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# **INVITED MEDICAL REVIEW**

# Nutrition and health: guidelines for dental practitioners

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Good nutrition is vital to overall health, and poor diet and a sedentary lifestyle are major causes of morbidity and mortality worldwide. Nutritional factors are implicated in many oral and systemic diseases and conditions, including obesity, hypertension, dyslipidemia, type II diabetes, cardiovascular disease, osteoporosis, dental caries and some cancers including oral cancers. This review focuses on the evidence for the relations between key nutritional factors and health. Energy intake is related to body weight and obesity, highlighting the importance of lowerenergy diets and regular physical activity for body weight maintenance and for preventing obesity. Evidence is presented for the health benefits of high quality carbohydrates, such as whole grain products, and fruits and vegetables, in reducing the risk of cardiovascular disease and cancer. The adverse effects of sugar, sweetened beverages, and trans and saturated fats on several diseases including caries, diabetes and cardiovascular disease are described. The health benefits of unsaturated fats, antioxidants, B vitamins and vitamin D in cardiovascular disease, periodontitis, cancer, and other conditions are documented. Both benefits and harmful effects of dairy product intake on health are discussed. Based on the evidence, nutritional guidelines are provided, as well as key recommendations for preventing obesity. Dentists can play a critical role in motivating and enabling healthy food choices.

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# Dentists contribute to knowledge about nutrition and health

Over 29,000 dentists have been directly contributing to research on the relation of diet to health since 1986, through their participation in the Health Professional's Follow-up Study (HPFS) conducted at Harvard School of Public Health. Participants have completed detailed dietary questionnaires every four years along with other questionnaires about lifestyle and medical and oral health history, and many have contributed blood samples. This research has provided unique information on the prevention of heart disease, many cancers, oral precancer, periodontal disease, and other important illness. Several manuscripts using the HPFS data are referenced in this review article. More details of the study and description of the research findings can be found at the Health Professional's Follow-up Study website: http://www.hsph.harvard.edu/hpfs

# Introduction

Good nutrition is vital to good health. Poor diet and a sedentary lifestyle are major causes of morbidity and mortality worldwide. Nutritional factors are implicated in several oral and systemic diseases and conditions, including overweight and obesity, hypertension, dyslipidemia, type 2 diabetes, cardiovascular disease, osteoporosis, dental caries, gastrointestinal disorders and most cancers including oral cancers.

One of the main challenges in the prevention of most chronic diseases is reducing obesity, which is now recognized as one of the major public health problems worldwide. It is estimated that one billion individuals in the world are overweight and at least 300 millions are obese (World Health Organization (WHO) (2003). Obesity is the leading cause of many chronic conditions (National Institute of Health, National Heart, Lung, and Blood Institute, 1998), and the mortality rate is 2–3 times higher in the obese individuals compared with

normal weight adults (Adams *et al*, 2006). A healthy diet is crucial for preventing and reducing obesity. Lack of physical activity is a major determinant of obesity and also a risk factor for many conditions.

This manuscript summarizes the evidence for the role of physical activity and intake of total energy (calories), specific foods and micronutrients in promoting oral as well as overall health and in preventing chronic diseases.

# Key relations between nutrition and health

Most people consume a wide variety of foods, with a combination of many nutrients and the dietary patterns often change over time. This makes it challenging to ascertain the health impact of any one nutrient or food. However, several foods and nutrients have well-established relationships to specific conditions as described below.

# Energy intake, physical activity and obesity

Maintaining body weight within normal ranges is important for well being, as obesity is a risk factor for hypertension, type 2 diabetes, coronary heart disease (CHD), stroke, gallbladder disease, osteoarthritis, sleep apnea, respiratory problems and some types of cancer (including endometrial, breast, prostate, pancreatic, kidney and colon) (National Institute of Health, National Heart, Lung, and Blood Institute, 1998). Obesity is also related to several aspects of oral health, such as xerostomia, caries and periodontitis (Mathus-Vliegen et al, 2007). Adipose tissue is also a source of inflammatory cytokines. Hence, increasing body fat may increase the likelihood of an active host inflammatory response in periodontitis (Shuldiner et al, 2001). Several cross-sectional studies have shown associations between obesity and periodontitis (Saito et al, 1998; Ritchie et al, 2002; Pischon et al, 2007).

Physical activity also independently reduces the risk of several diseases including cardiovascular disease, type 2 diabetes mellitus, osteoporosis, and colon and breast cancer (Haskell *et al*, 2007b), even among normal weight individuals. Increased physical activity has been also associated with lower levels of periodontal disease in two recent longitudinal studies, independent of obesity (Merchant *et al*, 2003; Al-Zahrani *et al*, 2005).

Body weight is determined primarily by the balance between energy (calorie) consumption and expenditure (activity); for weight maintenance, energy consumption should equal expenditure. Weight reduction can be achieved by decreasing energy intake and/or increasing physical activity, while the reverse would lead to weight gain. Most studies show that to maintain a healthy body weight (within normal ranges) and for weight-loss, individuals should consume low-energy diets and engage regularly in physical activity (Anderson et al, 2001; Wing and Phelan, 2005; Galani and Schneider, 2007; Haskell et al, 2007b). In a recent large randomized trial, differences in the percentage of energy from fat, carbohydrate and protein did not affect body weight over a 2-year period (Sacks et al, 2009). Recommendations for energy intake are based on each individual's weight, height and activity level [for calculating energy requirements, refer to the Dietary Reference Intakes report from the Institute of Medicine (Standing Committee on the Scientific Evaluation of Dietary Reference Intakes, Food and Nutrition board, Institute of Medicine, 2005)]. The American College of Sports Medicine and the American Heart Association's recommendation for healthy adults is to engage in at least 30 min of moderate-intensity aerobic activity on 5 days of every week or at least 20 min of vigorous-intensity aerobic activity on 3 days of the week (Haskell *et al*, 2007a). Even though this level of physical activity has clear benefits for disease prevention, some individuals will need greater amounts for weight control.

#### Carbohydrate intake

The specific percentage of calories from carbohydrate in the diet does not appear to be an important determinant of weight gain or risk of disease, but the quality of carbohydrates is important. High quality carbohydrates. such as whole grain products and fruits and vegetables, have positive health benefits. These foods are good source of fiber, minerals such as potassium, magnesium, zinc and selenium, and vitamins such as vitamin A, K, C and the B vitamins. Studies have found that diets rich in whole grain products reduce the risk for diabetes (Haskell et al, 2007b; de Munter et al, 2007; Barclay et al, 2008) and heart disease (Anderson et al, 2000; Barclay et al, 2008). In addition, whole grain products, especially if not milled into fine flour, and fruits and vegetables tend to have low glycemic indexes, which prevent a quick rise in blood sugar. Diets rich in highglycemic-index foods, such as white bread and potatoes, lead to a quick and strong increase in blood sugar levels, which then stimulates a large release of insulin, which in turn signals cells to absorb blood sugar for energy or storage. As a result, blood sugar declines suddenly and steeply and may lead to an increase in food intake. Meta-analyses of clinical trials also show that diets high in refined carbohydrates and sugar increase the levels of triglycerides and decrease the levels of HDL cholesterol (Anderson et al, 2004; Thomas et al, 2007). Several studies now show that such diets increase the risk for overweight (Maki et al, 2007; Thomas et al, 2007; Livesey et al, 2008), diabetes (Barclay et al, 2008) and heart disease (Beulens et al, 2007; Barclay et al, 2008). Some foods, low in glycemic index, include coarsely ground whole grain bread, beans and legumes, nuts, and high-fiber fruits and vegetables.

Total carbohydrate intake has not been related to periodontitis. However, one large cohort study of male US health professionals showed an inverse association between whole-grain intake and periodontitis (Merchant *et al.* 2006).

Fermentable carbohydrates result in acid production by cariogenic plaque bacteria and lowering of the plaque pH leads to demineralization. Sucrose is the most cariogenic sugar because it can form glucan that enables firm bacterial adhesion to teeth (Tinanoff and Palmer, 2003). The frequency of sugar intake is a major determinant of caries risk (Gustafsson *et al*, 1954).

Caries risk is the greatest if sugar is consumed at high frequency, more so when the sugar is in a form that is retained in the mouth for long periods of time. The effect of sugar or other carbohydrates on caries could be reduced by fluoride (ingested or topical) and good oral hygiene, but a reduction in sugar should be the primary approach as this has many health benefits.

Sugar and sweetened beverages intake. Regular consumption of sugar-sweetened beverages has been associated with increased energy consumption in children (O'Connor et al, 2006) and with the risk of obesity in children and adults (Ludwig et al, 2001; Malik et al, 2006, 2009; Wolff and Dansinger, 2008). Short intervention trials in children and adolescents to decrease sugar-sweetened consumption resulted in decreases in BMI in those with higher BMI at baseline (Ebbeling et al, 2006; Sichieri et al, 2009) and increases in lean body mass (Albala et al, 2008). Regular consumption of sugar-sweetened beverages has also been associated with increased risk of type II diabetes (Schulze et al, 2005; Palmer et al, 2008) and CHD (Fung et al, 2004, 2009). Consumption of soft drinks containing sugar is also related to increased caries prevalence and incidence (Lim et al, 2008).

Fiber intake. Several large, longitudinal epidemiological studies have found an inverse relation between dietary cereal fiber intake and risk of cardiovascular disease (Rimm et al, 1996; Pereira et al, 2004). In clinical trials, increased dietary fiber intake also reduces blood pressure (Streppel et al, 2005; Whelton et al, 2005) and total and LDL cholesterol levels (Brown et al, 1999). Also, in several large longitudinal studies, fiber intake from whole grain products has been associated with the risk of diabetes (Krishnan et al, 2007; de Munter et al, 2007; Schulze et al, 2007). Furthermore, clinical trials show that diets rich in whole grains result in improved insulin sensitivity in obese subjects (Rave et al. 2007). In addition, fiber consumption can reduce the occurrence of diverticular disease (Aldoori et al, 1998; Aldoori and Ryan-Harshman, 2002) and constipation (Hongisto et al, 2006). A pooled analysis of case-control studies supports an inverse association between dietary fiber and endometrial cancer (Bandera et al, 2007). However, for colon cancer, the evidence is not clear. Large epidemiological studies do not show any association of dietary fiber and colon cancer (Fuchs et al, 1999; Park et al, 2005; Schatzkin et al, 2007). Also, in clinical trials, higher intake of fiber has not reduced the occurrence of colorectal adenomas (Jacobs et al, 2006).

Fruits and vegetables intake. A diet rich in fruits and vegetables is associated with reduced risk of cardiovascular disease. Results from several large longitudinal studies show that individuals consuming five or more servings per day of fruits and vegetables significantly reduce their risk of cardiovascular disease (Hung et al, 2004; Dauchet et al, 2006; He et al, 2006, 2007). Clinical trials also show that diets rich in fruits and vegetables significantly reduced blood pressure (Appel et al, 1997).

With respect to cancer risk, the evidence is inconsistent. Case-control studies show an inverse association between fruits, vegetables and several other cancers (esophagus, lung, stomach and colorectum). Cohort studies have suggested a protective effect of fruits on bladder cancer (Riboli and Norat, 2003), while a very large cohort study did not find a protective effect of five or more servings of fruits and vegetables on total cancer risk (Hung et al, 2004). Both case-control and cohort studies show that high intake of fruits and vegetables are inversely associated with oral precancer (Gupta et al, 1998; Morse et al, 2000; Maserejian et al, 2006) and cancer (Gridley et al, 1990; Levi et al, 1998; Boeing et al, 2006; Pavia et al, 2006). Fruits show a stronger and more consistent inverse association with oral cancer; for vegetables, the strongest inverse association is seen for tomatoes (De Stefani et al, 2000). A metaanalysis of cohort studies on gastric cancer showed that fruits and vegetable intake was associated with lower incidence, but it was not associated with mortality (Lunet et al, 2005). These differences in results may be explained by the diversity of compounds, such as vitamins and minerals, present in different fruits and vegetables and their differential effects on certain types of cancer. This is discussed further in the Vitamins and minerals intake section.

#### Fat intake

Diets high in fats have been believed to be linked to obesity and chronic diseases (Bray and Popkin, 1998; Drewnowski, 2007; Mendoza et al, 2007), but the overall evidence does not support a relation between the percentage of fat in the diet and body weight or risks of cardiovascular disease or cancer (Hu and Willett, 2002; World Cancer Research Fund (WCRF)/American Institute for Cancer Research, 2007). A large randomized trial showed that total fat intake was unrelated to long-term control of body weight (Howard et al, 2006a), risk of cardiovascular disease (Howard et al, 2006b) or risk of cancer (Beresford et al, 2006; Prentice et al, 2006). Also, as noted above, in a large randomized trial, the percentage of fat was not related to body weight (Sacks et al, 2009). As with carbohydrates, the quality of fat is more important than the total amount of fat consumption.

Trans and saturated fat intake. Epidemiological studies show that the consumption of unsaturated fat decreases the risk of heart disease, whereas the consumption of trans fat increases the risk (Willett, 1993; Hu et al, 2001). The magnitude of this association is considerably stronger for trans fat compared with saturated fat consumption (Willett, 2006), and when saturated fat is compared with the same level of calories from carbohydrate, the difference in risk of heart disease is minimal. In controlled feeding studies, diets high in trans fat significantly increase LDL and VLDL cholesterol and significantly decrease HDL cholesterol compared with mono- and poly-unsaturated fats. Compared with carbohydrate, saturated fat increases both LDL and HDL cholesterol (Mensink et al, 2003). High levels of LDL

and VLDL cholesterol and low levels of HDL cholesterol are associated with increased risk of cardiovascular disease (Stamler et al, 2000; Anum and Adera, 2004). Furthermore, trans fat intake is positively associated with markers of systemic inflammation in women (Mozaffarian et al, 2004), suggesting a uniquely adverse cardiometabolic effect via pathways linked to insulin resistance (Mozaffarian and Willett, 2007). Red and processed meats are associated with an increased risk of hypertension (Steffen et al, 2005), CHD (Hu et al, 1999), and type 2 diabetes (Fung et al, 2004). It was estimated from the Nurses' Health Study that replacing only 5% of the energy from saturated fats and 2% of trans fat with unsaturated fats would reduce the risk of coronary disease by 42% and 53%, respectively (Hu et al, 1997).

There is not enough evidence to relate total fat, *trans* or saturated fat consumption with increased risk for most cancer. However, epidemiological evidence suggests that diets rich in animal fat increase the risk of breast cancer in young women (Cho *et al*, 2003), while diets rich in monounsaturated fats (mainly in the form of olive oil) may reduce this risk (Sieri *et al*, 2004). Similar results have been found with some other cancers. Red meat consumption has been associated with an increased risk of colorectal cancer (Larsson and Wolk, 2006; Sandhu *et al*, 2001; WCRF, 2007).

Unsaturated fat intake. While the consumption of trans and saturated fat may increase the risk of many chronic diseases, the opposite is true for the consumption of mono- and poly-unsaturated fat. Epidemiological and clinical studies show that unsaturated fat consumption can lower blood pressure, improve lipid levels (reduce LDL and total cholesterol and increase HDL cholesterol), and reduce cardiovascular risk (Willett, 1993, 2006; Ascherio et al, 1996; Hu et al, 2001, 2002, 2003; Mensink et al, 2003; Appel et al, 2005).

Several epidemiological studies also relate the consumption of fish to a lower risk of heart disease, especially sudden cardiac death (Albert *et al*, 2002; He *et al*, 2002). Fish is rich in the essential omega-3 fatty acids. Meta-analysis of observational studies showed that individuals consuming fish have a reduction of about 20% in the risk of fatal CHD and 10% in total CHD compared with those consuming little or no fish (Whelton *et al*, 2004), and eating fish once per week appears to reduce death from CHD by about 15% (He *et al*, 2004). Clinical trials using fish oil have also shown a reduction in fatal CHD and sudden death (Harper and Jacobson, 2005). Omega-3 fatty acids could also alter the inflammatory process in periodontitis (Kesavalu *et al*, 2006).

Nuts are also rich in mono- and poly-unsaturated fatty acids as well as fiber and micronutrients. Several epidemiological studies show that regular nut consumption is associated with lower risk of CHD (Hu and Stampfer, 1999). Based on the data from the Nurses' Health Study, substituting saturated fat with nuts would result in 45% reduction in the risk of CHD (Hu and Stampfer, 1999). Clinical trials have also shown that var-

ious forms of nuts decrease LDL cholesterol (Gebauer et al, 2008) and triglycerides (Alper and Mattes, 2003), and increase HDL cholesterol (Sheridan et al, 2007), with no effect on body weight if total calories are kept the same (Hollis and Mattes, 2007; Mattes et al, 2008).

#### Antioxidant intake

Antioxidants are substances that prevent or reduce damage caused by reactive oxygen species (ROS) or reactive nitrogen species. Certain vitamins (especially vitamins C and E) and minerals (selenium) act as antioxidants and may protect against tissue damage, thus, reducing the risk of heart disease and cancer. Many of these antioxidants are present in fruits and vegetables; therefore, the role of individual antioxidants on preventing chronic diseases is not very clear.

A pooled analysis of nine cohort studies that included information on intake of vitamin E, carotenoids and vitamin C found that only a high consumption of vitamin C (>700 mg day<sup>-1</sup>) was associated with lower risk of CHD (Knekt et al, 2004). A meta-analysis that included 11 randomized control trials of supplementation with vitamins E and C, beta-carotene, selenium, folate, vitamin B-6 or vitamin B-12 did not show a protective effect of any of these antioxidants on the progression of atherosclerosis (Blevs et al, 2006). The epidemiologic studies differ from the randomized trials in many ways because in the former, subjects with known CHD were excluded, whereas the randomized trials were almost all conducted among subjects with known CHD and on many medications. The Women's Health Study was conducted among primarily healthy people; although there was no overall effect of vitamin E supplements on the risk of CHD, the risk of total cardiovascular mortality was significantly reduced by 24% (Buring, 2006). Thus, the use of antioxidant supplements does not appear to be beneficial among patients with existing CHD, but the effects among apparently healthy people remain unclear.

Antioxidants could also alter the inflammatory process in periodontitis (Chapple *et al*, 2007). Vitamin C is clearly effective in preventing scurvy. Recent data from the NHANES III show an inverse association between dietary vitamin C and periodontal disease (Nishida *et al*, 2000b). In another study, serum levels of vitamin C were inversely associated with periodontitis prevalence, even among never smokers (Chapple *et al*, 2007). ROS have been associated with periodontitis, which if not buffered by sufficient antioxidants, could increase the damage to the periodontal tissue (Moynihan, 2005).

Antioxidants may also prevent certain cancer, by protecting cells from oxidative damage. However, the evidence from epidemiological and clinical studies is not clear. Several meta-analyses of prospective cohorts studies in North America and Europe have not found associations between carotenoid intake and risk of colorectal cancer (Mannisto *et al*, 2007), nor between vitamins A, C, E, and folate supplements and the risk of lung cancer (Cho *et al*, 2006), nor between alphacarotene, beta-carotene, beta-cryptoxanthin, lutein/zeaxanthin, and lycopene and the risk of invasive

epithelial ovarian cancer (Koushik et al, 2006). However, epidemiological studies seem to support a role of lycopene in reducing the risk of prostate cancer (Giovannucci et al. 2002: Etminan et al. 2004). Similarly, several meta-analyses of clinical trials have not found a protective effect of vitamin C and E supplements on the treatment and prevention of cancer in the doses evaluated (Coulter et al, 2006) nor in the consumption of antioxidant supplements in preventing colorectal adenoma (Bjelakovic et al, 2006) or gastrointestinal cancers (Bjelakovic et al, 2008). Furthermore, a recent meta-analysis of 12 randomized clinical trials found that beta-carotene supplementation appeared to increase cancer incidence and cancer mortality among smokers, whereas vitamin E supplementation had no effect (Bardia et al, 2008). However, trials using β-carotene supplements showed reduced risk of oral cancers and remission of precancers with an improvement of at least one grade dysplasia in 39% and no change in 61% of patients (Garewal et al, 1999). Cohort and case control studies also suggest reduced risk of oral cancer among people who consumed different carotenoids (WCRF, 2007).

Selenium, a mineral antioxidant, may have a preventive role on cancer. Several meta-analyses of case—control and cohort studies suggest that selenium intake may reduce the risk of prostate cancer (Etminan *et al*, 2005) and lung cancer (Zhuo *et al*, 2004). Clinical trials also suggest a protective role of selenium supplementation in primary cancer incidence in men (Bardia *et al*, 2008) and in gastrointestinal cancer occurrence (Bjelakovic *et al*, 2008). However, a recent report from the SELECT trial on more than 35 000 individuals with vitamin E and/or selenium supplementation followed for 5.5 years did not reduce prostate cancer risk (Lippman *et al*, 2009).

#### Vitamins and minerals intake

B-vitamins intake. B-vitamins may reduce the risk of cardiovascular disease by ways of reducing homocysteine levels. A meta-analysis of randomized clinical trial showed clear evidence that folate and vitamin B-12 supplementation reduce blood levels of homocysteine (Homocysteine Lowering Trialists' Collaboration, 2005). Observational studies also show a reduction in the risk of cardiovascular disease in individuals with high folate and B6 intake (Rimm et al, 1998; Bazzano et al, 2006). Recent randomized trials show little benefit of folate or B-12 on risks of cardiovascular disease (Bazzano et al, 2006). However, these recent trials have been performed primarily in patients with existing cardiovascular disease, of relatively short duration, and some were in populations where the food supply was fortified with folate. A meta-analysis published in Lancet 2 years ago showed a clear benefit of folic acid supplementation in reducing the risk of stroke, especially in populations with lower folate intake (Wang et al, 2007). In addition to the effects on cardiovascular health, folic acid supplementation reduces the risk of neural tube defects and other congenital anomalies (Goh et al, 2006).

Calcium and vitamin D intake. Calcium and vitamin D intakes are essential for maintaining bone mass. A metaanalysis of 33 clinical studies showed a positive effect of 1000 mg of calcium supplementation on bone mass in young and premenopausal women (Welten et al, 1995), while another meta-analysis showed that dairy products were only beneficial in young white women (Weinsier and Krumdieck, 2000). Around menopause, when bone resorption is greater than formation, calcium supplementation does not prevent bone loss (Dawson-Hughes et al, 1990). However, after about 5 years postmenopause, calcium supplementation prevents further bone loss and may increase bone mass by 1.6–2% (Shea et al, 2004). More recently, a large, randomized-control clinical trial in 36 282 healthy postmenopausal women, the Women's Health Initiative (WHI) study, indicated that prolonged calcium and vitamin D supplementation significantly reduced bone loss at the hip (Jackson et al, 2006). Despite these apparent benefits of calcium intake on bone mineral density, these studies can be misleading because the effects are small and temporary; once calcium supplementation is stopped, bone mineral density typically returns to that of the placebo group.

Calcium and vitamin D supplementation may prevent fracture risk. A meta-analysis of 15 clinical trials with calcium supplementation observed a 23% reduction in vertebral fractures and 14% in non-vertebral fractures (Shea et al, 2004). Two meta-analyses of calcium and vitamin D supplementation trials in 45 000 to 64 000 individuals 50 years and older found a 25% reduction in hip fracture risk (Tang et al, 2007), a 12% reduction of all fractures and 24% reduction of all fractures in those with high compliance to the treatment (Boonen et al., 2007). The WHI trial found a significant 29% decrease in the risk of hip fracture with 1000 mg day<sup>-1</sup> of calcium and 400 IU day<sup>-1</sup> of vitamin D supplementation among participants with good compliance to the treatment (Jackson et al, 2006). One of the difficulties in evaluating the effects of calcium on fracture risk is that in many studies it has been combined with vitamin D. In a metaanalysis of intervention trials using vitamin D supplementation, doses less than 700 IU day<sup>-1</sup> had no effect on risk of fractures, but higher doses of 700-800 IU day<sup>-1</sup> significantly reduced the risk of hip fractures by 26%, and reduced non-vertebral fractures by 23% (Bischoff-Ferrari et al, 2005). However, a recent metaanalysis found no relation between calcium intake and risk of hip fractures in prospective cohort studies, and no significant reduction in the risk of non-vertebral fractures in randomized trial of calcium supplements that did not include vitamin D, with a possible increase in risk (Bischoff-Ferrari et al, 2007). Thus, most people appear to be consuming sufficient calcium for bone health, and increases in vitamin D are likely to reduce risk of fractures, either by effects on bone health or reductions of falls.

Calcium intake has been hypothesized to reduce the risk of cardiovascular conditions, possibly by decreases in blood pressure. In the Nurses' Health Study, low calcium intake was associated with greater risk of ischemic stroke in middle-aged American women (Iso

et al, 1999); intake above 600 mg day<sup>-1</sup> did not appear to further reduce the risk. In the same population, calcium intake was not related to the risk of CHD (Al-Delaimy et al, 2003). Several meta-analyses of clinical trials suggest that calcium supplementation may reduce the risk for high blood pressure (Allender et al, 1996; Griffith et al, 1999; Hajjar et al, 2003; Dickinson et al, 2006; van Mierlo et al, 2006), but the effect appears to be small and may only benefit those with low dietary intakes. Although the evidence has been inconsistent, in some circumstances, higher calcium intake may reduce the risk of pre-eclampsia (Hofmeyr et al, 2006).

Vitamin D may also be related to systemic and local inflammation. A meta-analysis of 19 cross-sectional, 13 case-control and 12 prospective studies suggests that low serum 25-hydroxyvitamin D [25(OH)D] levels or vitamin D intake is associated with glucose intolerance,  $\beta$  cell function and insulin resistance, and with the risk of diabetes and metabolic syndrome in individuals from different populations and age groups (Pittas et al, 2007). The Nurses' Health Study also found that a high intake of calcium and vitamin D was associated with a reduce risk of type 2 diabetes compared with a low intake (Pittas et al, 2006). Furthermore, vitamin D and calcium deficiencies result in bone loss and increased inflammation, which are both components of periodontal disease (Hildebolt, 2005). Vitamin D also has immune-modulatory functions by which it may reduce periodontitis susceptibility (Dietrich et al, 2005). Serum levels of 25(OH)D have been associated with gingivitis (Dietrich et al, 2005) and periodontal disease (Dietrich et al, 2004). Low dietary calcium and dairy products intake are associated with increased levels of periodontal disease (Nishida et al, 2000a; Al-Zahrani, 2006). A clinical trial of calcium and vitamin D supplementation showed increased tooth retention after 3 years in the elderly people (Krall et al, 2001).

Few studies have assessed the protective effects of foods on caries. Recent studies have suggested that milk and cheese are related to reduced risk of caries; however, this is still not well established. Postulated mechanisms may involve buffering of acids, salivary stimulation, reduction of bacterial adhesion, reduction of enamel demineralization, and/or promotion of remineralization by casein and ionizable calcium and phosphorous (Moynihan *et al*, 1999; Kashket and DePaola, 2002).

Calcium and vitamin D intake may also be related to lower colon cancer risk. In a pooled analysis of 10 cohort studies, with more than 500 000 subjects followed for up to 16 years, those with the highest intake of milk and calcium had significantly reduced risk of colon cancer (Cho et al, 2004). This effect was stronger for calcium from foods compared with calcium from supplements (Hartman et al, 2005). Evidence from large cohort trials (the Cancer Prevention Study II Nutrition Cohort and the Multiethnic Cohort Study) also favors a protective effect among men (McCullough et al, 2003; Park et al, 2007) but not among women (Lin et al, 2005; Wactawski-Wende et al, 2006). Meta-analyses of clinical

trials also indicate that calcium supplementation may prevent colorectal adenomatous polyps (Weingarten et al, 2004) and recurrent colorectal adenomas (Shaukat et al, 2005). In addition, higher blood levels of vitamin D (25-OH D) have been consistently associated with reduced risk of colorectal cancer and higher intakes of vitamin D have also been related to lower risk (Gorham et al, 2005; Wei et al, 2008). Some evidences suggest that higher vitamin D levels may reduce the risk of breast cancer (Autier and Gandini, 2007; Garland et al, 2007; Gissel et al, 2008) and decrease total mortality rates (Autier and Gandini, 2007).

On the other hand, calcium and dairy product intake may be related to higher risk of prostate cancer. Metaanalyses of prospective studies indicate that high consumption of milk and dairy products (about >3 servings per day) increases the risk of prostate cancer (Gao et al, 2005; Qin et al, 2007; WCRF, 2007). A large prospective study found > 2.75 servings per day of dairy products, particularly low-fat types > 2000 mg day<sup>-1</sup> of calcium were modestly associated with increased risks only for non-aggressive prostate cancer (Ahn et al, 2007), but in most studies the associations have been primarily with metastatic or fatal prostate cancer. Although consumption of dairy products has been associated with the risk of ovarian cancer in some studies, in a large pooled analysis, the association with total dairy products and calcium intake was not significantly associated with the risk of this cancer (Genkinger et al, 2006). However, when expressed as lactose intake, a modest increase in risk was seen at levels equivalent to about three glasses of milk per day. Milk or dairy product consumption has not been associated with increased risk of breast cancer (Missmer et al, 2002; Shin et al, 2002; McCullough et al, 2005).

# Impact of tooth loss on nutrition

Several cross-sectional studies have shown associations between tooth loss and compromised diet (Joshipura et al, 1996; Krall et al, 1998; Walls and Steele, 2004). There are limited longitudinal studies investigating whether tooth loss leads to dietary changes. The association between self-reported tooth loss and concomitant dietary changes was investigated over a period of 8 years among 31 813 male US health professionals. Men who lost five or more teeth during the follow-up had significant reduction in intake of dietary fiber, whole fruit, increase in dietary cholesterol, and polyunsaturated fat than those who did not lose teeth (Hung et al, 2003). Subsequent to incident tooth loss, results from the Nurses' Health Study showed detrimental dietary changes over a 2-year period with a tendency for women who lost teeth to avoid hard foods such as raw carrot, fresh apple or pear (Hung et al, 2005). However, these differences were relatively small and their significance with respect to chronic disease risks uncertain (Hung et al, 2005). Nevertheless, a small impact on several foods and nutrients could still contribute to a modest overall impact on health.

#### **Guidelines for nutrition**

To maintain and promote good health, individuals should consume a variety of healthy foods as described below.

#### Folic acid

Women in the reproductive age should be taking multiple vitamins that include folic acid if there is any possibility that they might become pregnant. Folic acid has been shown to reduce the risk of birth defects if taken before or during the first few weeks of pregnancy.

#### Sugar and beverages

Pregnant women and mothers of infants should be given advice on preventing baby bottle tooth decay. They should be discouraged from giving their infants a bottle containing beverages with sugar. Milk should be consumed without sugar. It is extremely important for the prevention of baby bottle tooth decay to not allow infants to continue keeping bottles containing milk or other beverages in the mouth when sleeping, especially if these beverages contain added sugar.

Guidelines to reduce the exposure of children to sugar include avoidance of sodas, 'fruit drinks' or other high sugar beverages, limiting the consumption of fruit juice to one small glass (6 oz) per day, and avoiding sugar containing cough drops, chewing gum and snacks such as candy. Furthermore, good dental care should include appropriate use of fluorides, access to preventive and restorative dental care, limiting cariogenic foods during mealtimes, and rapidly clearing cariogenic foods from the child's oral cavity by tooth brushing.

# Fruits and vegetables

Fruits and vegetables provide a variety of micronutrients such as vitamins A, K, C, folate, potassium, magnesium, calcium, iron and fiber. Therefore, individuals should consume a wide variety of these on each meal. The US Dietary Guidelines recommend 4½ cups (nine servings) of fruits and vegetables (not counting starchy vegetables such as potato) for the reference 2000 calorie level (U.S. Department of Health and Human Services and U.S. Department of Agriculture, 2005). This level of intake may be challenging for many people; however, it is important to reassure individuals that even increases in one or two servings a day can have important benefits (Hung *et al*, 2004). Consumption of dark green, yellow and red vegetables should be emphasized to receive the health benefits described in the previous sections.

# Whole grains

Whole grains provide plenty of fiber as wells as micronutrients such as vitamin E, folate, some B vitamins, magnesium, copper, selenium and proteins. To increase the consumption of whole grains, individuals should replace white rice, bread and pasta with brown rice and whole grain products, as well as choose whole grain or high fiber cereals for breakfast. This has a double benefit because refined starches have adverse metabolic effects and increase the risks of obesity,

diabetes and heart disease, whereas whole grains have positive health benefits. The Dietary Reference Intake in the US for fiber is 25 g in women and 38 g in men (U.S. Department of Health and Human Services and U.S. Department of Agriculture, 2005). To obtain the recommended amount of fiber, individuals should consume whole grain products in most meals.

## Nuts

Nuts provide plenty of mono- and poly-unsaturated fatty acids, vitamin E, magnesium and copper, and protein. Individuals should include nuts in their daily diet. Although nuts are often considered as a snack and they are a healthy snack, they can also be considered as a protein source and replace meat in a mixed dish or with a salad. It is recommended that individuals consume 1–2 oz nuts per day (42 g day<sup>-1</sup>).

#### Fish

Fish and other seafood are rich sources of omega 3 fatty acids. Individuals should consume a variety of fish two or more times a week. Part of the benefit of fish can be due to replacement of foods high in saturated fat, such as red meats. Pregnant women should avoid the consumption of big fish, such as shark, swordfish, king mackerel or tilefish and tuna due to their high mercury content. Salmon and small fish, such as catfish, snapper, tilapia, sardines and others, are generally safe during pregnancy because they accumulate less mercury.

# Unsaturated oils

Some fats are good. Vegetable oils are rich in mono- and poly-unsaturated fatty acids, vitamins A, E and K. Double benefits are achieved by replacing butter and other animal fats with vegetable oils, such as canola, soybean olive, corn and sunflower for cooking, baking and for salads. Although margarines were high in trans fat, at this time, almost all margarines in the US contain zero grams of trans fat; although these usually have slightly more saturated fat than liquid oils, they are substantially superior to butter in their fatty acid composition. Until recently, there were concerns about fried foods as fats used for deep frying commercially were universally high in trans fats, but most restaurants are now using trans free oils that are high in unsaturated fats. There is no evidence that fried foods are inherently harmful; their health effects are related to the type of fat and their potential contribution to excessive caloric intake.

# Low-fat dairy products

Dairy products are rich in calcium, phosphorus, potassium, vitamins A and D, some B vitamins and protein. However, dairy products also provide saturated fat. Therefore, individuals should substitute whole-fat products with lower fat products. The US Dietary Guidelines recommend three daily servings of low fat dairy products to achieve calcium requirements for preventing bone loss later in life (U.S. Department of Health and Human Services and U.S. Department of Agriculture, 2005). However, as the evidence suggests both benefits

and harmful effects of dairy products intake on health, limiting dairy products to about two servings a day seems desirable. Other foods, such as cruciferous (brassica) vegetables (broccoli, spinach, cabbage, cauliflower, brussels sprouts, kale, collard greens, bok choi and kohlrabi) and tofu also provide absorbable calcium (2–5 servings of these products could substitute one serving of dairy product), as well as supplements. As the overall evidence suggests that vitamin D supplementation prevents bone loss, falls, fractures and other chronic diseases, most individuals would probably benefit by using a vitamin D supplement daily. The optimal dose is being actively investigated and is not yet established; 1000 IU day<sup>-1</sup> provides benefits, but optimal intakes may be several times this amount.

The revised USDA Pyramid 2005 recommends that individuals consume low amounts of fat and eat large amounts of carbohydrate-rich foods, but pays insufficient attention to the adverse effects of refined starches, and gives inadequate attention to the benefits of healthier oils. A food guide pyramid should encourage the consumption of healthy fats and whole grain foods, minimizing the refined carbohydrates, sugar, butter, and red meat.

# Key recommendations for preventing obesity

As stated above, obesity is a major growing problem with many health implications. Some key recommendations include:

- 1. pay attention to portion sizes and total caloric intake;
- 2. be physically active;
- 3. consume whole grain, high fiber carbohydrates as opposed to rapidly absorbed carbohydrates;
- 4. consume more fruits and vegetables;
- 5. reduce sugar consumption including beverages such as regular carbonated beverages and fruit drinks with added sugar;
- 6. limit fast foods.

# Role of dentists in influencing nutrition

Dentists should play a more active role in preventing and controlling dental caries and periodontal diseases, which are the major indicators for tooth loss. Advice on good dietary habits that help maintain both oral and general health could be included as part of routine preventive care. The importance of retaining teeth and the consequences of tooth loss should be stressed to patients before problems develop, and maintaining good nutrition should be emphasized before extractions are preformed. When dentists recommend extractions, they should inform the patient of the possible consequences of tooth loss on chewing, nutrition, and subsequently on systemic health (Joshipura, 2005). The visit during which the extractions are performed is an additional opportunity to inform the patient about the value of maintaining a healthy diet, for both dental and systemic health. If major chronic diseases are already present, referral to a dietitian may be needed. The need for a good prosthesis should be emphasized to enable patients to chew more efficiently after tooth loss. Follow-up visits for patients with new prosthesis should be scheduled to check its fit and comfort levels, but specifically to inquire about any difficulty encountered while eating when using the prosthesis, and to address this as soon as possible by adjusting the dentures. Even with a good prosthesis, it may be necessary to adjust the preparation of some foods by having hard-to-chew items processed in a way that makes them easier to eat with a compromised dentition, while retaining their nutritional value.

#### **Conclusions**

There is strong evidence that a diet rich in fruits, vegetables, whole grains, nuts, fish and unsaturated fats together with regular physical activity contributes greatly to overall health, including dental health. The appropriate role of dairy products in a healthy diet remains unsettled as both beneficial and harmful effects have been observed. At this point, prudence suggests an emphasis on low fat products and limiting intake to about two servings per day. Overweight and obesity is a huge and growing problem; low fat diets have little benefit, but elimination of sugar-sweetened beverages, attention to overall caloric intake, and regular physical activity will contribute to weight control.

#### **Author contributions**

All authors drafted and revised this review.

# References

- Adams KF, Schatzkin A, Harris TB *et al* (2006). Overweight, obesity, and mortality in a large prospective cohort of persons 50 to 71 years old. *N Engl J Med* **355**: 763–778.
- Ahn J, Albanes D, Peters U *et al* (2007). Dairy products, calcium intake, and risk of prostate cancer in the prostate, lung, colorectal, and ovarian cancer screening trial. *Cancer Epidemiol Biomarkers Prev* **16:** 2623–2630.
- Albala C, Ebbeling CB, Cifuentes M, Lera L, Bustos N, Ludwig DS (2008). Effects of replacing the habitual consumption of sugar-sweetened beverages with milk in chilean children. *Am J Clin Nutr* **88:** 605–611.
- Albert CM, Campos H, Stampfer MJ *et al* (2002). Blood levels of long-chain n-3 fatty acids and the risk of sudden death. *N Engl J Med* **346:** 1113–1118.
- Al-Delaimy WK, Rimm E, Willett WC, Stampfer MJ, Hu FB (2003). A prospective study of calcium intake from diet and supplements and risk of ischemic heart disease among men. *Am J Clin Nutr* 77: 814–818.
- Aldoori W, Ryan-Harshman M (2002). Preventing diverticular disease. review of recent evidence on high-fibre diets. *Can Fam Physician* **48**: 1632–1637.
- Aldoori WH, Giovannucci EL, Rockett HR, Sampson L, Rimm EB, Willett WC (1998). A prospective study of dietary fiber types and symptomatic diverticular disease in men. *J Nutr* **128**: 714–719.
- Allender PS, Cutler JA, Follmann D, Cappuccio FP, Pryer J, Elliott P (1996). Dietary calcium and blood pressure: a meta-analysis of randomized clinical trials. *Ann Intern Med* 124: 825–831.

- Alper CM, Mattes RD (2003). Peanut consumption improves indices of cardiovascular disease risk in healthy adults. *J Am Coll Nutr* **22:** 133–141.
- Al-Zahrani MS (2006). Increased intake of dairy products is related to lower periodontitis prevalence. J Periodontol 77: 289–294.
- Al-Zahrani MS, Borawski EA, Bissada NF (2005). Increased physical activity reduces prevalence of periodontitis. *J Dent* **33:** 703–710.
- Anderson JW, Hanna TJ, Peng X, Kryscio RJ (2000). Whole grain foods and heart disease risk. *J Am Coll Nutr* **19:** 291S–299S
- Anderson JW, Konz EC, Frederich RC, Wood CL (2001). Long-term weight-loss maintenance: a meta-analysis of US studies. Am J Clin Nutr 74: 579–584.
- Anderson JW, Randles KM, Kendall CW, Jenkins DJ (2004). Carbohydrate and fiber recommendations for individuals with diabetes: a quantitative assessment and meta-analysis of the evidence. *J Am Coll Nutr* **23:** 5–17.
- Anum EA, Adera T (2004). Hypercholesterolemia and coronary heart disease in the elderly: a meta-analysis. *Ann Epidemiol* **14:** 705–721.
- Appel LJ, Moore TJ, Obarzanek E *et al* (1997). A clinical trial of the effects of dietary patterns on blood pressure. DASH collaborative research group. *N Engl J Med* **336**: 1117–1124.
- Appel LJ, Sacks FM, Carey VJ et al (2005). Effects of protein, monounsaturated fat, and carbohydrate intake on blood pressure and serum lipids: results of the OmniHeart randomized trial. JAMA 294: 2455–2464.
- Ascherio A, Rimm EB, Giovannucci EL, Spiegelman D, Stampfer M, Willett WC (1996). Dietary fat and risk of coronary heart disease in men: cohort follow up study in the united states. *BMJ* **313:** 84–90.
- Autier P, Gandini S (2007). Vitamin D supplementation and total mortality: a meta-analysis of randomized controlled trials. *Arch Intern Med* **167:** 1730–1737.
- Bandera EV, Kushi LH, Moore DF, Gifkins DM, McCullough ML (2007). Association between dietary fiber and endometrial cancer: a dose-response meta-analysis. *Am J Clin Nutr* **86:** 1730–1737.
- Barclay AW, Petocz P, McMillan-Price J *et al* (2008). Glycemic index, glycemic load, and chronic disease risk a meta-analysis of observational studies. *Am J Clin Nutr* **87**: 627–637.
- Bardia A, Tleyjeh IM, Cerhan JR *et al* (2008). Efficacy of antioxidant supplementation in reducing primary cancer incidence and mortality: systematic review and meta-analysis. *Mayo Clin Proc* **83**: 23–34.
- Bazzano LA, Reynolds K, Holder KN, He J (2006). Effect of folic acid supplementation on risk of cardiovascular diseases: a meta-analysis of randomized controlled trials. *JAMA* 296: 2720–2726.
- Beresford SA, Johnson KC, Ritenbaugh C *et al* (2006). Lowfat dietary pattern and risk of colorectal cancer: the women's health initiative randomized controlled dietary modification trial. *JAMA* **295**: 643–654.
- Beulens JW, de Bruijne LM, Stolk RP *et al* (2007). High dietary glycemic load and glycemic index increase risk of cardiovascular disease among middle-aged women: a population-based follow-up study. *J Am Coll Cardiol* **50:** 14–21.
- Bischoff-Ferrari HA, Willett WC, Wong JB, Giovannucci E, Dietrich T, Dawson-Hughes B (2005). Fracture prevention with vitamin D supplementation: a meta-analysis of randomized controlled trials. *JAMA* **293**: 2257–2264.
- Bischoff-Ferrari HA, Dawson-Hughes B, Baron JA *et al* (2007). Calcium intake and hip fracture risk in men and women: a meta-analysis of prospective cohort studies and randomized controlled trials. *Am J Clin Nutr* **86:** 1780–1790.

- Bjelakovic G, Nagorni A, Nikolova D, Simonetti RG, Bjelakovic M, Gluud C (2006). Meta-analysis: antioxidant supplements for primary and secondary prevention of colorectal adenoma. Aliment Pharmacol Ther 24: 281–291.
- Bjelakovic G, Nikolova D, Simonetti RG, Gluud C (2008). Antioxidant supplements for preventing gastrointestinal cancers. *Cochrane Database Syst Rev* **3:** CD004183.
- Bleys J, Miller ER III, Pastor-Barriuso R, Appel LJ, Guallar E (2006). Vitamin-mineral supplementation and the progression of atherosclerosis: a meta-analysis of randomized controlled trials. *Am J Clin Nutr* **84:** 880–887; quiz 954–5.
- Boeing H, Dietrich T, Hoffmann K *et al* (2006). Intake of fruits and vegetables and risk of cancer of the upper aero-digestive tract: the prospective EPIC-study. *Cancer Causes Control* **17:** 957–969.
- Boonen S, Lips P, Bouillon R, Bischoff-Ferrari HA, Vander-schueren D, Haentjens P (2007). Need for additional calcium to reduce the risk of hip fracture with vitamin D supplementation: evidence from a comparative meta-analysis of randomized controlled trials. *J Clin Endocrinol Meta* **92**: 1415–1423.
- Bray GA, Popkin BM (1998). Dietary fat intake does affect obesity! *Am J Clin Nutr* **68:** 1157–1173.
- Brown L, Rosner B, Willett WW, Sacks FM (1999). Cholesterol-lowering effects of dietary fiber: a meta-analysis. *Am J Clin Nutr* **69:** 30–42.
- Buring JE (2006). Aspirin prevents stroke but not MI in women; vitamin E has no effect on CV disease or cancer. *Cleve Clin J Med* **73:** 863–870.
- Chapple IL, Milward MR, Dietrich T (2007). The prevalence of inflammatory periodontitis is negatively associated with serum antioxidant concentrations. *J Nutr* **137:** 657–664.
- Cho E, Spiegelman D, Hunter DJ *et al* (2003). Premenopausal fat intake and risk of breast cancer. *J Natl Cancer Inst* **95**: 1079–1085.
- Cho E, Smith-Warner SA, Spiegelman D *et al* (2004). Dairy foods, calcium, and colorectal cancer: a pooled analysis of 10 cohort studies. *J Natl Cancer Inst* **96**: 1015–1022.
- Cho E, Hunter DJ, Spiegelman D *et al* (2006). Intakes of vitamins A, C and E and folate and multivitamins and lung cancer: a pooled analysis of 8 prospective studies. *Int J Cancer* **118**: 970–978.
- Coulter ID, Hardy ML, Morton SC *et al* (2006). Antioxidants vitamin C and vitamin E for the prevention and treatment of cancer. *J Gen Intern Med* **21:** 735–744.
- Dauchet L, Amouyel P, Hercberg S, Dallongeville J (2006). Fruit and vegetable consumption and risk of coronary heart disease: a meta-analysis of cohort studies. *J Nutr* **136**: 2588–2593.
- Dawson-Hughes B, Dallal GE, Krall EA, Sadowski L, Sahyoun N, Tannenbaum S (1990). A controlled trial of the effect of calcium supplementation on bone density in postmenopausal women. *N Engl J Med* **323**: 878–883.
- De Stefani E, Oreggia F, Boffetta P, Deneo-Pellegrini H, Ronco A, Mendilaharsu M (2000). Tomatoes, tomato-rich foods, lycopene and cancer of the upper aerodigestive tract: a case–control in Uruguay. *Oral Oncol* **36:** 47–53.
- Dickinson HO, Nicolson DJ, Cook JV *et al* (2006). Calcium supplementation for the management of primary hypertension in adults. *Cochrane Database Syst Rev* 2: CD004639.
- Dietrich T, Joshipura KJ, Dawson-Hughes B, Bischoff-Ferrari HA (2004). Association between serum concentrations of 25-hydroxyvitamin D3 and periodontal disease in the US population. *Am J Clin Nutr* **80:** 108–113.
- Dietrich T, Nunn M, Dawson-Hughes B, Bischoff-Ferrari HA (2005). Association between serum concentrations of 25-hydroxyvitamin D and gingival inflammation. *Am J Clin Nutr* **82:** 575–580.

- Drewnowski A (2007). The real contribution of added sugars and fats to obesity. *Epidemiol Rev* **29:** 160–171.
- Ebbeling CB, Feldman HA, Osganian SK, Chomitz VR, Ellenbogen SJ, Ludwig DS (2006). Effects of decreasing sugar-sweetened beverage consumption on body weight in adolescents: a randomized, controlled pilot study. *Pediatrics* **117**: 673–680.
- Etminan M, Takkouche B, Caamano-Isorna F (2004). The role of tomato products and lycopene in the prevention of prostate cancer: a meta-analysis of observational studies. *Cancer Epidemiol Biomarkers Prev* 13: 340–345.
- Etminan M, FitzGerald JM, Gleave M, Chambers K (2005). Intake of selenium in the prevention of prostate cancer: a systematic review and meta-analysis. *Cancer Causes Control* **16:** 1125–1131.
- Fuchs CS, Giovannucci EL, Colditz GA *et al* (1999). Dietary fiber and the risk of colorectal cancer and adenoma in women. *N Engl J Med* **340**: 169–176.
- Fung TT, Schulze M, Manson JE, Willett WC, Hu FB (2004). Dietary patterns, meat intake, and the risk of type 2 diabetes in women. *Arch Intern Med* **164:** 2235–2240.
- Fung TT, Malik V, Rexrode KM, Manson JE, Willett WC, Hu FB (2009). Sweetened beverage consumption and risk of coronary heart disease in women. *Am J Clin Nutr* **89:** 1037–1042.
- Galani C, Schneider H (2007). Prevention and treatment of obesity with lifestyle interventions: review and meta-analysis. Int J Public Health 52: 348–359.
- Gao X, LaValley MP, Tucker KL (2005). Prospective studies of dairy product and calcium intakes and prostate cancer risk: a meta-analysis. *J Natl Cancer Inst* **97:** 1768–1777.
- Garewal HS, Katz RV, Meyskens F *et al* (1999). Beta-carotene produces sustained remissions in patients with oral leukoplakia: results of a multicenter prospective trial. *Arch Otolaryngol Head Neck Surg* **125**: 1305–1310.
- Garland CF, Gorham ED, Mohr SB *et al* (2007). Vitamin D and prevention of breast cancer: pooled analysis. *J Steroid Biochem Mol Biol* **103:** 708–711.
- Gebauer SK, West SG, Kay CD, Alaupovic P, Bagshaw D, Kris-Etherton PM (2008). Effects of pistachios on cardio-vascular disease risk factors and potential mechanisms of action: a dose-response study. *Am J Clin Nutr* **88:** 651–659.
- Genkinger JM, Hunter DJ, Spiegelman D *et al* (2006). Dairy products and ovarian cancer: a pooled analysis of 12 cohort studies. *Cancer Epidemiol Biomarkers Prev* **15:** 364–372
- Giovannucci E, Rimm EB, Liu Y, Stampfer MJ, Willett WC (2002). A prospective study of tomato products, lycopene, and prostate cancer risk. *J Natl Cancer Inst* **94:** 391–398.
- Gissel T, Rejnmark L, Mosekilde L, Vestergaard P (2008). Intake of vitamin D and risk of breast cancer a meta-analysis. *J Steroid Biochem Mol Biol* **111**: 195–199.
- Goh YI, Bollano E, Einarson TR, Koren G (2006). Prenatal multivitamin supplementation and rates of congenital anomalies: a meta-analysis. *J Obstet Gynaecol can* **28**: 680–689.
- Gorham ED, Garland CF, Garland FC et al (2005). Vitamin D and prevention of colorectal cancer. J Steroid Biochem Mol Biol 97: 179–194.
- Gridley G, McLaughlin JK, Block G *et al* (1990). Diet and oral and pharyngeal cancer among blacks. *Nutr Cancer* **14**: 219–225.
- Griffith LE, Guyatt GH, Cook RJ, Bucher HC, Cook DJ (1999). The influence of dietary and nondietary calcium supplementation on blood pressure: an updated metaanalysis of randomized controlled trials. *Am J Hypertens* **12**: 84–92.

- Gupta PC, Hebert JR, Bhonsle RB, Sinor PN, Mehta H, Mehta FS (1998). Dietary factors in oral leukoplakia and submucous fibrosis in a population-based case control study in Gujarat, India. *Oral Dis* **4:** 200–206.
- Gustafsson BE, Quensel CE, Lanke LS *et al* (1954). The vipeholm dental caries study; the effect of different levels of carbohydrate intake on caries activity in 436 individuals observed for five years. *Acta Odontol Scand* 11: 232–264.
- Hajjar IM, Grim CE, Kotchen TA (2003). Dietary calcium lowers the age-related rise in blood pressure in the United States: the NHANES III survey. J Clin Hypertens (Greenwich) 5: 122–126.
- Harper CR, Jacobson TA (2005). Usefulness of omega-3 fatty acids and the prevention of coronary heart disease. *Am J Cardiol* **96:** 1521–1529.
- Hartman TJ, Albert PS, Snyder K *et al* (2005). The association of calcium and vitamin D with risk of colorectal adenomas. *J Nutr* **135**: 252–259.
- Haskell WL, Lee IM, Pate RR *et al* (2007a). Physical activity and public health: updated recommendation for adults from the american college of sports medicine and the american heart association. *Med Sci Sports Exerc* **39:** 1423–1434.
- Haskell WL, Lee IM, Pate RR *et al* (2007b). Physical activity and public health: updated recommendation for adults from the American College of Sports Medicine and the American Heart Association. *Circulation* **116**: 1081–1093.
- He K, Rimm EB, Merchant A et al (2002). Fish consumption and risk of stroke in men. JAMA 288: 3130–3136.
- He K, Song Y, Daviglus ML *et al* (2004). Accumulated evidence on fish consumption and coronary heart disease mortality: a meta-analysis of cohort studies. *Circulation* **109**: 2705–2711.
- He FJ, Nowson CA, MacGregor GA (2006). Fruit and vegetable consumption and stroke: meta-analysis of cohort studies. *Lancet* **367**: 320–326.
- He FJ, Nowson CA, Lucas M, MacGregor GA (2007). Increased consumption of fruit and vegetables is related to a reduced risk of coronary heart disease: meta-analysis of cohort studies. *J Hum Hypertens* 21: 717–728.
- Hildebolt CF (2005). Effect of vitamin D and calcium on periodontitis. *J Periodontol* **76:** 1576–1587.
- Hofmeyr GJ, Atallah AN, Duley L (2006). Calcium supplementation during pregnancy for preventing hypertensive disorders and related problems. *Cochrane Database Syst Rev* 3: CD001059.
- Hollis J, Mattes R (2007). Effect of chronic consumption of almonds on body weight in healthy humans. *Br J Nutr* **98**: 651–656
- Homocysteine Lowering Trialists' Collaboration (2005). Dose-dependent effects of folic acid on blood concentrations of homocysteine: a meta-analysis of the randomized trials. *Am J Clin Nutr* **82:** 806–812.
- Hongisto SM, Paajanen L, Saxelin M, Korpela R (2006). A combination of fibre-rich rye bread and yoghurt containing lactobacillus GG improves bowel function in women with self-reported constipation. *Eur J Clin Nutr* **60**: 319–324.
- Howard BV, Manson JE, Stefanick ML *et al* (2006a). Low-fat dietary pattern and weight change over 7 years: the Women's Health Initiative Dietary Modification Trial. *JAMA* **295**: 39–49.
- Howard BV, Van Horn L, Hsia J *et al* (2006b). Low-fat dietary pattern and risk of cardiovascular disease: the Women's Health Initiative Randomized Controlled Dietary Modification Trial. *JAMA* **295**: 655–666.
- Hu FB, Stampfer MJ (1999). Nut consumption and risk of coronary heart disease: a review of epidemiologic evidence. *Curr Atheroscler Rep* **1:** 204–209.

- Hu FB, Willett WC (2002). Optimal diets for prevention of coronary heart disease. JAMA 288: 2569–2578.
- Hu FB, Stampfer MJ, Manson JE *et al* (1997). Dietary fat intake and the risk of coronary heart disease in women. *N Engl J Med* **337**: 1491–1499.
- Hu FB, Stampfer MJ, Manson JE *et al* (1999). Dietary saturated fats and their food sources in relation to the risk of coronary heart disease in women. *Am J Clin Nutr* **70**: 1001–1008.
- Hu FB, Manson JE, Willett WC (2001). Types of dietary fat and risk of coronary heart disease: a critical review. *J Am Coll Nutr* **20**: 5–19.
- Hu FB, Bronner L, Willett WC et al (2002). Fish and omega-3 fatty acid intake and risk of coronary heart disease in women. JAMA 287: 1815–1821.
- Hu FB, Cho E, Rexrode KM, Albert CM, Manson JE (2003). Fish and long-chain omega-3 fatty acid intake and risk of coronary heart disease and total mortality in diabetic women. *Circulation* 107: 1852–1857.
- Hung HC, Willett W, Ascherio A, Rosner BA, Rimm E, Joshipura KJ (2003). Tooth loss and dietary intake. J Am Dent Assoc 134: 1185–1192.
- Hung HC, Joshipura KJ, Jiang R *et al* (2004). Fruit and vegetable intake and risk of major chronic disease. *J Natl Cancer Inst* **96:** 1577–1584.
- Hung HC, Colditz G, Joshipura KJ (2005). The association between tooth loss and the self-reported intake of selected CVD-related nutrients and foods among US women. *Community Dent Oral Epidemiol* **33:** 167–173.
- Iso H, Stampfer MJ, Manson JE *et al* (1999). Prospective study of calcium, potassium, and magnesium intake and risk of stroke in women. *Stroke* **30:** 1772–1779.
- Jackson RD, LaCroixAZ, Gass M et al (2006). Calcium plus vitamin D supplementation and the risk of fractures. N Engl J Med 354: 669–683.
- Jacobs ET, Lanza E, Alberts DS *et al* (2006). Fiber, sex, and colorectal adenoma: results of a pooled analysis. *Am J Clin Nutr* **83**: 343–349.
- Joshipura KJ (2005). How can tooth loss affect diet and health, and what nutritional advice would you give to a patient scheduled for extractions? *J Can Dental Assoc* **71:** 421–422.
- Joshipura KJ, Willett WC, Douglass CW (1996). The impact of edentulousness on food and nutrient intake. J Am Dent Assoc 127: 459–467.
- Kashket S, DePaola DP (2002). Cheese consumption and the development and progression of dental caries. *Nutr Rev* 60: 97–103.
- Kesavalu L, Vasudevan B, Raghu B *et al* (2006). Omega-3 fatty acid effect on alveolar bone loss in rats. *J Dent Res* **85**: 648–652.
- Knekt P, Ritz J, Pereira MA et al (2004). Antioxidant vitamins and coronary heart disease risk: a pooled analysis of 9 cohorts. Am J Clin Nutr 80: 1508–1520.
- Koushik A, Hunter DJ, Spiegelman D *et al* (2006). Intake of the major carotenoids and the risk of epithelial ovarian cancer in a pooled analysis of 10 cohort studies. *Int J Cancer* **119:** 2148–2154.
- Krall E, Hayes C, Garcia R (1998). How dentition status and masticatory function affect nutrient intake. *J Am Dent Assoc* **129:** 1261–1269.
- Krall EA, Wehler C, Garcia RI, Harris SS, Dawson-Hughes B (2001). Calcium and vitamin D supplements reduce tooth loss in the elderly. *Am J Med* **111:** 452–456.
- Krishnan S, Rosenberg L, Singer M *et al* (2007). Glycemic index, glycemic load, and cereal fiber intake and risk of type 2 diabetes in US black women. *Arch Intern Med* **167**: 2304–2309.

- Larsson SC, Wolk A (2006). Meat consumption and risk of colorectal cancer: a meta-analysis of prospective studies. *Int* J Cancer 119: 2657–2664.
- Levi F, Pasche C, La Vecchia C, Lucchini F, Franceschi S, Monnier P (1998). Food groups and risk of oral and pharyngeal cancer. *Int J Cancer* 77: 705–709.
- Lim S, Sohn W, Burt BA *et al* (2008). Cariogenicity of soft drinks, milk and fruit juice in low-income African-American children: a longitudinal study. *J Am Dent Assoc* **139**: 959–967.
- Lin J, Zhang SM, Cook NR, Manson JE, Lee IM, Buring JE (2005). Intakes of calcium and vitamin D and risk of colorectal cancer in women. *Am J Epidemiol* **161:** 755–764.
- Lippman SM, Klein EA, Goodman PJ et al (2009). Effect of selenium and vitamin E on risk of prostate cancer and other cancers: the selenium and vitamin E cancer prevention trial (SELECT). JAMA 301: 39–51.
- Livesey G, Taylor R, Hulshof T, Howlett J (2008). Glycemic response and health systematic review and meta-analysis: relations between dietary glycemic properties and health outcomes. *Am J Clin Nutr* 87: 258S–268S.
- Ludwig DS, Peterson KE, Gortmaker SL (2001). Relation between consumption of sugar-sweetened drinks and child-hood obesity: a prospective, observational analysis. *Lancet* **357:** 505–508.
- Lunet N, Lacerda-Vieira A, Barros H (2005). Fruit and vegetables consumption and gastric cancer: a systematic review and meta-analysis of cohort studies. *Nutr Cancer* **53**: 1–10
- Maki KC, Rains TM, Kaden VN, Raneri KR, Davidson MH (2007). Effects of a reduced-glycemic-load diet on body weight, body composition, and cardiovascular disease risk markers in overweight and obese adults. *Am J Clin Nutr* **85**: 724–734.
- Malik VS, Schulze MB, Hu FB (2006). Intake of sugar-sweetened beverages and weight gain: a systematic review. *Am J Clin Nutr* **84:** 274–288.
- Malik VS, Willett WC, Hu FB (2009). Sugar-sweetened beverages and BMI in children and adolescents: reanalyses of a meta-analysis. *Am J Clin Nutr* **89:** 438–439; author reply 439–40.
- Mannisto S, Yaun SS, Hunter DJ *et al* (2007). Dietary carotenoids and risk of colorectal cancer in a pooled analysis of 11 cohort studies. *Am J Epidemiol* **165**: 246–255.
- Maserejian NN, Giovannucci E, Rosner B, Zavras A, Joshipura K (2006). Prospective study of fruits and vegetables and risk of oral premalignant lesions in men. *Am J Epidemiol* **164:** 556–566.
- Mathus-Vliegen EM, Nikkel D, Brand HS (2007). Oral aspects of obesity. *Int Dent J* **57:** 249–256.
- Mattes RD, Kris-Etherton PM, Foster GD (2008). Impact of peanuts and tree nuts on body weight and healthy weight loss in adults. *J Nutr* **138**: 1741S–1745S.
- McCullough ML, Robertson AS, Rodriguez C *et al* (2003). Calcium, vitamin D, dairy products, and risk of colorectal cancer in the cancer prevention study II nutrition cohort. *Cancer Causes Control* **14:** 1–12.
- McCullough ML, Rodriguez C, Diver WR et al (2005). Dairy, calcium, and vitamin D intake and postmenopausal breast cancer risk in the cancer prevention study II nutrition cohort. Cancer Epidemiol Biomarkers Prev 14: 2898–2904.
- Mendoza JA, Drewnowski A, Christakis DA (2007). Dietary energy density is associated with obesity and the metabolic syndrome in U.S. adults. *Diabetes Care* **30:** 974–979.

- Mensink RP, Zock PL, Kester AD, Katan MB (2003). Effects of dietary fatty acids and carbohydrates on the ratio of serum total to HDL cholesterol and on serum lipids and apolipoproteins: a meta-analysis of 60 controlled trials. *Am J Clin Nutr* 77: 1146–1155.
- Merchant AT, Pitiphat W, Rimm EB, Joshipura K (2003). Increased physical activity decreases periodontitis risk in men. Eur J Epidemiol 18: 891–898.
- Merchant AT, Pitiphat W, Franz M, Joshipura KJ (2006). Whole-grain and fiber intakes and periodontitis risk in men. *Am J Clin Nutr* **83:** 1395–1400.
- van Mierlo LA, Arends LR, Streppel MT *et al* (2006). Blood pressure response to calcium supplementation: a meta-analysis of randomized controlled trials. *J Hum Hypertens* **20**: 571–580.
- Missmer SA, Smith-Warner SA, Spiegelman D *et al* (2002). Meat and dairy food consumption and breast cancer: a pooled analysis of cohort studies. *Int J Epidemiol* **31:** 78–85.
- Morse DE, Pendrys DG, Katz RV *et al* (2000). Food group intake and the risk of oral epithelial dysplasia in a united states population. *Cancer Causes Control* **11:** 713–720.
- Moynihan PJ (2005). The role of diet and nutrition in the etiology and prevention of oral diseases. *Bull World Health Org* **83:** 694–699.
- Moynihan PJ, Ferrier S, Jenkins GN (1999). The cariostatic potential of cheese: cooked cheese-containing meals increase plaque calcium concentration. *Br Dent J* **187:** 664–667.
- Mozaffarian D, Willett WC (2007). Trans fatty acids and cardiovascular risk: a unique cardiometabolic imprint? *Curr Atheroscler Rep* **9:** 486–493.
- Mozaffarian D, Rimm EB, King IB, Lawler RL, McDonald GB, Levy WC (2004). Trans fatty acids and systemic inflammation in heart failure. *Am J Clin Nutr* **80:** 1521–1525.
- de Munter JS, Hu FB, Spiegelman D, Franz M, van Dam RM (2007). Whole grain, bran, and germ intake and risk of type 2 diabetes: a prospective cohort study and systematic review. *PLoS Med* **4:** e261.
- National Institute of Health, National Heart, Lung, and Blood Institute (1998). Clinical guidelines on the identification, evaluation, and treatment of overweight and obesity in adults: the evidence report. NIH Publication 98-4083. National Institute of Health, National Heart, Lung, and Blood Institute: Bethesda.
- Nishida M, Grossi SG, Dunford RG, Ho AW, Trevisan M, Genco RJ (2000a). Calcium and the risk for periodontal disease. J Periodontol 71: 1057–1066.
- Nishida M, Grossi SG, Dunford RG, Ho AW, Trevisan M, Genco RJ (2000b). Dietary vitamin C and the risk for periodontal disease. *J Periodontol* **71:** 1215–1223.
- O'Connor TM, Yang SJ, Nicklas TA (2006). Beverage intake among preschool children and its effect on weight status. *Pediatrics* **118:** e1010–e1018.
- Palmer JR, Boggs DA, Krishnan S, Hu FB, Singer M, Rosenberg L (2008). Sugar-sweetened beverages and incidence of type 2 diabetes mellitus in African American women. Arch Intern Med 168: 1487–1492.
- Park Y, Hunter DJ, Spiegelman D et al (2005). Dietary fiber intake and risk of colorectal cancer: a pooled analysis of prospective cohort studies. JAMA 294: 2849–2857.
- Park SY, Murphy SP, Wilkens LR, Nomura AM, Henderson BE, Kolonel LN (2007). Calcium and vitamin D intake and risk of colorectal cancer: the multiethnic cohort study. *Am J Epidemiol* **165:** 784–793.
- Pavia M, Pileggi C, Nobile CG, Angelillo IF (2006). Association between fruit and vegetable consumption and oral cancer: a meta-analysis of observational studies. *Am J Clin Nutr* **83:** 1126–1134.

- Pereira MA, O'Reilly E, Augustsson K *et al* (2004). Dietary fiber and risk of coronary heart disease: a pooled analysis of cohort studies. *Arch Intern Med* **164**: 370–376.
- Pischon N, Heng N, Bernimoulin JP, Kleber BM, Willich SN, Pischon T (2007). Obesity, inflammation, and periodontal disease. J Dent Res 86: 400–409.
- Pittas AG, Dawson-Hughes B, Li T *et al* (2006). Vitamin D and calcium intake in relation to type 2 diabetes in women. *Diabetes Care* **29:** 650–656.
- Pittas AG, Lau J, Hu FB, Dawson-Hughes B (2007). The role of vitamin D and calcium in type 2 diabetes. A systematic review and meta-analysis. *J Clin Endocrinol Metab* **92:** 2017–2029.
- Prentice RL, Caan B, Chlebowski RT *et al* (2006). Low-fat dietary pattern and risk of invasive breast cancer: the Women's Health Initiative Randomized Controlled Dietary Modification Trial. *JAMA* **295**: 629–642.
- Qin LQ, Xu JY, Wang PY, Tong J, Hoshi K (2007). Milk consumption is a risk factor for prostate cancer in western countries: evidence from cohort studies. *Asia Pac J Clin Nutr* **16:** 467–476.
- Rave K, Roggen K, Dellweg S, Heise T, tom Dieck H (2007). Improvement of insulin resistance after diet with a wholegrain based dietary product: results of a randomized, controlled cross-over study in obese subjects with elevated fasting blood glucose. *Br J Nutr* **98:** 929–936.
- Riboli E, Norat T (2003). Epidemiologic evidence of the protective effect of fruit and vegetables on cancer risk. *Am J Clin Nutr* **78:** 559S–569S.
- Rimm EB, Ascherio A, Giovannucci E, Spiegelman D, Stampfer MJ, Willett WC (1996). Vegetable, fruit, and cereal fiber intake and risk of coronary heart disease among men. *JAMA* **275**: 447–451.
- Rimm EB, Willett WC, Hu FB *et al* (1998). Folate and vitamin B6 from diet and supplements in relation to risk of coronary heart disease among women. *JAMA* **279:** 359–364.
- Ritchie CS, Joshipura K, Hung HC, Douglass CW (2002). Nutrition as a mediator in the relation between oral and systemic disease: associations between specific measures of adult oral health and nutrition outcomes. *Crit Rev Oral Biol Med* 13: 291–300.
- Sacks FM, Bray GA, Carey VJ *et al* (2009). Comparison of weight-loss diets with different compositions of fat, protein, and carbohydrates. *N Engl J Med* **360**: 859–873.
- Saito T, Shimazaki Y, Sakamoto M (1998). Obesity and periodontitis. *N Engl J Med* **339:** 482–483.
- Sandhu MS, White IR, McPherson K (2001). Systematic review of the prospective cohort studies on meat consumption and colorectal cancer risk: a meta-analytical approach. *Cancer Epidemiol Biomarkers Prev* **10:** 439–446.
- Schatzkin A, Mouw T, Park Y *et al* (2007). Dietary fiber and whole-grain consumption in relation to colorectal cancer in the NIH-AARP diet and health study. *Am J Clin Nutr* **85**: 1353–1360.
- Schulze MB, Hoffmann K, Manson JE *et al* (2005). Dietary pattern, inflammation, and incidence of type 2 diabetes in women. *Am J Clin Nutr* **82:** 675–684.
- Schulze MB, Schulz M, Heidemann C, Schienkiewitz A, Hoffmann K, Boeing H (2007). Fiber and magnesium intake and incidence of type 2 diabetes: a prospective study and meta-analysis. *Arch Intern Med* **167**: 956–965.
- Shaukat A, Scouras N, Schunemann HJ (2005). Role of supplemental calcium in the recurrence of colorectal adenomas: a meta-analysis of randomized controlled trials. *Am J Gastroenterol* **100:** 390–394.
- Shea B, Wells G, Cranney A *et al* (2004). Calcium supplementation on bone loss in postmenopausal women. *Cochrane Database Syst Rev* 1: CD004526.

- Sheridan MJ, Cooper JN, Erario M, Cheifetz CE (2007). Pistachio nut consumption and serum lipid levels. J Am Coll Nutr 26: 141–148.
- Shin MH, Holmes MD, Hankinson SE, Wu K, Colditz GA, Willett WC (2002). Intake of dairy products, calcium, and vitamin D and risk of breast cancer. J Natl Cancer Inst 94: 1301–1311.
- Shuldiner AR, Yang R, Gong DW (2001). Resistin, obesity and insulin resistance the emerging role of the adipocyte as an endocrine organ. *N Engl J Med* **345**: 1345–1346.
- Sichieri R, Paula Trotte A, de Souza RA, Veiga GV (2009). School randomised trial on prevention of excessive weight gain by discouraging students from drinking sodas. *Public Health Nutr* **12:** 197–202.
- Sieri S, Krogh V, Pala V *et al* (2004). Dietary patterns and risk of breast cancer in the ORDET cohort. *Cancer Epidemiol Biomarkers Prev* **13:** 567–572.
- Stamler J, Daviglus ML, Garside DB, Dyer AR, Greenland P, Neaton JD (2000). Relationship of baseline serum cholesterol levels in 3 large cohorts of younger men to long-term coronary, cardiovascular, and all-cause mortality and to longevity. *JAMA* **284**: 311–318.
- Standing Committee on the Scientific Evaluation of Dietary Reference Intakes, Food and Nutrition board, Institute of Medicine (2005). Dietary reference intakes for energy, carbohydrate, fiber, fat, fatty acids, cholesterol, protein, and amino acids. National Academy Press: Washington, DC.
- Steffen LM, Kroenke CH, Yu X et al (2005). Associations of plant food, dairy product, and meat intakes with 15-y incidence of elevated blood pressure in young black and white adults: the coronary artery risk development in young adults (CARDIA) study. Am J Clin Nutr 82: 1169–1177; quiz 1363–4.
- Streppel MT, Arends LR, van 't Veer P, Grobbee DE, Geleijnse JM (2005). Dietary fiber and blood pressure: a meta-analysis of randomized placebo-controlled trials. *Arch Intern Med* **165**: 150–156.
- Tang BM, Eslick GD, Nowson C, Smith C, Bensoussan A (2007). Use of calcium or calcium in combination with vitamin D supplementation to prevent fractures and bone loss in people aged 50 years and older: a meta-analysis. *Lancet* **370**: 657–666.
- Thomas DE, Elliott EJ, Baur L (2007). Low glycaemic index or low glycaemic load diets for overweight and obesity. *Cochrane Database Syst Rev* **3:** CD005105.
- Tinanoff N, Palmer CA (2003). Dietary determinants of dental caries and dietary recommendations for preschool children. *Refuat Hapeh Vehashinayim* **20:** 8–23.
- U.S. Department of Health and Human Services and U.S. Department of Agriculture (2005). *Dietary guidelines for Americans*, 2005. U.S. Department of Health and Human Services and U.S. Department of Agriculture: Washington.

- Wactawski-Wende J, Kotchen JM, Anderson GL *et al* (2006). Calcium plus vitamin D supplementation and the risk of colorectal cancer. *N Engl J Med* **354**: 684–696.
- Walls AW, Steele JG (2004). The relationship between oral health and nutrition in older people. *Mech Ageing Dev* **125**: 853–857.
- Wang X, Qin X, Demirtas H *et al* (2007). Efficacy of folic acid supplementation in stroke prevention: a meta-analysis. *Lancet* **369**: 1876–1882.
- Wei MY, Garland CF, Gorham ED, Mohr SB, Giovannucci E (2008). Vitamin D and prevention of colorectal adenoma: a meta-analysis. *Cancer Epidemiol Biomarkers Prev* 17: 2958–2969
- Weingarten MA, Zalmanovici A, Yaphe J (2004). Dietary calcium supplementation for preventing colorectal cancer and adenomatous polyps. *Cochrane Database Syst Rev* 1: CD003548.
- Weinsier RL, Krumdieck CL (2000). Dairy foods and bone health: examination of the evidence. *Am J Clin Nutr* **72:** 681–689.
- Welten DC, Kemper HC, Post GB, van Staveren WA (1995).
  A meta-analysis of the effect of calcium intake on bone mass in young and middle aged females and males. J Nutr 125: 2802–2813.
- Whelton SP, He J, Whelton PK, Muntner P (2004). Metaanalysis of observational studies on fish intake and coronary heart disease. *Am J Cardiol* **93:** 1119–1123.
- Whelton SP, Hyre AD, Pedersen B, Yi Y, Whelton PK, He J (2005). Effect of dietary fiber intake on blood pressure: a meta-analysis of randomized, controlled clinical trials. *J Hypertens* **23**: 475–481.
- Willett WC (1993). Dietary fat reduction among women with early breast cancer. *J Clin Oncol* 11: 2061–2062.
- Willett WC (2006). Trans fatty acids and cardiovascular disease-epidemiological data. *Atheroscler Suppl* 7: 5–8.
- Wing RR, Phelan S (2005). Long-term weight loss maintenance. *Am J Clin Nutr* **82:** 222S–225S.
- Wolff E, Dansinger ML (2008). Soft drinks and weight gain: how strong is the link? *Medscape J Med* **10**: 189.
- World Cancer Research Fund (WCRF)/American Institute for Cancer Research (2007). Food, nutrition, physical activity, and the prevention of cancer: a global perspective. World Cancer Research Fund (WCRF)/American Institute for Cancer Research: Washington.
- World Health Organization (WHO) (2003). *Global strategy on diet, physical activity and health*. World Health Organization (WHO): Washington, DC.
- Zhuo H, Smith AH, Steinmaus C (2004). Selenium and lung cancer: a quantitative analysis of heterogeneity in the current epidemiological literature. *Cancer Epidemiol Biomarkers Prev* **13:** 771–778.