REVIEW

Milk for Skeletal Muscle Health and Sarcopenia in Older Adults: A Narrative Review

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Abstract: Skeletal muscle aging manifests as a decline in muscle quantity and quality that accelerates with aging, increasing the risk of sarcopenia. Sarcopenia is characterized by a loss of muscle strength and mass, and contributes to adverse health outcomes in older adults. Intervention studies have shown that sarcopenia may be treated by higher protein intake in combination with resistance exercise (RE). In comparison, less is known about the role of whole protein-containing foods in preventing or treating sarcopenia. Liquid milk contains multiple nutrients and bioactive components that may be beneficial for muscle, including proteins for muscle anabolism that, alone or with RE, may have myoprotective properties. However, there is a lack of evidence about the role of milk and its effects on muscle aging. This narrative review considers evidence from three observational and eight intervention studies that used milk or fortified milk, with or without exercise, as an intervention to promote muscle health and function in older adults (aged 50–99 years). The observational studies showed no association between higher habitual milk consumption and muscle-related outcomes. The results of intervention studies using fortified milk in relation to elements of sarcopenia were also negative, with further inconclusive results from the studies using a combination of (fortified) milk and exercise. Although milk contains nutrients that may be myoprotective, current evidence does not show beneficial effects of milk on muscle health in older adults. This could be due to high habitual protein intakes (>1.0 g/kg BW/d) in study participants, differences in the type of milk (low-fat vs whole) and timing of milk consumption, length of interventions, as well as differences in the sarcopenia status of participants in trials. Adequately powered intervention studies of individuals likely to benefit are needed to test the effectiveness of a whole food approach, including milk, for healthy muscle aging.

Keywords: sarcopenia, muscle health, whole foods, milk, myoprotective properties, older adults

Introduction

The natural loss of skeletal muscle mass after the age of 30 accelerates in later life.[1] There are even steeper losses in muscle strength and power,[1,2] adversely affecting physical functioning in older adults and increasing the risk of sarcopenia. Sarcopenia—loss of skeletal muscle strength and mass[1,3]—is associated with adverse health outcomes in older adults, including frailty, falls, disability, hospitalization, and earlier death.[3–8] Sarcopenia has a complex pathogenesis and no pharmacological treatments have yet been shown to be effective. Several cellular processes including low-grade inflammation, oxidative stress, anabolic resistance, motor unit denervation and mitochondrial dysfunction are thought to contribute to reduced myofibre quantity and quality with aging.[9,10] Current recommendations for

treatment and prevention of sarcopenia focus on modifiable lifestyle factors such as diet and nutrition in combination with physical exercise[9,11] mostly using a single nutrient approach.[11–16] Using a whole food or whole diet approach, accounting for the combined effect of multiple food components on muscle has emerged as a treatment paradigm only recently.[17] For example, a protein intake higher than the recommended daily allowance of 0.8 g/kg body weight/day (g/kg BW/d) has been debated as optimal for maintaining and regaining muscle mass and better physical performance in healthy older adults[11–15]—preferably by consuming high-quality, protein-rich foods distributed across meals.[16] Dietary protein sources and isolated protein supplements such as whey that have a higher content of branched-chain amino acids (BCAA) (eg, 13.4% leucine in whey)—an established regulator of muscle protein anabolism[18]—have been regarded as superior for muscle mass and performance in younger and older adults.[19,20] However, less is known about how whole protein-containing foods, that are also rich in other essential nutrients and bioactive components, may be myoprotective and influence muscle health and function in older adults.[21] Protein-containing foods that are affordable and easy to prepare also need to be acceptable to older adults and a sustainable dietary source for the environment.[22,23] Bovine milk is an attractive candidate whole food for evaluation because of its nutrient composition and potential benefits for human health.

The purpose of this narrative review is threefold: (1) to discuss the potential myoprotective properties of liquid milk; (2) to summarize and discuss findings from observational and intervention studies of the effects of milk, with or without exercise, in relation to muscle-health outcomes and sarcopenia in older adults, and (3) to discuss the evidence needed to inform future interventions with whole foods, including dairy foods, for healthy muscle aging.

Potential Myoprotective and Health Benefits of Liquid Milk
Nutritional Composition of Milk

Milk and milk products (eg, yogurts and cheeses) are examples of whole foods dense in nutrients that may have potential for improving muscle mass[21] and performance,[24] and therefore increasing consumption could be a preventive strategy for sarcopenia.[25] Liquid milk, an important part of a healthy diet for over six billion people,[26] contains a range of nutrients[27] and bioactive components[28] that are potentially valuable for human health.[26] On average, whole (bovine) milk provides high-quality proteins (20% of whey and 80% of caseins), minerals (eg, calcium, phosphorus, magnesium, iodine), vitamins (eg, fat-soluble A and E and water-soluble B vitamins), carbohydrates (lactose and oligosaccharides), and fats—a mixture of 70% saturated (SFA), and 30% of mono- (MUFA) and polyunsaturated fatty acids (PUFA).[26]

Specifically, in addition to its average protein content of 32 g/L, daily consumption of 500 mL of whole milk would make a significant contribution to other daily nutrient intakes; approximating 15–20% of the dietary recommended intake for vitamin A (280 µg/L); 60–80% for riboflavin (1.83 mg/L); 90% for vitamin B_{12}; 40–50% for calcium (1.1 g/L); 18–25% for zinc (4 mg/L); 30% for selenium (37 µg/L); 12–16% for magnesium (100 mg/L) and 50% of the requirements for iodine (for an average of 160 µg/L) in US adults.[26,29]

Beyond protein, milk also provides bioactive peptides, which have several physiological effects, including antihypertensive, antithrombotic, antimicrobial and immunomodulatory effects.[28,30] Briefly, α- (13 g/L) and β-caseins (9.3 g/L) are precursors of several peptides, including those involved in the inhibition of angiotensin-converting enzyme (ACE),[31] whilst β-lactoglobulin (3.2 g/L) is recognized as a potential anti-oxidant and retinol carrier, and immunoglobulins (A, M, and G; 0.7 g/L) and lactoferrin (0.1g/L) have been linked to immunoprotection (reviewed in Mills et al).[28] In addition, milk contains essential fatty acids such as PUFA (2.3% of total FA or 2 g/L)[27,28]—although in low amounts—that can be manipulated by different farming practices (animal diet, management and season).[32] Specifically, milk PUFA include linoleic acid (LA 18:2 n–6; 18 g/kg of total FA), α-linoleic (ALA 18:3 n-3; 6.9 g/kg of total FA), and long-chain n-3 FA (LC n-3; 1.8 g/kg of total FA) that are metabolically active and have specific functions in cell membranes.[27,32]

In summary, milk is a complex food that contains a number of nutrients and other biologically active components that are beneficial for human health. Milk is also an established part of the diet for many older adults who are, in general, higher consumers compared with younger adults,[33] and thus may provide them with nutrients and bioactive components relevant for aging muscle.

Milk and Human Health

Epidemiological studies have reported associations between higher milk/dairy intake, and better health and functioning across the life course. For example, higher

milk/dairy intake has been associated with better bone mineralization in early life, and slower age-related bone loss and reduced risk of hip fractures in later life;[34–37] favorable or neutral associations with different cardiovascular clinical outcomes (coronary artery disease, stroke, type 2 diabetes, hypertension, and metabolic syndrome);[38] beneficial associations with body composition,[39] and reduced risk of obesity[40] and colorectal cancers in middle-aged and older adults.[41]

Higher intake of milk was positively associated with performance in the 20-m endurance test and negatively associated with BMI in children aged 9–12 years, thought to be mediated through greater dietary intakes of milk-derived vitamin B_2 and B_{12}, which contributed to higher cardio-respiratory fitness.[42] In addition, intervention studies with fat-free and skimmed milk in combination with RE in young adults and athletes have shown higher lean muscle, fat mass loss, increased muscle strength,[43,44] and greater muscle protein synthesis (MPS),[45] compared with an isocaloric carbohydrate or soy-based drink. Ingestion of milk (including chocolate milk) post-exercise has been shown to attenuate exercise-induced muscle damage and soreness, increase MPS, and rehydrate muscles in athletes and young, active adults as effectively or even better that any other commercially available recovery drink.[46,47] Taken together, studies in young adults highlight the potential health benefits of milk for muscle beyond anabolic effects of its main protein constituents (whey and caseins). However, little is known about how milk consumption may affect muscle health and function in older adults who are at high risk of sarcopenia and associated adverse health outcomes.

Potential Mechanisms of Milk Effects on Skeletal Muscle Health in Older Adults

A whole food approach in investigating the relationship between protein-rich foods and muscle with aging posits a number of myoprotective properties of unfortified milk, beyond those that are proanabolic (ie, MPS-promoting). These properties include anti-oxidative, anti-inflammatory, and immunomodulatory effects (Figure 1).

Proanabolic Effect of Milk-Based Proteins: An Example of Whey

Whey proteins in milk (such as β-lactoglobulin, α-lactalbumin, lactoferrin, and immunoglobulins) are characterized as a fast-digestible proteins giving rise to a high concentration of essential amino acids (EAA) available to support MPS post-digestion.[12,27,48] Whey is a rich source of the BCAA leucine (13.4% or 122 mg/g of whey),[30] which appears to be the most important EAA to stimulate MPS in skeletal muscle[18,48] through the rapamycin (mTOR) pathway—a key regulator of human MPS in response to increased EAA.[49] The ability of leucine to induce MPS has been explained by the "leucine threshold" hypothesis and demonstrated in numerous human and animal trials (discussed in Devries et al).[48] The hypothesis posits that, in order to increase MPS after protein ingestion, the intracellular leucine concentration in myofibres has to reach a desired level, which can be altered by other stimuli, lowered by RE and increased by aging and sedentary lifestyle.[48] For younger active men, the amount of leucine needed to exceed the threshold and induce maximal MPS has been estimated to be 1.7 to 2.4 g provided in 20 g of high-quality protein, or in a per meal feeding dose of 0.25 g protein/kg BW.[50] For older adults, a higher protein (leucine) intake for MPS has been suggested because of anabolic resistance, and estimated to be 0.38 g/kg BW per meal.[50] Numerous trials have shown a synergistic effect of protein supplementation and RE when protein ingestion follows a bout of RE, resulting in a greater MPS compared with either stimulus alone (discussed in Devries et al).[48] The whey (leucine) potential to augment the anabolic effect of prolonged exercise (>6 weeks) has been show for fat-free mass and one repetition maximum (1-RM) leg press strength in a meta-analysis that included six trials with older adults (aged >50 years), a mean difference of 0.91 kg ($p<0.0001$) and 20.7 kg ($p<0.005$), respectively.[51]

Anti-oxidative, Anti-inflammatory, and Immunomodulatory Properties of Milk Bioactive Components

Several lines of research have shown that oxidative stress and accumulation of reactive oxygen and nitrogen species (ROS/RNS) in aging muscle impairs cellular homeostasis and causes damage to key cell biomolecules (ie, proteins, nucleic acids, lipids) and organelles[52] contributing to sarcopenia.[9,10] A number of milk-derived bioactive peptides (eg, β-lactoglobulin, lactoferrin),[26–28,30] lipids and fatty acids (eg, α-linoleic acid, the milk fat globule membrane (MFGM) lipids and glycoproteins),[26–28,53] and minerals (eg, selenium, zinc) have been shown to have anti-oxidative properties, which may add to the exogenous antioxidant capacity of a balanced diet in neutralizing ROS/RNS in myofibres. An anti-inflammatory effect is
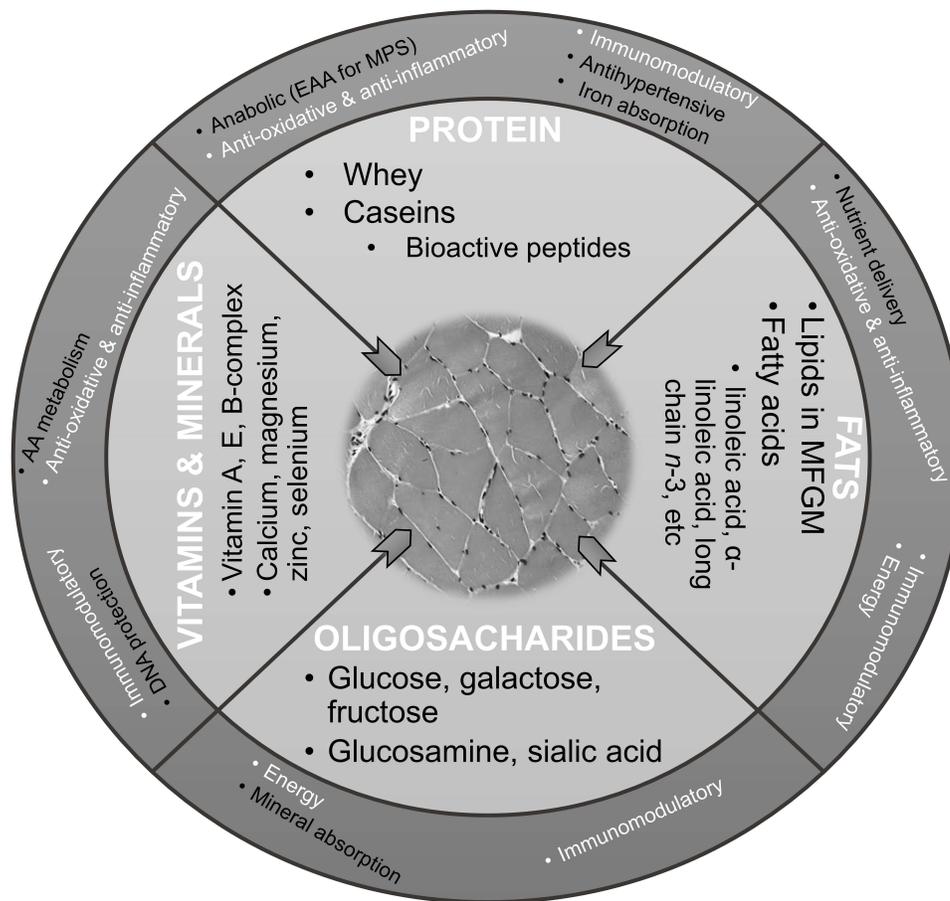
Figure 1 Hypothesized myoprotective properties of nutrients in liquid milk.

Notes: Hypothesized health effects and function of milk nutrients and bioactive components on muscle may include energy, minerals and vitamin delivery, anabolic, anti-oxidative, anti-inflammatory, and immunomodulatory pathways. Common pathways across the nutrients and non-nutrients are presented in the outer circle in white. © Newcastle University. This work is licensed under a Creative Commons Attribution-Noncommercial-Share Alike 3.0 Unported License https://creativecommons.org/licenses/by-nc-sa/3.0/.

Abbreviations: EAA, essential amino acids; MFGM, milk fat globule membrane; MPS, muscle protein synthesis.

the other bioactivity associated with milk constituents (such as casein-derived bioactive peptides[26,28] and *n*-3 PUFA) that may ameliorate inflammaging in muscle by reducing cytokine load (eg, by decreasing levels of interleukin 6 and 8 (IL-6 and IL-8) and tumor necrosis factor α (TNF-α)).[52] Inflammaging, the age-related chronic low-grade inflammation characterized by higher concentrations of pro-inflammatory mediators in serum and plasma, have been linked to worse age-related pathologies[9,10,54] and loss of muscle mass and function.[52,54] The n-3 PUFAs have been proposed as a therapeutic agent for sarcopenia because of their anti-inflammatory properties, anabolic effect on skeletal muscle metabolism through mTOR activation, and reduction of insulin resistance.[55] Although investigations about the role of immune function in sarcopenia lag behind other mechanistic studies, a decline of innate immunity and its link with inflammaging has been postulated in the pathogenesis of frailty and sarcopenia.[56]

Milk contains several immunomodulatory components (eg, immunoglobulins, lactoferrin, α-lactalbumin)[28] that may act against cytokine-derived inflammation.

Furthermore, antihypertensive bioactive peptides found in milk (eg, lactopeptides α-lactorphin and β-lactorphin released from α-lactalbumin and β-lactoglobulin, respectively) acting as ACE inhibitors[28,31] may have potential as a therapy for sarcopenia. ACE inhibitors (including those that are milk-derived) may have multiple beneficial effects on aging muscle,[57] including the ability to reduce inflammation, promote glucose uptake, and improve endothelial function, angiogenesis and muscle blood flow.[9,58] Synergistic and cumulative actions of milk-derived bioactive components through these pathways may enhance milk's proanabolic effects post-exercise. However, evidence is very limited and further research is needed to understand the bioactive potential of milk for healthy muscle aging. For example, future studies with older

adults are needed to determine whether the presence of fats (fatty acids) in whole milk enhances absorption of EAA for MPS as observed in young athletes post-exercise,[59] and in combination with other myoprotective effects, including the delivery of fat-soluble vitamins (eg, vitamin A, E, K) that may be relevant for muscle health.

In summary, myoprotective effects of milk may work through anabolic, anti-oxidative, anti-inflammatory and immunomodulatory pathways associated with the main nutrients in milk, including proteins, fats, vitamins and minerals, and milk sugars (Figure 1). Their synergistic and cumulative action may provide myoprotection beyond proanabolic effects of milk proteins such as whey.

Evidence About the Role of Liquid Milk in Skeletal Muscle with Ageing
Materials and Methods

For the narrative summary of evidence, systematic searches of four electronic databases (MEDLINE, Embase, Web of Science, CENTRAL) were conducted independently by two researchers (LD, CH) for articles published in English in the period from January 2010 until June 2019. Observational and intervention studies were searched using the following search terms and their combination: "liquid milk", "fluid milk", "whole milk", "skimmed milk", 'milk-based drinks', "fortified milk", "grip strength", "physical performance", "muscle mass'', "sarcopenia", "older adults", "elderly", "intervention", "randomized controlled trial" and "cohort study". Only full-text articles with clearly described populations (eg, community-dwelling older adults aged ≥65 years), exposure and study arms (milk, fortified milk with or without exercise; excluding reconstituted milk-based drinks from powder or those insufficiently defined), study duration, outcome (muscle mass, strength, power, physical performance, sarcopenia), and power to support the conclusions were critically evaluated (CH, LD) and selected for the review. Relevant information from the selected articles were extracted independently (CH, LD) and compared, including study name, duration, sex and age of participants, sample size, exposure, outcomes, main findings, and study limitations.

Results
Evidence from Observational Studies

Three observational studies (two articles) assessed evidence about the association between liquid milk consumption and muscle-related outcomes (Table 1). [60,61] One study aimed to address the association between habitual consumption and physical performance in old age using data from the Boyd Orr study and the Caerphilly Prospective Study (CaPS).[60] In the Boyd Orr study, a week-long dietary household inventory was completed by families, involving 4999 children (aged 0–19). Sixty-five years later, 405 men and women (mean age 70.7 years) took part in a detailed clinical examination involving physical performance tests. Individual consumption of whole milk (grams/day) was estimated from the household inventory. A higher childhood milk intake was associated with 5% faster walking times assessed by Timed Up-and-Go (TUG) test, and better balance in later life. However, the study found no association between adult milk intake and walking time, and a negative association between higher milk consumption and balance (Table 1). Contrasting results were observed in the CaPS study of 1195 men—a unit increase in whole milk intake (half a pint milk/day) at dietary assessment (age range: 59–73 years) was associated with 21% lower risk of poor balance at follow-up (age range: 66–86 years). The Boyd Orr findings suggest ensuring adequate milk consumption is a potentially modifiable health behavior in early years, to enhance physical performance in old age.

In the Helsinki Birth Cohort Study, a prospective study of over 1000 older adults (mean age 61 years at baseline) that assessed the association between a healthy Nordic Diet (ND) and physical performance at 10-year follow-up, low consumption of low-fat milk at baseline was independently associated with better overall Senior Fitness Test score in men but not in women.[61] Low-fat milk (skimmed and milk with fat content <2%) consumption, a favorable component of ND score, was estimated from food frequency questionnaires and assigned a value of 0–3 according to ascending sex-specific quartiles. The findings may have limited generalizability because those who were assessed at follow-up had healthier diets at baseline, were younger and more highly educated, and thus may not be representative of the general older adult population in Helsinki. In addition, uncontrolled confounding (eg, change in diet over time) and sex-specific differences in food choices (including milk) may have biased the results.

In summary, observational research investigating the association between liquid milk consumption, muscle strength and physical performance in older adults is scarce. We found only three studies focussed on older adults, and none of them have investigated the role of

Table I Observational Studies Investigating the Association between Milk Consumption and Muscle-Related Outcomes in Older Adults

Reference	Country	Study	Duration	Age M (SD)	Sex	Sample	Health	Exposure	Outcome	Comparator	Results
Birnie et al[60]	UK	Boyd Orr	65 years	Baseline (1937/39): 0–19 years	F, M	4999	BMI 27.5 (4.4)	Childhood daily milk intake (7-day household inventory)		Bottom third of milk intake (g)	(1) 5% ↑ walking speed at follow-up (2002) if in top third of childhood milk intake (2) 25% ↓ risk of being unable to balance for 5 s/SD ↑ in milk intake
				Follow-up (1997): 58–60 years	F, M	1648		Daily milk intake (FFQ)			
				Follow-up (2002): 70.7 (4.3) years	F, M	405		Daily milk intake (average FFQ 1997–2002)	(1) Walking time (get-up and go test) (2) Balance (flamingo test eyes open)	Lower milk group (by half pint)	(1) ↔ Between adult milk intake and walking time (2) 19% ↑ risk of poor balance with each increase in the milk group
		CaPS	25 years	Baseline (1979–83): 45–59 years	M	2512					
				Phase V (2002/04): 75.3 (4.3)	M	1195	BMI 27.8 (4.1)	Daily milk intake (average FFQ Phase III (1989–93) and Phase IV (1993–96))	(1) Walking time (get-up and go test) (2) Balance (flamingo test eyes open)	Lower milk group (by ½ pint)	(1) ↔ Between adult milk intake and walking time (2) 21% ↓ risk of poor balance with each increase in the milk group
Perala et al[61]	Finland	Helsinki Birth Cohort Study	12 years	Baseline (2001/04): 57–67 years	M, F	1094	BMI 27.2 (0.3) men BMI 27.1 (0.4) women	Low-fat milk (FFQ) as a part of the NDS (score 0–25)			
				Follow-up (2011/13): 67–77 years	M, F	1072			Senior Fitness Test (score 5–100)	Lower milk intake based on the lowest quartile of NDS (score <10)	Lower consumption of milk (NDS <10) associated with ↑ overall SFT

Notes: ↑, increase; ↓, decrease; ↔, no change or association
Abbreviations: BMI, body mass index; CaPS, Caerphilly Prospective Study; FFQ, food frequency questionnaire; NDS, Nordic Diet Score.

milk specifically in individuals with sarcopenia. Whilst higher childhood whole milk consumption might foster better physical performance and balance in later life, the results for the role of milk in muscle health and function in older adults were mixed, warranting further prospective research in older cohorts.

Evidence from Liquid Milk and Fortified Milk Intervention Studies with and without Exercise

Main Characteristic of the Studies: Participants, Intervention and Outcomes

In a recent meta-analysis of 14 randomized controlled trials (RCT) that investigated the efficacy of dairy protein supplementation on muscle strength, mass, and function in middle-aged and older adults (with or without sarcopenia), an increase in appendicular muscle mass (AMM) was observed with higher dairy protein intake, whilst the results for physical functioning were inconclusive.[21] However, the evidence was based mainly on isolated dairy-source proteins with only two trials using whole foods (cheese).

To date, only a limited number of intervention studies have evaluated the effects of liquid milk or fortified milk (ie, with additional nutrients) on measures of muscle health and physical performance in older adults. These studies typically involve either: (1) milk supplementation alone or (2) a combination of milk supplementation and exercise training and are summarized in Table 2. Briefly, we identified eight intervention studies[62–69] (two with milk supplementation alone[62,63]) and six with combined supplementation and exercise[64–69] involving healthy (three studies[65,67,68]) or (pre)sarcopenic community-dwelling older adults (two studies[63,66]), and older adults with physical impairments living in residential care facilities (three studies[63,64,69]). Seven were RCTs;[63–69] three included only men.[65–67] The studies enrolled older adults aged 50 to 99 years in samples ranging from 26 to 177 participants, with interventions lasting from 12 weeks to 18 months. Only one used fortified whole milk,[62] and the remainder used reduced fat milk/low fat milk ($\leq$1.6% fat)[63–69] of which two used chocolate milk.[66,67] Milk was fortified with several nutrients in all but one study,[67] including protein,[63,64,66,68,69] calcium and vitamin D[62,65] and EAA.[66] The amount of protein ingested via fortified milk varied from 10.5 g[68] to 40 g[63] consumed either every day or after exercise on the training days over the study period.

Muscle health and function-related outcomes also varied considerably across the studies, and included the measures of muscle mass (eg, total muscle mass, lean muscle mass, skeletal muscle index (SMI)),[63–66,68] strength (eg, leg press, chest press, grip strength),[62,63,65,67,68] and physical performance (eg, TUG, stair climb, chair rises, gait speed, balance).[62,63,65,66,68,69]

Interventions Involving Fortified Milk: Evidence

A study of 107 care residents (mean age 79.9±10.1 years) found no association between fortified whole milk intake, TUG and grip strength following a six-month intervention designed to promote milk consumption.[62] There was a trend for slower TUG time (mean (SD) −2.56 (15.6), p=0.07) over the study period, but no difference in GS pre- and post-intervention (17.4±0.9 kg vs 17.6±0.8 kg, p=0.7). However, the supply of milk to residents' diets (added to drinks and cereal) was left to the discretion of the nursing staff, aiming for a mean intake of 210 mL/day per resident. The median consumption was 160 mL/day and estimated based on self-reports. Consistent with this finding, a study of 50 community-dwelling older adults with reduced physical functioning, who were provided with 2x400 mL of protein-enriched milk (2x20 g protein) each day for 12 weeks, also showed no improvements in muscle mass, strength or functional performance when compared with a control group consuming an isocaloric, non-nitrogenous control drink.[63] Specifically, although chest press improved significantly in the protein (1.3 kg (0.1–2.5), p=0.03) and control group (1.5 kg (0.0–3.0), p=0.048), no difference between the groups (p=0.9) were observed. Furthermore, no significant change in leg press (p=0.9) or muscle mass (p=0.54), or the physical performance tests (ie, chair rise, stair climb, and GS test; p>0.05 for all tests) were observed between the groups after the 12-week intervention. Importantly, in both studies baseline protein intake was >0.8 g/kg BW/d, which would be expected to have affected the potential impact of (fortified) milk on muscle mass and function.

Interventions Involving Fortified Milk and Exercise: Evidence

In the longest duration intervention to date, an 18-month intervention, Kukuljan et al[65] found no beneficial effect of fortified milk consumption (2x200 mL of reduced fat UHT milk consumed daily supplemented with Ca and vitamin D), compared with resistance training alone on skeletal muscle size, strength or function in healthy older men with a higher

Table 2 Intervention Studies Investigating the Effect of Fortified Milk with or without Exercise on Muscle-related Outcomes in Older Adults

Reference	Country	Study Design	Duration	Participants			Intervention	Results	
				Age M (SD) Years	Sex	Sample		Compliance	Main Effects
Fortified milk interventions									
Grieger and Nowson[62]	Australia	Single group pre-postintervention	6 months	79.9 (10.1)	M, F	70 (in high-level care nursing, HLC) 37 (low-level care hostels, LLC)	(1) 1 L of fortified milk (190 mg Ca; 5 µg cholecalciferol; 75 µg folate/100 mL)/day added to residents' hot drinks, cereal and porridge by the nursing staff (2) Weekly investigators visits to encourage and record consumption	(1) Consumption over 6 months, mean (SD): 178 (132) mL (2) Higher in HLC compared with LLC (195 (17) mL vs 147 (17) mL) (3) No sex-specific differences in consumption (4) No difference in the estimated mean volume at the beginning (211 (16) mL) or end of the study (196 (15) mL)	↔ Between fortified milk intake and TUG or GS
Ottestad et al[63]	Norway	Double-blinded RCT	12 weeks	≥70	M, F	50	(1) 2×400 mL/day of protein-enriched milk (5.1 g protein; 4.9 g carbohydrate; 0.1 g fat/100 mL) with breakfast and evening meal (2) Isocaloric carbohydrate drink (2×400 mL/day) Groups: protein-enriched milk (n=24) placebo (n=26)	(1) 70% completed the study (2) 97.8% (3.8) in the protein-enriched group	(1) ↔ In muscle mass and total lean body mass within and between groups (2) ↑ In chest press in the protein-enriched milk group (1.3 kg (0.1 to 2.5), p=0.03) and the control group (1.5 kg (0.0 to 3.0), p=0.048) with no significant difference between the groups (p=0.85) (3) ↔ In leg press within or between the groups (p=0.93) (4) ↑ In stair climb test (without load) in the protein group (−0.4 s (−0.8 to 0.1), p=0.03), but not in the control group (0.0 s (−0.7 to 0.7), p=0.96); ↔ in the change in the protein group versus the control group (p=0.22) (5) ↔ Differences between the two groups in the changes in the functional performance tests: repeated chair rise, stair climb test (with 10 kg load), and hand grip strength

Fortified milk and exercise interventions							
Carlsson et al[64]	Sweden	Stratified cluster-RCT (double-blinded)	3 months	84.5 (6.4)	M, F	177	(1) 200 mL of milk-based protein enriched drink (7.4 g protein, 15.7 g carbohydrate and 0.43 g fat/100 g) five minutes after exercise programme (five sessions every two weeks) (2) 45 minutes exercise program (five sessions every two weeks) to improve lower-limb strength, balance and gait with exercises in functional weight-bearing positions (at least two lower-limb strength exercises and two balance exercises with progressing intensity) Groups and follow-up: enriched milk + exercise (n=36 at 3 months; n=37 at 6 months) placebo + exercise (n=30 at 3 months; n=26 at 6 months) enriched milk + control (n=40 at 3 months; n=37 at 6 months) placebo + control (n=43 at 3 months; n=39 at 6 months)	(1) The protein-enriched drink taken in 84% of all occasions and completely emptied on 82% of these occasions (2) 79% attendance level for the exercise group	(1) ↔ Difference in ICW and BW in any of the groups at 3-month follow-up in the exercise group compared to the control group, nor in the protein group compared to the placebo group. (2) ↔ Interaction effects (3) ↔ Within-group changes in ICW and BW in the exercise, control, protein or placebo groups

(Continued)

Table 2 (Continued).

Reference	Country	Study Design	Duration	Participants			Intervention	Results		Main Effects
				Age M (SD) Years	Sex	Sample		Compliance		
Kukuljan et al[65]	Australia	RCT	18 months	50–79	M	172	(1) 400 mL/day (2x200 mL) of reduced fat ultrahigh temperature milk (6.6 g protein, 2.2 g fat, 418 kJ/200 mL fortified with calcium (~500 mg calcium) and vitamin D3 (400 IU)) (2) Progressive resistance training performed on three nonconsecutive days/week for 18 months (session duration: 60–75 minutes) (3) Groups: fortified milk + exercise (n=43) exercise (n=44) fortified milk (n=43) control (n=42)	(1) ↔ Difference in exercise compliance between fortified milk + exercise (65%) and the exercise group (61%) (2) ↔ Difference in fortified milk compliance between the milk + exercise group (92%) and the milk group (89%)		(1) ↔ Effects of the fortified milk on muscle size, strength, or function after 18 months (2) ↑ In total body lean mass and mid-femur muscle CSA in the fortified milk + exercise group compared with either group alone or the control group; ↔ in the interaction terms for any muscle or functional parameter (3) 22–56% ↑ (all $p<0.001$) in upper and lower body muscle strength after 12 months of training in the exercise groups with a plateau thereafter, except in leg muscle strength (4) 11% ↑ in gait speed ($p<0.05$) after 18 months in the exercise group compared with the nonexercise group (5) ↔ Effects of exercise on any functional parameter (6) ↔ Within-group changes for lean mass, muscle CSA in the milk alone group (7) ↔ Main effects of milk on muscle strength or any functional measure

	Country	Design	Duration	Age	Sex	N	Intervention	Adherence	Outcomes
Maltais et al[66]	Canada	Double-blinded RCT	4 months	60–75	M	26	(1) Milk powder added to 1% fat chocolate cow milk (375 mL, 13.53 g protein, 7 g EAA (3.5 g from leucine), 37.5 g carbohydrate, 3.8 g fat; 270 kcal); EAA supplement from soy (12 g protein, 7g EAA); rice milk (nonprotein control); consumed immediately after exercise (2) Resistance training three on nonconsecutive days comprising of free weightlifting and RE equipment for leg press, bench press, leg extension and shoulder press, sit-ups, rowing extensions, bicep curls, and leg curls (3 8 repetitions); 6 to 8 repetitions at 80% of 1RM Groups: control + RE (n=10) EAA (soy) + RE (n=8) milk + RE (n=8)	>90%, on average, for all participants	(1) ↑ Lean body mass, muscle mass index and total muscle mass in all groups (2) ↔ Between-group changes (3) Each group improved at least one parameter of muscle strength (lateral pull down or bench press) (4) ↑ In the TUG test in the EAA group but not in the milk + RE and control + RE group
Mitchell et al[67]	Canada	RTC	12 weeks	74.4 (5.4) and 22.4 (2.1)	M	16 old adults 16 young adults	(1) 500 mL of 1% fat chocolate milk (320 kcal, 14 g protein, 5 g fat, 54 g carbohydrate) or placebo (310 kcal, 0.4 g protein, 5 g fat, 66 g carbohydrate)/day after exercise on training days and with breakfast on nontraining days (2) Resistance training 3x week for 12 weeks (2x lower-body exercise; 1x upper-body exercise) Groups: chocolate milk + RE placebo + RE for two age groups	Not reported	(1) ↑ I all muscle strength measure (MVC, leg press, leg extension, chest press) by the main effect of RE ($p<0.005$) (2) ↔ Effect of milk on any strength measure (3) ↔ Interactive effect of milk supplementation and RE for any strength measure ($p=0.30–0.86$) (4) ↑ In type I fiber area with RE ($p=0.008$) (5) ↔ Interactive effect of milk + RE on type I fiber area (6) ↔ Difference in type I fiber area between older and young men (7) ↑ In type II fiber area by the main effect of RE in young but not in old men ($p=0.048$) (8) ↔ Interactive effect of milk + RE on type II fiber area

(*Continued*)

Table 2 (Continued).

| Reference | Country | Study Design | Duration | Participants | | | Intervention | Results | |
				Age M (SD) Years	Sex	Sample		Compliance	Main Effects
Osuka et al[68]	Japan	RTC	12 weeks	65–79	M, F	56	(1) 250 mL fortified milk (10.5 g protein, 3.9 g fat, 9.3 g carbohydrate)/day after exercise on training days and as a snack between meals on nontraining days (2) Exercise intervention 2x week 12-weeks Groups: resistance training (RT) (chest press, leg extension, leg curl, leg press performed at progressive intensity (30–70% 1 RM) 3x10–12 reps) + fortified milk (n=28) aerobic training + resistance training (ART) (40–50% VO_{2peak}, cycle ergometer, 20–30 min duration + same RT) + fortified milk (n=28)	(1) Training attendance: 88.8% (RT) and 91.4% (ART) (2) Fortified milk intake: 95.1% (RT) and 95.4% (ART)	(1) ↑ In SMI, whole-body muscle mass, upper and lower extremity muscle mass, leg extension, leg curl, leg press, chest press, arm curl, and sit-to-stand in the RT group ($p<0.05$) (2) ↑ In lower extremity muscle mass, leg extension, leg curl, leg press, chest press, arm curl, and sit-to-stand ($p<0.05$) in the ART group (3) ↔ Differences in the changes in muscle mass and muscle strength between the two groups, (4)↑ in the changes in arm curl and sit-to-stand in the ART group compared with the RT group

| Rosendahl et al[69] | Sweden | Stratified cluster-RCT (double-blinded) | 3 months (29 sessions) | 84.7 (6.5) | 191 | (1) 200 mL of milk-based protein-enriched energy supplement (7.4 g protein, 15.7 g carbohydrate; 408 kJ/100 g) or 200 mL of placebo drink (0.2 g protein, 10.8 g carbohydrate; 191 kJ/100 g) consumed 5 minutes after exercise (2) The High-Intensity Functional Exercise Program (The HIF Program) 5x week every two weeks/3 months (45 minutes each) consisting of everyday tasks challenging leg strength, postural stability, and gait ability with progressive difficulty or the control activity program Groups and follow-up: The HIF Program + milk supplement (n=46) The HIF Program + placebo (n=45) Control Program + milk supplement (n=50) Control program + placebo (n=50) after 3 and 6 months | The HIF Program group attendance: 72% | (1) ↑ In self-paced gait speed in the exercise group compared with the control group (mean difference 0.04 m/s, p=0.02) at 3 months (2) ↑ In the Berg Balance Scale in the exercise compared with the control program group (1.9 points, p=0.05), self-paced gait speed (0.05 m/s, p=0.009), and lower-limb strength (10.8 kg, p=0.03) after 6 months (3) ↔ Interaction effects between the exercise and milk supplement |

Notes: ↑, increase or improvement; ↔, no change or association.
Abbreviations: BW, body weight; CSA, cross-sectional area; EAA, essential amino acids; GS, grip strength; HLC, high-level care; ICW, intracellular water; LLC, low-level care; MVC, maximal voluntary contraction; RCT, randomized controlled trial; RE, resistance exercise; SMI, skeletal muscle index (appendicular lean mass/height²); TUG, Timed Up-and-Go test, VO$_{2peak}$, peak oxygen uptake.

protein intake (>1.2 g/kg BW/d) at baseline. Whilst exercise significantly improved several muscle health outcomes—strength, lean mass (LM), muscle cross sectional area (CSA), and gait speed—compared with the no exercise group, no additional beneficial effects of milk beyond the effect of exercise were observed. Specifically, the gains in total body LM and mid-femur muscle CSA were two- to threefold greater in the exercise + fortified milk group compared with either group alone or the control group, but the interaction terms were not significant for any muscle or functional outcome after 18 months. The main effect analyses revealed that exercise significantly improved muscle strength (~20–52%, $p<0.001$), LM (0.6 kg, $p<0.05$), muscle CSA (1.8%, $p<0.001$), and gait speed (11%, $p<0.05$) compared with the no exercise group. Moreover, the fortified milk had no effect on muscle size, strength, or function.

Similar findings were noted in a study involving older adults living in a residential care facility with poor nutritional status and limitations in activities of daily living. Improvements in balance, gait speed and lower-limb strength were driven by a three-month high-intensity functional exercise program with no additional benefits from combining exercise with a milk-based protein supplement.[69] Specifically, there was a significant improvement in gait speed in the exercise compared with the control group (mean difference 0.04 m/s, $p=0.02$) at three months, and significant improvements in balance (Berg Balance Scale, 1.9 points, $p=0.05$), gait speed (0.05 m/s, $p=0.009$), and lower-limb strength (10.8 kg, $p=0.03$) after six months follow-up. However, no interaction effects were observed between the exercise and nutrition interventions (ie, milk-based protein-enriched energy supplement). This could be partially explained by the additional protein being oxidized to generate energy to compensate for a negative energy balance in malnourished older women.[69] In another study involving residential care residents with severe physical and cognitive impairment, no change in intracellular water (ICW), a proxy for muscle mass, was observed following a three-month intervention involving an exercise program and milk-based protein enriched drink providing ~15 g of protein after exercise.[64] For example, the between-group difference in ICW in exercise vs control activity group were not significant (mean 95% CI: −0.2 (−0.7 to 0.3), $p=0.37$), and no differences were observed between the protein drink vs placebo drink group (−0.2 (−0.7 to 0.3), $p=0.53$). In both studies all participants were assessed to be at risk of malnutrition (the Mini Nutritional Assessment score <24), which may have explained negative findings for muscle anabolism. Osuka et al[68] compared the

effect of a combination of aerobic and resistance training (ART) followed by consumption of fortified milk (10.5 g of protein) against RT with fortified milk on muscle mass, strength and function in healthy community-dwelling older adults after 12 weeks of intervention. No between-group differences in muscle mass were reported, but SMI, whole-body muscle mass, upper-extremity muscle mass were increased in the RT with the fortified milk group, whereas lower-extremity muscle mass was increased in both groups. Muscle strength (leg extension strength, leg curl, leg press, chest press, arm curl) and the time to complete chair stands also improved in both groups, with the ART + fortified milk group improving significantly more compared with the RT + fortified milk group (9.0±5.5 vs 5.3±3.8, $p=0.005$ arm curls per 30 s, and 5.9±3.9 vs 3.2±4.1, =0.01 chair rises per 30 s). In both intervention groups protein intake was >1.3 g/kg BW/d at baseline. Lack of a comparator group limits the interpretation of the study findings, to address whether fortified milk provided any benefits on muscle health outcomes.

Two studies[66,67] have evaluated the use of a chocolate milk-based drink, with Maltais et al[66] comparing EAA supplement (12 g of protein, 7 g of EAA from soy) against milk supplement (13.5 g of protein, 7 g of EAA) and nonprotein control (rice milk) in older men with low muscle mass index (MMI, muscle mass/high). All groups completed a 4-month resistance training program three times a week. All participants improved significantly in several parameters of muscle mass (ie, lean body mass, MMI and total muscle mass) and muscle strength (lateral pull down 1-RM), but no between-group differences were observed. For example, all groups experienced significant change in MMI (control group: 0.52 (0.32) kg/m^2; EAA group: 0.95 (0.55) kg/m^2, and milk group: 0.65 (0.47) kg/m^2, p for all ≤0.05). For physical capacity, only the EAA group improved in the TUG test. The study had only 8–10 participants per group and all had protein intake >0.8 g/kg BW/d. The authors concluded that RE was an effective way of improving muscle mass and strength regardless of protein supplementation. Conversely, a study that compared the effect of 500 mL non-supplemented chocolate milk (14 g of protein/day) with RE to placebo drink (0.4 g of protein) with RE in healthy young and old men after 12 weeks of intervention, found that milk did not enhance effects on skeletal muscle strength or hypertrophy following resistance training. Although a strong training effect was observed for all muscle strength measures (eg, 1-RM for leg press, leg extension, and chest press in kg; $p<0.005$), there was no significant interactive effect of

chocolate milk supplementation (p >0.3).[67] Protein intake pre-intervention was not reported.

In summary, several intervention studies have investigated how milk as a vehicle for protein and other nutrients may enhance the beneficial effect of resistance training on muscle mass, strength and function in older adults with and without functional impairments. Collectively, current evidence does not indicate any additional beneficial effects of milk supplementation beyond the positive effects of RE alone.

Discussion

Milk for Muscle Health: Summary of Evidence and Implications

Milk is a complex food constituted of nutrients and biologically active components with anabolic, anti-oxidative, anti-inflammatory, and immunomodulatory properties, which as a part of a balanced diet may provide myoprotection for aging muscle. However, there is limited evidence and current epidemiological and experimental research does not provide support for additional benefits of (unfortified or fortified) milk for muscle health and physical performance in older adults. Evidence from one observational study suggests that higher childhood intake of whole milk may have a beneficial effect on physical performance and balance in later life.[60] The associations between higher adult intake of milk and muscle health were inconclusive and based on only two studies, warranting further research.[60,61] Positive results observed for childhood milk intake and muscle function in later life need to be repeated in other populations.

In intervention studies (Table 2), (fortified) milk, alone or in the combination with resistance training, provided no benefits for muscle health. Specifically, two interventions with fortified milk showed no evidence for an independent effect of milk consumption on muscle strength and physical performance either in care residents[62] or community-dwelling older adults.[63] Although very limited, the results suggest that milk supplementation providing extra energy and protein above habitual consumption may not be effective or sufficient to improve muscle function in older adults with either higher dietary protein intake (ie, >0.8 g of protein/kg BW/d in both studies) or functional limitations. Of six intervention studies that combined exercise with (fortified) milk supplementation, five found the main effect of exercise on several parameters of muscle mass and function,[65–69] but no interaction effect of the exercise

and nutrition intervention in ether healthy community-dwelling older adults or in those residing in care homes. One study in care residents with functional impairments found no exercise or interaction effect on muscle mass,[64] which could be explained by malnutrition and negative protein energy for muscle anabolism, multimorbidity, and inadequate intervention duration for this population to observe significant change.

Promising evidence for an interaction effect of milk and exercise has been observed in several intervention studies with younger adults and athletes, both men and women.[19,43-45] A positive effect of (unfortified) milk immediately after exercise has been reported for lean muscle mass, strength, MPS, and loss of fat mass compared with control (isocaloric) or soy-based drink,[19,43-45] which the authors have contributed to the following mechanisms. Compared to other protein sources, milk proteins (especially whey) have greater ability to promote anabolism postexercise, resulting in lean muscle gain.[19,45,48,59] Decreases in fat mass have been explained by the interplay between parathyroid hormone, vitamin D metabolites, medium-chain fatty acids, bioactive peptides and serum calcium affecting cellular lipolysis (adipocytes lipid metabolism) and fatty acids absorption, thus resulting in loss of fat mass (discussed in Dougkas et al).[70]

However, the results from interventions studies combining fortified milk with exercise in older adults were inconclusive, and several limitations of the studies have been recognized by the authors. These include: (1) sample size and lack of power to detect change in muscle health outcomes affecting the validity of the study in the population of interest;[63–67] (2) lack of control group for proper comparison;[62,65] (3) insufficient study duration to observe an effect in the population under study;[62–64] (4) low milk (protein) dose to induce muscle fiber hypertrophy in older adults experiencing anabolic resistance;[66,67,69] (5) attrition and low compliance with the intervention;[63] (6) a small treatment effect, suggesting low clinical importance of (fortified) milk in the study population;[63,69] (7) selection bias toward healthy older adults;[65,68] (8) low generalizability to older adult population;[65,68] (8) no blinding of participants and assessors to the intervention;[68,69] (9) lack of mechanistic studies;[62,63,68] and (10) the timing of the nutrition intervention missing the "window of anabolic opportunity" (ie, the timing of the protein feeding exceeding 24 hours).[65]

Regardless of the differences across the studies (ie, participants, sample size, intervention type, duration, intensity, and muscle health outcomes), the results confirmed the

effectiveness of resistance training for better muscle aging in healthy and older adults with sarcopenia.[71,72] Importantly, the lack of the interaction effect between exercise and nutrition (milk) in all studies calls for a careful consideration of several factors that may have contributed to the null findings when designing future studies using a whole food approach (milk). These factors include: (1) nutrient (protein) dose in milk; (2) timing of milk (protein) consumption; (3) habitual protein intake; (4) insufficient understanding of the role of other nutrients in milk for MPS; and (5) difficulties recruiting older adults at risk of sarcopenia, who would likely benefit the most from these interventions.[73,74]

Milk for Muscle Health: Possible Reasons for Lack of Beneficial Effects in Studies with Older Adults

Provision of sufficient protein (ie, dose) within milk represents a key consideration, particularly for older adults who may exhibit anabolic resistance compared with younger adults.[30] Previous work has suggested that >20 g of protein per meal may be necessary to stimulate MPS in older adults,[11–13,22,48] yet all RCTs combining milk supplementation and exercise[64–69] (Table 2) have provided amounts below 20 g of protein/day either after exercise or every day during the study period. The two studies with milk supplementation intervention without exercise[62,63] provided >33 g of protein/day, but the absence of RE, a potent stimulus of MPS, may explain the negative results. A recent meta-analysis has indicated that there is no evidence to suggest that protein or EAA supplementation without concomitant exercise interventions increases muscle mass or strength in predominantly healthy older adults.[75] However, a meta-analysis with meta-regression of 49 studies in healthy adults has shown that the effect of protein supplementation in augmenting RE-induced change in fat-free mass was effective in young adults (<40 years) and less effective with advancing age (>40 years), but ineffective beyond total protein intake of ~1.6 g/kg BW/day.[76] This suggests that the protein dosage and the total protein intake may be the possible reasons for the negative results in the studies.

One challenge in collating this evidence is that the frequency of milk dose varied between the studies; three RCTs have provided nutritional supplement daily[65,67,68] and three on the training days only,[63,66,69] which makes the comparison between the studies challenging. Adherence to (fortified) milk drinking was >80% across

the studies, suggesting that milk drinking was feasible for older adults. However, a greater amount of milk is needed to provide a dose >20 g of protein, which needs to be assessed in this population.[77]

An important consideration for maximizing potential adaptation is the timing of protein consumption. For example, a study involving older men has shown that ingestion of protein supplement (10 g protein) immediately postexercise (within five minutes) stimulated greater skeletal muscle hypertrophy compared with the ingestion two hours post-training.[78] However, a study comparing the effect of protein supplementation pre- and post-RE between young (≤40 years) and older men (≥59 years) has found no effect of supplementation on muscle mass and strength in older adults.[79] This suggests that the dosages of protein may be more important than timing. The timing of the milk intake in the RCTs with exercise component described in this review varied from five minutes after exercise,[64,69] immediately after exercise[66–68] to non-specific,[65] the latter study possibly missing the "MPS window" post-RE stimulus for beneficial effect.

A key issue is the role of other nutrients present in milk, especially fats, which are rarely considered; the compositional differences in milk across the studies for MPS stimulation postexercise in older adults is unknown. There is some evidence that fat content may be important, as in a study of healthy young volunteers, Elliott et al[59] demonstrated that milk ingestion-stimulated net MPS following RE was more effective in the whole milk than in fat-free milk group—a difference that was explained by higher net amino acid uptake, hypothesized to be related to a higher fat content in whole milk. However, compositional difference in milk across the reviewed studies (eg, supplemented with protein,[63,64,66,68,69] vitamin D and Ca,[62,65] EAA,[66] chocolate;[66,67] reduced or low fat (≤1.6%) in seven out of eight studies) makes the synthesis of their findings difficult, and the implications for future trials of milk supplementation challenging.

Another important point to consider relates to participants' habitual dietary protein intake in these studies. In all the studies of older independent adults for whom there were baseline dietary data (Table 2), habitual protein intakes were above the recommended intakes (0.8 g/kg BW/d), reaching 1.2–1.3 g/kg BW/d in two studies.[65,68] Thus increased protein/milk consumption provided through the intervention was in populations that already consumed adequate amounts of protein for muscle health, and therefore unlikely to have a substantial additional

effect on muscle outcomes. For example, in a study of healthy older men and women with adequate habitual protein intake, protein supplementation of 15 g/day after a prolonged resistance-type exercise program did not augment beneficial effect of exercise on muscle mass, strength and function.[80] Similarly, in a recent RCT, the Liverpool Hope University—Sarcopenia Aging Trial, supplementation with a leucine-enriched whey protein isolate (1.5 g/kg BW/d) after resistance and functional exercise program did not provide any additional benefit in healthy older adults who already consumed sufficient amounts of dietary protein at study enrollment.[81] Interestingly, the authors suggest that future trials should use whole protein-containing foods instead of supplements because of low compliance (43±14%) to dietary-protein supplementation in this trial, which may have contributed to the null results.[81]

A big challenge to date for RCTs is the recruitment of older adults with sarcopenia, who may respond differently to the milk supplementation after RE treatment compared with those with lesser degrees of muscle dysfunction. In the present review only two trials included pre-sarcopenic older adults;[63,66] further research should determine whether those with different levels of skeletal muscle dysfunction respond differently to interventions.

Another challenge to consider in milk-based interventions with older adults relates to a substantial variation in milk consumption across the world regions,[33] explained in part by affordability and environmental concerns related to animal-based protein production and consumption,[22,23] and the presence of lactose intolerance (LI) and malabsorption in the populations.[82] Although much LI is genetically predetermined (ie, lactase nonpersistence, LNP), self-reported LI has been estimated at a greater prevalence then LNP in a number of studies, with links to female gender, advanced age, race, body size, dose of lactose, and genetic differences in LNP.[83,84] Also, the malabsorption of nutrients and gastrointestinal symptoms should be considered when designing nutritional interventions with whole foods, including milk and dairy in older adults. Nutrient absorption in older adults can be compromised by nutrient-drug interaction, atrophic gastritis leading to hypochlorhydria and altered acid-pepsin digestion, resulting in impaired absorption of vitamins and minerals, such as folate, vitamin B12, calcium, iron and β-carotene (discussed in Granic et al).[85] In a study of 400 Finnish adults (aged 18–64 years), only milk protein IgG, not IgA antibodies were associated with self-perceived gastrointestinal symptoms, which have been suspected to be caused by milk indigestion in over 40% of adults in primary care.[86]

In summary, synthesizing findings from existing intervention studies (Table 2) presents a challenge because of differences in study design (eg, sample size, study arms), participants (eg, baseline fitness, habitual protein intake, sarcopenia status, setting) and intervention characteristics (eg, exercise training program, protein supplementation protocol, adherence/compliance to intervention). Current evidence does not show benefit of milk supplementation of older adults for muscle health. However, inadequate dose of protein (via milk), high habitual protein intakes, fitness and sarcopenia status at baseline may be among the main reasons for the lack of demonstrated effects of milk.

Milk for Muscle Health: Areas for Future Aging Research

There is relatively little evidence about the role of milk in muscle health and functioning and sarcopenia in older adults from both observational and intervention studies. A whole food approach in testing the nutrition–muscle health hypothesis emphasizes nutrient-dense foods that are affordable, sustainable, easy to prepare, and palatable to older adults. Liquid milk may be such a food because it contains nutrients and other bioactive components that have multiple myoprotective properties, which may act against several pathogenic pathways implicated in sarcopenia, including inflammation and oxidative stress. However, there are several obstacles facing observational and intervention research of muscle aging in reaching a higher level of evidence about the role of whole foods (milk) in treatment and prevention of sarcopenia. These include: (1) the type and amount of milk (eg, low-fat vs whole; readily available vs fortified); (2) the type, frequency and intensity of exercise intervention (resistance training vs resistance and functional training); (3) the harmonization of the operational definition of sarcopenia[1,3] and muscle-related outcomes; (4) determination of target population (eg, sarcopenic vs at-risk population living in the community and residential care); (5) the timing of follow-up or intervention (weeks vs months); and (6) life stages (eg, a life course approach; mid vs late adulthood). Adequately powered RCTs of well characterized older adults likely to benefit from the interventions using a whole food approach are needed to test their effectiveness for healthy muscle ageing. In addition, milk may not be an appropriate functional food for mitigating loss of muscle mass and function for all older adults. For example, although higher milk and dairy intake has been associated with other positive health outcomes,[34–43,47] lactose intolerance should be considered

when designing RCTs with older adults with multimorbidities (eg, diabetes, obesity) and milk allergies.[82–84]

Conclusions

Milk contains biologically active nutrients and components that have anabolic, anti-inflammatory, anti-oxidative, and immunomodulating properties, and thus may be myoprotective. However, there is currently relatively little evidence from both observational and intervention studies with older adults about the benefits of milk as a functional food for muscle health. A limited number of studies summarized in this review included older adults that were mostly healthy, well-functioning, and well-nourished (ie, good protein intake), and varied greatly in the type, amount and timing of milk intake, length of intervention, and sarcopenia status. Sufficiently powered intervention studies in well characterized groups of older adults most likely to benefit from interventions are needed to test the effectiveness of a whole food approach, including milk, for healthy muscle aging.

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