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The Potential Nutrition-, Physical- and Health-Related Benefits of Cow's Milk for Primary School Aged Children

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28 Abstract

Cow's milk is a naturally nutrient-dense foodstuff. A significant source of many essential nutrients, its inclusion as a component of a healthy balanced diet has been long recommended. Beyond milk's nutritional value, an increasing body of evidence illustrates cow's milk may confer numerous benefits related to health. Evidence from adult populations suggests that cow's milk may have a role in overall dietary quality, appetite control, hydration and cognitive function. Although evidence is limited compared to the adult literature, these benefits may be echoed in recent paediatric studies. This article, therefore, reviews the scientific literature to provide an evidence-based evaluation of the associated health benefits of cow's milk consumption in primary-school aged children (4-11 years). We focus on seven key areas related to nutrition and health comprising nutritional status, hydration, dental and bone health, physical stature, cognitive function, and appetite control. The evidence consistently demonstrates cow's milk (plain and flavoured) improves nutritional status in primary-school aged children. With some confidence, cow's milk also appears beneficial for hydration, dental and bone health and beneficial to neutral concerning physical stature and appetite. Due to conflicting studies, reaching a conclusion has proven difficult concerning cow's milk and cognitive function therefore a level of caution should be exercised when interpreting these results. All areas, however, would benefit from further robust investigation, especially in free-living school settings, to verify conclusions. Nonetheless, when the nutritional-, physical- and health-related impact of cow's milk avoidance is considered, the evidence highlights the importance of increasing cow's milk consumption.

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62 **Introduction**

Commonly cited as nature's perfect food⁽¹⁾, milk is the foundation of life for all mammalian 63 neonates^(2, 3). Representing a complex and unique liquid, milk (which is produced by the mammary 64 glands⁽⁴⁾) is the only foodstuff designed by nature to serve as a complete food, at least during the 65 pre-weaning period. In this sense, milk contains numerous biological constituents and an assortment 66 of nutrients necessary for immunological protection and initial growth^(2, 5). Shortly after weaning, 67 most mammals stop consuming milk⁽⁶⁾. With domestication and milking of various animals, 68 however, some humans continue to drink milk throughout life⁽⁷⁾, mostly cow's milk. The value of 69 cow's milk in the human diet has been heavily debated for many years⁽⁸⁾ and has been portrayed by 70 some as 'white poison', a health hazard and promoter of Western chronic diseases⁽⁹⁾. This position 71 72 is based on numerous hypotheses including that cow's milk contains blood, pus, hormones and antibiotics, causes acne and cancer and has a high cholesterol and fat/saturated fat content 73 74 collectively contributing to cardiovascular disease risk, weight gain and obesity. Nevertheless, such perceptions of cow's milk being harmful to health are not supported by evidence. Conversely, cow's 75 milk has often been signified as the white $elixir^{(6)}$. 76

Childhood is a key stage for the development of healthy eating patterns⁽¹⁰⁾, and the school 77 environment provides a valuable setting to develop such behaviours. This is particularly true 78 79 considering children spend much of their time at school. Indeed, dietary habits shaped throughout the childhood years might carry forth and track into adulthood. Between the ages of 4-11 years 80 (primary-school age), children grow and mature at a rapid rate preceding the onset of puberty⁽¹¹⁾. 81 Childhood is therefore a critical transition period preceding adolescence, characterised by growing 82 83 independence and marked physical development. Good nutrition for the childhood period is therefore particularly important⁽¹²⁾. Not only may good nutrition support proper growth and 84 85 maturation, but it may therefore act as a base for immediate and lasting health, well-being and disease prevention. This is best achieved by consuming a balanced and varied diet that provides all 86 87 the nutrients needed. The inclusion of cow's milk as a staple component of a healthy balanced diet has been long recognised and is central in most public dietary recommendations⁽¹³⁾. Milk is naturally 88 nutrient-dense and is a significant source of many essential macro and micronutrients. Indeed, cow's 89 milk may confer nutrition-, physical- and health-related benefits beyond that of helping children to 90 simply meet nutrient targets. Based on the available literature, it appears that many of these effects 91 are a product of milk's nutritional composition. It therefore seems relevant to provide readers with 92 a brief description of the nutritional composition of cow's milk. 93

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96 Nutritional Composition of Cow's Milk and Patterns of Consumption

The nutritional composition of cow's milk is influenced by factors including genes (species and 97 breed), physical state (age and stage of lactation) and environment (available nutrition and climatic 98 conditions)⁽¹⁴⁾. The composition of whole cow's milk is approximately 87 % water and 13 % 99 solids⁽¹⁾. The percentage split of water and solids of cow's milk, as well as energy, is primarily 100 determined by the amount of fat⁽¹⁵⁾ but is also influenced by added sugars and sweeteners. Cow's 101 milk is approximately 4.9 % carbohydrate. The primary carbohydrate portion of milk is lactose, a 102 disaccharide comprising glucose and galactose. The lipid component of cow's milk and other milk-103 104 based dairy foods contribute numerous properties including the provision of fat-soluble micronutrients and essential fatty acids, as well as influencing flavour, texture and appearance⁽¹⁵⁾. Of the 105 lipid content within cow's milk, roughly 64 % are saturated fatty acids, with a considerable amount 106 (~26 %) from monounsaturated fatty acids and a small contribution from trans and polyunsaturated 107 fatty acids (both $\sim 3 \%$)⁽¹⁶⁾. Additionally, cow's milk provides high-quality proteins, namely casein 108 and whey. Casein and whey constitute approximately 82 % and 18 % of the total protein found in 109 cow's milk and provide an abundance of essential amino acids⁽¹⁷⁾. Aside from the macronutrient 110 111 content, cow's milk contains essential micronutrients that contribute to dietary quality and overall nutritional status. As shown in Table 1, with the exception of vitamin C (which is broken down 112 113 during pasteurisation) and vitamin D (unless fortified), cow's milk is a good source of all vitamins⁽¹⁸⁾. Calcium and phosphorus, crucial to healthy growth and maturation, as well as other 114 biological processes, are the most prominent minerals present in cow's milk. Cow's milk also makes 115 significant contributions to intakes of other major minerals⁽¹⁹⁾. In this sense, cow's milk makes a 116 substantial contribution to, and is the main dietary source of, calcium (26 %), iodine (37 %), 117 riboflavin (25 %), magnesium (10 %) and potassium (14 %) in the diets of primary-school aged 118 children⁽²⁰⁾. While volume specific data are limited, in the UK, the National Diet and Nutrition 119 Survey (NDNS) provides a nationally representative assessment concerning dietary habits of 120 individuals, aged 1.5 v and older, living in private households and remains the only surveillance 121 programme to do so. Temporal and age-related trends of cow's milk to average daily total energy, 122 macro and micronutrient intake from 2009/10 through to 2015-16 are presented in Table 2 for 123 children (4-10 years) and adolescents (11-18 years). Despite the clear value of cow's milk in the 124 everyday diet it is clear intakes steadily decline as children age. This trend is not only restricted to 125 the UK, it is also true in the US^(21, 22), Australia⁽²³⁾ other European countries⁽²⁴⁾. This is of great 126 concern among children, especially as dietary habits may track into adulthood, as milk avoidance 127 may have detrimental implications over time, leaving populations vulnerable to micronutrient 128 deficiencies (e.g. calcium, iodine and riboflavin deficiencies to name a few⁽²⁵⁾) and lasting nutrition-129

and health-related complications (e.g. cardiovascular disease⁽²⁶⁾, metabolic syndrome⁽²⁷⁾,
 hypertension^(27, 28), poor weight management and bone health^(25, 29)).

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133 Scope and Methodology of the Narrative Review

Evidence from adult populations suggests that adequate cow's milk consumption is associated with 134 a reduced risk of cardiovascular disease⁽³⁰⁾, metabolic syndrome^(28, 31) and obesity⁽³²⁻³⁴⁾. Emerging 135 data also suggests that cow's milk may have a role in overall dietary quality, appetite control, 136 hydration and cognitive function. Although the evidence to date is limited compared to the adult 137 138 literature, these benefits appear echoed in recent paediatric studies. There is a need to review the literature to assess whether there is sufficient evidence of a beneficial or detrimental effect of cow's 139 milk in the diet and health of children. The aim of this narrative review is therefore to summarise 140 and appraise the scientific literature to form an accurate, evidence-based evaluation of the associated 141 nutrition-, physical- and health-related benefits of cow's milk consumption in primary-school aged 142 143 children (4-11 years). Based on recent publications, it is not the intention of this narrative review to focus on body weight and body composition as with some confidence it is known cow's milk 144 145 consumption is inversely (or not) associated with body weight and body composition in children and adolescents⁽³⁵⁾. Where possible, this review focuses solely on literature in children aged 4-11 years, 146 147 however, where there is no applicable literature, data from general child studies will be reviewed (and possibly some adult studies). This is particularly true for mechanistic considerations as much 148 of this evidence comes from adult studies, therefore any adult-specific data should be interpreted 149 with some caution as the observations cited may not always be replicated in children. The narrative 150 review will also highlight current knowledge gaps in this field and suggest directions for future 151 research. 152

To identify research articles to facilitate this narrative review, PubMed (US National Library 153 of Medicine National Institutes of Health) and Web of Science (Thomson Reuters, UK), were 154 searched using various combinations of keywords relevant to the scope of the review up to August 155 2020. Search terms included: humans, child, milk, flavoured milk, animal milk, cow milk, infant, 156 adolescent, preschool, primary-school, paediatric, diet, energy intake, nutritional status, diet quality, 157 obesity, school-age children, food habits, dairy products, milk beverages, height, stature, 158 159 anthropometry, calcium, health association, health benefits, dental health, caries, decay, bone health, appetite, cognition, memory, mental performance, school milk, insulin, glucose, hydration, snack. 160 Studies were considered eligible for inclusion in the review if (i) they were conducted in children 4-161 11 years; (ii) participants received cow's milk, and not supplemental forms of dairy minerals (unless 162 a milk group was included); (iii) the study had been peer-reviewed. Results from studies that grouped 163

data over a wider age-range were also considered where this was deemed appropriate. This was especially true for prospective and intervention studies, yet the mean age of participants had to fall between the stipulated age range. Studies were excluded if (i) human participants were not used; (ii) participants lay directly outside of the stipulated age-range; (iii) studies used supplements only (iv) studies used different dairy foods (i.e. yogurt, cheese, dairy desserts) (v) the milk consumed included different sources of animal milk; (vi) data reporting was poor and/or not published in English.

In this narrative review we have summarised the evidence base, which we identified by related 170 benefits; nutrition-, physical- and health-related and provide a summary for the studies identified. 171 172 As this is not a systematic review, we did not perform a quality assessment of each study (based on design and implementation) or conduct a meta-analysis by study design but summarise the findings 173 for each study and sub-section in Tables 3 (nutrition-related benefits), 4 (physical-related benefits) 174 and 5 (health-related benefits). Symbols \uparrow and \downarrow indicate a statistically significant positive and 175 negative effect/relationship, respectively, between cow's milk and related subsections, with \leftrightarrow 176 177 indicating no statistically significant effect/relationship or a neutral effect. This approach has successfully been used in a recent narrative review in this journal⁽³⁵⁾, and therefore direct readers to 178 179 this paper for justifications and interpretations for this approach. In brief, however, a tabulated summary allowed us to draw overall conclusions for this narrative review, although we did not 180 181 undertake a detailed assessment of quality of individual studies.

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183 Nutrition-Related Benefits of Cow's Milk

184 Nutritional Status

The importance of regular cow's milk consumption in children has been recognized for over 40 185 years⁽³⁶⁾. As described, cow's milk is a naturally nutrient-dense foodstuff, providing a rich source of 186 many essential nutrients. While the potential of cow's milk to improve nutritional status may be 187 unsurprising, the effect of this in primary-school aged children has only been investigated by 188 numerous observational studies⁽³⁷⁻⁴²⁾ and three intervention-based study⁽⁴³⁻⁴⁵⁾ (see Table 3). All 189 reported improved nutritional status with increased cow's milk consumption, yet, due to the study 190 designs utilised, findings must be interpreted with caution. In the few intervention-based studies⁽⁴³⁻ 191 ⁴⁵⁾, which ranged from 16 weeks to 6 months, primary school children were provided with flavoured 192 milk or no drink (control)⁽⁴⁴⁾, 200 mL⁽⁴³⁾ and 250 mL⁽⁴⁵⁾ of multi-micronutrient fortified milk or 193 unfortified milk to establish whether unfortified and fortified cow's milk influences micronutrient 194 status in primary-school aged children. In those consuming fortified versus unfortified cow's milk 195 significant increases in blood riboflavin and vitamin B₁₂ were observed, with all other analytes 196 (selenium, ferritin, and vitamin D) comparable between drinks⁽⁴³⁾. Additionally, significant 197

increases in serum ferritin and zinc increased have been observed after 6 months with both fortified 198 versus unfortified cow's milk compared to a control drink^{(45).} With respect to nutritional status, 199 dietary patterns characterised by high cow's milk consumption (both plain and flavoured milk) 200 resulted in greater intakes of energy, protein, phosphorus, magnesium, calcium, potassium, vitamin 201 A, zinc, riboflavin (vitamin B₂), and niacin compared to children who seldom consumed cow's milk. 202 Notably, diets high in cow's milk may also limit intakes of foods and beverages high in fat and 203 sugar. In studies of Dutch⁽³⁷⁾ and American primary-school aged children⁽⁴⁶⁾, for example, cow's 204 milk intake was inversely related with the intake of sugar-sweetened beverages. In these studies, 205 206 children with low cow's milk intakes had lower protein, fibre, calcium, magnesium, potassium and 207 phosphorus intakes. Low cow's milk consumption has significant implications for intakes of several key nutrients that are of importance in childhood. Based on the nutritional contribution to dietary 208 intakes, it is important to note that children who drink cow's milk regularly are more likely to meet 209 dietary recommendations for many nutrients, and thus have a better nutritional status⁽¹³⁾. It could 210 therefore be argued that milk intake might be a marker for healthier eating habits(37). 211

With the above in mind, in the UK, it should be noted that dietary intakes of vitamin D in 212 213 primary-school aged children are low. Considering UK cows' milk is generally not a good source of vitamin D because it is not fortified, as it is in other countries, this narrative review may provide 214 215 justification for UK policy makers to reconsider widespread fortification. There is evidence (albeit from a theoretical modelling perspective) from Northern Ireland that cows' milk can be used as a 216 successful vehicle for vitamin D fortification⁽⁴⁷⁾. Fortification of cows' milks with 1 µg, 1.5 µg and 217 2.0 µg/100g, theoretically increased median vitamin D intakes from 2.0 µg/day to 4.2 µg, 5.1 µg and 218 5.9 µg/day, respectively and may therefore provide strong evidence for the efficacy of widespread 219 fortification. This modelling appears to translate to human studies. In support of this, a recent review 220 comprising of 20 studies showed positive associations between the consumption of vitamin D 221 fortified milk and 25(OH)D status in different population groups⁽⁴⁸⁾. Furthermore, in Finland, 222 Canada, United States who exercise a national vitamin D fortification policy (covering various fluid 223 milk products), milk products contributed 28-63 % to vitamin D intake, while in countries without 224 a fortification policy, or when the fortification covered only some dairy products (Sweden, Norway), 225 the contribution was much lower or negligible⁽⁴⁸⁾. Based on the above, widespread fortification of 226 milk seems helpful to help bolster vitamin D intakes and could be adopted by the UK to increase 227 vitamin D intakes in primary-school children. 228

Concerns about added sugars in foods and beverages has increased recently. Accordingly, many schools have limited access to foods and beverages high in added sugars, included flavoured cow's milk. In some studies, removal of flavoured cow's milk from the school environment reduces

energy and sugar intake, but negatively impacts essential nutrient intake and even reduced the 232 overall intake of non-flavoured cow's milk^(49, 50). In this sense, total milk consumption (both plain 233 and flavoured) decreased by 12.3 %⁽⁵⁰⁾. Contrariwise, children who drink more flavoured cow's milk 234 generally have higher non-flavoured milk intakes and display nutritional intakes similar to children 235 who only consume non-flavoured milk⁽⁵¹⁾. While greater cow's milk consumption (both plain and 236 flavoured) is generally associated with higher daily energy intake with some confidence we know 237 this does not impact body mass or composition⁽³⁵⁾. If daily energy intake is of concern for children 238 predisposed to overweight and obesity, results of a recent study suggest that replacement of whole 239 cow's milk or semi-skimmed cow's milk with skimmed cow's milk may help reduce total energy 240 intake, without impacting on nutrient provision⁽⁵²⁾. 241

When taken together, these results suggest that cow's milk consumption (both plain and 242 flavoured) might serve as a useful strategy to boost nutritional status in primary-school age children 243 and may act as a surrogate marker of diet quality. There may be a need for widespread vitamin D 244 fortification of cow's milk, however, and based on the above seems justified and should be reviewed 245 by UK policy makers. Cow's milk is a readily available, accessible, and affordable means of 246 providing valuable essential nutrients to the diets of primary-school children. The nutritional 247 implications of cow's milk provision and/or removal in the school environment must be considered 248 249 and is especially relevant in children suffering with nutritional inadequacies. The data are, however, primarily limited to observational investigations and require verification in more controlled 250 intervention-based studies covering differing populations. 251

252

253 *Hydration*

Whole cow's milk is approximately 87 % water⁽¹⁾ and may therefore be a beneficial choice for 254 hydration. For children, maintenance of euhydration is important for good health⁽⁵³⁾ but may also 255 increase concentration and mental performance (cognitive function)⁽⁵⁴⁾ while helping reduce 256 instances of headaches, constipation and other disorders⁽⁵³⁾. This is important as children are at a 257 greater risk of dehydration compared to adults, having relatively greater fluid losses at rest and 258 during exercise and being less able to recognise thirst⁽⁵⁴⁾. Given that water is continually being lost 259 260 from the body water pool, it is important to constantly replenish fluid losses to prevent dehydration. Establishing beverages that encourage longer-term fluid retention and maintenance of euhydration 261 has real clinical and practical implications⁽⁵⁵⁾. This is particularly true in situations where free access 262 to fluids is limited or when frequent breaks for urination are not desirable⁽⁵⁵⁾, such as in in the school 263 setting where many children arrive at school in an already hypo-hydrated state⁽⁵⁶⁾. There is little 264 research concerning the impact of cow's milk on hydration in primary-school aged children (see 265

Table 3). To date, three studies have been conducted⁽⁵⁷⁻⁵⁹⁾. Two of these were exercise-based intervention studies conducted by the same research group in Canada^(58, 59), and the remaining study was cross-sectional⁽⁵⁷⁾. Nonetheless, all reported improved hydration status with cow's milk in comparison to water or alternative beverages. While these findings show promise, additional controlled-intervention studies should be performed in different settings to examine repeatability of these effects.

272 In the two exercise-based intervention studies, researchers examined the influence of postexercise cow's milk consumption on rehydration in 7-11-year-old⁽⁵⁹⁾ and 10-12-year-old children⁽⁵⁸⁾. 273 The rehydration potential of cow's milk (or milk-protein) was compared to water or a carbohydrate-274 electrolyte drink, zero or a low cow's milk protein beverage and were given at 100 % and 150 % of 275 the children's body mass losses following exercise, respectively. In both studies, children exercised 276 in the heat (~34.5 °C) and consumed the test beverages immediately following exercise. The children 277 were subsequently observed for a period of 2 hours⁽⁵⁹⁾ and 4 hours⁽⁵⁸⁾, respectively, during which 278 measures of hydration status were collected (urine output and fluid balance). Findings over 2 hours 279 showed cow's milk was more effective than both water and the carbohydrate-electrolyte drink at 280 replacing fluid loss during exercise⁽⁵⁹⁾. This was similar over 4 hours. The authors suggested that the 281 protein in cow's milk may be a factor responsible for the rehydrating properties⁽⁵⁸⁾. 282

Montenegro-Bethancourt and colleagues⁽⁵⁷⁾ conducted a cross-sectional study designed initially to establish the contribution of fruit and vegetable intake on hydration status. Children (4-10-year-old) recorded all food and beverages consumed over 3 days using a weighed food diary alongside 24 hour urine samples. Regular intake of fruit and vegetables made a substantial contribution to hydration status, but notably, cow's milk consumption was also a strong dietary predictor of hydration status. In particular, cow's milk increased free water reserve by 25 mL in boys and 33 mL in girls per 100 g of intake. These findings corroborate studies in adult populations⁽⁶⁰⁻⁶²⁾.

Based solely from adult studies, there are a number of potential mechanisms explaining the 290 greater fluid retention (and thus hydration potential) with cow's milk⁽⁶³⁾. Firstly, milk contains 291 modest amounts of sodium (~20 mmol/L, similar to most commercial sports drinks) and large 292 amounts of potassium (~40 mmol/L). Sodium, as the main cation in the extracellular fluid plays a 293 major role in fluid retention⁽⁶⁴⁾, whilst potassium may exert some beneficial effects⁽⁶⁵⁾, although this 294 is not a consistent finding. Secondly, the protein content of cow's milk may help facilitate greater 295 fluid retention^(66, 67). While water and alternative beverages meet the basic intentions of rehydration, 296 they do not offer the abundance of nutrients present in cow's milk that will also aid in normal growth 297 and maturation. Nevertheless, it should be considered that while these findings suggest that cow's 298 milk helps improve the hydration status of children, there remains room for further studies to clarify 299

300 301 the role of cow's milk in hydration, especially in a free-living school setting. While the interventionbased studies show promise, many primary-school children do not exercise in such conditions.

302

303 **Physical-Related Benefits of Cow's Milk**

304 Dental Health

Milk contains multiple nutrients that may offer anticariogenic properties, protecting against the 305 development of dental caries, and thus supporting dental health in children^(68, 69). The nutrients 306 principally believed to play a role in dental health include calcium, phosphorus and protein⁽⁶⁸⁻⁷⁰⁾. As 307 reported earlier, cow's milk intake has been shown to be inversely related with the intake of sugar-308 sweetened beverages^(37, 46). Although speculative, one could argue increased cow's milk intake 309 might indirectly improve dental health. Calcium is required for bone and teeth formation, whereas 310 phosphorus ions works alongside calcium to maintain tooth strength⁽⁷⁰⁾. Cow's milk proteins 311 (particularly α_{s1} -, α_{s2} - and β -casein) may act to prevent tooth enamel erosion and demineralisation 312 of the tooth surface by producing casein phosphopeptides⁽⁷¹⁾. Based on the available evidence, 313 inverse associations between cow's milk intake and the incidence of tooth decay have frequently 314 been reported⁽⁷²⁻⁷⁵⁾ (see Table 4). The evidence, however, has been derived from cross-sectional 315 research, so causal conclusions cannot to be justified and caution must be exercised. 316

317 The findings of the cited studies were generated from recall methods which have obvious 318 shortcomings. Nevertheless, all studies (following visual examinations accompanied with food frequency questionnaires) reported inverse associations with cow's milk and the development of 319 dental caries. Interestingly, though sugar is suggested to possess acidogenic and cariogenic 320 potential⁽⁷⁶⁾, one study⁽⁷⁴⁾ reported the association between cow's milk consumption and protection 321 against of dental caries was stronger for children with diets high in sucrose. This might suggest that 322 cow's milk offers protection against the harmful effects of sugar, though this is speculative. To this 323 end, cow's milk intake during primary-school years has been reported as a predictor of incidences 324 of caries later in childhood⁽⁷²⁾. In this sense, greater cow's milk intake is inversely associated with 325 indices of dental caries. Collectively, these studies support the suggestion that dietary habits 326 established and maintained during primary-school years could have longer-term effects on health 327 328 outcomes.

Although no controlled trials have been conducted in primary-school children, the available literature appears to suggest that cow's milk could help reduce the incidence of dental caries and contribute to dental health in children. To reduce the occurrence of tooth decay, it is recommended primary-school children limit their consumption of sugary beverages (especially when not consumed with a meal) and increase consumption of cow's milk. The exact mechanism by which cow's milk

reduces the incidence of dental caries remains uncertain, though calcium, phosphate and casein 334 phosphopeptides may all play a role⁽⁶⁸⁻⁷⁰⁾. Casein phosphopeptides are phosphorylated casein-335 derived peptides produced by tryptic digestion of casein in the duodenum⁽⁷¹⁾. The anticariogenic 336 activity of casein phosphopeptides is due to their ability to stabilise high levels of amorphous 337 calcium phosphate on tooth surface, preventing demineralisation and enhancing remineralisation of 338 enamel caries⁽⁷⁷⁾. In addition, milk fat could be adsorbed onto the enamel surface and may have a 339 protective role; and thirdly, milk enzymes may have a role in reducing the growth of acidogenic 340 plaque bacteria⁽⁷⁸⁾. Where prior observational research provides a solid foundation, any future work 341 342 should seek to implement robust randomised clinical trials (RCT) to confirm any causal relationships between cow's milk consumption and dental health in primary-school aged children. 343

344

345 Bone Health and Physical Stature

Childhood is a critical time for bone growth and lasting bone health^(79, 80) and the nutritional 346 composition of cow's milk has evolved to stimulate and support this^(2, 5). The scientific opinion that 347 regular cow's milk consumption is associated with greater physical stature has a long history, dating 348 back to the 1920s^(81, 82). Cow's milk contains multiple nutrients that may support childhood 349 growth⁽⁸⁰⁾ and lasting bone health⁽⁷⁹⁾. The beneficial effects of cow's milk consumption on bone 350 351 health in children may include increased bone mineral content (BMC) and bone mineral density (BMD), characteristics necessary for the prevention of bone-related diseases later in life 352 (osteoporosis). Evidence of these benefits, however, is equivocal showing a beneficial to neutral 353 effect of cow's milk on these constructs^(25, 29, 83-88). In this sense, of the cited studies, two reported 354 greater total body BMC^(85, 88), while one reported no effect. One study reported no effect on total 355 body BMD⁽⁸⁷⁾, yet three studies reported increased regional BMC^(29, 83, 86) and another two studies 356 reported increased regional BMD^(25, 84) (see Table 4). The mixed findings reported throughout these 357 studies may be due to a lack of consistency in methodological approaches, length of study, location 358 and measures of bone health, all of which confound comparisons and prevent a clear conclusion. 359 Furthermore, age and sex differences must be considered, especially as puberty and bone 360 mineralisation typically occurs earlier in girls compared to boys⁽⁸⁹⁾. 361

- In two recent studies^(85, 88), school milk interventions increased total body and regional (forearm) BMC and BMD compared to children who seldom drank cow's milk. In these studies, the beneficial effects of cow's milk on bone health were observed in n = 435-757 children (mean age 11 years) following daily school milk intake for 1-2 years.
- 366 On a comparative basis with other animal sources, whole cow's milk is the richest source of 367 calcium and represents the biggest contributor of dietary calcium during childhood⁽⁹⁰⁾. In addition,

considering beef meat and eggs for example, cow's milk is the cheapest source of protein, calcium, 368 phosphorus, and vitamin D⁽⁹¹⁾. During childhood, the beneficial effects of cow's milk on bone health 369 in children are commonly attributed to calcium⁽⁹²⁾. However, many other nutrients including 370 phosphorus and protein are needed for normal growth and lasting bone health⁽⁹³⁾. In 5-11-year-old 371 children (n = 99), for example, calcium supplementation for 10 months did not influence total body 372 or regional BMC⁽⁸⁷⁾. In contrast, in an earlier study where cow's milk (and dairy) was supplemented 373 374 daily for 1 year (distributed to deliver 1200 mg calcium daily), lumbar (lower back) BMD increased compared to children who maintained their usual eating habits⁽⁸⁴⁾. This may illustrate that calcium 375 and other nutrients work together for bone growth, and thus lasting bone health⁽⁹³⁾. It is important to 376 note, however, that children in the calcium supplementation study were already consuming near 377 daily recommended amounts, which may illustrate intakes exceeding calcium recommendations 378 (from either supplemental calcium or cow's milk) offers no further benefit to bone health in children. 379

In several studies, it has been observed that children who avoid consuming cow's milk characteristically exhibit low calcium intakes, short statures, increased fatness and lower BMC (and thus exhibit reduced bone health) compared to their cow's milk-drinking counterparts^(25, 86). In children (mean age = 8 years) who previously avoided cow's milk, the introduction of cow's milk to the diet increased not only habitual cow's milk consumption but also increased total body BMC⁽²⁹⁾. This may suggest it is never too late for children to introduce cow's milk into their diet for bone health benefits.

From a stature perspective, available data appear to suggest that cow's milk consumption 387 almost certainly has a positive effect on growth in children. To date, fourteen studies in primary-388 school aged children have evaluated this aspect of cow's milk consumption^(25, 29, 36, 44, 81-83, 85, 94-99). 389 Seven were intervention-based, four were observational, and the remaining three were prospective 390 designs. Based on these studies, the evidence suggests that cow's milk intake positively influences 391 physical stature in primary-school aged children. All six intervention studies showed increased 392 stature with increased cow's milk (one study included milk calcium). In addition, all prospective 393 studies reported increased stature with increased cow's milk. With regards to the cross-sectional 394 studies, two illustrated cow's milk avoiders displayed stunted growth compared to cow's milk 395 drinkers while the remaining study showed increased adult stature with increased cow's milk 396 consumption in childhood. Although trends are consistent across studies, there remains a need for 397 evidence from robust controlled-intervention trials in primary-school aged children to verify 398 causality. 399

400 Beneficial effects on physical stature were observed with both whole and reduced fat cow's 401 milk, distributed at a range of 190 mL - 568 mL daily. In several of these studies, it was reported

that childhood cow's milk intake was associated with higher skeletal development (BMD of the hip 402 and the forearm), bone growth and periosteal bone expansion. These were likely established earlier 403 during growth periods and maintained into late adolescence and young adulthood⁽⁹⁶⁻⁹⁹⁾, supporting 404 the notion that dietary habits established and maintained during the primary- and secondary-school 405 years may not only induce short-term effects but offer lasting benefits. Indeed, in children with 406 407 prolonged cow's milk avoidance, stunted growth and physical stature is observed compared to children who habitually consume cow's milk, and this is maintained into adulthood^(25, 29, 86). During 408 a pubertal growth spurt, about 37 % of the entire skeletal mass is accumulated⁽¹⁰⁰⁾. Therefore, 409 410 inadequate calcium intake during this period may compromise volumetric bone density and overall stature attained. Notably, in a study that explored both cow's milk and supplemental calcium, those 411 children in the cow's milk (and dairy) group were 3 cm taller (166 cm) compared to the supplemental 412 calcium (163 cm) and placebo (163 cm) group⁽⁹⁶⁾. This may indicate that cow's milk has more of a 413 beneficial effect on physical stature and growth than single cow's milk constituents (i.e. calcium). 414

415 The precise mechanisms or nutrients in cow's milk responsible for stimulating growth and lasting bone health are not yet clear. Evidence suggests that maintaining adequate calcium intake 416 417 during childhood might be advantageous for the attainment of peak bone mass, which may be crucial in reducing the risk of bone-related diseases later in $life^{(101)}$. Interestingly, it appears that whole foods 418 419 may offer greater benefits than the equivalent amount of calcium in supplemental form. It has also 420 been suggested that the growth-stimulating effect of cow's milk is likely attributed to hormonal effects that can be influenced by ingested cow's milk proteins (predominantly whey protein and the 421 release of amino acids during digestion but also casein), micronutrients and also energy⁽¹⁰²⁻¹⁰⁴⁾. 422 423 Observations from child studies show these nutrients stimulate the secretion of insulin-like growth factor-1 and insulin, both of which are anabolic hormones that play an essential role in the regulation 424 of growth and accrual of bone mass during childhood⁽¹⁰²⁻¹⁰⁴⁾ though there is some controversy that 425 cow's milk intake upregulates insulin and insulin-like growth factor-1 signalling and thus promotes 426 chronic diseases such cancer (prostate, breast and colorectal) and cardiovascular disease⁽⁹⁾, though 427 these perceptions are hypothetical at present and not supported by evidence. 428

Nonetheless, when taken together, it appears that cow's milk consumption promotes both health and may increase physical stature in primary-school aged children. This suggests that cow's milk consumption during childhood might be important to ensure full growth potential is achieved. While there is some suggestion about the improved bone health with increased cow's milk consumption and the mechanisms responsible for the growth-stimulating properties of cow's milk, more intervention studies are needed to elucidate the components responsible for these effects and to prove and/or disprove the chronic diseases hypotheses. This is especially prudent when 436 considering bone health as the methodological approaches previously employed have been diverse437 in nature.

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439 Health-Related Benefits of Cow's Milk

440 *Appetite Regulation*

441 Appetite comprises numerous regulatory processes associated with the initiation and termination of eating and the selection and amount of food consumed. The regulation of appetite and eating 442 behaviour depend on the detection and integration of signals relaying nutritional status, and their 443 444 interaction with signals associated with food palatability and gastrointestinal handling, in addition to circadian, social, emotional, habitual and other situational influences⁽¹⁰⁵⁾. Consequently, appetite 445 and the regulation of eating behaviour are complex processes, regulated by homeostatic and non-446 homeostatic influences⁽¹⁰⁶⁾. Cow's milk contains a host of nutrients that might exert a favourable 447 effect on appetite and eating behaviour⁽¹⁰⁷⁾, yet in primary-school children there is limited evidence 448 concerning the short-term effect of cow's milk on these measures. There are also no data on the 449 moderate- and longer-term effects of daily cow's milk consumption on these outcomes in primary-450 451 school children.

From the available studies⁽¹⁰⁸⁻¹¹²⁾, cow's milk has principally been given as a mid-morning 452 453 snack or alongside breakfast, with the volume of cow's milk offered to children ranging from 160-250 mL. Based on these studies, the evidence concerning cow's milk and appetite regulation is 454 inconclusive. Three of these studies found a decrease in energy intake after cow's milk consumption 455 yet three reported no effect. Two studies reported cow's milk consumption reduced subjective 456 appetite, one reported increased subjective appetite compared to a fruit-based snack, while two 457 studies did not measure subjective appetite. Only one study measured hormonal indicators of 458 appetite and reported cow's milk consumption stimulated the secretion of glucagon-like peptide-1. 459

In two studies of 34 overweight and obese boys (mean age = 11 years)^(109, 110), when compared 460 with volume and energy matched servings of water or fruit-juice, 240 mL of low-fat cow's milk 461 with breakfast reduced energy intake at an ad-libitum lunchtime meal. In another study comprising 462 of 48 obese children (mean age = 11 years), girls reported higher satiety scores 4 hours after drinking 463 whole cow's milk compared to skimmed milk, and low-fat cow's milk significantly reduced appetite 464 compared to apple juice 2 hours after consumption. These differences, however, did not transpire to 465 changes in energy intake at an *ad-libitum* lunchtime meal across all conditions in girls, boys and the 466 group as a whole⁽¹⁰⁸⁾. As mentioned, only one investigation (comprising two experiments) has 467 sought to establish the effect of cow's milk (and other dairy products) on appetite and feeding 468 behaviour in normal weight and overweight/obese children (mean age = 11.5), where subjective 469

appetite and appetite-related analytes were measured⁽¹¹²⁾. In both experiments, preloads (experiment 470 1: 1 % fat (1 g/100g) chocolate cow's milk, 2 % (2 g/100g) fat cow's milk, 1.5 % (1.5 g/100g) fat 471 yogurt drink, fruit punch or a water drink; experiment 2: 2 % (2 g/100g) fat cow's milk or a fruit 472 punch) were provided 60 min preceding and during an *ad libitum* pizza meal. All preloads were 473 474 matched for volume (250 mL) and energy content (130 kcal, 543.9 kJ; except water in experiment 475 1). The first experiment comprised measures of subjective appetite, whereas the second experiment 476 included measures of subjective appetite together with appetite-related analytes (serum glucose, insulin and plasma GLP-1 and peptide YY). Reduced energy intake was observed following 477 478 chocolate cow's milk and yogurt consumption compared to a water drink in the first experiment. Consistent with a reduction in energy intake, subjective appetite (combined appetite score) was 479 significantly lower following 2 % (2 g/100g) fat cow's milk compared with the yogurt drink only. 480 No additional effects were observed concerning energy intake following the consumption of 2 % (2 481 g/100g) fat cow's milk and fruit punch or on subjective measures of appetite after 1 % fat chocolate 482 cow's milk, 1.5 % (1.5 g/100g) fat yogurt drink, fruit punch or water. Compared with the fruit punch 483 preload, cow's milk consumption resulted in a significantly greater GLP-1 area under the curve. 484 485 Nonetheless, ad libitum energy intake, insulin and glucose AUC were comparable between trials. Considering all aforementioned studies, it is important to consider that in these studies, energy intake 486 487 was principally assessed via ad libitum assessments which may not be reflective of free-living eating behaviour. 488

The mechanism(s) by which cow's milk consumption might influence eating behaviour are 489 not fully understood, but there are several plausible suggestions from the adult literature and 490 491 constituents of cow's milk that may act synergistically to explain possible actions. In the studies 492 where cow's milk reduced appetite and subsequent eating behaviour, it is probably unsurprising that cow's milk consumption suppressed energy intake at *ad-libitum* assessment meals, considering the 493 macronutrient composition of cow's milk compared to fruit-juice drinks and water. Cow's milk 494 contains considerably more protein than fruit-juice drinks and water. Although it is not a universal 495 finding, it is widely recognised that dietary proteins are more satiating than energetic equivalents of 496 carbohydrate and fat under most conditions, commonly suppressing eating behaviour at the next 497 meal⁽¹¹³⁻¹¹⁵⁾. Consequently, cow's milk proteins (whey and casein, and their products of digestion) 498 499 may act to potentiate peptides of gastrointestinal, pancreatic and adipose tissue origin, increasing perceptions of satiety and acutely reducing energy intake (anorexigenic behaviours)⁽¹¹³⁾. Moreover, 500 medium-chain triglycerides, conjugated linoleic acid (CLA) and lactose may also be implicated in 501 the reduction of energy intake after cow's milk intake⁽¹⁰⁷⁾. Medium-chain triglycerides are absorbed 502 directly into the portal circulation and transported to the liver for rapid oxidation. A combined action 503

of increased energy expenditure, decreased fat deposition and increased satiety may reduce of energy 504 intake via pre-absorptive signals, post-absorptive changes in metabolites and appetite-related 505 analytes⁽¹⁰⁷⁾ and similar appetite-related analyte responses have been observed with lactose⁽¹¹⁶⁾. 506 When considering CLA intake and its potential implications in appetite regulation, cow's milk (and 507 dairy) and cattle meat (cows and lamb) represents the almost exclusive dietary sources⁽¹¹⁷⁾ where 508 the predominant isomer of CLA (accounting for more than 90 % of the total CLA intake) is the cis-509 9, trans-11-CLA. It is, however, strongly proposed that other isomers, such as trans-10, cis-12-CLA 510 may influence body-weight and fat changes⁽¹¹⁸⁾. In agreeance with an earlier narrative review⁽³⁴⁾, it 511 remains unknown whether cow's milk (and dairy), when providing physiological doses of CLA, can 512 elicit any meaningful impact on appetite and body weight regulation in humans. This is especially 513 prudent when considering experimental design, age, sex, energy intake and CLA metabolism of the 514 participants, and the dose and chemical form of the CLA isomer administered as differences may 515 arise solely from these methodological differences⁽¹¹⁹⁾. From a child perspective, potential 516 underlying mechanisms of CLA on appetite regulation are poorly understood though evidence from 517 adult studies suggested that CLA can inhibit fatty acid synthase and stearoyl-CoA desaturase-1⁽¹¹⁸⁾, 518 519 enhance fat oxidation and thermogenesis and reduce lipogenesis and preadipocyte differentiation and proliferation⁽¹²⁰⁾.</sup> 520

521 In summary, evidence suggests that cow's milk may have a unique potential to influence elements of energy balance. Macro and micronutrients and other bioactive constituents might act 522 individually, though probably synergistically, to impart beneficial effects on energy balance and 523 body mass regulation through actions related to appetite, eating behaviour and metabolism. 524 However, there is little mechanistic exploration of cow's milk consumption and appetite regulation 525 from a paediatric perspective. Understanding the relationship between cow's milk consumption and 526 appetite regulation could provide novel nutritional interventions to contribute toward the fight 527 against childhood overweight and obesity⁽¹²¹⁾, whilst bolstering nutritional status and improving 528 elements of cognitive function and hydration. Controlled intervention studies are necessary to 529 determine the best possible timings to administer cow's milk and establish whether consumption 530 delivers these benefits when administered during the school day. 531

532 Collectively, the effects of cow's milk intake on appetite regulation in primary-school aged 533 children are unclear and could be dependent on BMI. The studies summarised suggest that mid-534 morning milk consumption influences eating behaviour at the next meal in overweight and obese 535 children, showing that cow's milk could be beneficial for reducing body mass. In lean children, the 536 evidence suggests there is no effect of cow's milk consumption on appetite and eating behaviour, 537 but cow's milk may boost the nutritional quality of the diet. There is much scope for further studies to clarify the role of cow's milk on appetite and eating behaviour, especially in a free-living school
environment where methodological approaches are more reflective of habitual eating behaviours. In
addition, it will be important to fully distinguish the effects of cow's milk on appetite- and on
metabolism-related peptides and subsequent eating behaviour.

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543 *Cognitive Function*

Compared to children with better dietary quality, those with nutritional inadequacies demonstrate 544 decreased attention and academic performance⁽¹²²⁾. Aside from improving nutritional status and 545 dietary quality, emerging evidence illustrates that cow's milk may positively influence cognitive 546 function in primary-school children^(43, 123, 124). Varying in duration from 2 hours to 9 months, studies 547 of an intervention-based nature generally report that cow's milk consumption increases elements of 548 cognitive function, though some of the specific outcome measures demonstrate no effect or in some 549 cases a negative effect of cow's milk. These studies highlight that consideration should be given to 550 potential moderators such as sex and to the use of cow's milk as a non-stigmatised method for 551 providing nutrients through fortification⁽¹²⁵⁾. One non-interventional study considered the adherence 552 of n = 1595 children to Canadian nutrition recommendations in Grade 5 (10-11 years) and in relation 553 to academic achievement in the provincial achievement tests taken approximately one year later⁽¹²⁶⁾. 554 555 A positive association was identified for boys, with those who met the nutrition recommendations for milk and alternatives (at the time of the study: 3-4 servings per day) scoring 3.45 % better on an 556 average measure of Math and Language Arts tests than those who did not meet the 557 recommendations⁽¹²⁶⁾. An account of the intervention studies follows, but overall, the varying 558 559 methodological approaches used suggests caution is needed in making firm conclusions about potential links between cow's milk consumption and improved cognitive function. 560

In a study⁽¹²⁴⁾ involving n = 469 boys and girls (mean age = 8 years), evaluating the effects of 561 daily mid-morning cow's milk consumption on physical, mental and school performance, 562 researchers found that a school feeding scheme focusing on daily cow's milk intake had beneficial 563 effects on school performance for girls. In this study, children received a serving of cow's milk (250 564 mL) daily for 12 weeks. Assessments of physical, mental and school performance were conducted 565 prior to and at the end of the 12-week supplementation period. Similarly, in a smaller study⁽¹²³⁾ 566 involving 40 children (mean age 11 years), the effects of a carbohydrate drink, a cow's milk drink 567 or a combination of both on subsequent cognitive function (processing speed, memory, attention 568 and perceptual speed) was assessed over a 3 hour period⁽¹²³⁾. Findings showed cow's milk 569 consumption improved short-term memory. Children were able to recall 0.7-0.8 more words 570 compared with 0.5 fewer words after the carbohydrate drink. However, this effect was only observed 571

in girls and not boys^(123, 124). There were slightly more mixed findings from a crossover study in 572 which 84 children (mean age 10 years) were given 237 mL of cow's milk or apple juice⁽¹²⁷⁾. While 573 the beverages were not standardised for temperature, participants were asked to complete 574 computerised tasks of inhibitory control, speeded working memory and sustained attention at 575 baseline and 30-, 90- and 120-min following beverage consumption. Although the significant results 576 following cow's milk compared to juice consumption were, again, only apparent in girls, there was 577 a negative effect (decreased working memory accuracy) alongside the positive one (improved 578 reaction time on the sustained attention task). There were non-significant trends in the opposite 579 580 direction for boys. No significant effects of the beverages were observed for on-task behaviour during the testing. 581

The mechanisms responsible for the beneficial effects of cow's milk on improved cognitive 582 function are unclear. One suggestion is a sustained blood glucose response following 583 consumption^(128, 129). Findings from Anderson et al.⁽¹²⁷⁾, an adult-based study, suggest that 584 glucoregulation may play a role as participants with higher fasting glucose levels demonstrated 585 faster reaction times on an inhibition task following cow's milk compared to juice. There were 586 587 however no sex differences in initial glucose elevation or change in glucose levels over time to explain the apparent sex differences in cognition. Such findings suggest a likelihood that factors 588 589 other than glucoregulation will explain why cow's milk differentially affects the cognition and behaviour of boys and girls. The micronutrient content of cow's milk is another potential mechanism 590 for the observed effects⁽⁴³⁾. Low intakes of vitamin B_1 , folate and vitamin B_{12} affect short-term 591 memory and impair learning, causing low cognitive scores and development in primary-school age 592 children⁽¹³⁰⁾. In addition, low iron and riboflavin intake may adversely affect motor skill 593 development and psychomotor performance⁽¹³¹⁾. All of these micronutrient are heavily present in 594 cow's milk. In one of the longer intervention-based studies available (5 months), Kuriyan and 595 colleagues⁽⁴³⁾ attempted to establish if fortification of cow's milk with multiple micronutrients 596 597 influenced the mental and physical performance of children (7-10 years) compared to an unfortified cow's milk drink. The children were randomised into groups and were provided with cow's milk (2 598 x 200 mL) 6 days a week for 5 months, with assessments of attention and executive function 599 conducted at baseline and 5 months. The findings showed improved cognitive and physical 600 performance in both groups, illustrating that further fortification of cow's milk provided no 601 additional benefits to cognitive and physical performance but did improve some elements of 602 nutritional status⁽³⁹⁾. Finally, in a double-blind RCT (9 months), the learning potential of 7-9-year-603 olds (31.9 % with a moderate or severe iodine deficiency at baseline) improved to a greater degree 604

following 200 mL daily of fortified cow's milk (fortified with 45 μ g iodine (given as potassium iodide) and other micronutrients) than following nonfortified milk (20.8 μ g iodine)⁽¹³²⁾.

Some of the components of cow's milk may be detrimental for the cognitive performance of 607 lactase deficient children. In a study involving children of 5-6 years (85 % lactase deficient), 608 information processing efficiency was assessed after 5 days' consumption of 150mL twice daily of 609 conventional cow's milk (containing A1 and A2 β-casein) or cow's milk containing A2 β-casein 610 only⁽¹³³⁾. While post-intervention response times were significantly improved from baseline for both 611 cow's milk products, error rates reduced in the A2 β-casein only condition. Furthermore, consuming 612 conventional cow's milk in the second phase of the crossover study appeared to undo the positive 613 614 effects on error rate obtained from the A2 β-casein only milk in the first phase, which were maintained through the 9 day washout period. 615

It is prudent to highlight that none of the studies of cognition in primary-school children have 616 compared cow's milk to a control beverage. It is therefore difficult to ascertain whether these studies 617 simply show less detrimental effects of cow's milk than comparator beverages. Furthermore, where 618 no comparators have been employed this could reflect practice effects. Nonetheless, based on the 619 published studies, and bearing in mind the varying methodological approaches employed, it is 620 difficult to ascertain the role of cow's milk in cognitive function for primary-school aged children 621 622 but it may at the very least be a useful conduit for nutrient fortification. This could be particularly meaningful within the school environment. The intervention studies did not measure academic 623 achievement directly, but their findings have relevant scholastic implications that warrant further 624 625 investigation using RCTs to establish if increased milk consumption influences academic achievement in primary-school aged children. This is especially so given the identification of a 626 positive association for boys between academic achievement and meeting the recommendations for 627 consumption of milk and alternatives (126). 628

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630 Future Directions and Conclusions

The aim of this narrative review was to evaluate evidence for the potential nutritional-, physicaland health-related benefits of cow's milk consumption for primary-school aged children (4-11 years). Cow's milk consumption (both plain and flavoured) improves nutritional status without adversely impacting on body mass and body composition⁽³⁵⁾. With some confidence, cow's milk also appears beneficial for hydration, dental and bone health and to have a beneficial to neutral effect concerning physical stature and appetite. Due to conflicting studies, reaching a conclusion has proven difficult concerning cow's milk and cognitive function therefore a level of caution should be exercised when interpreting these results. All areas, however, would benefit from further robust investigation, especially in free-living school settings, to verify conclusions. Improving elements of cognitive function, hydration and appetite regulation could have important long-term health and scholastic implications that should certainly be explored further. Indeed, recent research involving adolescent populations illustrates that high intakes of cow's milk are positively associated with academic performance⁽¹³⁴⁾, increased motivation for learning⁽¹³⁴⁾ and impact favourably on eating behaviour following acute and chronic consumption⁽¹³⁵⁾.

Despite a growing body of scientific literature exploring the potential benefits of cow's milk 645 646 consumption, there are several pertinent knowledge gaps that would benefit from further investigation. This is particularly true for cognitive function, hydration status and appetite 647 regulation, where there is little research available, especially in free-living school settings. Further 648 research, especially of a robust and methodologically sound nature (such as double-blind 649 randomised controlled trials), should seek to explore the mechanisms responsible for any effects 650 651 observed. At present, few studies on this population have attempted to establish mechanisms. Most purported mechanisms given throughout this narrative review come from adult-specific data and 652 653 should therefore be interpreted with some caution as the observations cited may not always be replicated in children. When working with child populations there are various considerations that 654 655 must be accounted for. The methodological approaches deemed most appropriate for the study of cognitive function, hydration status and appetite regulation in children will differ according to the 656 objective of the study, type of data required, and resources available⁽¹³⁶⁾. In children, it is of great 657 importance to adopt approaches that are non-invasive with a low level of participant stress. Current 658 techniques available to assess cognitive function, hydration status and appetite are arguably invasive, 659 elicit a moderate level of participant stress and increased ethical risk. This might explain the current 660 lack of studies and mechanistic insight from a child perspective. In recent years, however, research 661 groups have been pursuing non-invasive techniques that offer an opportunity to conduct 662 comprehensive mechanistic work in vulnerable populations. For example, developments in appetite 663 and metabolism research have identified fingertip-capillary blood sampling as an efficacious, 664 comparable and reproducible alternative to antecubital-venous blood sampling for the quantification 665 of appetite-related peptides^(137, 138). These developments will certainly help provide a better 666 understanding of mechanisms that influence appetite and eating behaviour in younger populations, 667 where traditional methods of venous blood sampling might be contraindicated. Furthermore, 668 fingertip-capillary blood sampling offers many advantages including simplistic application, 669 cost/time effectiveness and reduced volume of blood required for analysis⁽¹³⁹⁾. 670

671	Considering the nutritional-, physical- and health-related impact of cow's milk avoidance, the
672	evidence begins to highlight the importance of increasing cow's milk consumption. Cow's milk is a
673	naturally nutrient-dense foodstuff, providing a significant contribution of several essential nutrients
674	and bioactive constituents that potentially impact human health favourably. Establishing and
675	shaping healthy eating behaviours during the primary-school years is vital. Dietary behaviours
676	shaped throughout the childhood years progress through to adolescence and adulthood ⁽¹⁴⁰⁾ , making
677	healthy eating environments crucial. For primary-school aged children, the school setting may be an
678	ideal environment to promote cow's milk consumption, and school milk schemes are a great place
679	to start developing healthy eating behaviours given 35 % to 40 % of children's daily nutritional
680	needs are met at school ⁽¹⁴¹⁾ . Cow's milk is a readily available, accessible and affordable means of
681	providing valuable essential nutrients to the diets of primary-school children. Based on the evidence
682	presented in this manuscript, there appears no reason for primary-school children to limit cow's milk
683	consumption and there may, in fact, be many potential benefits to milk consumption.
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705 Author Contributions

- Benjamin Green and Penny Rumbold conceived and planned this narrative review. Penny Rumbold,
 Nicola McCullogh, Ruth Boldon and Benjamin Green sourced the relevant literature. All authors
 contributed to the writing and critically reviewed and approved the final manuscript.

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Conflicts of Interest

Penny Rumbold has previously received funding from The Dairy Council (UK), Nourishmenow and Cool Milk Ltd. Benjamin Green has previously been supported by The Dairy Council (UK) in the award of a PhD studentship and has received funding from Cool Milk Ltd and is presently an employee of Danone Specialised Nutrition. Lewis James has previously received funding from Volac International Ltd. Emma Stevenson has previously received funding from The Dairy Council (UK) and Arla Food Ingredients. Collectively, the above (with the exception of Cool Milk Ltd) was not related in any way to the work presented in this article. Nicola McCullogh, Ruth Boldon and Crystal Haskell-Ramsay declare no conflicts of interest.

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Per 100 g		Whole	Semi-Skimmed	Skimmed	Flavoured	
0		milk	milk	milk	milk <mark>*</mark>	
Energy (kJ)	274	195	144	270	
Protein (g)		3.3	3.5	3.5	3.6	
Carbohydr	ate (g)	4.6	4.7	4.8	9.6	
of which s	ugars (g)	4.6	4.7	4.8	8.9	
Fat (g)		3.9	1.7	0.3	1.5	
	of which saturates	2.5	1.1	0.1	1	
	monounsaturates	1	0.4	0.1	0.3	
	polyunsaturates	0.1	Trace	Trace	0.1	
	trans fatty acids	0.1	0.1	Trace	Trace	
Thiamin (r	ng)	0.03	0.03	0.03	0.03	
Niacin (mg	g)	0.2	0.1	0.1	0.1	
Niacin from	n Tryptophan (mg)	0.6	0.6	0.7	0.8	
Calcium (r	ng)	118	120	125	120	
Riboflavin	(mg)	0.23	0.24	0.22	0.17	
Vitamin B	6 (mg)	0.06	0.06	0.06	0.03	
Vitamin B	12 (µg)	0.6	0.9	0.8	0.1	
Folate (µg))	8	9	9	2	
Vitamin D	(µg)	Trace	Trace	Trace	Trace	
Biotin (µg))	2.5	3.0	2.5	2.2	
Pantothena	ite (mg)	0.6	0.7	0.5	0.3	
Vitamin C	(mg)	2	2	1	Trace	
Retinol (µ	z)	30	19	1	20	
Carotene	ug)	19	9	Trace	8	
Sodium (m	1g)	43	43	44	52	
Potassium	(mg)	155	156	162	168	
Magnesiur	n (mg)	11	11	11	12	
Phosphoru	s (mg)	93	94	96	102	
Zinc (mg)		0.4	0.4	0.5	0.4	
Copper (m	g)	Trace	Trace	Trace	Trace	
Selenium ((ug)	1	1	1	N/A	
Manganes	e(mg)	Trace	Trace	Trace	Trace	
Iodine (ug)	31	30	30	N/A	
1118 Note: N/A	, = values not available fo	or this food	: Trace = nutrient is pr	resent in less than 0	.1g per 100g. *Data taken	from a
1119 sample of s	trawberry and banana fl	avoured su	gar sweetened milk.			
1120 Adapted fro	om the Dairy UK. Avail	able at: htt	p://www.milk.co.uk/p	ublications/default.	aspx	
1	5		1 1		1	
1121						
1122						
1177						
1123						
1124						
1125						
1100						

1117 Table 1.0 Nutritional composition of cow's milk

	NDN Y 4-10	8 Rollin ′ears 1-2 years	g Progr 2; 2009/ 11-18	amme 10 years	NDNS Y 4-10	S Rollin ′ears 3-4 years	g Progr 4; 2011/ 11-18	amme 12 years	NDNS Y 4-10	5 Rollin ears 5-6 years	g Progr 5; 2013/ 11-18	amme 14 years	NDNS Rollin Years 7-8 4-10 years	g Programme 3; 2015/16 11-18 years
	Boys	Girls	Boys	Girls	Boys	Girls	Boys	Girls	Boys	Girls	Boys	Girls	Combined	Combined
Energy	8	8	4	4	7	7	5	4	8	7	4	4	7	4
Protein	14	12	8	7	13	12	9	8	13	13	10	7	11	8
Carbohydrate	5	5	4	3	5	5	4	3	5	5	4	3	5	4
Fat	9	10	6	5	9	10	6	4	10	11	6	4	9	6
Saturated fat	17	16	8	9	15	14	9	7	16	17	9	8	14	9
trans fatty acids	14	12	8	8	19	19	14	11	20	21	14	12	17	13
Vitamin A	11	10	7	7	10	10	9	4	12	12	9	7	11	9
Calcium	31	29	21	22	29	27	22	19	29	29	23	20	26	22
Riboflavin	32	29	22	21	28	29	23	20	29	31	23	18	25	19
Vitamin D	0	0	0	0	1	0	0	0	0	1	0	0	1	1
Sodium	5	4	3	4	6	5	3	2	6	6	3	4	6	4
Potassium	16	15	9	9	14	13	11	10	15	16	11	8	14	9
Magnesium	12	10	8	8	11	10	8	7	11	12	8	7	10	7
Zinc	14	13	8	9	13	13	9	8	13	13	10	8	12	8
Iron	1	1	0	1	1	0	1	0	1	1	1	1	1	1
Selenium	6	5	3	3	5	5	3	3	5	5	3	3	4	3
Folate	9	8	4	5	8	7	5	5	8	9	6	4	7	5
Iodine	40	38	30	30	37	37	32	28	39	28	34	27	37	30

Table 2. Percentage contribution of cow's milk to average daily total energy, macro- and micro-nutrient intake overtime by sex and age 1130

1131 Note: Percentage contribution information taken from National Diet and Nutrition Survey (NDNS) rolling programme. Percentage contributions for Vitamin B₆, Vitamin B₁₂, 1132 Vitamin C and Phosphorus are not reported in the NDNS. * The NDNS rolling programme for 2015/16 (https://www.gov.uk/government/statistics/ndns-results-from-years-7-

1133 and-8-combined) stopped providing sex specific intake data so the data are presented as boys and girls combined.

Reference	Details	Design	Methodological approach	Results and conclusion	Effect
Nutritional Statu	s				
Cook et al. (1975) ⁽³⁸⁾	n 312 (159 boys) Age: 8-11 years United Kingdom	Observational	7 day diet record	Nutrient intake was higher in children drinking school milk every day than non-consumers.	1
				Boys: energy (P < 0.05), protein (P < 0.01), fat (P < 0.05), calcium (P < 0.001), thiamin (P < 0.01) and riboflavin (P < 0.001) intakes higher than non-consumers	Ţ
				Girls: energy (P < 0.001), protein (P < 0.001), fat (P < 0.01), calcium (P < 0.001), thiamin (P < 0.001) and riboflavin (P < 0.001) intakes higher than non-consumers	↑
LaRowe et al. (2007) ⁽³⁹⁾	n 793 (410 boys) Age: 6-11 years United States	Observational	24 hour dietary recalls collected during NHANES 2001-2002.	General linear models showed children clustered as high-fat milk consumers had higher ($P < 0.05$ for all) intakes of energy, protein, riboflavin, folate, vitamin A, vitamin C and calcium compared to clusters of water, SSB, soda and mix/light drinks	Î
Albala et al. (2008) ⁽⁴⁴⁾	n 98 (56 boys) Age: 8-10 years Chile	Randomised, control trial	16 week intervention. Children drank 3 servings/d of flavoured milk or control (no drink)	Protein and calcium intakes increased ($P = 0.0001$) and energy intake decreased ($P = 0.009$) with milk compared to controls	↑
Murphy et al. (2008) ⁽⁴⁰⁾	n 2097 (1061 boys) Age: 6-11 years United States	Observational	24 hour dietary recalls collected during NHANES 1999-2002. Participants grouped by age (2-5 years; 6-11 years; 12-18 years).	Calcium, phosphorus, magnesium, potassium, and vitamin A intakes comparable between plain and flavoured milk consumers	\leftrightarrow
				Calcium, phosphorus, magnesium, potassium, and vitamin A intakes significantly higher ($P < 0.05$ for all) with milk consumption (plain and flavoured) compared to non-consumers	↑

Table 3 Studies in primary-school aged children that measured nutritional status and hydration with cow's milk consumption

Lien et al. (2009) ⁽⁴⁵⁾	n 454 (217 boys) Age: 7-8 years Vietnam	Double-blind, intervention	Volume matched drink (250 mL) served twice daily (morning), 6 days per week for 6 months: (a) multi-micronutrient fortified milk (b) unfortified milk (c) control (no drink)	Intakes of energy, protein, fat, sugar and vitamin A increased (P < 0.05 for all) after 6 months in conditions (a) and (b) and compared with condition (c)Serum ferritin and zinc increased (P < 0.05 for all) after 6 months in conditions (a) and (b) and compared with condition (c)Zinc levels however also increased in condition (c) (P < 0.05)	↑ ↑
Wang et al. (2012) ⁽⁴²⁾	n 632 Age: 8-10 years Canada	Observational	3 24 hour diet recalls	FM drinkers had a higher mean intake for calcium (930 vs. 837 mg; P = 0.010), Vitamin D (6.9 vs. 5.9 μ g; P = 0.021) and total sugar (99 g vs. 90 g, P = 0.015) than non-consumers	1
Rangan et al. (2013) ⁽⁴¹⁾	n 222 (121 boys) Age: 8-10 years Australia	Observational	3 non-consecutive 24 hour diet recalls	Milk (and dairy) intake significantly associated with increased intake of energy (P < 0.001), protein (P = 0.02), calcium (P < 0.001), phosphorus (P < 0.001), magnesium (P < 0.001), potassium (P = 0.009), zinc (P = 0.019), vitamin A(P < 0.001), riboflavin (P < 0.001), and niacin (P = 0.03)	Î
Campmans- Kuijpers et al. (2016) ⁽³⁷⁾	n 1007 (504 boys) Age: 7-13 years Netherlands	Observational	Two non-consecutive 24 hour dietary recalls with an interval of 2 to 6 weeks	Higher milk consumption was associated with significantly higher intakes of energy (P = 0.003), protein (P < 0.0001), fat (P = 0.03), fibre (P = 0.02), calcium (P < 0.0001), folate (P < 0.0001), iodine (P < 0.0001), potassium (P < 0.0001), magnesium (P < 0.0001), phosphorus (P < 0.0001), selenium (P = 0.002), zinc (P < 0.0001) and vitamins B ₁ (P < 0.0001), B ₂ (P < 0.0001) and B ₁₂ (P < 0.0001)	↑
Kuriyan et al. (2016) ⁽⁴³⁾	n 225 (52 boys) Age: 7-10 years India	Double-blind, randomised placebo-controlled	Volume matched drink (200 mL) served twice daily (morning and afternoon), 6 days per week for 5 months: (a) multi-micronutrient fortified milk	At the end of the intervention, levels of blood Vitamin B_{12} , and riboflavin were significantly different (P < 0.001) between the study groups, in favour of the fortified milk group	↑ ↔

			(b) unfortified milk	Vitamin D, selenium and body iron showed no difference with either group	
Hydration					
Montenegro- Bethancourt et al. (2013) ⁽⁵⁷⁾	n 442 (309 boys) Age: 4-10 years, Germany	Observational	DONALD Study 3 day weighed dietary records	Milk consumption significantly increased FWR (P < 0.05) by 25 mL in boys and 33 mL in girls per 100 g of intake	1
Volterman et al. (2014) ⁽⁵⁹⁾	n 38 (19 boys) Age: 7-11 years and 14-17 years, Canada	Randomised, repeated-measures cross-over	3 exercise sessions in 34.5 °C (2 × 20-min cycling bouts at 60% peak oxygen uptake) followed by consumption of: (a) plain water (W) (b) a carbohydrate/electrolyte solution (CES) (c) skimmed milk (SM)	Fraction of ingested beverage retained at 2 hour of recovery was greater with SM (74% \pm 18%) than W (47% \pm 26%) and CES (59% \pm 20%, p < 0.001 for both), and greater in CES than water (p < 0.001)	↑
Volterman et al (2016) ⁽⁵⁸⁾	n 15 (7 boys) Age: 10-12 Canada	Randomised, counterbalanced crossover design	 45 min of alternating running and cycling exercise in 34.5 °C followed by consumption (150% of body mass loss) of isocaloric and electrolyte-matched beverages containing: (a) 0 g (control), (b) 0.76 g (Lo-PRO) (c) 1.5 g (Hi-PRO) of milk protein/100 mL 	Less negative fluid balance at 2 hour of recovery after consumption of Hi-PRO vs. CONT (P = 0.01) Compared with CONT, beverage retention was enhanced by Hi-PRO at 2 h (P < 0.05)	↑ ↑

Abbreviations in order of appearance: FM = flavoured milk; NHANES = National Health and Nutrition Examination Survey; SSB = sugar sweetened beverage;

FWR = free water reserve; W = water; CES = carbohydrate/electrolyte solution; SM = skimmed milk; Lo-PRO = low protein; Hi-PRO = high protein.

Reference	Details	Design	Methodological approach	Results and conclusion	Effect
Dental Health					
Petti et al. (1997) ⁽⁷⁴⁾	n 439 (217 boys) Age: 6-11 years Italy	Descriptive cross- sectional	24 hour diet diary, oral examination using Plaque Index and number of decayed, extracted or filled teeth	Milk consumption significantly reduced the probability of carries ($P < 0.05$); greater significance in high-sucrose (<4/day) consuming children ($P < 0.01$)	Î
Levine et al. (2007) ⁽⁷²⁾	n 315 Age: 7-11 years England	Prospective cohort/cross- sectional	Three-day dietary diary, tooth brushing habits, dental examination using BASCD survey	Significantly ($P < 0.05$) less caries associated with moderate consumption (1-2/day) of milk (and dairy) products by children aged 11-15 years	1
Llena & Forner (2008) ⁽⁷³⁾	n 369 (220 boys) Age: 6-10 years Spain	Descriptive cross- sectional	Food frequency questionnaire, one dental examination for caries and fillings	No significant impact of milk on the incidence of carries	\leftrightarrow
Curtis et al. (2018) ⁽⁷⁵⁾	n 392 (183 boys) Age: 0-19 years United States	Longitudinal, observational	Dietary questionnaires at six- month intervals, cavities assessed age 5, 9, 13 and 17 years	Higher milk intake associated with lower expected adjusted decayed and filled surface increments, however, was not significant	1
Bone Health and	Physical Stature				
Cook et al. (1979) ⁽³⁶⁾	n 1210 (581 boys) Age 6-7 years United Kingdom	Prospective cohort	Free school milk consumed (190 mL/d at school) vs control; height gain	Significantly greater height gain in the milk group in Scottish females ($P < 0.05$)	1
_				No difference in males or any groups in England	\leftrightarrow
Rona & Chinn (1989) ⁽⁹⁸⁾	n 670 Age 5-11 years United Kingdom	Prospective cohort	Free school availability (190 mL/d at school) vs unavailable; change in standardised height score	Significantly greater increments in standardised height scores in 'milk available' group after 2 years in Scotland ($P < 0.05$)	↑
				No difference in England	\leftrightarrow
Baker et al. (1980) ⁽⁹⁴⁾	n 581 (267 boys) Age 7-8 years United Kingdom	Randomised, control trial	12 month intervention; 190 mL/d of milk at school vs control; height gain	Significantly greater height gain in milk group than control ($p < 0.05$)	1
Chan et al. (1995) ⁽⁸⁴⁾	n 48 females Age: 9-13 years USA	Randomised control trial	12 month intervention; ≥ 1200 mg/d calcium from dairy vs control; DXA of total body bone mineral content	Greater gain in total body bone mineral content in dairy calcium group (P < 0.001)	1

1142	Table 4 Studies in primar	y-school aged children that m	easured dental health, bone health and	l physical stature with cow's milk consum	ption
	1				

Bonjour et al. (1997) ⁽⁸³⁾	n 149 females Age: 6-10 years Switzerland	Randomised control trial	12 month intervention; 850 mg/d milk-extracted calcium vs placebo; DXA of regional bone mineral content, height gain and change in lumbar spine length	Greater increment in six-site mean bone mineral content in supplementation group (P < 0.05). At 1 year follow-up greater increment in femoral shaft bone mineral content (P < 0.02)	$\uparrow \\ \leftrightarrow \\ \uparrow$
Black et al. (2002) ⁽²⁵⁾	n 250 (120 boys) Age 3-10 years New Zealand	Cross-sectional	Food frequency questionnaire, and DXA of total body and regional bone mineral content and height	Milk avoiders had significantly lower total body bone mineral content than age, sex-matched controls ($P < 0.01$) Milk avoiders significantly shorter than control children of the same age and sex ($P < 0.01$)	↑ ↑
Du et al. (2004) ⁽⁸⁵⁾	n 757 (girls) Age 10-12 years China	Randomised control trial	24 month intervention; 330 mL fortified milk (560 mg calcium) / school day vs 300mL fortified milk (560 mg calcium) and 5 or 8 µg cholecalciferol / school day vs control group. DXA of regional and total body bone mineral content with stature assessed pre, mid and post-trial	Greater increase in total-body bone mineral content (> 1.2%) and bone mineral density (> 3.2%) in milk groups compared to control Significantly greater % change in height in milk supplement groups compared to control group (P < 0.005)	↑ ↑
Goulding et al. (2004) ⁽⁸⁶⁾	n 50 (20 boys) Age: 3-10 years New Zealand	Observational longitudinal	Milk avoiders vs birth cohort population, DXA of total body and regional bone mineral content, history of bone injuries, estimation of calcium intake	Greater number of children reporting fractures and increased total fractures in milk-avoidance compared to control ($P < 0.001$)	Ť
Rockell et al. (2005) ⁽²⁹⁾	n 46 (18 boys) Age 5-12 years New Zealand	Cross-sectional, longitudinal	Food frequency questionnaire, four-day food diary and DXA of total body bone mineral content and stature	Increase in total body bone mineral content in prior milk avoiders when milk consumption had increased ($P < 0.05$) but remained lower than none milk avoiders	1

Wiley	n 2592	Cross-sectional	24 hour recall, milk frequency in	Milk intake not associated with height at age 5-11	\leftrightarrow
$(2005)^{(99)}$	Age 5-11 year		past 30 day; height and adult	years ($P = 0.385$)	
	United States		height		
Iuliano-Burns et al.	n 99 (58 boys)	Randomised	10 month intervention;	Greater gain in pelvic bone mineral content in pre-	\leftrightarrow
(2006) ⁽⁸⁷⁾	Age 5-11 yearrs	control trial	800 mg/d of calcium from CaCO ₃	pubertal children vs. controls at 10 months in the	
	Austria		vs 800 mg/d of calcium from milk	milk mineral group ($P < 0.02$) but not biologically	
			vs placebo, DXA or total and	meaningful	
			regional bone mineral content		
Albala et al.	n 98 (56 boys)	Randomised,	16 week intervention;	Significantly greater increase $(P = 0.01)$	1
$(2008)^{(44)}$	Age: 8-10 years	control trial	Children drank 3 servings/d of		
	Chile		flavoured milk or control (no	Smaller increase in height in females in	\leftrightarrow
			drink)	intervention group vs control ($P = 0.10$)	
Zhou et al.	n 435 (188 boys)	Retrospective	School milk group (1 year, 4/wk or	Greater ulna bone mineral content in school milk	1
$(2011)^{(88)}$	Age 10-12 years	cohort	3 year, 1-3/wk) vs. 'seldom milk'	group (P < 0.001)	
	China		group, ulna bone mineral content		
Guo et al.	n 41,439	Cross-sectional	Validated questionnaire assessing	Milk intake not associated with stature ($P > 0.05$)	\leftrightarrow
(2020) ⁽⁹⁵⁾	(19,618 boys)		milk intake classified as low,		
	Age 6-17 years		medium and high; stature		
	China				

1143 Abbreviations in order of appearance: DXA = dual energy X-ray absorptiometry; CaCO₃ = calcium carbonate.

Reference	Details	Design	Methodological approach	Results and conclusion	Effect
Appetite Regulati	on				
Rumbold et al. (2013) ⁽¹¹¹⁾	n 25 (11 boys) Age: 9-10 years United Kingdom	Randomised controlled cross- over	Energy matched (0.3 MJ) preloads during school mid-morning break: (a) 160 mL of semi-skimmed milk (b) 153 g of apple	Boys and girls felt less hunger and expressed a lower desire to eat when apple was consumed compared to semi-skimmed milk ($P = 0.02$) Energy intake comparable at lunch between semi- skimmed milk and apple ($P > 0.05$)	$\begin{array}{c} \downarrow \\ \leftrightarrow \end{array}$
Mehrabani et al. (2014) ⁽¹¹⁰⁾	n 34 (obese boys) Age: 10-12 years Iran	Three-way repeated measure randomised controlled cross- over	Volume matched preloads with breakfast (all 240 mL): (a) low fat milk (1.5 %) (b) apple juice (isoenergetic control) (c) water (control)	Energy intake lower at lunch following low fat milk consumption compared to water (P <0.005) and apple juice (P <0.05)	↑
Vien et al. (2014) ⁽¹¹²⁾	Normal and overweight children Mean age: 11.5 years Av age: 11.5 years Canada	Not stated	Volume (250 mL) and energy matched (130 kcal) with the exception of water given pre- and within meals: (a) 1 % fat (1 g/100 g) flavoured milk (b) 2 % fat (2 g/100 g) milk (c) 1.5 % fat (1.5 g/100 g) yogurt drink (d) fruit punch (e) water	 Experiment 1 Energy intake lower after chocolate milk and yogurt (P < 0.01) compared to water Post-meal SA was lower after milk than yogurt (P < 0.01) Post-meal SA was lower after milk than fruit punch (P < 0.01) Experiment 2 GLP-1 AUC was higher after milk than fruit punch and in OWOB compared to NW children (P < 0.03). Post-prandial drop in ghrelin was greater after milk than fruit punch in OWOB children (-24 vs 14%) but was not significant (P = 0.06) 	\uparrow \uparrow \uparrow \leftrightarrow
					\leftrightarrow

Table 5 Studies in primary-school aged children that measured appetite and cognitive function with cow's milk consumption

				Energy intake, insulin and glucose AUC were comparable between all preloads ($P > 0.05$)	
Mehrabani et al. (2016) ⁽¹⁰⁹⁾	n 34 obese boys Age: 10-12 years Iran	Three-way repeated measure randomised controlled cross- over	Volume matched preloads with breakfast (all 240 mL): (a) low fat milk (1.5 %) (b) apple juice (isoenergetic control)	Higher satiety scores after drinking low fat milk with breakfast compared with water and apple juice (P < 0.05) Energy intake lower at lunch following low fat	↑ ↑
			(c) water (control)	milk compared to water ($P < 0.001$) and apple juice ($P = 0.03$)	1
Cognitivo Eurotion					
Rahmani et al. (2011) ⁽¹²⁴⁾	n 469 (230 boys) Av age: 8 years	Case-control population-based	Drink at morning school break: (a) tetra-pack sterilised and	Grade-point average increased from pre- to post- test in girls with milk ($P < 0.05$)	1
	Iran	intervention	(b) control (no milk supplementation)	No change for the control group, nor for either of the groups of boys.	\leftrightarrow
				Mean IQ after milk was better in boys compared to boys at post-test in the control ($P < 0.05$)	↑
Brindal et al. (2013) ⁽¹²³⁾	n 40 (19 boys) Age: 10-12 years, Australia	Double-blind, randomised three- way repeated measures crossover	Volume matched (455 mL) morning drink following ≥ 8 hours fasting: (a) glucose beverage	No effect of beverage type in speed of processing, working memory, short-term memory, attention switching, perceptual speed and inspection time	\leftrightarrow
			(b) full milk beverage(c) half milk/glucose beverage	No interactions between beverage type and timing of the cognitive testing (60, 120 and 180 minutes post-drink)	\leftrightarrow
				In the conditions with milk, girls recalled $0.7-0.8$ more words but in the glucose condition they recalled 0.5 fewer words (P = 0.014).	Ť
				For boys, no difference between beverages was found (P >0.09).	\leftrightarrow
				Sex differences identified for change in word recall after full milk ($P < 0.001$) and half	\leftrightarrow

				milk/glucose ($P < 0.001$) conditions: girls showed an increase and boys showed a decrease	
Kuriyan et al. (2016) ⁽⁴³⁾	n 225 (52 boys) Age: 7-10 years India	Double-blind, randomised placebo-controlled design	Volume matched drink (200 mL) served twice daily (morning and afternoon), 6 days per week for 5 months: (a) multi-micronutrient fortified milk (b) unfortified milk	No group × time interaction for short-term memory and executive functions from baseline	\leftrightarrow
Zahrou et al. (2016) ⁽¹³²⁾	n 200 Age: 7-9 years Morocco	Double-blind, controlled design	Volume matched (200 mL) drink served daily at school for 9 months, including weekends: (a) fortified milk; added potassium iodide and other micronutrients (b) non-fortified milk	Fortified milk group performed significantly better than the non-fortified milk group ($P=0.02$) on a dynamic testing procedure designed to assess children's learning potential	\leftrightarrow
Faught et al. (2017) ⁽¹²⁶⁾	n 1595 (732 boys) Age: 10-11 years Canada	Observational	Analysis of secondary data (food frequency questionnaire) from a 2012 population-based survey using a stratified random sampling design	Boys meeting the nutrition recommendations for milk and alternatives (3-4 servings per day) scored 3.45 % better on an average measure of Math and Language Arts standardised provincial achievement tests than those not meeting the recommendations (P =0.02)	Î
Petrova et al. (2019) ⁽¹²⁵⁾	n 103 (52 boys) Age: 8-14 years Spain	Double-blind, randomised controlled design	Volume matched (200 mL) drink served three times per day for 5 months: (a) fortified milk beverage; added micronutrients and essential fatty acids (b) regular full milk	Greater baseline to post-intervention increases in working memory capacity in the fortified milk group (32 % increase) compared to the regular milk group (13 % increase) ($P = 0.027$)	\leftrightarrow
Sheng et al. (2019) ⁽¹³³⁾	n 75 (42 boys) Age: 5-6 years China	Double-blind, multicentre randomised controlled parallel crossover	 Volume matched (150 mL) drinks consumed twice daily following meals for 5 days: (a) conventional milk, containing A1 and A2 β -casein (b) milk containing only A2 β- casein 	Subtle Cognitive Impairment Test improved following both conventional milk (P < 0.014) and milk containing only A2 β -casein (P < 0.002) No difference in Subtle Cognitive Impairment Test between drinks	\leftrightarrow
					\leftrightarrow

				Consumers of conventional milk in phase 1 and milk containing only A2 β -casein in phase 2, no effect on error rates in phase 1 (P = 0.101) but a decrease in error rates in phase 2 (P < 0.001) Consumers of milk containing only A2 β -casein in phase 1 and conventional milk in phase 2, there was a decrease in the error rate in phase 1 which continued through the washout period (P < 0.001), but error rates then increased in phase 2 (P < 0.001)	Ļ
Anderson et al. (2020) ⁽¹²⁷⁾	n 84 (39 boys) Age: 8-12 years, United States	Randomised counterbalanced crossover	Volume matched (237 mL) morning drink following ≥ 8 hours' fasting: (a) 1% fat milk (b) apple juice	Inhibitory control Reaction time significantly faster after milk compared to apple juice (P <0.05)	↑ ↓
			(0) apple juice	No effect of beverage on accuracy Speeded working memory No difference between the beverages on reaction time ($P = 0.45$)	\leftrightarrow
				Sex \times beverage interaction for accuracy (P = 0.003); compared to apple juice, milk decreased performance accuracy for females whereas non-significant increase in accuracy for males	↑↔
				Sustained attention Sex \times beverage interaction (P = 0.02); compared to apple juice, milk significantly improved reaction time for females whereas reaction time decreased in males (though not significantly)	↑↔

Abbreviations in order of appearance: SA = subjective appetite; GLP-1 = glucagon like peptide-1; OWOB = overweight and obese; NW = normal weight; AUC = area under the curve; IQ = intelligence quotient