

Dairy and cardiovascular health: friend or foe?

Article

Published Version

Creative Commons: Attribution 3.0 (CC-BY)

Open Access

Markey, O., Vasilopoulou, D., Givens, I. ORCID: <https://orcid.org/0000-0002-6754-6935> and Lovegrove, J. ORCID: <https://orcid.org/0000-0001-7633-9455> (2014) Dairy and cardiovascular health: friend or foe? Nutrition Bulletin, 39 (2). pp. 161-171. ISSN 1467-3010 doi: 10.1111/nbu.12086 Available at <https://centaur.reading.ac.uk/36671/>

It is advisable to refer to the publisher's version if you intend to cite from the work. See [Guidance on citing](#).

To link to this article DOI: <http://dx.doi.org/10.1111/nbu.12086>

Publisher: Wiley

All outputs in CentAUR are protected by Intellectual Property Rights law, including copyright law. Copyright and IPR is retained by the creators or other copyright holders. Terms and conditions for use of this material are defined in the [End User Agreement](#).

www.reading.ac.uk/centaur

CentAUR

Central Archive at the University of Reading

Reading's research outputs online

Dairy and cardiovascular health: Friend or foe?

O. Markey*, D. Vasilopoulou*†, D. I. Givens† and J. A. Lovegrove*

*Hugh Sinclair Unit of Human Nutrition and Institute for Cardiovascular and Metabolic Research (ICMR), Department of Food and Nutritional Sciences, University of Reading, UK;

†Food Production and Quality Research Division, School of Agriculture, Policy and Development, Faculty of Life Sciences, University of Reading, UK

Abstract

Cardiovascular disease (CVD) prevalence at a global level is predicted to increase substantially over the next decade due to the increasing ageing population and incidence of obesity. Hence, there is an urgent requirement to focus on modifiable contributors to CVD risk, including a high dietary intake of saturated fatty acids (SFA). As an important source of SFA in the UK diet, milk and dairy products are often targeted for SFA reduction. The current paper acknowledges that milk is a complex food and that simply focusing on the link between SFA and CVD risk overlooks the other beneficial nutrients of dairy foods. The body of existing prospective evidence exploring the impact of milk and dairy consumption on risk factors for CVD is reviewed. The current paper highlights that high milk consumption may be beneficial to cardiovascular health, while illustrating that the evidence is less clear for cheese and butter intake. The option of manipulating the fatty acid profile of ruminant milk is discussed as a potential dietary strategy for lowering SFA intake at a population level. The review highlights that there is a necessity to perform more well-controlled human intervention-based research that provides a more holistic evaluation of fat-reduced and fat-modified dairy consumption on CVD risk factors including vascular function, arterial stiffness, postprandial lipaemia and markers of inflammation. Additionally, further research is required to investigate the impact of different dairy products and the effect of the specific food matrix on CVD development.

Keywords: arterial stiffness, blood pressure, cardiovascular disease, dairy products, milk, saturated fatty acids

Introduction

Although mortality from cardiovascular disease (CVD) is now falling in most European countries, CVD is ranked

as the leading cause of mortality in the UK and world-wide (BHF 2010; WHO 2011; Nichols *et al.* 2012). It is envisaged that the global prevalence of CVD will continue to increase and is expected to be responsible for more than 23.6 million deaths by 2030 (Smith *et al.* 2012), a figure that is largely attributable to today's dramatic demographic changes with increasing proportions of ageing and obese groups. The outcome of today's obesity and ageing trends may, if not moderated, result in unsustainable costs to global society. Currently, CVD costs the European Union (EU) economy approximately €196 billion per annum in direct and indirect charges,

Correspondence: Professor Julie A. Lovegrove, Professor of Human Nutrition, Head of the Hugh Sinclair Unit of Human Nutrition and Deputy Director of the Institute for Cardiovascular and Metabolic Research (ICMR), Hugh Sinclair Unit of Human Nutrition, Department of Food and Nutritional Sciences, University of Reading, Whiteknights, PO Box 266, Reading, RG6 6AP, UK.
E-mail: j.a.lovegrove@reading.ac.uk

with the cost per person in the UK exceeding the EU average (Nichols *et al.* 2012). There is growing pressure to reduce risk factors for CVD at a population level. An atherogenic diet, characterised by a high intake of dietary saturated fatty acids (SFA), is a key modifiable risk factor for CVD. While the effects of the amount and type of dietary fat have been examined in relation to CVD, less focus has been placed on the role of animal-derived staple foods such as milk and dairy products, which are significant dietary sources of SFA.

Saturated fat consumption

In order to reduce the population health burden of CVD, a number of strategies can be addressed; these include increasing physical activity, improving weight profiles and reducing tobacco and alcohol intake. However, this paper will focus on strategies for combating the overconsumption of dietary SFA.

The adverse effect of SFA on CVD risk is well established; this is primarily mediated via increases in serum lipids, particularly low-density lipoprotein (LDL)-cholesterol (Mensink *et al.* 2003). Current dietary recommendations include an intake of dietary SFA of less than 10% of total energy (DH 1991; WHO 2008). The current rolling *National Diet and Nutrition Survey* (NDNS), as outlined in Table 1, highlights that this dietary target is exceeded by the majority of men, women and children in the UK (DH 2012).

Substitution of dietary saturated fat with carbohydrate or unsaturated fatty acids

The question of whether replacement of dietary SFA with carbohydrate (CHO), *cis*-polyunsaturated fatty acids (*cis*-PUFA) or *cis*-monounsaturated fatty acids (*cis*-MUFA) can have beneficial effects on CVD mortality and risk has received considerable attention (Astrup *et al.* 2011; Hooper *et al.* 2012; Vafeiadou *et al.* 2012).

There would seem to be no clear evidence for a benefit of substituting CHO for SFA in the prevention of CVD (Astrup *et al.* 2011; Hooper *et al.* 2012) but there is some recent evidence for the benefit of replacing SFA with unsaturated fatty acids (Jakobsen *et al.* 2009; Micha & Mozaffarian 2010; Hooper *et al.* 2012). There is good evidence that replacing SFA with *cis*-PUFA will reduce CVD mortality and risk markers, although there is less information on the effects of replacing SFA with *cis*-MUFA, mainly because few relevant randomised controlled trials have been performed (Mozaffarian *et al.* 2010). Overall, the Cochrane meta-analysis of Hooper *et al.* (2012) identified that a reduction of dietary SFA intake (where it is replaced by unsaturated fat) and/or reduction of total dietary fat lowered the risk of cardiovascular events by 14% [relative risk (RR) 0.86; 95% confidence interval (CI) 0.77–0.96] but had no effect on total mortality.

The contribution of dairy products to saturated fat intake

Milk and dairy products are one of the most significant contributors to SFA intake in the UK diet. In the context of this review, we are defining dairy as cows' milk or any food product derived from cows' milk. However, it should be noted that some dairy products (including butter and cream) do not fall into the 'milk and dairy' section of the *eatwell* plate because they are predominantly viewed as significant sources of fat (FSA 2011). Recent NDNS data (Table 2) suggest that milk and dairy products are responsible for 22–25% of SFA consumption in adults (DH 2011); however, it should be noted that this analysis failed to take into account the intake of composite dairy dishes (*e.g.* pizza and lasagne) as well as butter consumption, the latter contributing around 5% to daily average saturated fat intakes; thus SFA intake from milk and dairy product consumption may be underestimated.

Table 1 Average intake of saturated fatty acids in absolute terms and as a percentage of food/total energy, by age and gender (NDNS 2008/2009–2010/2011)

	Boys/Men			Girls/Women		
	4–18 years	19–64 years	≥65 years	4–18 years	19–64 years	≥65 years
SFA (g/day)	25.8 ± 8.6	28.8 ± 12.1	29.3 ± 10.7	22.7 ± 8.1	22.0 ± 9.8	23.2 ± 9.0
% food energy	12.9 ± 2.6	12.8 ± 3.3	14.4 ± 3.5	12.9 ± 2.7	12.6 ± 3.4	14.0 ± 3.7
% total energy	12.9 ± 2.6	12.0 ± 3.4	13.6 ± 3.4	12.9 ± 2.8	12.0 ± 3.3	13.7 ± 3.3

Source: DH (2012).

Mean ± SD. NDNS, National Diet and Nutrition Survey; SFA, saturated fatty acids.

Table 2 Percentage contribution of food groups to average daily total saturated fatty acid intake, by age and gender (NDNS 2008/2009–2009/10)

Food groups (%)	Boys/Men			Girls/Women		
	4–18 years	19–64 years	≥65 years	4–18 years	19–64 years	≥65 years
Cereals and cereal products	25	18	17	24	20	18
Milk and milk products	25	22	28	27	22	24
Eggs and egg dishes	2	3	3	2	4	4
Fat spreads	9	10	13	8	10	16
Meat and meat products	22	27	22	20	23	19
Fish and fish dishes	2	3	4	1	3	5
Vegetables and potatoes	5	7	5	6	7	6
Savoury snacks	2	1	0	2	1	0
Nuts/seeds/fruits	0	1	1	0	2	0
Sugars/preserves/confectionary	7	4	2	7	5	3
Drinks	0	1	1	0	1	0
Miscellaneous	2	3	3	2	4	3

Source: DH (2011).

NDNS, National Diet and Nutrition Survey.

Effects of saturated fat in dairy products on plasma lipids and lipoproteins

A detailed review by Huth and Park (2012) revealed that diets higher in SFA from whole milk and butter increase LDL-cholesterol but may also have a beneficial impact on high-density lipoprotein (HDL)-cholesterol concentrations, resulting in a neutral or beneficial effect on the total cholesterol:HDL-cholesterol ratio. Fermented dairy products including cheese and yogurt appear to have a different impact on circulating levels of plasma lipid and lipoproteins. For example, compared to butter matched for milk fat intake, consumption of hard cheese for a 6-week period led to significantly lower concentrations of total cholesterol, LDL-cholesterol and HDL-cholesterol (Hjerpsted *et al.* 2011). However, it should be noted that butter is not necessarily a good comparator. There are a number of confounding variables that negate the drawing of firm conclusions in relation to cheese and CVD risk due to the differential effect of individual cheese varieties that differ in macronutrient content, degree of fermentation and food matrix (Huth & Park 2012). Furthermore, it highlights that the impact of dairy products on CVD risk factors may be dependent on the specific dairy food, even when supplying the same mass of dairy fat. However, further long-term studies are necessary before firmer conclusions can be drawn on the relative impact of milk and milk-derived food consumption on plasma lipids and lipoproteins.

The consumption of milk and milk-derived products and cardiovascular disease

Potential benefits of milk and dairy consumption

Dairy fats are high in SFA; however, ruminant milk is a complex food and there is much debate as to whether nutrients within dairy foods act independently or synergistically in relation to chronic disease development. The association of SFA with CVD development may be dependent on other nutrients/macronutrients in the matrix of the SFA-containing food (de Oliveira Otto *et al.* 2012). Milk is a significant source of a number of essential micronutrients including calcium, potassium and iodine. Calcium, for example, has a higher bioavailability in milk compared with that present in some other foods (Weaver *et al.* 1999). Furthermore, mineral bioavailability is enhanced by the lack of inhibitors present in milk, including phytates and oxalates, and by the presence of lactose and certain amino acids that may promote mineral absorption (FAO 2013). Moreover, lipids mediate the delivery of essential fat-soluble vitamins, including vitamins D and A, and fatty acids associated with dairy products, namely conjugated linoleic acids, may also have cardio-protective properties, although the data are inconsistent and require confirmation in further human studies (Dilzer & Park 2012). Furthermore, emerging evidence suggests that plasma phospholipid *trans*-palmitoleic acid (*trans* 16:1 *n*-7), a circulating fatty acid biomarker positively corre-

lated with self-reported intakes of dairy fat intake (whole-fat dairy products and butter), is associated with a more favourable metabolic profile and incident diabetes rate (Micha & Mozaffarian 2010; Mozaffarian *et al.* 2013). However, the aforementioned findings regarding *trans* 16:1 *n*-7 should be interpreted with caution as they do not necessarily prove cause and effect; thus, oversimplifying the relationship between dairy product intake (in terms of SFA content) and CVD risk may be misleading (Givens 2012).

Changing trends in dairy consumption

UK trends in dairy product consumption have changed markedly over recent years; this could be partly due to the negative connotations surrounding dairy products and SFA content. *Family Food* statistics published by the Department for Environment, Food and Rural Affairs (Defra) over the past 20 years show a decline in total liquid milk consumption, largely brought about by a decrease in whole milk consumption, which now accounts for around 20% of milk consumed in the UK today. Butter consumption has decreased markedly from the 1970s by about 70%, and fell around 2% lower in the decade between 2001 and 2011, while cheese and yogurt consumption increased by around 5% and 30%, respectively, over the same time period (AHDB-DairyCo 2012).

Dietary patterns associated with milk and dairy product consumption

Analysis of dietary patterns is recognised as an alternate and complementary strategy for investigating the association between diet and disease risk. As dietary patterns are more representative of overall food consumption, they may facilitate a more valid prediction of CVD risk compared with assessment of an individual nutrient or food (Hu 2002). Prospective data from 88 517 middle-aged women indicated that adherence to the Dietary Approaches to Stop Hypertension (DASH) diet, characterised by a moderate intake of low-fat dairy, legumes and nut products, high intake of fruit, vegetables and wholegrains, and low intake of red meat, was inversely associated with risk of coronary heart disease (CHD) and stroke during a 24-year follow-up period (Fung *et al.* 2008). This finding was in agreement with the *Atherosclerosis Risk in Communities Study* which illustrated that a dietary pattern rich in dairy and nut products, but less meat, is associated with a lower risk of incident hypertension in middle-aged adults (Weng *et al.* 2013). However, many international and national

dietary guidelines recommend the reduced intake of full-fat dairy products as one aspect of a dietary pattern linked to reducing risk of CHD (Erlinger & Appel 2005).

Dairy consumption and cardiovascular disease risk: Prospective evidence

A number of studies have investigated the effect of milk and dairy products on different CVD events. As previously mentioned, there is a lack of robust evidence on the potential differential effects of individual dairy foods on CVD risk as most observational studies combine dairy products as a single food group, although milk is better studied. A meta-analysis of prospective cohort studies reported that, overall, high milk consumption does not increase the RR of CHD (Elwood *et al.* 2008). A second meta-analysis, which combined prospective cohort and clinical studies, revealed that there was insufficient evidence for an association between milk consumption and CHD (RR 0.94; 95% CI 0.75–1.13) (Mente *et al.* 2009). A later more extensive meta-analysis on milk and dairy consumption and CVD events concluded that high consumption of milk was related to a significant reduction in risk of stroke development (Elwood *et al.* 2010). Table 3 is largely based on Elwood *et al.* (2010) but has been updated by the addition of data from six recently published studies (Bonhuis *et al.* 2010; Goldbohm *et al.* 2011; Sonestedt *et al.* 2011; Soedamah-Muthu *et al.* 2012; Avalos *et al.* 2013; van Aerde *et al.* 2013). A recent study showed that compared with the lowest quintile of dairy consumption, total dairy intake was inversely related to myocardial infarction (MI) risk following an 11.6-year follow-up period (HR 0.77; 95% CI 0.63–0.95). Further analysis revealed that butter used on bread was positively correlated with MI risk, while total cheese intake had an inverse risk association (Patterson *et al.* 2013). This evidence supports the hypothesis that dairy products, excluding butter, are associated with no detrimental effect, and in some cases a significant reduced CVD risk.

Hypertension is one of the key risk factors for CVD development and is influenced by gene polymorphisms, nutrition, the environment and interactions between these factors. Milk and milk-derived products provide essential micronutrients (such as calcium, potassium and iodine) and protein (whey, casein and specific bioactive peptides), some of which have been associated with beneficial hypotensive effects, either independently or synergistically (Kris-Etherton *et al.* 2009). There are a number of proposed mechanisms by which milk and its components could reduce blood pressure (BP; for a

Table 3 Summary of the relative risk for milk and dairy consumption and CVD events

Disease outcome	Number of cohort studies (number used in analyses)	Adjusted RR (95% CI) for milk/dairy consumption*	Significance of heterogeneity between studies
Ischaemic heart disease	22 (17)	0.92 (0.86, 0.99)	$P=0.765$
All strokes	12	0.79 (0.68, 0.91)	$P=0.001$
Haemorrhagic stroke	5	0.75 (0.60, 0.94)	$P=0.014$
Subarachnoid bleed	3	0.93 (0.84, 1.02)	$P=0.004$
All-cause mortality	12 (9)	0.91 (0.78, 1.05)	$P=0.070$

Source: Adapted from Givens *et al.* (2014).

*Estimate of the risk of each disease in individuals with the highest consumption of milk/dairy products compared to the risk in individuals with the lowest consumption.

CI, confidence interval; CVD, cardiovascular disease; RR, relative risk.

detailed review, see Fekete *et al.* 2013). Bioactive peptides present in casein and whey proteins have been observed to play a role in controlling BP by inhibiting the action of angiotensin-I-converting enzyme, resulting in vasodilation (FitzGerald & Meisel 2000), by modulating the release of endothelin-1 by endothelial cells (Maes *et al.* 2004) and acting as opioid receptor ligands increasing nitric oxide production which mediates arterial tone (Kris-Etherton *et al.* 2009). An important consideration is the potential impact of a threshold dependency mechanism whereby benefit is conferred in those at low nutrient status, such as calcium, whereas in individuals with adequate baseline status, little effect is observed (McCarron *et al.* 1991; Wennersberg *et al.* 2009; Park & Cifelli 2013). This has important considerations in respect of public health advice on dairy consumption within population groups with different nutritional status. Although the *Rotterdam Study* found an inverse association between low-fat dairy intake and hypertension risk in older adults (Engberink *et al.* 2009), limited evidence exists as to the potential additional benefit of low-fat dairy foods and the type of dairy products in relation to BP reduction. Low-fat dairy is the product of choice in most trials, yet both low- and high-fat alternatives appear to have an overall beneficial effect in relation to BP (Ralston *et al.* 2012). Fumeron *et al.* (2011) reported that consumption of either a variety of dairy products excluding cheese, or cheese alone, and the calcium density of the diet were associated with a lower 9-year diastolic BP after analysing data from the *Epidemiological Study on the Insulin Resistance Syndrome*. Moreover, data from the *Caerphilly Prospective Study* illustrated that when compared to non-milk consumers, men who consumed >586 ml/day had a 10.4 mmHg lower systolic BP after a 22.8-year follow-up (Livingstone *et al.* 2013). Unsurprisingly, greater hypotensive effects of dairy consumption are

observed in those with hypertension or who present with calcium sensitivity. In normotensive subjects, dairy consumption is often related to retaining BP homeostasis rather than hypotensive effects (Park & Cifelli 2013).

Elasticity of the blood vessels can be influenced by chronic dietary patterns (Kesse-Guyot *et al.* 2010). Cardiovascular events and all-cause mortality are independently predicted by carotid-femoral pulse wave velocity (PWV), the gold standard measurement of arterial stiffness (Vlachopoulos *et al.* 2010; Van Bortel *et al.* 2012). Further evidence from the *Caerphilly Prospective Study* highlighted that, with the exception of butter consumption, dairy product intake does not impact negatively on PWV (Livingstone *et al.* 2013). Furthermore, augmentation index, another indicator of arterial stiffness, was 1.8% lower in men with the highest quartiles of dairy food consumption (Livingstone *et al.* 2013). Similarly, cross-sectional study findings illustrated that dairy food intake was inversely correlated with PWV (Crichton *et al.* 2012a).

A low-grade systemic inflammation is recognised as a major factor contributing to the development and progression of a number of disorders related to CVD (Labonte *et al.* 2013). Cross-sectional studies that have investigated the relationship between dairy intake and low-grade systemic inflammation have found an inverse association (Salas-Salvado *et al.* 2008; Esmailzadeh & Azadbakht 2010). However, a review that grouped several studies involving overweight or obese subjects found a degree of heterogeneity which hinders any definite conclusions (Labonte *et al.* 2013). Although there is evidence supporting a beneficial association between dairy consumption and inflammation, the mechanisms are still unclear and studies are either underpowered or use more than one type of dairy product as an intervention, making it difficult to distinguish between dairy products (Labonte *et al.* 2013).

Dietary strategies for lowering consumption of saturated fat and the implication for cardiovascular disease

Low-fat milk and dairy products

When compared with low-fat alternatives, there is no established nutritional benefit of whole-fat dairy consumption, except in young children; therefore, the intake of low-fat milk and milk-related products may be considered an effective strategy to lower SFA intake. However, there is currently no consensus on whether fat-reduced dairy foods are associated with a reduced risk of CVD (Benatar *et al.* 2013). Observational studies have indicated that low-fat dairy consumption is an effective strategy to promote lower BP levels (Engberink *et al.* 2009; Toledo *et al.* 2009; van Meijl & Mensink 2011), circulating markers of inflammation (Esmailzadeh & Azadbakht 2010), the ratio of total cholesterol:HDL-cholesterol (Mensink *et al.* 2003) and LDL-cholesterol concentration (Kai *et al.* 2013), as well as aid in weight maintenance or reduction (Abargouei *et al.* 2012). The *Nurses' Health Study* cohort illustrated that the associated RR of CHD varied according to consumption of high-fat (RR 1.12; 95% CI 1.05–1.20) or low-fat dairy consumption (RR 0.80; 95% CI 0.73–0.87) (Hu *et al.* 1999). On the contrary, findings from a prospective population-based cohort of 33 636 women suggested there were no significant differences between consumption of specific low-fat and high-fat dairy products and MI risk (Patterson *et al.* 2013). Furthermore, findings from a 12-month randomised crossover trial concluded that inclusion of reduced-fat dairy products in the diets of overweight adults had no impact on cardio-metabolic outcomes, including blood lipids, BP and arterial compliance (Crichton *et al.* 2012b). However, before it can be clearly established whether or not removal of milk fat is beneficial to overall cardio-metabolic health, further evidence from well-controlled human intervention studies is required.

Altering the dairy cow diet to manipulate the fatty acid profile of milk

As an alternative to promoting low-fat dairy product consumption, modification of the fatty acid profile of bovine milk offers a strategy for lowering the population's intake of SFA, by removing SFA from the food chain, while preserving the beneficial contributions that dairy products make to the protein and micronutrient content of the human diet (Shingfield *et al.* 2008). Over 100 studies have explored the potential of partially replacing milk SFA with *cis*-MUFA or *cis*-PUFA through

supplementation of the bovine diet with plant oils or oilseeds (Givens & Shingfield 2006; Glasser *et al.* 2008). Through a reduced synthesis of short- and medium-chain SFA by the mammary gland, this feeding strategy enhances the long-chain (>C18) unsaturated fatty acid concentration in the milk (Doreau *et al.* 1999). Inclusion of 49 g/kg of dry matter of rapeseed oil in the ruminant diet for a 28-day period increased *cis*-MUFA from 20 to 33 g/100 g FA, while reducing SFA from 70 to 55–60 g/day FA (Givens *et al.* 2009). Although a more substantial decrease in SFA (~20 g/100 g of FA) has been documented, the alteration to milk FA composition was adversely linked to voluntary bovine nutrient intake and milk yield compared to the control diet (Givens *et al.* 2003). In order for modification of the composition of ruminant-derived foods to be recognised as a sustainable strategy for reducing SFA intake at a population level, it is essential to find an optimal balance between maximising the unsaturated FA profile of the milk and minimising the impact of the supplementation on animal performance (Givens 2008). Furthermore, it should be considered that ruminal biohydrogenation of PUFA results in the formation of intermediates including *trans* 18:1 and leads to small increases in PUFA relative to MUFA concentrations and therefore, it might be more feasible to supplement the bovine diet with MUFA (Shingfield *et al.* 2013).

In addition to the reductions in SFA and increases in *cis*-MUFA, inclusion of unsaturated fatty acids into the bovine-feeding regimen can lead to increased concentrations of naturally produced ruminant *trans* fatty acid (rTFA), namely linoleic acid isomers and *trans* MUFA, in the milk. The intake of *trans* fatty acids (TFA) from industrially hydrogenated vegetable oils is known to have a negative impact on cardiovascular health (Mozaffarian *et al.* 2006; Brouwer *et al.* 2010) and, accordingly, there has been a significant reduction in the level of 'industrial' TFA (iTFA) in the food chain (Hulshof *et al.* 1999). Conversely, the association between rTFA and CVD remains inconclusive (Gebauer *et al.* 2011; Brouwer *et al.* 2013) with some studies showing a cardio-protective effect of ruminant sources of *trans* fats (Mozaffarian *et al.* 2006; Jakobsen *et al.* 2008). In an attempt to resolve the conflicting reports, a systematic review and meta-analysis was undertaken by Bendsen *et al.* (2011). They reported that the RR for high vs. low quintiles of total TFA intake (2.8 to approximately 10 g/day) was 1.22 (95% CI 1.08–1.38; $P = 0.002$) for CHD events and 1.24 (95% CI 1.07–1.43; $P = 0.003$) for fatal CHD. rTFA intake (0.5–1.9 g/day) was not significantly associated with CHD risk (RR 0.92; 95% CI 0.76–1.11; $P = 0.36$) although neither

was iTFA. There was, however, a trend towards a positive association (RR 1.21; 95% CI 0.97–1.50; $P = 0.09$) for iTFA intake. The authors concluded that while iTFA may be positively related to CHD, rTFA is not, but the limited number of studies available prevented a firm conclusion concerning whether the source of TFA is important. The lack of an association of rTFA with CHD risk may be due to lower intake levels (Bendsen *et al.* 2011). However, at levels currently consumed in the UK diet, there is no evidence of risk from rTFA.

Over the past 10–15 years, the total TFA intake in the UK diet has decreased substantially as a result of voluntary action by the UK food industry; this has led to a greater proportion of the total dietary *trans* fats originating from rTFA (SACN 2007). However, while the proportion of dietary rTFA has increased, the absolute intake of ruminant fat is unchanged. The current dietary intake of TFA in the UK (0.7% of food energy in adults) (DH 2012) is below the recommended population maximum (2% of food energy intake) (DH 1991), with milk and milk products contributing to around 25% of this intake (DH 2011). Consequently, TFA intake from ruminant sources is not seen as a major cause of concern to cardiovascular health at a population level (Tardy *et al.* 2011; Brouwer *et al.* 2013). However, it has yet to be determined whether increasing rTFA intake, through manipulation of the fatty acid profile of milk and dairy products to decrease SFA content, impacts on cardiovascular health (Livingstone *et al.* 2012).

Impact of modified dairy products on cardiovascular disease risk factors

A review of the current evidence suggests that consumption of modified feed-reduced SFA milk and milk products may be beneficial to CVD risk in healthy and hypercholesterolaemic populations when compared to commercially available whole milk dairy products (Livingstone *et al.* 2012). However, it should be noted that there is a distinct lack of human intervention-based research (Givens 2012; Livingstone *et al.* 2012) and the studies that have been performed have relied on plasma lipid levels as a predictor of CVD risk and on butter as the main test food. Some selected data illustrate that, in comparison to conventional milk, cheese, butter and ice cream (70 g/100 g SFA, 28 g/100 g *cis*-MUFA), total cholesterol and LDL-cholesterol were significantly lowered following a 3-week period of consuming matched fat-modified dairy products (51 g/100 g SFA, 39 g/100 g *cis*-MUFA) (Noakes *et al.* 1996). Dairy products are complex, nutrient-dense foods and focusing on a single outcome measure could lead to misleading

conclusions by failing to establish the impact on other CVD risk factors.

Evidence suggests that some of the effects of SFA on CVD risk are mediated by impairment in endothelial function and subsequent establishment of atherosclerosis (Nicholls *et al.* 2006; Blumenthal *et al.* 2010; Vafeiadou *et al.* 2012) and by influencing postprandial lipaemia (Berry & Sanders 2005). Endothelial dysfunction, an early modifiable event in the coronary atherosclerotic process, is positively associated with increased risk of CVD (De Caterina 2000; Schachinger *et al.* 2000). Flow-mediated dilation, which measures the vasodilatory response of the brachial artery to an increase in blood flow-associated shear stress and carotid intima-media thickness, can be used to non-invasively assess endothelial function and arterial structural changes, respectively (Anderson 2006). As previously discussed, it is recommended that arterial stiffness, a surrogate marker of central arterial function, should also be evaluated when exploring the impact of modified dairy consumption on cardiovascular risk factors (Givens 2012). Postprandial lipaemia, characterised by elevated and prolonged triacylglycerol concentrations in the fed state, is influenced by the type and quantity of the meal fat (Chong *et al.* 2010; Jackson & Lovegrove 2012; Jackson *et al.* 2012). Postprandial lipaemia is a significant independent risk marker of CVD that requires attention in future modified fat studies (Nordestgaard *et al.* 2007). Further research is warranted to examine the impact of fat-modified or any total low-fat or full-fat dairy consumption on holistic measures of CVD including vascular function, arterial stiffness, postprandial lipaemia and inflammation.

This is currently being addressed at the University of Reading in the *RESET (REplacement of SaturatEd fat in dairy on Total cholesterol) Study* (ClinicalTrials.gov NCT02089035), a 3-year Medical Research Council funded project that is investigating the impact of reducing SFA intake by using modified milk and dairy products on vascular function and CVD risk biomarkers, without limiting dairy product consumption. This will be achieved by producing milk and dairy products that have a substantial proportion of the SFA replaced with *cis*-MUFA. In a randomised, crossover, double-blind, controlled study, it will be determined whether modified dairy product consumption improves vascular function and other CVD risk biomarkers relative to typical commercially available products in both acute and chronic settings. The project, which started in late 2013, will provide unique evidence to inform public health policy on food-based dietary recommendations for CVD risk reduction.

Conclusion

Much of the UK population currently exceeds the dietary SFA recommendation of <11% of food energy intake, with milk and dairy products as a group making a considerable contribution to SFA intake in the average UK diet. While it is intuitive to consider further reducing dairy consumption as a means of decreasing SFA intake, epidemiological data suggest that this strategy may be counterproductive, given the array of cardio-metabolic benefits that milk products appear to offer to human health. Fat-reduced or fat-modified dairy product consumption may offer a more feasible option for reducing intake of SFA with minimal change to habitual eating patterns and, hence, CVD risk at a population level. Nevertheless, before the impact of both fat-reduced and modified dairy consumption on CVD risk reduction can be evaluated, it is essential to conduct more robust, controlled human intervention-based research using both traditional and novel assessments of cardiovascular risk. Additionally, more research is required to differentiate between dairy food matrices.

Acknowledgements

O. Markey and D. Vasilopoulou are supported by the Medical Research Council (MR/K020218/1).

Conflict of interest

The authors declare no conflict of interest.

References

- Abargouei AS, Janghorbani M, Salehi-Marzijarani M *et al.* (2012) Effect of dairy consumption on weight and body composition in adults: a systematic review and meta-analysis of randomized controlled clinical trials. *International Journal of Obesity* **36**: 1485–93.
- AHDB-DairyCo (Agriculture and Horticulture Development Board) (2012) Purchases of Milk and Dairy. Available at: <http://www.dairyco.org.uk/market-information/dairy-sales-consumption/#.Uz6-X61g6Fc> (accessed 7 March 2014).
- Anderson TJ (2006) Arterial stiffness or endothelial dysfunction as a surrogate marker of vascular risk. *Canadian Journal of Cardiology* **22**: 72B–80B.
- Astrup A, Dyerberg J, Elwood P *et al.* (2011) The role of reducing intakes of saturated fat in the prevention of cardiovascular disease: where does the evidence stand in 2010? *American Journal of Clinical Nutrition* **93**: 684–8.
- Avalos EE, Barrett-Connor E, Kritiz-Silverstein D *et al.* (2013) Is dairy product consumption associated with the incidence of CHD? *Public Health Nutrition* **16**: 2055–63.
- Benatar JR, Sidhu K & Stewart RA (2013) Effects of high and low fat dairy food on cardio-metabolic risk factors: a meta-analysis of randomized studies. *PLoS ONE* **8**: 0076480.
- Bendsen NT, Christensen R, Bartels EM *et al.* (2011) Consumption of industrial and ruminant trans fatty acids and risk of coronary heart disease: a systematic review and meta-analysis of cohort studies. *European Journal of Clinical Nutrition* **65**: 773–83.
- Berry SE & Sanders TA (2005) Influence of triacylglycerol structure of stearic acid-rich fats on postprandial lipaemia. *Proceedings of the Nutrition Society* **64**: 205–12.
- BHF (British Heart Foundation) (2010) Coronary heart disease statistics 2010. Available at: <http://www.bhf.org.uk/publications/view-publication.aspx?ps=1001546> (accessed 5 December 2013).
- Blumenthal JA, Babyak MA, Hinderliter A *et al.* (2010) Effects of the DASH diet alone and in combination with exercise and weight loss on blood pressure and cardiovascular biomarkers in men and women with high blood pressure: the ENCORE study. *Archives of Internal Medicine* **170**: 126–35.
- Bonthuis M, Hughes MC, Ibiebele TI *et al.* (2010) Dairy consumption and patterns of mortality of Australian adults. *European Journal of Clinical Nutrition* **64**: 569–77.
- Brouwer IA, Wanders AJ & Katan MB (2010) Effect of animal and industrial trans fatty acids on HDL and LDL cholesterol levels in humans—a quantitative review. *PLoS ONE* **5**: 0009434.
- Brouwer IA, Wanders AJ & Katan MB (2013) Trans fatty acids and cardiovascular health: research completed? *European Journal of Clinical Nutrition* **67**: 541–7.
- Chong MF, Lockyer S, Saunders CJ *et al.* (2010) Long chain n-3 PUFA-rich meal reduced postprandial measures of arterial stiffness. *Clinical Nutrition* **29**: 678–81.
- Crichton GE, Elias MF, Dore GA *et al.* (2012a) Relations between dairy food intake and arterial stiffness: pulse wave velocity and pulse pressure. *Hypertension* **59**: 1044–51.
- Crichton GE, Howe PR, Buckley JD *et al.* (2012b) Dairy consumption and cardiometabolic health: outcomes of a 12-month crossover trial. *Nutrition & Metabolism* **9**: 19.
- de Oliveira Otto MC, Mozaffarian D, Kromhout D *et al.* (2012) Dietary intake of saturated fat by food source and incident cardiovascular disease: the Multi-Ethnic Study of Atherosclerosis. *American Journal of Clinical Nutrition* **96**: 397–404.
- De Caterina R (2000) Endothelial dysfunctions: common denominators in vascular disease. *Current Opinion in Lipidology* **11**: 9–23.
- DH (Department of Health) (1991), Dietary Reference Values for Food Energy and Nutrients for the United Kingdom. Vol. 41: Report on Health and Social Subjects. London: Her Majesty's Stationery Office.
- DH (Department of Health) (2011) National Diet and Nutrition Survey: Headline results from Years 1 and 2 (combined) of the rolling programme 2008/9–2009/10. Available at: <https://www.gov.uk/government/publications/national-diet-and-nutrition-survey-headline-results-from-years-1-and-2-combined-of-the-rolling-programme-2008-9-2009-10> (accessed 6 December 2013).
- DH (Department of Health) (2012) National Diet and Nutrition Survey: Headline results from Years 1, 2 and 3 (combined) of the rolling programme 2008/9–2010/11. Available at: <http://webarchive.nationalarchives.gov.uk/20130402145952/http://transparency.dh.gov.uk/2012/07/25/ndns-3-years-report/> (accessed February 2014).
- Dilzer A & Park Y (2012) Implication of conjugated linoleic acid (CLA) in human health. *Critical Reviews in Food Science and Nutrition* **52**: 488–513.

- Doreau M, Chillard Y, Rulquin H *et al.* (1999) Manipulation of milk fat in dairy cows. In: *Recent Advances in Animal Nutrition*, pp. 81–109. Nottingham University Press: Nottingham.
- Elwood PC, Givens DI, Beswick AD *et al.* (2008) The survival advantage of milk and dairy consumption: an overview of evidence from cohort studies of vascular diseases, diabetes and cancer. *Journal of the American College of Nutrition* 27: 723S–34S.
- Elwood PC, Pickering JE, Givens DI *et al.* (2010) The consumption of milk and dairy foods and the incidence of vascular disease and diabetes: an overview of the evidence. *Lipids* 45: 925–39.
- Engberink MF, Geleijnse JM, de Jong N *et al.* (2009) Dairy intake, blood pressure, and incident hypertension in a general Dutch population. *Journal of Nutrition* 139: 582–7.
- Erlinger T & Appel LJ (2005) Dietary patterns and coronary heart disease risk. In: *Coronary Heart Disease Epidemiology*, 2nd edn. Available at: <http://dx.doi.org/10.1093/acprof:oso/9780198525738.003.0014> (accessed 4 April 2014).
- Esmailzadeh A & Azadbakht L (2010) Dairy consumption and circulating levels of inflammatory markers among Iranian women. *Public Health Nutrition* 13: 1395–402.
- FAO (Food and Agriculture Organization) (2013) Milk and dairy products in human nutrition. Available at: <http://www.fao.org/publications> (accessed December 2013).
- Fekete AA, Givens DI & Lovegrove JA (2013) The impact of milk proteins and peptides on blood pressure and vascular function: a review of evidence from human intervention studies. *Nutrition Research Reviews* 26: 177–90.
- FitzGerald RJ & Meisel H (2000) Milk protein-derived peptide inhibitors of angiotensin-I-converting enzyme. *British Journal of Nutrition* 84: S33–7.
- FSA (Food Standards Agency) (2011) The Eatwell Plate. Available at: <http://www.Eatwell.gov.uk/healthydiet/Eatwellplate/> (accessed 2 February 2014).
- Fumeron F, Lamri A, Abi Khalil C *et al.* (2011) Dairy consumption and the incidence of hyperglycemia and the metabolic syndrome: results from a French prospective study, Data from the Epidemiological Study on the Insulin Resistance Syndrome (DESIR). *Diabetes Care* 34: 813–17.
- Fung TT, Chiuve SE, McCullough ML *et al.* (2008) Adherence to a DASH-style diet and risk of coronary heart disease and stroke in women. *Archives of Internal Medicine* 168: 713–20.
- Gebauer SK, Chardigny JM, Jakobsen MU *et al.* (2011) Effects of ruminant trans fatty acids on cardiovascular disease and cancer: a comprehensive review of epidemiological, clinical, and mechanistic studies. *Advances in Nutrition* 2: 332–54.
- Givens DI (2008) Session 4: Challenges facing the food industry in innovating for health. Impact on CVD risk of modifying milk fat to decrease intake of SFA and increase intake of cis-MUFA. *Proceedings of the Nutrition Society* 67: 419–27.
- Givens DI (2012) Milk in the diet: good or bad for vascular disease? *Proceedings of the Nutrition Society* 71: 98–104.
- Givens DI & Shingfield KJ (2006) Optimising dairy milk fatty acid composition. In: *Improving the Fat Content of Foods*, pp. 252–80. Woodhead Publishing Ltd: Cambridge.
- Givens DI, Allison R & Blake JS (2003) Enhancement of oleic acid and vitamin E concentrations of bovine milk using dietary supplements of whole rapeseed and vitamin E. *Animal Research* 52: 531–42.
- Givens DI, Kliem KE, Humphries DJ *et al.* (2009) Effect of replacing calcium salts of palm oil distillate with rapeseed oil, milled or whole rapeseeds on milk fatty-acid composition in cows fed maize silage-based diets. *Animal* 3: 1067–74.
- Givens DI, Livingstone KM, Pickering JE *et al.* (2014) Milk: White elixir or white poison? An examination of the associations between dairy consumption and disease in human subjects. *Animal Frontiers* 4: 8–15.
- Glasser F, Ferlay A & Chilliard Y (2008) Oilseed lipid supplements and fatty acid composition of cow milk: a meta-analysis. *Journal of Dairy Science* 91: 4687–703.
- Goldbohm RA, Chorus AM, Galindo Garre F *et al.* (2011) Dairy consumption and 10-y total and cardiovascular mortality: a prospective cohort study in the Netherlands. *The American Journal of Clinical Nutrition* 93: 615–27.
- Hjerpsted J, Leedo E & Tholstrup T (2011) Cheese intake in large amounts lowers LDL-cholesterol concentrations compared with butter intake of equal fat content. *American Journal of Clinical Nutrition* 94: 1479–84.
- Hooper L, Summerbell CD, Thompson R *et al.* (2012) Reduced or modified dietary fat for preventing cardiovascular disease. *Cochrane Database of Systematic Reviews* 5: CD002137.
- Hu FB (2002) Dietary pattern analysis: a new direction in nutritional epidemiology. *Current Opinion in Lipidology* 13: 3–9.
- 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. *American Journal of Clinical Nutrition* 70: 1001–8.
- Hulshof KF, van Erp-Baart MA, Anttolainen M *et al.* (1999) Intake of fatty acids in western Europe with emphasis on trans fatty acids: the TRANSFAIR Study. *European Journal of Clinical Nutrition* 53: 143–57.
- Huth PJ & Park KM (2012) Influence of dairy product and milk fat consumption on cardiovascular disease risk: a review of the evidence. *Advances in Nutrition* 3: 266–85.
- Jackson KG & Lovegrove JA (2012) Impact of probiotics, prebiotics and synbiotics on lipid metabolism in humans. *Nutrition and Aging* 1: 181–200.
- Jackson KG, Walden CM, Murray P *et al.* (2012) A sequential two meal challenge reveals abnormalities in postprandial TAG but not glucose in men with increasing numbers of metabolic syndrome components. *Atherosclerosis* 220: 237–43.
- Jakobsen MU, Overvad K, Dyerberg J *et al.* (2008) Intake of ruminant trans fatty acids and risk of coronary heart disease. *International Journal of Epidemiology* 37: 173–82.
- Jakobsen MU, O'Reilly EJ, Heitmann BL *et al.* (2009) Major types of dietary fat and risk of coronary heart disease: a pooled analysis of 11 cohort studies. *American Journal of Clinical Nutrition* 89: 1425–32.
- Kai SH, Bongard V, Simon C *et al.* (2013) Low-fat and high-fat dairy products are differently related to blood lipids and cardiovascular risk score. *European Journal of Preventive Cardiology* 3: 3.
- Kesse-Guyot E, Vergnaud A-C, Fezeu L *et al.* (2010) Associations between dietary patterns and arterial stiffness, carotid artery intima-media thickness and atherosclerosis. *European Journal of Cardiovascular Prevention & Rehabilitation* 17: 718–24.

- Kris-Etherton PM, Grieger JA, Hilpert KF *et al.* (2009) Milk products, dietary patterns and blood pressure management. *American College of Nutrition* **28**: 1035–195.
- Labonte ME, Couture P, Richard C *et al.* (2013) Impact of dairy products on biomarkers of inflammation: a systematic review of randomized controlled nutritional intervention studies in overweight and obese adults. *American Journal of Clinical Nutrition* **97**: 706–17.
- Livingstone KM, Lovegrove JA & Givens DI (2012) The impact of substituting SFA in dairy products with MUFA or PUFA on CVD risk: evidence from human intervention studies. *Nutrition Research Reviews* **25**: 193–206.
- Livingstone KM, Lovegrove JA, Cockcroft JR *et al.* (2013) Does dairy food intake predict arterial stiffness and blood pressure in men?: Evidence from the Caerphilly Prospective Study. *Hypertension* **61**: 42–7.
- McCarron DA, Morris CD, Young E *et al.* (1991) Dietary calcium and blood pressure: modifying factors in specific populations. *American Journal of Clinical Nutrition* **54**: 215S–219S.
- Maes W, Van Camp J, Vermeirssen V *et al.* (2004) Influence of the lactokinin Ala-Leu-Pro-Met-His-Ile-Arg (ALPMHIR) on the release of endothelin-1 by endothelial cells. *Regulatory Peptides* **118**: 105–9.
- Mensink RP, Zock PL, Kester AD *et al.* (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. *American Journal of Clinical Nutrition* **77**: 1146–55.
- Mente A, de Koning L, Shannon HS *et al.* (2009) A systematic review of the evidence supporting a causal link between dietary factors and coronary heart disease. *Archives of Internal Medicine* **169**: 659–69.
- Micha R & Mozaffarian D (2010) Saturated fat and cardiometabolic risk factors, coronary heart disease, stroke, and diabetes: a fresh look at the evidence. *Lipids* **45**: 893–905.
- Mozaffarian D, Katan MB, Ascherio A *et al.* (2006) Trans fatty acids and cardiovascular disease. *New England Journal of Medicine* **354**: 1601–13.
- Mozaffarian D, Micha R & Wallace S (2010) Effects on coronary heart disease of increasing polyunsaturated fat in place of saturated fat: a systematic review and meta-analysis of randomized controlled trials. *PLoS Medicine* **7**: 1000252.
- Mozaffarian D, de Oliveira Otto MC, Lemaitre RN *et al.* (2013) Trans-palmitoleic acid, other dairy fat biomarkers, and incident diabetes: the Multi-Ethnic Study of Atherosclerosis (MESA). *American Journal of Clinical Nutrition* **97**: 854–61.
- Nicholls SJ, Lundman P, Harmer JA *et al.* (2006) Consumption of saturated fat impairs the anti-inflammatory properties of high-density lipoproteins and endothelial function. *Journal of the American College of Cardiology* **48**: 715–20.
- Nichols M, Townsend N, Scarborough P *et al.* (2012) European Cardiovascular Disease Statistics. European Heart Network, Brussels, European Society of Cardiology, Sophia Antipolis.
- Noakes M, Nestel PJ & Clifton PM (1996) Modifying the fatty acid profile of dairy products through feedlot technology lowers plasma cholesterol of humans consuming the products. *American Journal of Clinical Nutrition* **63**: 42–6.
- Nordestgaard BG, Benn M, Schnohr P *et al.* (2007) Nonfasting triglycerides and risk of myocardial infarction, ischemic heart disease, and death in men and women. *Journal of the American Medical Association* **298**: 299–308.
- Park KM & Cifelli CJ (2013) Dairy and blood pressure: a fresh look at the evidence. *Nutrition Reviews* **71**: 149–57.
- Patterson E, Larsson SC, Wolk A *et al.* (2013) Association between dairy food consumption and risk of myocardial infarction in women differs by type of dairy food. *Journal of Nutrition* **143**: 74–9.
- Ralston RA, Lee JH, Truby H *et al.* (2012) A systematic review and meta-analysis of elevated blood pressure and consumption of dairy foods. *Journal of Human Hypertension* **26**: 3–13.
- Salas-Salvado J, Garcia-Arellano A, Estruch R *et al.* (2008) Components of the Mediterranean-type food pattern and serum inflammatory markers among patients at high risk for cardiovascular disease. *European Journal of Clinical Nutrition* **62**: 651–9.
- SACN (Scientific Advisory Committee on Nutrition) (2007) Update on trans fatty acids and health. Position statement. Available at: http://www.sacn.gov.uk/pdfs/sacn_trans_fatty_acids_report.pdf (accessed 2 March 2014).
- Schachinger V, Britten MB & Zeiher AM (2000) Prognostic impact of coronary vasodilator dysfunction on adverse long-term outcome of coronary heart disease. *Circulation* **101**: 1899–906.
- Shingfield KJ, Chilliard Y, Toivonen V *et al.* (2008) Trans fatty acids and bioactive lipids in ruminant milk. *Advances in Experimental Medicine and Biology* **606**: 3–65.
- Shingfield KJ, Bonnet M & Scollan ND (2013) Recent developments in altering the fatty acid composition of ruminant-derived foods. *Animal* **1**: 132–62.
- Smith SC, Collins A, Ferrari R *et al.* (2012) Our time: a call to save preventable death from cardiovascular disease (heart disease and stroke). *Circulation* **126**: 2769–75.
- Soedamah-Muthu SS, Masset G, Verberne L *et al.* (2012) Consumption of dairy products and associations with incident diabetes, CHD and mortality in the Whitehall II study. *British Journal of Nutrition* **7**: 1–9.
- Sonestedt E, Wirfalt E, Wallstrom P *et al.* (2011) Dairy products and its association with incidence of cardiovascular disease: the Malmö diet and cancer cohort. *European Journal of Epidemiology* **26**: 609–18.
- Tardy AL, Morio B, Chardigny JM *et al.* (2011) Ruminant and industrial sources of trans-fat and cardiovascular and diabetic diseases. *Nutrition Research Reviews* **24**: 111.
- Toledo E, Delgado-Rodriguez M, Estruch R *et al.* (2009) Low-fat dairy products and blood pressure: follow-up of 2290 older persons at high cardiovascular risk participating in the PREDIMED study. *British Journal of Nutrition* **101**: 59–67.
- van Aerde MA, Soedamah-Muthu SS, Geleijnse JM *et al.* (2013) Dairy intake in relation to cardiovascular disease mortality and all-cause mortality: the Hoorn Study. *European Journal of Nutrition* **52**: 609–16.
- van Meijl LE & Mensink RP (2011) Low-fat dairy consumption reduces systolic blood pressure, but does not improve other metabolic risk parameters in overweight and obese subjects. *Nutrition, Metabolism & Cardiovascular Diseases* **21**: 355–61.
- Vafeiadou K, Weech M, Sharma V *et al.* (2012) A review of the evidence for the effects of total dietary fat, saturated, monounsaturated and n-6 polyunsaturated fatty acids on vascular function, endothelial progenitor cells and microparticles. *British Journal of Nutrition* **107**: 303–24.

- Van Bortel LM, Laurent S, Boutouyrie P *et al.* (2012) Expert consensus document on the measurement of aortic stiffness in daily practice using carotid-femoral pulse wave velocity. *Journal of Hypertension* **30**: 445–8.
- Vlachopoulos C, Aznaouridis K & Stefanadis C (2010) Prediction of cardiovascular events and all-cause mortality with arterial stiffness: a systematic review and meta-analysis. *Journal of American College of Cardiology* **55**: 1318–27.
- Weaver CM, Proulx WR & Heaney R (1999) Choices for achieving adequate dietary calcium with a vegetarian diet. *American Journal of Clinical Nutrition* **70**: 543s–548s.
- Weng L-C, Steffen L, Szklo M *et al.* (2013) A diet pattern with more dairy and nuts, but less meat is related to lower risk of developing hypertension in middle-aged adults: the Atherosclerosis Risk in Communities (ARIC) Study. *Nutrients* **5**: 1719–33.
- Wennergren MH, Smedman A, Turpeinen AM *et al.* (2009) Dairy products and metabolic effects in overweight men and women: results from a 6-mo intervention study. *American Journal of Clinical Nutrition* **90**: 960–8.
- WHO (World Health Organization) (2008) Interim summary of conclusions and dietary recommendations on total fat and fatty acids. Available at: http://www.who.int/nmh/publications/ncd_report2010/en/ (accessed 5 December 2013).
- WHO (World Health Organization) (2011) Global status report on noncommunicable diseases 2010. Available at: http://www.who.int/nmh/publications/ncd_report_full_en.pdf (accessed 5 December 2013).