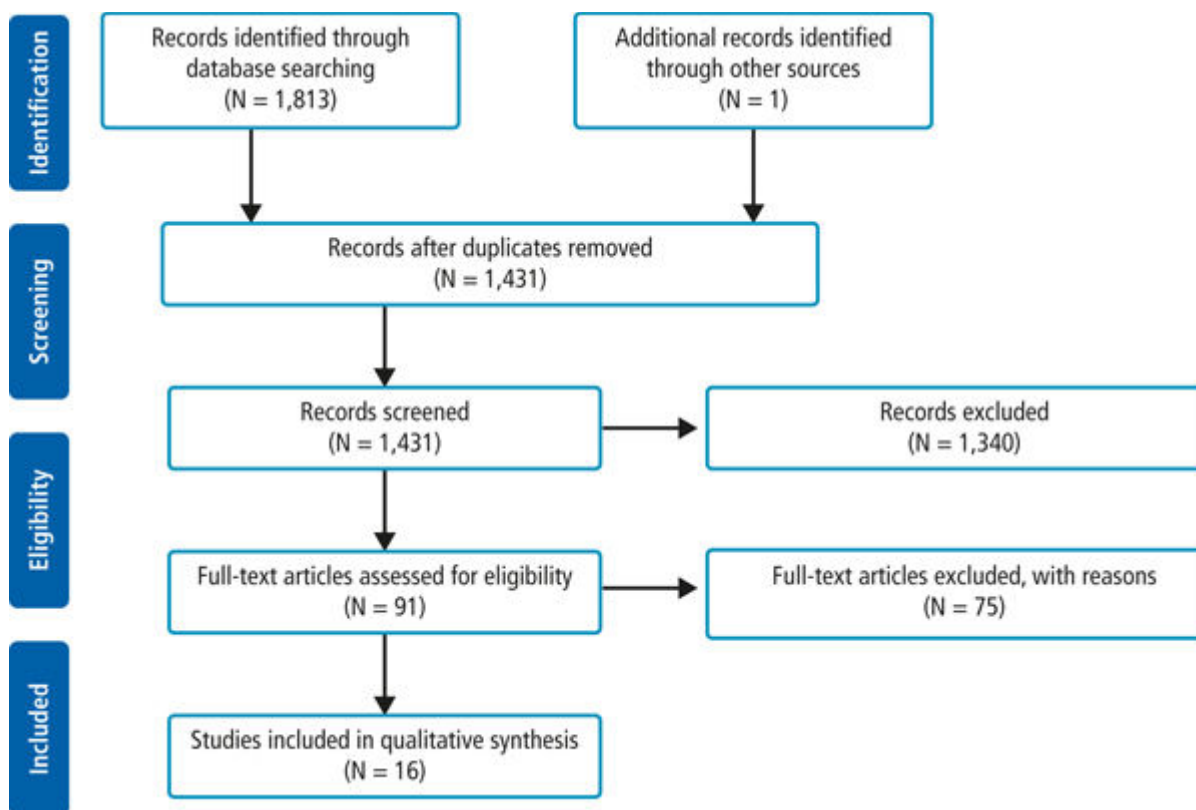
- 1 dementia.mp. (af)
- 2 alzheimer\$.mp. (af)
- 3 lewy-body.mp. (af)
- 4 vascular dementia.mp. (af)
- 5 cognitive impairment.mp. (af)
- 6 hippocampus.mp. (af)
- 7 cognition.mp. (af)
- 8 neurodegen\$.mp. (af)
- 9 cognitive reasoning.mp. (af)
- 10 cognitive function.mp. (af)
- 11 oral health.mp. (af)
- 12 dentistry.mp. (af)
- 13 dental health.mp. (af)
- 14 dental disease.mp. (af)
- 15 dental treatment.mp. (af)
- 16 dental therapy.mp. (af)
- 17 dental care.mp. (af)
- 18 oral medicine.mp. (af)
- 19 dry mouth.mp. (af)
- 20 xerostomia.mp. (af)
- 21 periodon\$.mp. (af)
- 22 gum disease.mp. (af)
- 23 saliva\$.mp. (af)
- 24 caries.mp. (af)
- 25 dental restoration.mp. (af)
- 26 dental an\*sthesia.mp. (af)
- 27 randomi\$ controlled trials.mp. (af)
- 28 controlled trial.mp. (af)

- 29 controlled clinical trial.mp. (af)
- 30 placebo.mp. (af)
- 31 single blind.mp. (af)
- 32 double blind.mp. (af)
- 33 triple blind.mp. (af)
- 34 rct.mp. (af)
- 35 cohort.mp. (af)
- 36 case contro\$.mp. (af)
- 37 case-contro\$.mp. (af)
- 38 systematic review.mp. (af)
- 39 meta analy\$.mp. (af)
- 40 meta-analy\$.mp. (af)
- 41 cross-sectional.mp. (af)
- 42 cross sectional.mp. (af)
- 43 cognitive decline.mp. (af)
- 44 DMFT.mp. (af)
- 45 CPITN.mp. (af)
- 46 DMFS.mp. (af)
- 47 1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9 or 10
- 48 11 or 12 or 13 or 14 or 15 or 16 or 17 or 18 or 19 or 20 or 21 or 22 or 23 or 24 or 25 or 26 or 44 or 45 or 46
- 49 27 or 28 or 29 or 30 or 31 or 32 or 33 or 34 or 35 or 36 or 37 or 38 or 39 or 40 or 41 or 42
- 50 47 and 48 and 49
- 51 remove duplicates from 50.

**Table 1-2**

In total, 16 papers were included in the final synthesis: 11 cohort and five case-control studies. They ranged from a sample size of 59 up to 11,140 participants, with data drawn from 28 countries. Most studies were small and there was considerable variation in how oral health outcomes were assessed, and the presence of a diagnosis of dementia or cognitive impairment was generally not reliably determined. A flow diagram of the process is shown in Figure 2.

**Figure 2: PRISMA flowchart of paper selection**



Critical appraisal was undertaken using the features of the Newcastle-Ottawa tool for cohort and case-control studies (15). Studies were rated out of 10, with those scoring three and under considered low and those scoring seven and above rated as high quality (Table 1-2).

The evidence presented here is based on a synthesis of disparate primary studies (n=16) of varying quality: high (n=5), medium (n=8) and low (n=3). Common bias included the lack of measure of exposure or outcome at baseline, failure to account for possible confounders, unclear definition of periodontal disease and lack of criteria for diagnosis of dementia. Additionally, proxy measures of oral health were used such as dental attendance and self-reported oral health.



## Results: evidence synthesis

The results are reported in two main sections following the questions examined by the review group. First, whether the presence of dementia (to include cognitive decline) had an association with the oral health status of people living with the condition in a range of settings (community, care home, hospital settings). Second, whether poor oral health was a risk factor for the development of dementia (to include cognitive decline).

### Part 1] Is dementia a risk factor for poor oral health?

This section of the results addresses the question of whether patients with dementia were at a higher risk of poor oral health. It is reported in six parts, each highlighting the association of dementia with different aspect(s) of oral health.

#### *1] Dental plaque, gingival bleeding and gingivitis*

Five case-control studies reporting on people with dementia were reviewed (Table 1A) (10, 16-19). The effectiveness of oral hygiene was assessed using measures of dental plaque accumulation (on tooth and denture surfaces), gingival bleeding and gingivitis. There was variation across the studies in relation to the recording of plaque scores and gingival bleeding, with the latter measured through self-report or clinical assessment.

#### **Table 1A**

Older people with dementia, assessed in care homes had **higher plaque** scores recorded on all tooth surfaces compared with controls (19). Community dwelling people with dementia attending hospital neurology departments had higher average plaque scores compared to controls ( $p < 0.001$ ) (17). In one small high quality prospective case-control study, an increase in MMSE score (a measure of cognitive decline) in patients with AD was correlated with an increase in dental plaque accumulation ( $p = 0.015$ ) (18); in this study plaque accumulation was recorded on tooth and denture surfaces using a composite score.

In another case-control study Chu et al., found that **twice daily toothbrushing** was *less commonly practised* by people with mild-level late-onset AD compared with matched controls (67% cf. 83%;  $p = 0.045$ ), and this group was six times more likely to have received support with their toothbrushing ( $p < 0.001$ ) (16).

In relation to **gingivitis** there was contrasting evidence; two case-control studies of community dwellers suggested that people with dementia had *higher* bleeding scores ( $p < 0.001$ ) (10, 11), whilst one small Chinese study did not report a difference (16).

In summary, the studies in this section suggested that people with dementia in community and institutional settings had higher levels of plaque present and worse oral hygiene compared with controls. Regular oral hygiene was less commonly practised in people with dementia and they were more likely to require support to

clean their teeth or dentures. In relation to gingivitis, there was conflicting evidence with the higher quality studies suggesting that people with dementia were more likely to have gingivitis compared with controls.

### *II] Periodontal disease*

Three case-control studies were included (Table 1B). The presence of periodontal disease was recorded using the following measures: Community Periodontal Index [CPI], attachment loss [AL], clinical attachment loss [CAL], and probing pocket depth [PD].

#### **Table 1B**

These three studies investigated community dwelling people (10, 16, 17). In a high quality study Gil-Montoya et al (17), suggested that **periodontitis** (AL >3mm; 67% of sites) appeared to be associated with cognitive impairment after controlling for confounders such as age, sex, and education level. The proportion of sites with AL >3mm was greater in patients with cognitive impairment compared to those with no impairment (p=0.002). When adjusted for age, sex, education, teeth present, oral hygiene habits, and hyperlipidaemia, the odds ratio of moderately extensive AL in adults with moderate cognitive impairment was 2.64 (1.18-5.92) and severe impairment was 2.31 (1.15-4.66) (p=0.04). In a study of patients with mild AD, de Souza Rolim et al (10), using the American Academy of Periodontology [AAP] definition of periodontal disease, reported that gingivitis and severe periodontal disease were more frequent in patients with mild AD compared with healthy controls (p=0.002). Chu et al (16), in a study involving older adults with mild level late-onset AD, found no evidence of an increased prevalence of advanced periodontal disease (assessed as CPITN  $\geq$ 3mm) between patients with dementia and matched controls.

In summary, whilst there are few studies examining whether people with dementia experience greater levels of periodontal disease, one high quality study suggests that the proportion of sites affected is significantly greater in patients with cognitive impairment compared with those without evidence of impairment.

### *III] Caries levels and saliva flow*

Five papers, comprising two cohort and three case-control studies, were included (Table 1C) (16, 18-21). Dental caries experience was defined as mean numbers of decayed, missing and filled teeth (DMFT), or surfaces (DMFS) or caries increment over a defined period. Decayed coronal and root surfaces were presented, separately if appropriate.

#### **Table 1C**

Chen et al, (21), in a high quality cohort study showed that adults with a diagnosis of dementia attending a dental clinic had significantly higher levels of **dental caries** compared to controls. Ellefson et al, (20), in a medium quality cohort study showed that older people attending a memory clinic (other than patients with AD) were at elevated risk of developing high levels of coronal and root surface caries during their first year after referral. Additionally, those who received a dementia diagnosis, other than AD, appeared to be at a particularly high risk of developing multiple carious lesions in the first year after diagnosis (20).

Evidence from three case–control studies in community dwelling individuals showed no statistically significant difference in decay experience between people with dementia compared to those without (16, 18, 19). One study, however, did show a correlation between a decline in MMSE and increase in DMFT scores ( $p=0.022$ ) (18). Chu et al., showed that people with dementia had lower unstimulated saliva flow rates (ml/min) compared with a control group (0.30 V 0.41) ( $p=0.043$ ) with reduced salivary flow being a risk factor for dental caries (16); however, there is no evidence presented of controlling for other causes of dry mouth.

In summary, there are very few studies exploring caries experience and those that exist are of variable quality (one high, three medium and one low). Whilst there is evidence that people with dementia are at increased risk of presenting to dental services with caries and may be at higher risk of developing caries once dementia is established, the results overall are conflicting.

#### *IV] Tooth Loss*

Five studies (10, 16, 18, 19, 22) examine the link between dementia and tooth loss, one cohort, and four case-control as presented in Table 1D.

#### **Table 1D**

In a high quality cohort study Chen et al, (22) found that amongst affected community dwellers dementia was not associated with **tooth loss**, although all participants (with and without dementia), with more teeth ( $n_{\geq 25}$ ) at first assessment, were at higher risk of tooth loss during follow-up ( $p<0.001$ ). A medium quality study in Hong Kong found no significant difference in the number of missing teeth for people with AD and matched controls (16); similarly patients with AD and matched controls had similar rates of edentulism in a small low quality case-control study in Brazil (10). Adam and Preston (19), found that 70% ( $n=38$ ) of those with mild or no dementia and 63% ( $n=51$ ) of those with moderate or severe dementia were edentulous. One small medium quality case-control study in Turkey found a positive correlation between teeth present with MMSE scores in patients with AD (18).

In summary, one high quality cohort study and one medium quality case-control study found no association between dementia and tooth loss, whilst two low quality case-controls found similar rates of edentulism between people over 59 years old with and without dementia. Just one small medium quality case-control study found a positive correlation between the stage of dementia, as determined using MMSE, and the number of teeth present; i.e. in later stages of dementia patients had fewer remaining teeth.

#### *V] Denture related conditions:*

Three case-control studies of variable quality (two low and one medium) examined denture-related conditions (Table 1E) (10, 18, 19). The findings are equivocal but highlight a range of issues relating to **wearing and caring for dentures**. In edentulous individuals, Adam and Preston (2006) conducted a study of care homes and showed that 60% of those with moderate to severe dementia wore no dentures at all compared with only 10% of those with no or mild dementia ( $p=0.04$ ) (19). Whilst one small case-control suggested no difference

in denture wearing between those with and without mild dementia (10), other evidence from one case-control study by Hatipoglu et al, suggests that community living people with moderate or severe dementia were less likely to remove their denture at night and were more likely to have denture related stomatitis in both the maxilla and mandible ( $p=0.001$  for all) (18).

### Table 1E

In summary, there is low quality evidence that people tend to stop wearing their dentures as cognitive decline increases, while the presence of denture stomatitis increases for those who continue to wear dentures.

## Part 2] Is poor oral health a risk factor for developing dementia (including cognitive decline)?

This section of the results addresses the question of whether patients or communities with poor oral health were at a higher risk of developing dementia (to include cognitive decline). It is reported in five parts, each highlighting the association of dementia with the relevant aspect(s) of oral health.

### *1] Dental plaque, gingival bleeding and gingivitis*

Six studies (11, 17, 18, 23-25) of variable quality were reviewed (Table 2A); four cohort and two case-control studies.

### Table 2A

One high quality five-year prospective multi-centre cohort study found a higher than two-fold cognitive decline (modified MMSE score) in people with **gingival inflammation** denoted by higher gingival index scores (OR 2.54; CI 1.75-3.70), after correcting for potential confounders (23). In a medium quality large community-based cohort study Paganini-Hill in the US found that *not* performing daily toothbrushing amongst residents of a large retirement community was associated with an increased risk of developing dementia over an 18-year period compared with those brushing three times a day (males 22%; females 65%), although this was only statistically significant in women (11).

A medium quality case-control study of patients with AD reported evidence of a correlation between **oral hygiene status** and mini-mental state examination (MMSE) scores, but affected individuals were not significantly worse than controls (18). A four year prospective cohort study in Japan of low quality, suggested an association between not taking care of dental health and risk of dementia was partly explained by co-factors such as socio-demographics, health behaviours and forgetfulness as an early symptom of mild cognitive impairment (25). In a high quality case-control study Gil-Montaya et al reported the odds of cognitive impairment were 15.7-fold greater in patients with a higher plaque accumulation measured by a plaque score (2.51-3 cf 0 - 1);  $p<0.001$ ; however, the direction of the relationship remained uncertain (17).

In summary, there is evidence from cohort and case-control studies of variable quality that failure to perform toothbrushing and the presence of gingival inflammation may be risk predictors associated with developing dementia.

### *III] Periodontal disease*

Six studies in total, five cohort and one case-control study examined periodontal disease (Table 2B) (17) (23, 26) (25, 27) (28).

#### **Table 2B**

Kaye et al (27), in a 32-year longitudinal cohort study showed that risk of cognitive decline over a decade increased by 2-5% for each tooth that had progression of **alveolar bone loss** or **probing pocket depth** (27). Furthermore, the tendency to have a lower MMSE score (denoting increased cognitive impairment) was consistently higher in older men compared with younger men and thus rates of tooth loss and periodontal disease progression during adulthood independently predicted performance on the MMSE cognitive test (27). Gil-Montoya et al (17), reporting a case-control study, suggested that **periodontitis** appeared to be associated with cognitive impairment after controlling for age, sex, education level and oral hygiene habits. Risk of cognitive impairment was more than three times higher in patients with severe periodontitis compared to those with mild or no periodontitis;  $p < 0.001$ .

In contrast to the above, three medium to high quality cohort studies did not show a relationship between cognitive decline and periodontal disease. Stewart et al. (23), in their 5-year cohort analysis examined periodontal health by calculating the probing depth and loss of attachment for 954 participants; their findings showed no prediction of cognitive decline as associations were substantially confounded by education and race. Arrive (28), in a 15-year cohort study showed that periodontal status was not associated with the risk of dementia before, and after, correcting for confounders. Finally, Okamoto et al (26), in a large Japanese cohort study found no evidence of an association between the community periodontal index (CPI) and cognitive impairment.

In summary, the results are equivocal: three cohort studies (one high and two of medium quality) found no association between measures of periodontal disease and cognitive decline, whilst one very long cohort study of medium quality and two case-control studies of medium/high quality suggest an association.

### *IV] Caries*

Only one study of medium quality was reviewed (Table 2C). Kaye et al (27), in a 32-year long cohort study of male veterans showed that development of new caries or need for new restorations was associated with greater risk of poor performance in MMSE and was greater in men aged over 45.5 years of age at baseline.

#### **Table 2C**

## VJ Tooth loss

Ten studies examine the link between tooth loss (total and number of remaining teeth) and dementia, nine cohort studies (two high, five medium and two low quality) (11, 23-30) and one high quality case-control study (17), were included (Table 2D).

### Table 2D

A 37-year medium quality cohort study found that dementia was twice as common in those with fewer than nine teeth in comparison to those with 25 or more teeth (29). A 32-year cohort study, of medium quality, found that with each tooth lost per decade, the risk of having a low cognitive test score increased by 9% to 12% (27). A five-year cohort study, also of medium quality, found those participants with one to eight remaining teeth and progressing to total tooth loss were associated with having mild memory impairment ( $p=0.008$ ); however, lower tooth count was not associated with dementia (26).

Two cohort studies (one low and one medium quality) considered masticatory function by using the number of teeth remaining and denture wear. Yamamoto (25), found that those people with fewer teeth and no replacement dentures had a 1.85-fold increased risk for dementia. Paganini-Hill et al (11), found that participants with fewer than ten upper teeth, fewer than six lower teeth, and not wearing dentures, had a higher risk of dementia compared to those with adequate natural masticatory function; the risk was significant in men (Hazard ratio (HR) = 1.91) but not in women (HR = 1.22) (11).

In a high quality cohort study, edentulous participants from the English Longitudinal Study of Ageing (ELSA) were found to recall fewer words than dentate participants; however, the association was attenuated when corrected for socioeconomic status (30). One low quality cohort study also suggests a trend in the development of dementia or cognitive decline for those participants who were edentulous (24). Conversely one medium quality cohort study found the risk of dementia was significantly lower in people with eleven or more missing teeth and a lower educational attainment (HR=0.30; 95% CI 0.11-0.79), compared with higher educational attainment (HR=1.07; 95% CI 0.57-2.02). Two high quality studies (one cohort and one case-control) found that neither number of teeth present nor numbers of occluding pairs of teeth were associated with cognitive decline when other risk factors were considered (17, 23).

In summary, whilst there are a number of studies relating to this aspect of oral health, there are conflicting results as evidenced by the fact that several studies of medium/high quality suggest that progressive tooth loss was associated with increasing risk of cognitive decline, while others do not. Studies researching edentulous participants have not been able to provide conclusive evidence to support an association with an increased risk of cognitive decline; however, there is sufficient evidence to suggest that this is an important area for further research.

## *Vij Dentures*

Studies involving investigation of dentures are limited to two cohort studies of low (25), and medium quality (11) (Table 2E).

### **Table 2E**

An 18-year cohort study in a retirement community in the US suggest that for denture wearers, adequate masticatory function involving 10 or more upper, and six or more lower teeth, was associated with a lower risk of dementia (11). Furthermore, the study also suggests that cleaning dentures was not significantly related to dementia (11). In contrast, a four-year cohort study in Japan showed that people with fewer teeth and no dentures were found to be at greater risk for dementia (25). Compared with people having 20 or more teeth, those with fewer teeth and no dentures were at almost twice the risk of dementia [1.85 (CI 1.04 – 3.31);  $p=0.04$ ]; however, general health status, health behaviour and forgetfulness attenuated the association (11, 25).

Thus, in summary whilst the findings, of medium to low quality, regarding dentures and dementia are equivocal, there is a suggestion that masticatory function may be important and further research is required.

## **Discussion**

This rapid review highlights the paucity of evidence on the relationship between dementia and oral health over the 10-year review period. The evidence available is of mixed quality and covers a wide range of oral diseases and conditions. Given the extent of heterogeneity meta-analysis by outcome has not been possible.

This review has a number of strengths and limitations which should be acknowledged. First, the review process conducted by a multidisciplinary team containing medical, dental, and public health professionals was considered a strength. Second, this is a “rapid review”, and so was intended to summarise existing evidence, rather than undertake quantitative synthesis. The restriction to the 10 year review period however meant that some important landmark papers may have been missed in the searches. Third, given the nature of oral disease in this population, there is a wide range of oral conditions to be examined. Fourth, there was large heterogeneity in the studies reviewed; nonetheless, there is important learning to inform research in this field.

Certain general limitations relating to research in this field need to be acknowledged. First, there are thought to be different pathological mechanisms for the various different dementias so the potential relationships between oral health/diseases and dementias could be different for different pathologies. This makes trying to identify relationships complicated, particularly when many of the dementia diagnoses are made post mortem. Second, studies in subjects with dementia that rely on self-reported activity such as toothbrushing, are subject to considerable risk of error due to reduced, and often variable, cognitive performance in this population.

Is dementia a risk factor for poor oral health? The evidence suggests that people with dementia in community and institutional settings may have higher plaque accumulation, more teeth and surfaces affected by dental caries and more extensive periodontal disease compared to controls. A large proportion of this association may be explained by increasing age and fragility, greater functional dependence and the presence of other confounding co-morbidities. This is not however the finding across all the studies we explored and there is

some additional learning and research warranted in communities where people's oral health appears to be less affected by dementia. The differences may well be related to the quality and conduct of the studies and further confounded by access to health and social care; nevertheless, more research using large well designed longitudinal prospective cohort studies is required.

Is poor oral health a risk factor for dementia? There is some limited evidence of the impact of oral health (oral hygiene, caries and number of teeth) on dementia (cognitive impairment, dementia onset and progression), including plaque accumulation, gingivitis and notably tooth loss. Risk factors and risk predictors for dementia are of interest in current research, particularly the early detection and identification of disease. Research into tooth loss and its putative role as either a predictor or as a co-factor in cognitive decline may become increasingly relevant as we move towards earlier recognition and diagnosis of dementia. There is very weak and limited evidence that in the absence of regular toothbrushing, the presence of gingivitis and reduced masticatory function may be risk factors for dementia, but there is no evidence of a direct effect on caries and periodontal disease. It is important to acknowledge that changes in MMSE could be predicted by a change in oral health status such as a loss of teeth, or vice versa. Alternatively, both may be a feature of ageing with no plausible biological link between the two phenomena.

We considered it helpful to examine further evidence from two recently published systematic reviews of cohort studies shown in Table 3. Cerrutti-Kaplin et al. (31), reviewed 10 studies, five of which are considered in our review (11, 24, 25, 27, 28). Whilst the authors report that the relationship between periodontal disease and cognitive impairment showed conflicting results, individuals with tooth loss and suboptimal dentitions (<20 teeth) were considered at a 20% higher risk of developing cognitive decline (HR = 1.26, 95% CI = 1.14 to 1.40) and dementia (HR = 1.22, 95% CI = 1.04 to 1.43) compared to those with an optimal dentition (≥20 teeth) lending some support to the hypothesis that tooth loss is associated with an increased risk of cognitive impairment. Wu et al. (32), considered the number of teeth, dental caries, periodontal disease and dentures and highlighted conflicting and inconsistent findings in relation to oral health and cognitive status. The authors attributed the conflicting results to methodological weaknesses concluding that there was uncertainty about an association between oral health status and cognitive decline.

### **Table 3**

In relation to the questions examined overall, and given the range of conditions investigated, the findings from our rapid review can only suggest the presence of potential associations with oral hygiene, gingivitis, periodontal disease, dental caries, saliva flow, tooth loss, and denture related conditions including masticatory function. In common with Cerruti-Kaplin et al and Wu et al, our review was unable to draw firm conclusions relating to cause and effect due to the limitations of the underpinning primary research which included: small sample size, unclear inclusion and exclusion criteria and appropriateness of experimental design. There is, however, sufficient emerging evidence to suggest that this is an important area for further research given demographic changes and the fact that older adults have already experienced significant oral disease and its sequelae. Given the problems with existing evidence, future research should include the use of standardised criteria for planning, undertaking fieldwork and assessing outcomes as shown in Table 4.

### **Table 4**



On a practical level, daily self-care and/or assisted care is vitally important for oral health (33), whether or not people have experienced cognitive decline or dementia. It has particular relevance for those with dementia as they are often reliant on others for support to maintain oral hygiene. Therefore, for those carers and health care workers (HCWs) who provide direct personal care, mandatory training should be provided in oral health and hygiene, to ensure carers and HCWs have a basic competence in assessing the mouth for oral hygiene status and abnormalities. HCWs and social care professionals are often unfamiliar with pain assessment in cognitively impaired patients; therefore, training is needed to help identification and management of dental pain, which is often inaccurately attributed to other causes. Additionally training should include practical oral hygiene measures, including techniques for those patients who may resist support for oral hygiene and denture care. This training may be more easily implemented in long term care facilities; however, the evidence suggests that people living with dementia in the community are just as likely as those in long-term care facilities to have poor oral hygiene. It is clear that every effort must be made to ensure that those living in the community should also receive the necessary oral health care support to secure optimal oral health.

While it has not been possible to establish poor oral health predicts dementia, there is more concrete evidence that oral health declines in the presence of increasing cognitive impairment and dementia. Dental care must therefore be included in the wider care plan for people with dementia, in order to improve oral health throughout each individual's life course, and minimise impact on the quality of life. There must be consideration of the progressive nature of cognitive decline and dementia, and also the challenges of dental care in later stages of the disease when formulating individual oral care plans. Implementation of delivering better oral health guidelines and tailored support at various stages of dementia are also important (34).

Looking to the future, there is a clear need for primary research into links between oral health and dementia, such as the link between periodontal disease and systemic inflammatory loading (24, 28), along with research into whether increased inflammatory loading has an impact on the progression or development of dementia.

It is important to explore the possibility of multidisciplinary research approaches, using standardised globally acceptable assessments and definitions (4, 5, 35). For example there is potential for collaboration with physiotherapists and occupational therapists to focus on the activities of daily living concurrently such as manual dexterity, gait and toothbrushing.

Dieticians with their expertise in nutrition, could ensure diet plans consider oral health and speech and language therapists could contribute with their assessment of concomitant swallowing difficulties, as these are prevalent in people with dementia and may affect oral hygiene practices and conversely, oral microflora which could have implications for pulmonary health. This approach will also help to explore the holistic relationships of dementia, oral health, and general health.

## **Conclusion**

This rapid review of evidence suggests that people with dementia may experience worse oral health, with poor oral hygiene and dementia (cognitive decline/impairment) showing the most consistent association. There is conflicting evidence relating to tooth loss and the risk of cognitive decline and dementia associated with poor oral health, though professional management and satisfactory treatment of oral conditions remains important

in this patient group. Overall, given the quality and paucity of research in this field, the findings should be treated with caution and require more rigorous testing in well designed and conducted studies, preferably longitudinal in design.

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**Table 1: Cohort and case-control studies: Is dementia (inc. cognitive decline) a risk factor for poor oral health?**

Author and year	Study type	Population	Results	Authors' conclusions	Quality
<b>A. Dementia impact on plaque, gingival bleeding and gingivitis</b>					
Chu et al 2015 16	Case-control  Country: Hong Kong	<p><b>Cases:</b> 59 patients in day care centres of Hong Kong Alzheimer's Disease Association. All with mild level late-onset Alzheimer's disease. All <math>\geq 60</math> years of age.</p> <p><b>Controls:</b> 59 healthy controls without dementia and matched for age and gender, from Prince Philip Dental Hospital (not under current dental treatment).</p>	<p><b>1. Percentage of people performing brushing twice daily</b> Dementia (67%); control (83%); <math>p=0.045</math>.</p> <p><b>2. Percentage of people receiving assistance when brushing</b> Dementia (31%) control (5%); <math>p&lt;0.001</math></p> <p><b>3. Percentage of people with reversible gingivitis (as indicated by a community periodontal index score maximum of 1)</b> Dementia (N = 5/47) 11%; Control (N = 7/50) 14%</p> <p><b>4. Assistance with toothbrushing</b> Dementia (31%); Control (5%); <math>p&lt;0.001</math></p>	<p>Compared to those without dementia, fewer elderly Chinese with dementia practiced tooth brushing twice daily <i>And they were more likely to have assistance with toothbrushing</i></p>	Medium
Gil-Montoya et al 2015 17	Case-control  Country: Spain	<p><b>Cases:</b> Patients treated in neurology departments of two hospitals, who had a firm diagnosis of AD, Dementia, or MCI (mild cognitive impairment)</p> <p><b>Controls:</b> patients being seen in primary healthcare, for problems other than dental or neurological problems. Excluded schizophrenic, depressive, personality disorder, diseases not under treatment, drug abuse, and having previous periodontal treatment in the last 6 months</p>	<p><b>1. OR of cognitive impairment in relation to oral hygiene habits</b> (reference group those who brushed twice or more a day) i. mouthwash only OR=11.37 CI(5.46 - 23.68) <math>p&lt;0.001</math> ii. No brushing OR=7.21 CI(3.48 - 14.96) iii. Brushing once a day OR=3.34 CI(2.03 - 5.49)</p> <p><b>2. OR of cognitive impairment in relation to dental variables</b> i. Plaque index (reference group; 0.00 - 1.00). 1.01 to 2.00; OR=1.92 CI(0.94 - 3.90) 2.01 to 2.50=6.33 CI(3.15 - 12.72) 2.51 to 3.00=15.70 CI(7.72 - 31.92) ii. Bleeding index (reference group was 0-25% BOP) 25.1 to 50% BOP OR=2.32 CI(1.21 - 4.48) 50.1 to 90% BOP OR=2.67 CI (1.47 - 4.85) 90.1 to 100% BOP OR=3.53 CI (1.88 - 6.63)</p> <p><b>3. Association between dental variables and levels of cognitive impairment</b> i. Mean Plaque index MCI; 2.26.(SD=0.49) Mild Dementia; 2.27 (SD=0.76) Moderate Dementia; 2.4 (SD 0.57) Severe Dementia 2.5 (SD=0.62) <math>p=0.18</math> ii. Mean Bleeding index MCI; 45.4 (SD=20.9) Mild Dementia; 62 (SD=29.5) Moderate Dementia; 61.5 (SD=34.2) Severe Dementia; 77.6 (SD: 27.3) <math>p=0.001</math></p>	<p>Periodontitis appears to be associated with cognitive impairment after controlling for age, sex, education level and oral hygiene habits</p> <p><i>Average plaque and bleeding scores were significantly higher in the patients with a diagnosis of AD, Dementia or MCI</i></p> <p><i>There is evidence of increased bleeding score with cognitive decline; however, there wasn't significantly more plaque present with cognitive decline</i></p> <p><i>The present results evidence a significant worsening of oral hygiene habits (reduction or cessation of toothbrushing) with more advanced cognitive disease. The direction of the association is yet to be established</i></p>	High

			<p>iii. Oral hygiene habits Brushed twice or more a day cf brushed once a day (n) cf used only mouthwash cf no oral hygiene habits (n) in relation to MCI, mild-moderate- and severe-dementia; p&lt;0.001</p> <p><b>4. Mean plaque scores (PI)</b> Mean plaque 2.37 (SD =0.65) cases cf 1.55 (SD = 0.89) controls p&lt;0.001.</p> <p><b>5. Mean bleeding scores (BI)</b> Mean bleeding scores 63 (SD = 31.1) cases cf 50.6 (SD = 34.2) controls p&lt;0.001.</p>		
de Souza Rolim et al 2014  10	Case-control  Country: Brazil	<p><b>Cases:</b> 29 patients with mild AD from Cognitive Neurology and Behaviour Group of the Neurology Department of the School of Medicine of the University of Sao Paulo, Brazil</p> <p><b>Controls:</b> 30 elderly without AD (relatives of the patients: wives, husbands, brothers, and/or sisters) Mix of dentate (36) and edentate (23) patients</p>	<p><b>1. Difference in dental characteristics between case and control groups</b> (AAP Definitions of periodontal disease) <i>Gingivitis:</i> Case 31% c.f. Control =10% <i>Moderate perio disease:</i> Case=6.9% c.f. Control =10% <i>Severe perio disease:</i> Case=20.7% c.f. Control =6.7% p=0.002</p>	Periodontal infections were more frequent in patients with mild AD than in healthy subjects	Low
Hatipoglu et al 2011  18	Case-control  Country: Turkey	<p><b>Cases:</b> patients with Alzheimer's disease (n=31)</p> <p><b>Controls:</b> patients of a similar age and gender without known systemic disorders (n=47) Cases and controls were from the Neurology Department of an Institute in Turkey.</p>	<p><b>1. Oral hygiene status</b> cases c.f. control patients: Good; 1 (3.2%) c.f. 9 (19.1%) p&gt;0.510 Fair; 9 (29.0%) c.f. 15 (31.9%) p&gt;0.510 Poor; 21 (67.7%) c.f. 23 (48.9%) p&gt;0.510</p> <p><b>2. Correlation between clinical and oral findings:</b> MMSE scores were significantly correlated with oral hygiene status p=0.015</p>	The results suggest that cognitive status (MMSE) of patients is significantly correlated with oral hygiene status	Medium

Adam and Preston 2006 19	Case-control Country: United Kingdom	135 individuals, aged 65 years and over, residing in four nursing homes in Cheshire, UK <b>Cases:</b> 81 people with moderate to severe dementia  <b>Controls:</b> 54 people with no or mild dementia	<p><b>1. Mean Plaque score</b> Greene and Vermillion index:</p> <ul style="list-style-type: none"> <li>i. All surfaces; No/mild dementia 1.33 (SD=0.60), Moderate/severe dementia 2.14 (SD=0.73)</li> <li>ii. Buccal surfaces, No/mild dementia 1.42 (SD=0.72), Moderate/severe dementia 2.14 (SD=0.73)</li> <li>iii. Lingual surfaces, No/mild dementia 1.24 (SD=0.47), Moderate/severe dementia 2.14 (SD=0.72)</li> </ul> <p><b>2. Mean calculus score</b> Greene and Vermillion index</p> <ul style="list-style-type: none"> <li>i. All surfaces; No/mild dementia 1.34 (SD=0.60), Moderate/severe dementia 2.04 (SD=0.75)</li> <li>ii. Buccal surfaces; No/mild dementia 1.38 (SD=0.69), Moderate/severe dementia 2.05 (SD=0.76)</li> <li>iii. Lingual surfaces, No/mild dementia 1.29 (SD=0.51), Moderate/severe dementia 2.04 (SD=0.73)</li> </ul> <p>Note. No evidence of significance provided</p>	For individuals' resident in nursing homes, moderate to severe dementia might have a deleterious effect on oral health (plaque and calculus).	Low
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B. Dementia impact on periodontitis					
Author and year	Study type	Population	Results	Authors' conclusions	Quality
Gil-Montoya et al 2015  17	Case-control  Country: Spain	<p><b>Cases:</b> Patients treated in a behavioural and cognitive neurology department, with a firm diagnosis of AD, Dementia, or MCI (n=180)</p> <p><b>Controls:</b> patients being seen in primary healthcare, for problems other than dental or neurological problems (n=229). Excluded schizophrenic, depressive, personality disorder, diseases not under treatment, drug abuse, and previous periodontal treatment in the last 6 months</p>	<p><b>1. OR of cognitive impairment in relation to dental variables</b> (Periodontitis: % of sites with AL &gt;3mm where moderate 33-66%; severe 67-100%) (reference group absent/mild periodontitis) Moderate; OR=2.58 (1.31 - 5.09) Severe; OR=3.04 (1.69 - 5.46) p&lt;0.001 <i>Adjusted for age, sex, present teeth, OH habits, hyperlipidaemia, studies:</i> Moderate; OR=2.64 (1.18-5.92) Severe; OR=2.31: (1.15-4.66) p=0.04</p> <p><b>2. Association between dental variables and levels of cognitive impairment</b></p> <p>i. Mean Pocket Depth; MCI; 2.8 (S.D=0.5) Mild Dementia; 3.0 (S.D=0.6) Moderate Dementia; 3.1 (S.D=0.6) Severe Dementia; 2.0 (S.D=0.9) p=0.47</p> <p>ii. Mean Attachment loss (mm) MCI; 4.6(S.D=1.0) Mild Dementia; 4.8 (S.D=1.4) Moderate Dementia 5.1 (S.D=2.0) Severe Dementia 4.7 (S.D=1.3) p=0.42</p> <p>iii. Attachment loss (percentage &gt;3mm) MCI; 75.2 (S.D=19.8) Mild Dementia; 76.6 (S.D=27.4) Moderate Dementia; 73.3 (S.D=33.8) Severe Dementia; 75.0 (S.D=27.1) p=0.94</p> <p><b>iv. Severe periodontitis (% with 67-100% AL &gt;3mm)</b> MCI; 66.7% Mild Dementia; 67.7% Moderate Dementia; 70.5% Severe Dementia; 66.7 p=0.98</p> <p><b>3. Adjusted OR of cognitive impairment in relation to periodontitis</b> <i>Periodontitis severity; reference group absent/mild periodontitis;</i> i. Moderate periodontitis OR=2.64 CI(1.18 - 5.92), ii. Severe periodontitis OR=2.31 CI(1.15 - 4.66) p=0.04</p>	Periodontitis appears to be associated with cognitive impairment after controlling for confounders such as age, sex, and education level	High



Chu et al 2015 16	Case-control Country: Hong Kong	<p><b>Cases:</b> 59 Patients in day care centres of Hong Kong Alzheimer's Disease Association</p> <p><b>Controls:</b> 59 generally healthy controls matched for age and gender, from Prince Philip Dental Hospital (not under current dental treatment)</p>	<p><b>1. Prevalence of advanced periodontal disease (CPI of 3 or more)</b> Dementia group=78% c.f. Control group=74%; p= 0.64</p>	<p>Compared to those without dementia, fewer elderly Chinese with dementia practiced tooth brushing twice daily <i>And they were more likely to have assistance with toothbrushing</i></p>	Medium
de Souza Rolim et al 2014 10	Case-control Country: Brazil	<p><b>Cases:</b> 29 patients from Cognitive Neurology and Behavior Group of the Neurology Department of the School of Medicine of the University of Sao Paulo, Brazil with mild AD</p> <p><b>Controls:</b> 30 elderly people without AD (relatives of the patients: wives, husbands, brothers, and/or sisters) Mix of dentate (36) and edentate (23) patients</p>	<p><b>1. Difference in periodontal disease between case and control groups</b> (AAP Definitions of periodontal disease) <i>Gingivitis:</i> Case 31% c.f. Control =10% <i>Moderate perio disease:</i> Case=6.9% c.f. Control =10% <i>Severe perio disease:</i> Case=20.7% c.f. Control =6.7% p=0.002</p>	<p>Periodontal infections were more common in patients with mild AD than in healthy subjects</p>	Low

C. Dementia impact on caries and salivary flow					
Author and year	Study type	Population	Results	Authors' conclusions	Quality
Chu et al 2015  16	Case-control	<p><b>Cases:</b> 59 Patients in day care centres of Hong Kong Alzheimer's Disease Association with dementia.</p> <p><b>Controls:</b> 59 generally healthy controls matched for age and gender, from Prince Philip Dental Hospital (not under current dental treatment).</p>	<p><b>1. Mean DMFT, DT, FT of cases vs controls</b> DMFT cases 22.3 (SD=8.2), controls 21.5 (SD=8.2); p=0.59 DT cases 1.2 (SD=1.9), controls 0.8 (SD=1.4); p=0.28 FT cases 2.5 (SD=3.3), controls 2.4 (SD=2.5); p=0.88</p> <p><b>2. Unstimulated salivary flow rate (ml/min)</b> Dementia group - 0.3 (SD=0.17) c.f. control group - 0.41 (SD=0.28); p=0.0043</p>	No significant difference in dental caries experience (DMFT) between AD patients and controls. Resting salivary secretion was reduced in dementia patients; however, their caries experience was not significantly different	Medium
Chen et al 2013  21	Retrospective longitudinal cohort study  Country: United States of America	<p>491 older adults who received dental care as new patients from the Wilder Senior Dental Clinic</p> <p>119 with diagnosis of Dementia, AD or chronic brain syndrome of no diagnosis</p> <p>- control group was not matched</p>	<p><b>1. Number of decayed and broken teeth at baseline</b> Dementia; 4.2 ± SD 4.5 (range 0-20) Without dementia; 3.1 ± SD 3.8 (range 0-18) p=0.006</p>	Patients with dementia present with higher rates of caries at baseline	High
Hatipoglu et al 2011  18	Prospective Case-control	<p><b>Cases:</b> Patients with AD (31)</p> <p><b>Controls:</b> (47) matched for similar age and gender without known systemic disorders</p> <p>Cases and control from Neurology Department of the Institution.</p>	<p><b>1. Mean DMFT score (case c.f. control)</b> 24.19 (SD=6.8) c.f. 19.68 (SD=9.5); p= 0.126</p> <p><b>2. Correlation between clinical and oral findings</b> MMSE scores were significantly correlated with DMFT; p=0.022</p>	<p>No significant difference in dental caries experience (DMFT) between patients with AD and the control group</p> <p>Decreased MMSE scores were significantly correlated with increased DMFT.</p>	Medium

<p>Ellefson et al 2009</p> <p>20</p>	<p>Cohort</p>	<p>77 patients with Alzheimer's disease (AD), other dementia (OD), no dementia (ND) referred to the memory clinics of two hospitals in Copenhagen, Denmark</p>	<p>Crude caries (coronal and root) increments (CCI), net caries increments (NCI), and adjusted caries and filling increments (ADJCI), for AD, OD and ND patients</p> <p><b>1. Change in average number of decayed surfaces (DS) in first year after referral to memory clinic</b>  All patients: increased from 5.7 (SD=6.7) to 7.9 (SD=8.8); <math>p \leq 0.005</math>  OD: increased from 5.0 (SD=6.6) to 10.2 (SD=9.8); <math>p \leq 0.005</math>  AD: increased from 6.7 (SD=7.2) to 8.0 (SD=9.2) NS  ND but referred to memory clinic: increased from 2.8 (SD 3.0) to 4.8 (SD 5.1) NS</p> <p><b>2. Adjusted OR of ADJCI after one year</b>  <i>(reference group; no dementia)</i>  OR of having a higher coronal caries increment (&gt;2) in OD was 4.13 CI(0.44-39.05) c.f. ND  OR of having a higher coronal caries increment (&gt;2) in AD was 0.64 CI(0.09-4.5) c.f. ND</p>	<p>Elderly people referred to a memory clinic were at an elevated risk of developing high levels of coronal and root surface caries during the first year after referral (independent of whether or not they fulfilled the criteria for a dementia diagnosis), and that elderly people with a dementia diagnosis other than AD appeared to be at a particularly high risk of developing multiple carious lesions during the first year after diagnosis.</p>	<p>Medium</p>
<p>Adam and Preston 2006</p> <p>19</p>	<p>Case-control</p>	<p>135 individuals, aged 65 years and over, residing in four nursing homes in Cheshire, UK  <b>Cases:</b> 81 people with moderate to severe dementia  <b>Controls:</b> 54 people with no or mild dementia</p>	<p><b>1. Mean number decayed teeth DT</b>  No/mild dementia; 0.11 (SD=3.42); Moderate/severe dementia; 0.80 (SD=1.87) NS</p> <p><b>2. Mean number of missing teeth MT</b>  No/mild dementia 28.22 (SD=6.64), Moderate/severe dementia 27.28 (SD=7.73) NS</p> <p><b>3. Mean number of filled teeth</b>  No/mild dementia 0.69 (SD=1.30); Moderate/severe dementia 0.90 (SD=2.39) NS</p>	<p>In an institutionalised population there were no statistical differences between the two groups (no/mild dementia and moderate/severe dementia) in terms of the DMFT</p>	<p>Low</p>

D. Dementia impact on tooth loss/number					
Author and year	Study type	Population	Results	Authors' conclusions	Quality
Chu et al 2015 16	Case-control Country: Hong Kong	<b>Cases:</b> 59 patients in day care centres of Hong Kong Alzheimer's Disease Association. All with mild level late-onset Alzheimer's disease. All $\geq 60$ years of age.  <b>Controls:</b> 59 healthy controls without dementia and matched for age and gender, from Prince Philip Dental Hospital (not under current dental treatment).	<b>Missing teeth, cases c.f. controls</b> Cases 18.9 (SD=9.4), controls 18.3 (SD=8.9) p=0.75	No significant difference in number of missing teeth between cases and controls	Medium
de Souza Rolim et al 2014 10	Case-control Country: Brazil	<b>Cases:</b> 29 patients with mild AD from Cognitive Neurology and Behaviour Group of the Neurology Department of the School of Medicine of the University of Sao Paulo, Brazil  <b>Controls:</b> 30 elderly without AD (relatives of the patients: wives, husbands, brothers, and/or sisters) Mix of dentate (36) and edentate (23) patients	<b>Difference in dental characteristics between case and control groups</b> Completely edentulous (% of group); case 32.3% c.f. control 43.3% p=0.614	Patients and controls had no statistical difference in edentulism	Low
Hatipoglu et al 2011 18	Case-control Country: Turkey	<b>Cases:</b> patients with Alzheimer's disease (n=31)  <b>Controls:</b> patients of a similar age and gender without known systemic disorders (n=47) Cases and controls were from the Neurology Department of an Institute in Turkey.	<b>1. DMFT score</b> Cases $24.10 \pm 6.8$ c.f. controls $19.68 \pm 9.5$ p = 0.126  <b>2. Mean number of present teeth</b> Cases $5.07$ (SD $\pm 7.7$ ) c.f. controls $10.55$ (SD $\pm 10.6$ ) p=0.060  <b>3. Correlations between clinical features and oral findings in the study groups</b> MMSE and PT 0.011 (correlations were significant at the 0.05 and 0.01 level (one-tailed) with Pearson correlation test)	In this study, teeth present (PT) was found to be positively correlated with MMSE scores in AD patients. In our study, we observed similar oral and dental findings in the patients with AD and control group. Groups were not comparable for age and gender. We would like to conclude that the relation between dementia or AD and dental health should be further evaluated.	Medium
Chen et al 2010 22	Retrospective longitudinal cohort study Country: United States of America	491 older adults who received dental care as new patients from the Wilder Senior Dental Clinic	<b>1. Number of teeth at baseline</b> Dementia; $19.6 \pm SD 7.4$ (range 2-31) Without dementia; $18.1 \pm SD 7.7$ (range 2-30); p=0.06  <b>2. Number of total teeth lost according to dementia status (cases c.f. controls)</b> <i>Zero teeth lost; 71.4% c.f. 73.1%, one tooth lost; 13.4% c.f. 15.6% two teeth lost; 4.2% c.f. 4.8%, three-four teeth lost; 5.9% c.f. 2.4%, five to six teeth lost; 2.5% c.f. 1.6%, seven to eight teeth lost; 1.7% c.f. 1.6%, edentulous; 0.8% c.f. 0.8%; p=0.61</i>	Based on data available in a community-based geriatric dental clinic, dementia was not associated with tooth loss. Although their oral health was poor at arrival, participants with dementia maintained their dentition as well as participants without dementia	High

			<p><b>3. Percentage of participants with tooth loss events during follow-up according to number of teeth at arrival</b></p> <p>i. 1-6 teeth arrival Dementia; 5.9% c.f. Without dementia; 5%</p> <p>ii. 7-12 teeth Dementia; 20.6% c.f. Without dementia; 15.0%</p> <p>iii. 13-18 teeth Dementia; 23.5% c.f. Without dementia; 26%</p> <p>iv. 19-24 teeth Dementia; 26.5% c.f. Without dementia; 33%</p> <p>v. 25-32 teeth Dementia; 23.5% c.f. Without dementia; 21%</p> <p><b>4. Comparison of tooth loss events per 100 patient years</b> Dementia; 14.9 (Standard error, SE=2.04) CI(11.4-19.5) Without dementia; 14.9 (SE=1.36) CI(12.4-17.8) Fully adjusted p=0.52</p> <p><b>5. Comparison of number of teeth lost per patient per 5 years</b> Dementia; 1.21 (SE=0.25) CI(0.80-1.82) Without dementia; 1.01 (SE=0.15) CI(0.76-1.34) Fully adjusted p=0.19</p>	<p>when dental treatment was provided.</p> <p>Participants with and without dementia did not differ significantly in the association between tooth loss and number of teeth remaining at arrival, although all participants with more teeth remaining at arrival had a higher risk of losing teeth during follow-up (p&lt;0.001).</p>	
Adam and Preston 2006 19	Case-control Country: United Kingdom	<p>135 individuals, aged 65 years and over, residing in four nursing homes in Cheshire, UK</p> <p><b>Cases:</b> 81 people with moderate to severe dementia</p> <p><b>Controls:</b> 54 people with no or mild dementia</p>	<p><b>1. Mean number of missing teeth (MT)</b> No/mild dementia 28.22 (SD=6.64), Moderate/severe dementia 27.28 (SD=7.73)</p> <p>Total cohort = 135 Number of edentulous subjects = 89/135 No/mild dementia and edentulous = 38/89 Moderate/severe dementia and edentulous = 51/89</p> <p>Number of subjects with some natural teeth = 46/135 Average number of teeth present was 11.7 in both groups.</p>	<p>When considering the teeth present or absent for the two populations, it was shown that most of the individuals in each group had a large number of teeth missing. There were no statistical differences between the two groups in terms of the oro-dental parameters used, except that for denture stability.</p>	Low

E. Dementia impact on dentures					
Author and year	Study type	Population	Results	Authors' conclusions	Quality
de Souza Rolim et al 2014 10	Case-control Country: Brazil	<p><b>Cases:</b> 29 patients with mild AD from Cognitive Neurology and Behaviour Group of the Neurology Department of the School of Medicine of the University of Sao Paulo, Brazil</p> <p><b>Controls:</b> 30 elderly without AD (relatives of the patients: wives, husbands, brothers, and/or sisters) Mix of dentate (36) and edentate (23) patients</p>	<p><b>1. Difference in dental characteristics between case and control groups</b> Complete Denture lower and upper (% of group); Control=43.3% Case=25.8%; p=0.334</p>	No significant differences in the wearing of denture between patients with or without (mild) dementia	Low
Hatipoglu et al 2011 18	Prospective Case-control Country:	<p><b>Cases;</b> Patients with AD (31)</p> <p><b>Controls;</b> (47) matched for similar age and gender without known systemic disorders</p> <p>Cases and control from Neurology Department of the Institution.</p>	<p><b>1. Presence of maxillary denture</b> Cases; 30/31 (97%) c.f. Controls; 27/47 (57%); p= 0.348</p> <p><b>2. Presence of mandibular denture</b> Cases: 31/31 (100%) c.f. Controls; 26/47 (55%); p= 0.291</p> <p><b>3. Maxillary denture status (cases c.f. controls)</b> i. No denture; 1/31 (3.2%) c.f. 4 (8.5%); p&gt;0.05 ii. Denture owned but not worn 6/31 (19.3%) c.f. 15 (31.9%); p&gt;0.05 iii. Fixed prosthesis 2/31 (6.5%) c.f. 1 (2.1%); p&gt;0.05 iv. Complete prosthesis 17/31 (54.8%) c.f. 24 (51.1%); p&gt;0.05 v. Removable partial prosthesis 5/31 (16.1%) c.f. 3 (6.4%); p&gt;0.05 vi. Not removing maxillary denture at night 15/22 (68.2%) c.f. 6/27 (22.2%); p=0.001 vii. Denture-related stomatitis 13/22 (59.1%) c.f. 2/27 (7.4%) p= 0.001</p> <p><b>4. Mandibular denture status (cases c.f. controls)</b> i. No denture 0/31 (-) c.f. 5 (10.1%); p&gt;0.05 ii. Denture owned but not worn 8/31 (25.8%) c.f. 14 (29.8%); p&gt;0.05 iii. Fixed prosthesis 1/31 (3.2%) c.f. 2 (4.3%); p&gt;0.05 iv. Complete prosthesis 14/31 (45.2%) c.f. 20 (42.6%); p&gt;0.05 v. Removable partial prosthesis 8/31 (25.8%) c.f. 6 (12.8%); p&gt;0.05 vi. Not removing mandibular denture at night 14/22 (63.6%) c.f. 6/26 (23.0%); p=0.001 vii. Denture-related stomatitis 4/22 (18.1%) c.f. 0/26 (-); p=0.001</p> <p><b>5. Correlation between clinical and oral findings</b> MMSE scores were significantly correlated with denture removal at night (DRN) and related stomatitis p&lt;0.001</p>	<p>Oral hygiene status is closely related with cognitive functions of the patients with AD.</p> <p>Especially, denture-related problems may be obvious due to impaired denture care in these patients</p> <p><i>Oral hygiene status and maxillary and mandibular denture status were similar in the patients with AD and in the control group. However, the patients with AD were found to have denture-related stomatitis due to the lack of denture removal at night</i></p>	Medium

<p>Adam and Preston 2006 19</p>	<p>Case-control Country: United Kingdom</p>	<p>135 individuals, aged 65 years and over, residing in four nursing homes in Cheshire, UK <b>Cases:</b> 81 people with moderate to severe dementia  <b>Controls:</b> 54 people with no or mild dementia</p>	<p><b>Satisfactory denture characteristics</b> i. Denture stability satisfactory No/mild dementia 93% Moderate/severe dementia 76% p=0.04 ii. Denture retention satisfactory No/mild dementia 76.7% Moderate/severe dementia 76%; iii. Occlusion satisfactory No/mild dementia 91.4% Moderate/severe dementia 66.7%; iv. Vertical height of complete dentures satisfactory No/mild dementia; 82.1%, Moderate/severe dementia; 71.4%</p>	<p>If the wearing of dentures in edentulous individuals is considered, 60% of those with moderate to severe dementia wore no dentures at all compared with only 10% of those with no or mild dementia. The figures for the no/mild dementia group are comparable with other studies for rates of denture wear. It could be suggested that the high rate of not wearing dentures in the moderate/severe dementia group may in part be due to the dementia itself.</p>	<p>Low</p>
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**Table 1: Cohort and case-control studies: Is dementia (inc. cognitive decline) a risk factor for poor oral health?**

Author and year	Study type	Population	Results	Authors' conclusions	Quality
<b>A. Dementia impact on plaque, gingival bleeding and gingivitis</b>					
Chu et al 2015 16	Case-control  Country: Hong Kong	<b>Cases:</b> 59 patients in day care centres of Hong Kong Alzheimer's Disease Association. All with mild level late-onset Alzheimer's disease. All $\geq 60$ years of age.  <b>Controls:</b> 59 healthy controls without dementia and matched for age and gender, from Prince Philip Dental Hospital (not under current dental treatment).	<b>1. Percentage of people performing brushing twice daily</b> Dementia (67%); control (83%); $p=0.045$ .  <b>2. Percentage of people receiving assistance when brushing</b> Dementia (31%) control (5%); $p<0.001$  <b>3. Percentage of people with reversible gingivitis (as indicated by a community periodontal index score maximum of 1)</b> Dementia (N = 5/47) 11%; Control (N = 7/50) 14%  <b>4. Assistance with toothbrushing</b> Dementia (31%); Control (5%); $p<0.001$	Compared to those without dementia, fewer elderly Chinese with dementia practiced tooth brushing twice daily <i>And they were more likely to have assistance with toothbrushing</i>	Medium
Gil-Montoya et al 2015 17	Case-control  Country: Spain	<b>Cases:</b> Patients treated in neurology departments of two hospitals, who had a firm diagnosis of AD, Dementia, or MCI (mild cognitive impairment)  <b>Controls:</b> patients being seen in primary healthcare, for problems other than dental or neurological problems. Excluded schizophrenic, depressive, personality disorder, diseases not under treatment, drug abuse, and having previous periodontal treatment in the last 6 months	<b>1. OR of cognitive impairment in relation to oral hygiene habits</b> (reference group those who brushed twice or more a day) i. mouthwash only OR=11:37 CI(5.46 - 23.68) $p<0.001$ ii. No brushing OR=7.21 CI(3.48 - 14.96) iii. Brushing once a day OR=3.34 CI(2.03 - 5.49)  <b>2. OR of cognitive impairment in relation to dental variables</b> i. Plaque index (reference group; 0.00 - 1.00). 1.01 to 2.00; OR=1.92 CI(0.94 - 3.90) 2.01 to 2.50=6.33 CI(3.15 - 12.72) 2.51 to 3.00=15.70 CI(7.72 - 31.92) ii. Bleeding index (reference group was 0-25% BOP) 25.1 to 50% BOP OR=2.32 CI(1.21 - 4.48) 50.1 to 90% BOP OR=2.67 CI (1.47 - 4.85) 90.1 to 100% BOP OR=3.53 CI (1.88 - 6.63)  <b>3. Association between dental variables and levels of cognitive impairment</b> i. Mean Plaque index MCI; 2.26.(SD=0.49) Mild Dementia; 2.27 (SD=0.76) Moderate Dementia; 2.4 (SD 0.57) Severe Dementia 2.5 (SD=0.62) $p=0.18$  ii. Mean Bleeding index MCI; 45.4 (SD=20.9) Mild Dementia; 62 (SD=29.5) Moderate Dementia; 61.5 (SD=34.2) Severe Dementia; 77.6 (SD: 27.3) $p=0.001$	Periodontitis appears to be associated with cognitive impairment after controlling for age, sex, education level and oral hygiene habits  <i>Average plaque and bleeding scores were significantly higher in the patients with a diagnosis of AD, Dementia or MCI</i>  <i>There is evidence of increased bleeding score with cognitive decline; however, there wasn't significantly more plaque present with cognitive decline</i>  <i>The present results evidence a significant worsening of oral hygiene habits (reduction or cessation of toothbrushing) with more advanced cognitive disease. The direction of the association is yet to be established</i>	High



			<p>iii. Oral hygiene habits Brushed twice or more a day cf brushed once a day (n) cf used only mouthwash cf no oral hygiene habits (n) in relation to MCI, mild-moderate- and severe-dementia; p&lt;0.001</p> <p><b>4. Mean plaque scores (PI)</b> Mean plaque 2.37 (SD =0.65) cases cf 1.55 (SD = 0.89) controls p&lt;0.001.</p> <p><b>5. Mean bleeding scores (BI)</b> Mean bleeding scores 63 (SD = 31.1) cases cf 50.6 (SD = 34.2) controls p&lt;0.001.</p>		
de Souza Rolim et al 2014  10	Case-control  Country: Brazil	<p><b>Cases:</b> 29 patients with mild AD from Cognitive Neurology and Behaviour Group of the Neurology Department of the School of Medicine of the University of Sao Paulo, Brazil</p> <p><b>Controls:</b> 30 elderly without AD (relatives of the patients: wives, husbands, brothers, and/or sisters) Mix of dentate (36) and edentate (23) patients</p>	<p><b>1. Difference in dental characteristics between case and control groups</b> (AAP Definitions of periodontal disease) <i>Gingivitis:</i> Case 31% c.f. Control =10% <i>Moderate perio disease:</i> Case=6.9% c.f. Control =10% <i>Severe perio disease:</i> Case=20.7% c.f. Control =6.7% p=0.002</p>	Periodontal infections were more frequent in patients with mild AD than in healthy subjects	Low
Hatipoglu et al 2011  18	Case-control  Country: Turkey	<p><b>Cases:</b> patients with Alzheimer's disease (n=31)</p> <p><b>Controls:</b> patients of a similar age and gender without known systemic disorders (n=47) Cases and controls were from the Neurology Department of an Institute in Turkey.</p>	<p><b>1. Oral hygiene status</b> cases c.f. control patients: Good; 1 (3.2%) c.f. 9 (19.1%) p&gt;0.510 Fair; 9 (29.0%) c.f. 15 (31.9%) p&gt;0.510 Poor; 21 (67.7%) c.f. 23 (48.9%) p&gt;0.510</p> <p><b>2. Correlation between clinical and oral findings:</b> MMSE scores were significantly correlated with oral hygiene status p=0.015</p>	The results suggest that cognitive status (MMSE) of patients is significantly correlated with oral hygiene status	Medium

<p>Adam and Preston 2006 19</p>	<p>Case-control Country: United Kingdom</p>	<p>135 individuals, aged 65 years and over, residing in four nursing homes in Cheshire, UK <b>Cases:</b> 81 people with moderate to severe dementia <b>Controls:</b> 54 people with no or mild dementia</p>	<p><b>1. Mean Plaque score</b> Greene and Vermillion index: i. All surfaces; No/mild dementia 1.33 (SD=0.60), Moderate/severe dementia 2.14 (SD=0.73) ii. Buccal surfaces, No/mild dementia 1.42 (SD=0.72), Moderate/severe dementia 2.14 (SD=0.73) iii. Lingual surfaces, No/mild dementia 1.24 (SD=0.47), Moderate/severe dementia 2.14 (SD=0.72) <b>2. Mean calculus score</b> Greene and Vermillion index i. All surfaces; No/mild dementia 1.34 (SD=0.60), Moderate/severe dementia 2.04 (SD=0.75) ii. Buccal surfaces; No/mild dementia 1.38 (SD=0.69), Moderate/severe dementia 2.05 (SD=0.76) iii. Lingual surfaces, No/mild dementia 1.29 (SD=0.51), Moderate/severe dementia 2.04 (SD=0.73) Note. No evidence of significance provided</p>	<p>For individuals' resident in nursing homes, moderate to severe dementia might have a deleterious effect on oral health (plaque and calculus).</p>	<p>Low</p>
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B. Dementia impact on periodontitis					
Author and year	Study type	Population	Results	Authors' conclusions	Quality
Gil-Montoya et al 2015  17	Case-control  Country: Spain	<p><b>Cases:</b> Patients treated in a behavioural and cognitive neurology department, with a firm diagnosis of AD, Dementia, or MCI (n=180)</p> <p><b>Controls:</b> patients being seen in primary healthcare, for problems other than dental or neurological problems (n=229). Excluded schizophrenic, depressive, personality disorder, diseases not under treatment, drug abuse, and previous periodontal treatment in the last 6 months</p>	<p><b>1. OR of cognitive impairment in relation to dental variables</b> (Periodontitis: % of sites with AL &gt;3mm where moderate 33-66%; severe 67-100%) (reference group absent/mild periodontitis) Moderate; OR=2.58 (1.31 - 5.09) Severe; OR=3.04 (1.69 - 5.46) p&lt;0.001 <i>Adjusted for age, sex, present teeth, OH habits, hyperlipidaemia, studies:</i> Moderate; OR=2.64 (1.18-5.92) Severe; OR=2.31: (1.15-4.66) p=0.04</p> <p><b>2. Association between dental variables and levels of cognitive impairment</b></p> <p>i. Mean Pocket Depth; MCI; 2.8 (S.D=0.5) Mild Dementia; 3.0 (S.D=0.6) Moderate Dementia; 3.1 (S.D=0.6) Severe Dementia; 2.0 (S.D=0.9) p=0.47</p> <p>ii. Mean Attachment loss (mm) MCI; 4.6(S.D=1.0) Mild Dementia; 4.8 (S.D=1.4) Moderate Dementia 5.1 (S.D=2.0) Severe Dementia 4.7 (S.D=1.3) p=0.42</p> <p>iii. Attachment loss (percentage &gt;3mm) MCI; 75.2 (S.D=19.8) Mild Dementia; 76.6 (S.D=27.4) Moderate Dementia; 73.3 (S.D=33.8) Severe Dementia; 75.0 (S.D=27.1) p=0.94</p> <p><b>iv. Severe periodontitis (% with 67-100% AL &gt;3mm)</b> MCI; 66.7% Mild Dementia; 67.7% Moderate Dementia; 70.5% Severe Dementia; 66.7 p=0.98</p> <p><b>3. Adjusted OR of cognitive impairment in relation to periodontitis</b> <i>Periodontitis severity; reference group absent/mild periodontitis;</i> i. Moderate periodontitis OR=2.64 CI(1.18 - 5.92), ii. Severe periodontitis OR=2.31 CI(1.15 - 4.66) p=0.04</p>	Periodontitis appears to be associated with cognitive impairment after controlling for confounders such as age, sex, and education level	High

<p>Chu et al 2015 16</p>	<p>Case-control Country: Hong Kong</p>	<p><b>Cases:</b> 59 Patients in day care centres of Hong Kong Alzheimer's Disease Association  <b>Controls:</b> 59 generally healthy controls matched for age and gender, from Prince Philip Dental Hospital (not under current dental treatment)</p>	<p><b>1. Prevalence of advanced periodontal disease (CPI of 3 or more)</b> Dementia group=78% c.f. Control group=74%; p= 0.64</p>	<p>Compared to those without dementia, fewer elderly Chinese with dementia practiced tooth brushing twice daily <i>And they were more likely to have assistance with toothbrushing</i></p>	<p>Medium</p>
<p>de Souza Rolim et al 2014 10</p>	<p>Case-control Country: Brazil</p>	<p><b>Cases:</b> 29 patients from Cognitive Neurology and Behavior Group of the Neurology Department of the School of Medicine of the University of Sao Paulo, Brazil with mild AD  <b>Controls:</b> 30 elderly people without AD (relatives of the patients: wives, husbands, brothers, and/or sisters) Mix of dentate (36) and edentate (23) patients</p>	<p><b>1. Difference in periodontal disease between case and control groups</b> (AAP Definitions of periodontal disease) <i>Gingivitis:</i> Case 31% c.f. Control =10% <i>Moderate perio disease:</i> Case=6.9% c.f. Control =10% <i>Severe perio disease:</i> Case=20.7% c.f. Control =6.7% p=0.002</p>	<p>Periodontal infections were more common in patients with mild AD than in healthy subjects</p>	<p>Low</p>

C. Dementia impact on caries and salivary flow					
Author and year	Study type	Population	Results	Authors' conclusions	Quality
Chu et al 2015  16	Case-control	<p><b>Cases:</b> 59 Patients in day care centres of Hong Kong Alzheimer's Disease Association with dementia.</p> <p><b>Controls:</b> 59 generally healthy controls matched for age and gender, from Prince Philip Dental Hospital (not under current dental treatment).</p>	<p><b>1. Mean DMFT, DT, FT of cases vs controls</b> DMFT cases 22.3 (SD=8.2), controls 21.5 (SD=8.2); p=0.59 DT cases 1.2 (SD=1.9), controls 0.8 (SD=1.4); p=0.28 FT cases 2.5 (SD=3.3), controls 2.4 (SD=2.5); p=0.88</p> <p><b>2. Unstimulated salivary flow rate (ml/min)</b> Dementia group - 0.3 (SD=0.17) c.f. control group - 0.41 (SD=0.28); p=0.0043</p>	No significant difference in dental caries experience (DMFT) between AD patients and controls. Resting salivary secretion was reduced in dementia patients; however, their caries experience was not significantly different	Medium
Chen et al 2013  21	Retrospective longitudinal cohort study  Country: United States of America	<p>491 older adults who received dental care as new patients from the Wilder Senior Dental Clinic</p> <p>119 with diagnosis of Dementia, AD or chronic brain syndrome of no diagnosis</p> <p>- control group was not matched</p>	<p><b>1. Number of decayed and broken teeth at baseline</b> Dementia; 4.2 ± SD 4.5 (range 0-20) Without dementia; 3.1 ± SD 3.8 (range 0-18) p=0.006</p>	Patients with dementia present with higher rates of caries at baseline	High
Hatipoglu et al 2011  18	Prospective Case-control	<p><b>Cases:</b> Patients with AD (31)</p> <p><b>Controls:</b> (47) matched for similar age and gender without known systemic disorders</p> <p>Cases and control from Neurology Department of the Institution.</p>	<p><b>1. Mean DMFT score (case c.f. control)</b> 24.19 (SD=6.8) c.f. 19.68 (SD=9.5); p= 0.126</p> <p><b>2. Correlation between clinical and oral findings</b> MMSE scores were significantly correlated with DMFT; p=0.022</p>	<p>No significant difference in dental caries experience (DMFT) between patients with AD and the control group</p> <p>Decreased MMSE scores were significantly correlated with increased DMFT.</p>	Medium

<p>Ellefson et al 2009</p> <p>20</p>	<p>Cohort</p>	<p>77 patients with Alzheimer's disease (AD), other dementia (OD), no dementia (ND) referred to the memory clinics of two hospitals in Copenhagen, Denmark</p>	<p>Crude caries (coronal and root) increments (CCI), net caries increments (NCI), and adjusted caries and filling increments (ADJCI), for AD, OD and ND patients</p> <p><b>1. Change in average number of decayed surfaces (DS) in first year after referral to memory clinic</b>  All patients: increased from 5.7 (SD=6.7) to 7.9 (SD=8.8); <math>p \leq 0.005</math>  OD: increased from 5.0 (SD=6.6) to 10.2 (SD=9.8); <math>p \leq 0.005</math>  AD: increased from 6.7 (SD=7.2) to 8.0 (SD=9.2) NS  ND but referred to memory clinic: increased from 2.8 (SD 3.0) to 4.8 (SD 5.1) NS</p> <p><b>2. Adjusted OR of ADJCI after one year</b>  <i>(reference group; no dementia)</i>  OR of having a higher coronal caries increment (&gt;2) in OD was 4.13 CI(0.44-39.05) c.f. ND  OR of having a higher coronal caries increment (&gt;2) in AD was 0.64 CI(0.09-4.5) c.f. ND</p>	<p>Elderly people referred to a memory clinic were at an elevated risk of developing high levels of coronal and root surface caries during the first year after referral (independent of whether or not they fulfilled the criteria for a dementia diagnosis), and that elderly people with a dementia diagnosis other than AD appeared to be at a particularly high risk of developing multiple carious lesions during the first year after diagnosis.</p>	<p>Medium</p>
<p>Adam and Preston 2006</p> <p>19</p>	<p>Case-control</p>	<p>135 individuals, aged 65 years and over, residing in four nursing homes in Cheshire, UK  <b>Cases:</b> 81 people with moderate to severe dementia  <b>Controls:</b> 54 people with no or mild dementia</p>	<p><b>1. Mean number decayed teeth DT</b>  No/mild dementia; 0.11 (SD=3.42); Moderate/severe dementia; 0.80 (SD=1.87) NS</p> <p><b>2. Mean number of missing teeth MT</b>  No/mild dementia 28.22 (SD=6.64), Moderate/severe dementia 27.28 (SD=7.73) NS</p> <p><b>3. Mean number of filled teeth</b>  No/mild dementia 0.69 (SD=1.30); Moderate/severe dementia 0.90 (SD=2.39) NS</p>	<p>In an institutionalised population there were no statistical differences between the two groups (no/mild dementia and moderate/severe dementia) in terms of the DMFT</p>	<p>Low</p>

D. Dementia impact on tooth loss/number					
Author and year	Study type	Population	Results	Authors' conclusions	Quality
Chu et al 2015 16	Case-control Country: Hong Kong	<b>Cases:</b> 59 patients in day care centres of Hong Kong Alzheimer's Disease Association. All with mild level late-onset Alzheimer's disease. All $\geq 60$ years of age.  <b>Controls:</b> 59 healthy controls without dementia and matched for age and gender, from Prince Philip Dental Hospital (not under current dental treatment).	<b>Missing teeth, cases c.f. controls</b> Cases 18.9 (SD=9.4), controls 18.3 (SD=8.9) p=0.75	No significant difference in number of missing teeth between cases and controls	Medium
de Souza Rolim et al 2014 10	Case-control Country: Brazil	<b>Cases:</b> 29 patients with mild AD from Cognitive Neurology and Behaviour Group of the Neurology Department of the School of Medicine of the University of Sao Paulo, Brazil  <b>Controls:</b> 30 elderly without AD (relatives of the patients: wives, husbands, brothers, and/or sisters) Mix of dentate (36) and edentate (23) patients	<b>Difference in dental characteristics between case and control groups</b> Completely edentulous (% of group); case 32.3% c.f. control 43.3% p=0.614	Patients and controls had no statistical difference in edentulism	Low
Hatipoglu et al 2011 18	Case-control Country: Turkey	<b>Cases:</b> patients with Alzheimer's disease (n=31)  <b>Controls:</b> patients of a similar age and gender without known systemic disorders (n=47) Cases and controls were from the Neurology Department of an Institute in Turkey.	<b>1. DMFT score</b> Cases $24.10 \pm 6.8$ c.f. controls $19.68 \pm 9.5$ p = 0.126  <b>2. Mean number of present teeth</b> Cases $5.07$ (SD $\pm 7.7$ ) c.f. controls $10.55$ (SD $\pm 10.6$ ) p=0.060  <b>3. Correlations between clinical features and oral findings in the study groups</b> MMSE and PT 0.011 (correlations were significant at the 0.05 and 0.01 level (one-tailed) with Pearson correlation test)	In this study, teeth present (PT) was found to be positively correlated with MMSE scores in AD patients. In our study, we observed similar oral and dental findings in the patients with AD and control group. Groups were not comparable for age and gender. We would like to conclude that the relation between dementia or AD and dental health should be further evaluated.	Medium
Chen et al 2010 22	Retrospective longitudinal cohort study Country: United States of America	491 older adults who received dental care as new patients from the Wilder Senior Dental Clinic	<b>1. Number of teeth at baseline</b> Dementia; $19.6 \pm SD 7.4$ (range 2-31) Without dementia; $18.1 \pm SD 7.7$ (range 2-30); p=0.06  <b>2. Number of total teeth lost according to dementia status (cases c.f. controls)</b> <i>Zero teeth lost; 71.4% c.f. 73.1%, one tooth lost; 13.4% c.f. 15.6% two teeth lost; 4.2% c.f. 4.8%, three-four teeth lost; 5.9% c.f. 2.4%, five to six teeth lost; 2.5% c.f. 1.6%, seven to eight teeth lost; 1.7% c.f. 1.6%, edentulous; 0.8% c.f. 0.8%; p=0.61</i>	Based on data available in a community-based geriatric dental clinic, dementia was not associated with tooth loss. Although their oral health was poor at arrival, participants with dementia maintained their dentition as well as participants without dementia	High

			<p><b>3. Percentage of participants with tooth loss events during follow-up according to number of teeth at arrival</b></p> <p>i. 1-6 teeth arrival Dementia; 5.9% c.f. Without dementia; 5%</p> <p>ii. 7-12 teeth Dementia; 20.6% c.f. Without dementia; 15.0%</p> <p>iii. 13-18 teeth Dementia; 23.5% c.f. Without dementia; 26%</p> <p>iv. 19-24 teeth Dementia; 26.5% c.f. Without dementia; 33%</p> <p>v. 25-32 teeth Dementia; 23.5% c.f. Without dementia; 21%</p> <p><b>4. Comparison of tooth loss events per 100 patient years</b> Dementia; 14.9 (Standard error, SE=2.04) CI(11.4-19.5) Without dementia; 14.9 (SE=1.36) CI(12.4-17.8) Fully adjusted p=0.52</p> <p><b>5. Comparison of number of teeth lost per patient per 5 years</b> Dementia; 1.21 (SE=0.25) CI(0.80-1.82) Without dementia; 1.01 (SE=0.15) CI(0.76-1.34) Fully adjusted p=0.19</p>	<p>when dental treatment was provided.</p> <p>Participants with and without dementia did not differ significantly in the association between tooth loss and number of teeth remaining at arrival, although all participants with more teeth remaining at arrival had a higher risk of losing teeth during follow-up (p&lt;0.001).</p>	
Adam and Preston 2006 19	Case-control Country: United Kingdom	<p>135 individuals, aged 65 years and over, residing in four nursing homes in Cheshire, UK</p> <p><b>Cases:</b> 81 people with moderate to severe dementia</p> <p><b>Controls:</b> 54 people with no or mild dementia</p>	<p><b>1. Mean number of missing teeth (MT)</b> No/mild dementia 28.22 (SD=6.64), Moderate/severe dementia 27.28 (SD=7.73)</p> <p>Total cohort = 135 Number of edentulous subjects = 89/135 No/mild dementia and edentulous = 38/89 Moderate/severe dementia and edentulous = 51/89</p> <p>Number of subjects with some natural teeth = 46/135 Average number of teeth present was 11.7 in both groups.</p>	<p>When considering the teeth present or absent for the two populations, it was shown that most of the individuals in each group had a large number of teeth missing. There were no statistical differences between the two groups in terms of the orodental parameters used, except that for denture stability.</p>	Low



E. Dementia impact on dentures					
Author and year	Study type	Population	Results	Authors' conclusions	Quality
de Souza Rolim et al 2014 10	Case-control Country: Brazil	<p><b>Cases:</b> 29 patients with mild AD from Cognitive Neurology and Behaviour Group of the Neurology Department of the School of Medicine of the University of Sao Paulo, Brazil</p> <p><b>Controls:</b> 30 elderly without AD (relatives of the patients: wives, husbands, brothers, and/or sisters) Mix of dentate (36) and edentate (23) patients</p>	<p><b>1. Difference in dental characteristics between case and control groups</b> Complete Denture lower and upper (% of group); Control=43.3% Case=25.8%; p=0.334</p>	No significant differences in the wearing of denture between patients with or without (mild) dementia	Low
Hatipoglu et al 2011 18	Prospective Case-control Country:	<p><b>Cases;</b> Patients with AD (31)</p> <p><b>Controls;</b> (47) matched for similar age and gender without known systemic disorders</p> <p>Cases and control from Neurology Department of the Institution.</p>	<p><b>1. Presence of maxillary denture</b> Cases; 30/31 (97%) c.f. Controls; 27/47 (57%); p= 0.348</p> <p><b>2. Presence of mandibular denture</b> Cases: 31/31 (100%) c.f. Controls; 26/47 (55%); p= 0.291</p> <p><b>3. Maxillary denture status (cases c.f. controls)</b> i. No denture; 1/31 (3.2%) c.f. 4 (8.5%); p&gt;0.05 ii. Denture owned but not worn 6/31 (19.3%) c.f. 15 (31.9%); p&gt;0.05 iii. Fixed prosthesis 2/31 (6.5%) c.f. 1 (2.1%); p&gt;0.05 iv. Complete prosthesis 17/31 (54.8%) c.f. 24 (51.1%); p&gt;0.05 v. Removable partial prosthesis 5/31 (16.1%) c.f. 3 (6.4%); p&gt;0.05 vi. Not removing maxillary denture at night 15/22 (68.2%) c.f. 6/27 (22.2%); p=0.001 vii. Denture-related stomatitis 13/22 (59.1%) c.f. 2/27 (7.4%) p= 0.001</p> <p><b>4. Mandibular denture status (cases c.f. controls)</b> i. No denture 0/31 (-) c.f. 5 (10.1%); p&gt;0.05 ii. Denture owned but not worn 8/31 (25.8%) c.f. 14 (29.8%); p&gt;0.05 iii. Fixed prosthesis 1/31 (3.2%) c.f. 2 (4.3%); p&gt;0.05 iv. Complete prosthesis 14/31 (45.2%) c.f. 20 (42.6%); p&gt;0.05 v. Removable partial prosthesis 8/31 (25.8%) c.f. 6 (12.8%); p&gt;0.05 vi. Not removing mandibular denture at night 14/22 (63.6%) c.f. 6/26 (23.0%); p=0.001 vii. Denture-related stomatitis 4/22 (18.1%) c.f. 0/26 (-); p=0.001</p> <p><b>5. Correlation between clinical and oral findings</b> MMSE scores were significantly correlated with denture removal at night (DRN) and related stomatitis p&lt;0.001</p>	<p>Oral hygiene status is closely related with cognitive functions of the patients with AD.</p> <p>Especially, denture-related problems may be obvious due to impaired denture care in these patients</p> <p><i>Oral hygiene status and maxillary and mandibular denture status were similar in the patients with AD and in the control group. However, the patients with AD were found to have denture-related stomatitis due to the lack of denture removal at night</i></p>	Medium

<p>Adam and Preston 2006 19</p>	<p>Case-control Country: United Kingdom</p>	<p>135 individuals, aged 65 years and over, residing in four nursing homes in Cheshire, UK <b>Cases:</b> 81 people with moderate to severe dementia  <b>Controls:</b> 54 people with no or mild dementia</p>	<p><b>Satisfactory denture characteristics</b> i. Denture stability satisfactory No/mild dementia 93% Moderate/severe dementia 76% p=0.04 ii. Denture retention satisfactory No/mild dementia 76.7% Moderate/severe dementia 76%; iii. Occlusion satisfactory No/mild dementia 91.4% Moderate/severe dementia 66.7%; iv. Vertical height of complete dentures satisfactory No/mild dementia; 82.1%, Moderate/severe dementia; 71.4%</p>	<p>If the wearing of dentures in edentulous individuals is considered, 60% of those with moderate to severe dementia wore no dentures at all compared with only 10% of those with no or mild dementia. The figures for the no/mild dementia group are comparable with other studies for rates of denture wear. It could be suggested that the high rate of not wearing dentures in the moderate/severe dementia group may in part be due to the dementia itself.</p>	<p>Low</p>
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**Table 3 Systematic review on tooth loss and periodontal disease and cognitive impairment**

Author, date and title	Populations included in studies reviewed	Studies Types in Review	Oral disease and diag criteria or oral disease intervention and description	Cognitive diagnostic criteria of	Intervention, comparison: Including covariates adjusted for	Results	Author's conclusions	Amstar Score and Justification for quality score
<p>Cerutti-Kopplin et al., 2016</p> <p>Tooth loss increases the risk of diminished cognitive function: A systematic review and meta-analysis</p>	<p>Individuals aged ≥18 years with cognitive status/impairment as the outcome of interest.</p>	<p>10 cohort studies - 8 were included in meta-analysis</p>	<p><b>I)</b> Tooth loss primary oral health outcomes <b>II)</b> 5/10 looked at periodontal disease</p> <p>Tooth loss and periodontal disease were assessed by clinical examination/ full-mouth radiographs in (4/10) and by self-administrated questionnaires in (6/10)</p>	<p>Cognitive function was ascertained during follow-ups ranging from 4yrs to 32 yrs</p> <p>(4/10) measured and reported cognitive status at baseline and at the last follow-up</p> <p>(1/10) reported the cognitive status at each annual follow-up and (5/10) studies assessed this outcome every 2 yrs</p> <p>The assessment of cognitive function and diagnosis of cognitive impairment or dementia was based on the patient's medical record (3/10) or via clinical or standard neuropsychological examination by a neurologist, psychiatrist, or other health professional (7/10)</p> <p>Genetic predisposition to dementia (3/10) as well as depressive symptoms (3/10) were assessed</p>	<p>The relationship between periodontal disease and cognitive impairment ( 5/10)</p> <p>The relationship between tooth loss and cognitive impairment (10/10)</p>	<p>The relationship between periodontal disease and cognitive impairment shows conflicting results</p> <p>Random effects analysis showed, with statistically low heterogeneity, that individuals with suboptimal dentition (&lt;20 teeth) were at a 20% higher risk for developing cognitive decline (HR = 1.26, 95% CI = 1.14 to 1.40) and dementia (HR = 1.22, 95% CI = 1.04 to 1.43) than those with optimal dentition (≥20 teeth) and dementia</p>	<p>Studies on the association between periodontal disease and cognitive status showed conflicting results. Within the limits of the quality of published evidence, this meta-analysis lends further support to the hypothesis that tooth loss is associated with an increased risk of cognitive impairment</p>	<p>11</p> <p>High All Amstar questions were satisfied</p>

<p>Wu et al .,2016 Association between oral health and cognitive status: a systematic review</p>	<p>Older adults</p>	<p>16 longitudinal studies</p>	<p>Tooth loss is the primary oral health outcome (14/16) studies. Number of decayed, missing, filled teeth; plaque; periodontal disease (4/16) to minimal (self-reported approximate number of teeth currently or when younger) (4/16)</p> <p>Subject-provided information included pain or discomfort (1/16) bleeding gums (14/16) and denture use (3/16)</p> <p>(2/16) relied on up to 40 years of records</p> <p>(9/16) studies used oral health evaluation and (7/16) dental records</p> <p>In examining periodontal disease, attention was paid to extent of gingival bleeding on probing (ratio of examined sites), proportion of examined sites with loss of attachment greater than 3 mm, and mean pocket depth</p>	<p>The standard diagnostic criteria for dementia included Diagnostic and Statistical Manual of Mental Disorders, Third Edition, Revised (DSM-III-R), DSM-IV, International Classification of Diseases, Ninth Revision, Clinical Modification, Revised (ICD-9-CM-R), and ICD-10, reflecting the date of evaluation</p>	<p>The association between oral health and cognitive status</p>	<p>Some studies found that oral health measures, such as number of teeth and periodontal disease, were associated with risk of cognitive decline or incident dementia, whereas others did not find an association.</p> <p>Similarly, cognitive decline was not consistently associated with greater loss of teeth or number of decayed teeth. It is likely that methodological limitations play a major role in explaining the inconsistent findings.</p>	<p>It is unclear how or whether oral health and cognitive status are related. Additional research is needed in which there is greater agreement on how oral health and cognitive states are assessed to better examine the linkages between these two health outcomes.</p>	<p>11 High All Amstar questions were satisfied</p>
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