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2	INVITED REVIEW: ADVANCES IN GOAT MILK RESEARCH
3	Subtitle: One hundred years of advancing goat milk research through JDS
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15	ABSTRACT
16	In the century of research chronicled between 1917 and 2017, dairy goats have gone from
17	simply serving as surrogates to cows, to serving as transgenic carriers of human enzymes. Goat
18	milk has been an important part of human nutrition for millennnia, in part because of the
19	greater similarity of goat milk to human milk, softer curd formation, higher proportion of small
20	milk fat globules and different allergenic properties compared to cow milk; however key
21	nutritional deficiencies limit its suitability for infants. Great attention has been given not only
22	to protein differences between goat and cow milk, but also fat and enzyme differences, and
23	their impact on the physical and sensory properties of goat milk and milk products.
24	Physiological differences between the species necessitate different techniques for analysis of
25	somatic cell counts, which are naturally higher in goat milk. The high value of goat milk
26	throughout the world has generated a need for a variety of techniques to detect adulteration of
27	goat milk products with cow milk. Advances in all of these areas have been largely documented
28	in the Journal of Dairy Science (JDS); this review summarizes such advances.
29	

30 key words: adulteration, composition, nutrition, somatic cells, safety

31

32 INTRODUCTION

33 Previously considered the "poor man's cow", goats and goat milk products began gaining 34 attention in the US in the 1960s because of health and nutritive values attributed to goat milk 35 and milk products. Touted for its easy digestibility and lower allergenic properties compared to 36 cow milk, goat milk has been considered a nutraceutical for decades, but many initial reports 37 were anecdotal. The JDS played a large role in documented the true differences between cow 38 and goat milk. Haenlein (1980) even credited JDS as "a major US research organ on dairy goats 39 as well as on dairy cows". In the 100-year period since 1917, JDS has published more than 850 40 research manuscripts related to goat milk and milk products. However, these numbers do not 41 reflect the full scope of research related to dairy goats, or the role that goat milk and milk products have played in advancing the global dairy industry in the past century. With particular 42 43 focus on JDS publications, this manuscript is dedicated to those discoveries.

44

45 ADVANCES IN GOAT MILK RESEARCH FROM 1917 TO 2017

46 Goat milk and human nutrition

47 Likely since the beginning of domestication, the importance of goats for human nutrition has 48 been recognized. Indeed, the first publications related to goat milk, published in *The Lancet*, 49 tended to focus on infant feeding, and some of the risks and benefits associated with it (Blackham 1906; Cahill 1906; Dalebrook, 1902; Wright 1906). One letter to the editor of The 50 51 *Lancet* claimed that "goats practically never have tubercle, therefore their milk can be given 52 without pasteurizing... their milk is said to be better for infants than cow's milk because the 53 curd is finer" (Edmunds, 1914). Prompted by the observation that goat milk rarely forms a 54 cream layer, though fat content was similar to that of cow milk, Schultz and Chandler (1921) 55 reported that 91% of goat milk fat globules were under 4 µm in diameter. Previous work by 56 Bitting (1902) reported that 90% of cow milk fat globules were more than 4 μ m in diameter. 57 Although it soon became clear that goat milk was also susceptible to microbial contamination, the softer curd and higher proportion of small fat globules have been selling points of goat milk
ever since these early works.

60

61 In the early 1900s, vitamins and minerals were almost exclusively studied in rats, chicks, and 62 monkeys. Approximately 15 years before the early "Our Industry Today" report by Elvehjem (1953), work in his lab revealed that rats grew more slowly on goat milk than cow milk. By 63 64 then, several cases of severe anemia had been associated with goat milk feeding of human infants, and the term "goat's milk anemia" was coined. Elvehjem (1953) reported that goat 65 66 milk provided inferior amounts of vitamin B_{12} and that levels of folic acid in goat milk and cow 67 milk were "about equal" (which has been since shown untrue). However, since improvement in 68 rat growth was seen with folic acid supplementation, a sparing effect of folic acid on vitamin B₁₂ 69 was indicated. Still in the early days of understanding the role of folic acid and B₁₂ in human 70 health, Collins et al. (1953a and 1953b) published two companion papers in JDS, the former 71 related to cow colostrum and milk, and the latter related to goat colostrum and milk. Because 72 vitamin B₁₂ levels in sheep milk could be increased by the addition of cobalt or trace-minerals 73 (containing cobalt; Harper et al. 1951), they wanted to evaluate the impact of such diet 74 supplementation in goats. Goats receiving trace-mineralized salt (containing cobalt) had a 75 higher level of vitamin B₁₂ in their colostrum and milk during the first week post-partum 76 compared to those receiving only iodized salt. Trace-mineralized salt or a 50 mg supplement of 77 cobalt per goat per day had no influence upon the level of B₁₂ in goat milk after this time. The 78 addition of trace-minerals to the diet of the goat did not influence the free folic acid level of the 79 goat milk. The authors admitted that the information reported in the JDS work was "more 80 accurate" than what the reported in their previous work (Collins et al. 1951).

81

It was not realized until later that goat milk was deficient, with respect to human nutrition, in
folic acid, and vitamins B₁₂ and B₆, nutrients that are essential for normal human baby
development (Ford and Scott 1968; Parkash and Jenness 1968). Nonetheless, goat milk
products gained considerable attention in the 1970s because of perceived health and nutritive
value. Jenness (1980) provided a good review of goat milk nutritive value based upon literature

87 of the time. Similar to cow milk, goat milk is an adequate to excellent source of protein, 88 calcium, niacin, pantothenic acid phosphorus, potassium, riboflavin, thiamin and vitamin A to 89 the human diet (Parkash and Jenness 1968; Jenness 1980). Neither cow nor goat milk is a good 90 source of iron, vitamin C or vitamin D (unless fortified). In contrast to cow milk, goat milk 91 contains less than adequate levels of vitamins B₆, vitamin B₁₂, and folic acid than cow milk for 92 infant nutrition (Ford and Scott 1968; Parkash and Jenness 1968; Jenness 1980). Folic acid and 93 vitamin B₁₂ deficiencies became a focus of research in the 1970s, regarding megaloblastic 94 anemia in children exclusively fed goat milk (Davidson and Townley 1977), and continue to be 95 of concern today (Basnet et al. 2010; Ziegler et al. 2005).

96

97 One of the main characteristics of goat milk that has contributed to its appeal as an alternative 98 to cow milk is its lower allergenic properties compared to cow milk. Even today, families are 99 known to switch to goat milk or to buy a dairy goat to avoid cow milk consumption. Yet mostly 100 antecdotal evidence for the lower allergenicity of goat milk was reported until the 1990s 101 (Haenlein 2001; Loewenstein et al. 1980). With an incidence of 2 to 3% in the first year of life, 102 cow milk allergy is the most common food allergy in early childhood, but the remission rate is 103 approximately 85 to 90% by adulthood (Høst 2002). In an outstanding review published in JDS, 104 Jenness (1980) noted that in many cases, allergy to cow milk proteins was not improved by 105 shifting patients to goat milk, and he recognized that α_{s1} -casein may play a role. It was not until 106 Ballabio et al. (2011) published in JDS that the clear relationship was established. By running 107 individual milk samples from 25 goats with different α_{s1} -casein genotypes through SDS-PAGE 108 and immunoblotting using monoclonal antibodies specific for bovine α -casein and sera from 109 children allergic to cow milk, Ballabio et al. (2011) showed that goat milk allergenicity is a 110 function of α_{s1} -casein genetic polymorphism. Lower reactivity was shown for samples with null 111 α_{s1} -casein genotypes (0₁0₁ or 0₁F). Their work confirmed that caution must be taken before 112 goat milk is suggested as an alternative to cow milk for patients with cow milk allergy. They 113 went further to indicate that goat milk from particular α_{s1} -casein genotypes could possibly 114 serve as protein sources for hypoallergenic formulas (Ballabio et al. 2011). The findings were 115 echoed by Lisson et al. (2014), who confirmed that although genetic variants of caseins differ in

their allergenicity, cross-reactivity of IgE antibodies of goats and buffaloes with cow milk

117 caseins limit feeding goat or buffalo products to cow milk-allergic patients.

118

119 In "Past, present, and future perspectives of small ruminant dairy research", Haenlein (2001) 120 provided an outstanding review of over 135 manuscripts related to, primarily, goats and sheep. 121 Haenlein noted that research prior to 2001 was scarce on the unique qualities of goat and 122 sheep milk compared to cow milk; largely it had been assumed that technical research on cows 123 could be extrapolated to small ruminants. Haenlein summarized differences in anatomy, 124 physiology, nutrition, metabolism, and pathology of goats and sheep, as well as differences in 125 their milk and milk products and economic profitability. Although not mentioned in his 126 manuscript, perhaps a dairy goat check-off program could help narrow the gap of disparity in 127 research dollars spent on cows and dairy goats. Particularly compelling was Haenlein's 128 statement regarding the potential of goat and/or sheep milk to combat under- and malnutrition 129 of people in poor areas and countries. Only 21 out of the 24 countries Haenlein included in his 130 summary met the recommended level of calcium intake (1,000 mg/day). All but five countries 131 met the recommended level of protein consumption (50 g/day) in the form of animal protein 132 (developed countries); six countries had below or borderline levels of protein consumption 133 even after plant sources of protein were added in. The bottom line is that many countries have 134 room to improve animal protein and milk utilization. In his conclusion, Haenlein (2001) urged 135 continued research, extension service, and public support to improve the productivity of small ruminant dairy animals, particularly in developing nations that rely on these animals to a much 136 137 greater extent than developed nations.

138

In recent years, epidemiological studies have led investigators to consider estrogen a factor that
may contribute to reproductive system cancers (Farlow et al. 2009; Yager and Davidson, 2006).
The World Cancer Research Fund/American Institute for Cancer Research (2007) demonstrated
no relationships of importance for consumption of milk and cancer, with the exception of
colorectal (decreased risk) and prostate (increased risk) cancers. Because of concerns about
estrogen metabolites in milk, and consumption being associated with cancers of the

reproductive system (Farlow et al. 2009), Farlow et al. (2012) compared estrone (E_1) and 17 β estradiol (E_2) levels in commercial goat and cow milk. Goat milk exhibited a lower combined concentration of E_1 and E_2 than cow milk.

148

149 Goats serve as surrogates to cows

150 The earliest JDS manuscript that specifically mentioned dairy goat milk was published in 151 Volume 15, in May 1932, entitled "Fat Metabolism in the Lactating Goat" (Bender and 152 Maynard, 1932). Similar to the sister manuscript, published in Volume 17, in March 1934, 153 entitled "The Effect of Specific Dietary Fats on the Blood Lipids of Lactating Goats" (Williams 154 and Maynard, 1934), authors of both manuscripts stated that dairy goats were selected for the 155 research to save expense, rather than to study the dairy goat metabolism in particular. The 156 authors explicitly stated an assumption that, physiologically, dairy goats and dairy cows would 157 perform similarly. It is surprising that in those early works, adequate sample size and 158 replication were not required for publication. Findings for four goats, who received four 159 different dietary treatments, were reported in the Bender and Maynard (1932) manuscript. 160 Cunningham and Addington (1935) destined goats to be "used more and more in fundamental 161 research problems" because of their convenient size-equating them with five to seven dairy 162 cows-and greater offspring potential. Since these early works, it has been realized that caution 163 must be exercised when using the goat as a model for the dairy cow (Larson 1978), and dairy 164 goats and their milk are worthy of study in their own right.

165

166 Advances in goat milk composition research

Bergman and Turner (1937) were among the first to report on the composition of dairy goat colostrum, in particular, the globulins (importantly associated with immune bodies). They reported a rapid transition of colostrum (characterized by high total solids, fat and total protein) from six Toggenburg does into nearly normal milk by the third and fourth day after parturition. At the time, total protein was composed of four groups of protein, namely casein, casein globulin, albumin, and globulin. They used the "newer methods of protein analysis", including precipitation with 8% trichloroacetic acid for determination of total protein, and casein precipitation with an acetate buffer solution, to quantitatively determine total protein,
free from non-protein nitrogen and casein. By salting out with MgSO₄, Bergman and Turner
(1937) were also able to track globulin and albumin separately, for the first time. The most
rapid change was seen in globulin, which was reported to decrease from 1.76% on day one to
0.40% on day two, and 0.11% by day nine. Since albumin did not decrease to the same extent,
globulin was determined to be the driver in protein transition between colostrum and normal
milk.

181

182 Until 1940, only four research manuscripts reported goat milk composition data (Bosworth and 183 Van Slyke 1916a, b, c; Lythgoe 1940). Lythgoe (1940) conducted proximate analysis on 335 184 samples from individual goats from 21 herds in MA, across a 16-month period. The results are 185 summarized in Table 1. The work confirmed the high individual and seasonal variability in total 186 solids (driven primarily by high variability of fat), which was more pronounced in goats than in 187 cows. Fifteen years later, in another early JDS "Our Industry Today" literature reviews (Rusoff 188 1955) included a table comparing milk composition of various mammals (Table 2). The information was from the 2nd edition of a McGraw-Hill book, *The Market Milk Industry* 189 190 (Roadhouse and Henderson, 1950). In subsequent years, a few manuscripts related to goat 191 milk composition were published. Jenness (1980) compiled the mean total solids, fat, crude 192 protein, lactose and ash from milk of international goat breeds from 11 references reported in 193 nine countries between 1968 and 1979. At the time Jenness (1980) wrote his review, the 194 composition of milk from individual US goat breeds had still not been reported; oddly, milk 195 from pygmy goats was used to represent US goat milk composition. It wasn't until Alderson 196 and Pollack (1980) summarized 3,481 milk and fat yield records of Alpine, LaMancha, Nubian, 197 Saanen and Toggenburg goats from a cooperating herd in CA, that we gained an appreciation 198 for the differences in milk composition and milk production of US dairy goat breeds. Milk and 199 fat yield were influenced by age, month, and year of freshening, and Nubians had the lowest 200 yields but highest fat content (3.8%). Haenlein (1981), who evaluated the production records 201 of US dairy goats, also showed that milk from Nubians had the highest fat content (4.6%) and 202 lowest yield (806 kg/305-day record).

203

204 [Table 1 near here]

205

206 [Table 2 near here]

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208 More recently, Guo et al. (2001) collected commingled commercial goat milk shipments for an 209 entire year to provide fundamental information for cheese making and milk cheese yield 210 potential and pricing. Samples were collected weekly from bulk milk shipments to a 211 commercial cheese plant from April, 1996 to March, 1997. The bulk milk was composed of milk 212 from 12 dairy farms, composed of Saanen, Nubian, LaMancha, Alpine, and Toggenburg breeds, 213 in New Hampshire and Vermont. Total solids (TS) and fat (F) contents decreased over the first 214 20 weeks from 12.7 and 3.6% to 11.3 and 3.0%, respectively, then increased to peak values of 215 13.4 and 4.4 in January. The contents of crude protein and casein also decreased in the first 20 216 weeks, from 3.5 and 2.7% to 3.2 and 2.3%, respectively, then increased gradually to 3.8 and 217 2.9% in February. The physicochemical properties of commingled goat milk, Table 3, was 218 adapted from Table 1 from the manuscript (Guo et al. 2001). Summer milk had the highest 219 yield potential per kg of protein, due to a higher proportion of casein in crude protein; late 220 lactation milk from does that freshened in the summer had the lowest yield potential. Guo et 221 al. (2001) also concluded that, because of the high lactational and seasonal variability, milk 222 standardization, especially in February, will enable greater uniformity in cheese composition 223 and functionality.

224

[Table 3 near here]

226

A viral disease that causes animal and economic losses in goat production throughout the world is caprine arthritis encephalitis (CAE) (The Center for Food Security and Public Health, 2007). The lentivirus responsible for CAE disease is caprine arthritis encephalitis virus (CAEV), which affects animals in the form of chronic progressive arthritis, pneumonia, chronic weight loss, encephalomyelitis, and indurative mastitis (Kaba et al. 2012). Because contradictory results

232 have been shown for milk production studies, Kaba et al. (2012) investigated the influence of 233 CAEV on milk yield, somatic cell count (SCC), and percent fat, protein and lactose in a 12-year 234 cohort study with 177 does. No significant differences were found between infected and 235 uninfected animals for daily milk yield or SCC (non-leukocytic epithelial cell-like particles). 236 However, the milk of uninfected goats contained more total protein, fat and lactose than that 237 of the infected goats. Martínez-Navalón et al. (2013) studied the Marciano-Granadina breed, 238 which commonly carries CAEV. Longer lactations, higher milk yield, fat, normalized mean SCC 239 and lactose content were found in seronegative goats. According to their findings, CAEV 240 infection could be a major cause for decreased milk production in dairy goats, however they 241 mentioned that transmission routes and potential causes of this disease are still unclear and 242 need more research (Martínez-Navalón et al. 2013).

243

244 Studies with somatic cells

245 Similar to dairy cows, mastitis is the primary, and most costly infection of dairy goats, and dairy 246 goat mastitis research published in JDS has been extensive in the past four decades (Contreras 247 et al. 1997; Contreras et al. 2003; Gelasakis et al. 2016; Koop et al. 2010; Koop et al. 2012; 248 Moroni et al. 2005; Timms and Schultz 1985). Summary of such work is beyond the scope of 249 this manuscript. However, although high SCC are strongly associated with mastitis in cows, that 250 is not always the case with goats (Dulin et al. 1983; Koop et al. 2012; Park and Humphrey 1986). 251 It has long been known that the milk of goats naturally contains elevated levels of somatic cells 252 compared to cows, because of the apocrine secretory system in the mammary gland, and that 253 for cows, elevated SCC are associated with cheese quality defects (Dulin et al. 1982; Dulin et al. 254 1983; Poutrel and Lerondelle 1983; Park 1991; Zeng and Escobar 1996). But since bacterial cell 255 counts do not explain high SCC in goat milk (Park and Humphrey 1986), the impact of SCC on 256 goat cheese has been debated. Dulin et al. (1982) studied the differentiation and enumeration 257 of SCC in goat milk. Results indicated that cytoplasmic particles that are similar in size to milk 258 somatic cells, and commonly found in goat milk, can be mistakenly counted as somatic cells by 259 milk quality machines, therefore it was recommended to use counting methods that are 260 specific for DNA for estimation of somatic cells in goat milk to have an accurate differentiation

261 of cells from cell-like material. Zeng (1995) compared SCC and chemical composition of goat 262 milk using Fossomatic-300 and Dairylab II, calibrated either with goat milk or cow milk. In both 263 machines, SCC estimation in goat milk was higher when cow milk was used as a standard than 264 with goat milk as a standard. Moreover, results significantly exceeded legal limits established 265 by the FDA (1,000,000 cells/mL for goat milk at the time, as opposed to the 750,000 cells/mL 266 limit for cows (HHS, PHS, FDA, 2011). Zeng (1995) indicated that the natural differences 267 between cow and goat milk can lead to SCC, protein, and fat reading errors by milk quality 268 equipment when they are set up with cow milk as a standard. Therefore, he recommended to 269 use goat milk as a standard when testing goat milk quality to collect reliable data.

270

271 Somatic cells, and their impact on cow milk quality, have been extensively studied. Factors 272 such as milking methods, breed, age, stage of lactation, season, and management have been 273 reported to affect SCC in cow milk, however it is not always the case in goat milk (Zeng and 274 Escobar, 1996). Milk from Nubian and Alpine dairy goats and three milking methods (pipeline, 275 bucket and hands) were tested to compare SCC, standard plate count (SPC) and chemical milk 276 composition during a complete lactation (Zeng and Escobar 1996). There was no significant 277 difference among milking methods, but SCC increased as lactation advanced and SPC was 278 higher in Nubian milk than Alpine milk. Milk fat and protein of both breeds increased during 279 the first 60 days of lactation and then decreased. Some milk samples contained over 1 million 280 somatic cells/mL, which exceeded the legal limit for Grade "A"; nevertheless does did not 281 experience mastitis symptoms such as swelling or redness of udder. A pathology test indicated 282 that *Staphylococci* were the predominant bacteria, but there was no mastitis condition. 283 Therefore, there was an indication that healthy does could produce milk with more than 1 284 million SCC/mL and the Grade "A" SCC rule should be reviewed to truly reflect goat udder 285 health. In a more recent study with sixty Alpine goats not exhibiting clinical mastitis, Chen et al. 286 (2010) demonstrated that milk composition did not change when SCC varied from 214,000 to 287 1,450,000 cells/mL. Milk with higher SCC actually had lower standard plate count. Coliform and 288 psychrotrophic bacteria counts, milk components (fat, protein, lactose, casein and total solids), 289 and yield of semisoft goat cheese, did not differ among low, medium and high SCC goat milk.

290 However, body and texture scores provided by trained panelists were lower and FFA were 291 higher for high cheeses made with milk with highest SCC (Chen et al. 2010). Today, the Grade 292 "A" Pasteurized Milk Ordinance allows 1,500,000 cells/mL for goat milk, 750,000 cells/mL for 293 cow, sheep and camel milk (HHS, PHS, FDA, 2015). Albenzio et al. (2015) went deeper into the 294 goat physiology with their research into activities of indigenous proteolytic enzymes in goat 295 milk of different SCC. They identified 700,000 cells/mL as the threshold for changes in the 296 immune status of the goat mammary gland. Similar to cow and sheep, plasmin appeared to be 297 the predominant enzyme activity in goat milk, which was correlated to SCC, and macrophages 298 in particular (Albenzio et al. (2015).

299

300 It is important to note that the other temperature, chemical, physical and bacteriological 301 standards for Grade "A" raw milk and Grade "A" pasteurized milk and/or milk products do not 302 differ for cow and goat milk (HHS, PHS, FDA, 2015). Goats, and goat milk and milk products are 303 held to the same high standards for safety and quality that the dairy industry is known for. 304

305 Findings with fatty acids

306 By 1964, the overall significance of short-chain fatty acids (FA) of ruminant milk fat was not fully 307 known, but it was recognized that goat milk had unique flavor properties. Efthymiou and 308 Mattick (1964) developed a domestic feta cheese in order to provide uniformity to the 309 unpredictable quality of feta cheeses that were being produced in the US at the time. Although 310 feta cheese is traditionally made from goat and/or sheep milk, their method was developed to 311 produce characteristics of "typical Greek Feta" using cow milk. The authors concluded that 312 characteristic (desirable) rancid flavor, specifically from free fatty acids (FFA) C₂ to C₁₀, could be 313 consistently produced using a mixed culture of Streptococcus (now Lactococcus) lactis and 314 either Lactobacillus casei or Lactobacillus acidophilus, and lipase powder (either Capalase-KL 315 (kid and lamb-derived) or Capalase-L (lamb-derived pregastric esterase)). Bitter, atypical rancid, 316 and unclean flavors were associated with the use of Capalase-K or Italase, and FA of C₁₂ or 317 higher chain length predominated.

318

319 As the recognition of unique properties of goat milk grew, in particular FA, Dimick and Patton 320 (1965) set out to understand the role of butyric acid, and its function in milk fat synthesis. It 321 had previously been reported in JDS articles (Jack et al. 1963; Jensen et al. 1961), that butyrate 322 was esterified in sn-1 and sn-3 positions in triglycerides. For their experiments, they utilized 323 fresh raw milk from cows (herd milk) and goat (one animal) milk. Their work demonstrated 324 similarities in mole percent distribution of FA from each of the triglyceride fractions, with 325 butyrate concentrations topping out at 20.4 mole percent (goat) and 20.0 mole percent (cow), 326 yet they determined few, if any, dibutyryl triglycerides exist in either type of milk. Ultimately, 327 the authors concluded that butyrate exists predominantly as one mole per mole of triglyceride 328 in both cow and goat milk. Freeman et al. (1965) later reported, in JDS, the distribution of FA in 329 goat, sheep, Indian buffalo, cow and human milk using methyl esters via gas-liquid 330 chromatography. Short-chain FA, $C_{4:0}$ and $C_{6:0}$, were determined to be esterified predominantly 331 in the sn-1 and sn-3 positions in all species. The C_{14:0} and C_{15:0} FA were preferentially esterified 332 at the sn-2 position, while C_{18:0} was primarily esterified at sn1 or sn3 positions. Not long later, 333 Breckenridge and Kuksis (1967), utilizing butyl esters in gas-liquid chromatography, revealed 334 the FA distribution of milk from seven species (Table 4). Differences in the findings between 335 Freeman et al. (1965) and Kuksis (1967) were attributed to methodology (methyl vs. butyl 336 esterification) and milk sources.

337

338 Attaie and colleagues (1993) were the first to investigate FA profiles of cow and goat colostrum. 339 Simultaneous distillation extraction was used to separate short-chain from long-chain FA, and 340 the n-butyl esters of FA were quantified by gas chromatography and identities confirmed by gas 341 chromatography-mass spectrometry. Table 5 displays the concentration of total FA in goat and 342 cow colostrum. Similar to milk of goats and cows, significant differences in colostrum fatty acid 343 profile were generally found between species. However, the amounts of hexanoic, octanoic, 344 decanoic, 9-Decenoic, and dodecanoic acids also differed between goat breeds, with Nubians 345 presenting more of each of the aromatic compounds.

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347 [Table 4 near here]

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349 [Table 5 near here]

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351 Chilliard and colleagues (2003) provided an excellent review on the nutritional and physiological 352 factors affecting goat milk lipid synthesis and lipolysis. In contrast to cows, goat milk fat 353 content increases with almost all studied fat supplements, however, the response of fatty acid 354 composition is similar in the two species. Although the LPL system of goat milk is lower than 355 that of cow milk, it is more bound to the fat globules (vs. casein micelles in cows) and more 356 strongly correlated to spontaneous lipolysis (lipolysis at 4°C) in goat milk. LPL activity is 357 influenced by stage of lactation, milking frequency, fasting, and lipid supplementation (Chilliard 358 et al. 2003). The lipolysis and characteristic goat milk flavor were attributed to a combination 359 of goat milk fatty acid (FA) composition, triglyceride structure (i.e., high proportion of C_6 to C_{10} 360 FA esterified on carbon 3) and LPL characteristics. The authors also suggested fat 361 supplementation of diets to improve goat milk composition for greater control of cheese 362 processing and sensory properties (Chilliard et al. 2003).

363

364 Bouattour et al. (2008) showed that feeding a moderate level of soybean oil (6% as fed in the 365 concentrate) to dairy goats increased total milk fat, conjugated linoleic acid (*cis*-9, *trans*-11 $C_{18:2}$ 366 CLA), and *trans*-vaccenic acid (trans-11 C_{18:1} VA) in milk without negative effects on intake, milk 367 yield or protein content. In the same issue of JDS, Luna et al. (2008) reported increases in α -368 linolenic acid, CLA, VA, as well as minor conjugated linoleic acid isomers, in the milk of goats fed 369 whole linseed and sunflower oil. Subsequently, Chen et al. (2009) demonstrated that feeding of 370 a dietary supplement containing *trans*-10, *cis*-12 conjugated linoleic acid (3 to 6g/d/goat) 371 reduced milk fat synthesis in dairy goats and decreased cheese moisture and yield. Martínez 372 Marín (2012) fed increasing amounts of 3 plant oils (linseed oil, LO; high oleic sunflower oil, 373 HOSFO; and regular sunflower oil, RSFO) to dairy goats. Oil supplementation decreased the 374 level of saturated FA in milk fat (especially $C_{16:0}$) and increased mono-and polyunsaturated FA in 375 a linear manner. LO supplementation appeared to be the most favorable alternative of the 376 three because of the positive impact on rumenic acid and vaccenic acid and decrease in the

377 omega-6 to omega-3 FA ratio in milk fat (Martínez Marín 2012). Even more recently, Toral et al. 378 (2015) set out to compare lipid metabolism of goats and cows. Animals were fed diets 379 containing no additional oil (control), or supplements of fish oil, sunflower oil and wheat starch, 380 in a 3 X 3 Latin square design, with 26-d experimental periods. Their work demonstrated 381 interspecies differences in mammary lipogenesis, suggesting a lower sensitivity to the inhibitory 382 effects of trans-10, cis-12 CLA in goats and that ruminal biohydrogenation pathways are more 383 stable and less prone to diet-induced shifts toward *trans*-10-containing intermediates in goats 384 than cows.

385

386 With the emergence of biorenewable sources of fuel has come the production of by-products 387 such as dried distillers grains with solubles (DDGS), a by-product of the ethanol industry. A 388 good amount of literature is available on the impact of DDGS feeding on poultry, swine, beef, 389 dairy cows, and even cow milk and cheese (Sankarlal et al. 2015, Testroet et al. 2015). Cais-390 Sokolińska et al. (2015) were the first to report on the impact of DDGS on goat and sheep milk 391 and milk products, when they evaluated formation of volatile compounds in the fermented 392 beverage, kefir. Their work showed that the increased polyunsaturated fats resulting from 393 DDGS feeding resulted in significant changes to the fermentation process and aroma profile of 394 the resulting kefirs Cais-Sokolińska et al. 2015).

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396 Evolution of goat milk enzyme research

397 The enzyme composition of ruminant milk was not completely characterized by 1968, and is 398 probably still not. Chandan et al. (1964) reported the lysozyme content of human milk in 399 *Nature*, then proceeded to report on the composition of lysozyme, lipase and ribonuclease in 400 the milk of five species in JDS (Chandan et al. 1968). There was interest in lysozyme due to the 401 discovery that human milk had nearly 3,000 times the amount of lysozyme than that of cow 402 milk, and potential implications to infant feeding and keeping quality of milk. The investigators 403 confirmed the great discrepancy in lysozyme content of human milk (40,000 μ g/100 mL) 404 compared to cow (13 μ g/100 mL), goat (25 μ g/100 mL), sheep (10 μ g/100 mL), and sow (0 405 μ g/100 mL). Differences in lipase (13, 132, 39, 9, and 141 μ M/min/100 mL, respectively) and

406 ribonuclease (305, 1,100, 425, 300 and 30 µg/100 mL, respectively) were also notable, but not 407 as extreme (Chandan et al. 1964). Later, with the emergence of genetic engineering, transgenic 408 goats were developed to express human lysozyme at least 67% of the concentration found in 409 human milk that enhanced the antimicrobial properties of goat milk to select mastitis and 410 pathogenic microorganisms (Brundige et al. 2008; Maga et al. 2006). Maga et al. (2006) 411 demonstrated that milk from the five transgenic goats had lower somatic cell count, but the 412 overall component composition of the milk and milk production were not different from 413 controls. Additional benefits included that milk from the transgenic goats had a shorter rennet 414 clotting time and increased curd strength.

415

416 Milk xanthine oxidase (XO) was also a hot topic in the 80s. Oster (1971) proposed an 417 association between XO and atherosclerosis. Because of such concerns, Zikakis and Wooters 418 (1980) evaluated a total of 195 commercially processed dairy products, polarographically, for 419 XO activity. Fluid milk, cream, powdered and evaporated milk, yogurt and ice cream, cheese, 420 butter, as well as goat and sheep products, were evaluated. The authors reported XO activity 421 of raw milk increased with storage, particularly frozen storage, and that commercial processing 422 destroyed about 82% of XO activity compared to raw milk. Commercial processing allowed the 423 release of XO from the milk fat globule membrane, enabling destruction. Cheeses made from 424 goat and sheep milk (Feta and imported blue) were reported to contain low to no XO activity 425 (Zikakis and Wooters 1980).

426

427 DeFeo et al. (1982) were among the first to distinguish differences in the lipoprotein lipase (LPL) 428 system between goats and cows. The importance of the research lies in the fact that hydrolytic 429 rancidity (lipolysis) aromas and flavors from volatile FFA are influenced by native LPL, and 430 acceptability of goat milk products are largely influenced by rancid flavors. To characterize 431 components of the lipolytic system in goats, in part because of the unique flavor characteristics 432 of goat milk, Chilliard et al. (1984) activated spontaneous lipolysis in goat milk. Unlike for cow 433 milk, LPL activity is correlated with spontaneous lipolysis in goat milk. Goat milk LPL was found 434 to be distributed primarily in the cream (46%) and serum (46%), with little activity in the caseins

(8%), in comparison to cows (6, 17 and 78%, respectively). It has been shown that the LPL
activity differs among several breeds of goat, with evidence of genetic polymorphism
influencing the functional properties of this enzyme (Badaoui et al. 2007).

438

439 Plasmin is likely the most important proteases in milk because of its influence upon milk and 440 cheese quality. Although a lot of research into the plasmin enzyme system had previously been 441 conducted in cows, it was not until the early 1990s that anyone reported on the plasmin system 442 in goats. Like in cow milk, the complex plasmin enzyme system, composed of plasmin (PL), 443 plasminogen (PG), plasminogen activators (PA), plasminogen activator inhibitors, and plasmin 444 inhibitors, is present in goat milk (Politis et al. 1994). For the first time, Politis et al. (1994) 445 demonstrated that tissue plasminogen activators (t-PA) were present in the casein and serum 446 fractions of goat milk; urokinase plasminogen activators (u-PA) were present in all fractions 447 (i.e., casein, serum, and somatic cells). Electrophoretic studies by Trujillo et al. (1997) 448 demonstrated that plasmin hydrolyzed the same regions in β -casein in cow and goat milk. The 449 plasmin system is also involved in mammary involution, with higher PL and PA activity in late 450 lactation cows (Baldi et al. 1996). Fantuz et al. (2001) evaluated the plasminogen activation 451 system in goat milk and its relation with composition and coagulation properties toward the 452 end of lactation. Compared to cow and sheep milk, goat milk PG activity was low, but 453 consistent with the high activity of PA. The high PL and PA activity in goat milk was negatively 454 correlated with coagulating properties in late lactation, which was likely related to degradation 455 of casein (Fantuz et al. 2001).

456

The quality of cheeses largely depends on the rate, extent and nature of the two main biochemical processes involved in cheese aging, proteolysis and lipolysis. With the growing popularity of goat milk cheeses, and paucity of information regarding proteolysis and lipolysis specific to goat milk cheeses, Park (2001) published a review on the topic in JDS. Regarding proteolysis, one of the distinguishing differences between cow and goat milk is the ratio of caseins. Because of its naturally lower content of α_{s1} -casein, goat milk has a higher proportion of β -, α_{s2} - and κ -casein than cow milk. As a consequence, goat cheeses tend to be less firm,

and less resistant to enzymatic degradation than cow cheeses (Park 2001). Earlier work
published in JDS, by Fontecha et al. (1990), Ha and Lindsay (1991), Attai and Richter (1996a),
and Jin and Park (1995), were also cited as being important for the characterization of lipolysis
in goat milk cheeses.

468

Because the majority of goat milk cheese sold in the US are fresh, soft cheese (chevre), and because goats are largely seasonal breeders, availability is variable. Thus, Van Hekken et al. (2005) evaluated the impact of frozen storage on the proteolytic and rheological properties of soft goat cheese. The creation and removal of ice crystals in the cheese matrix and the limited proteolysis of caseins resulted in only slight changes to cheese texture. Thus, authors concluded that frozen storage of soft cheeses may be appropriate to enable year-round supply of soft goat cheese, but consumer evaluation was not conducted to confirm this.

476

477 Genetic variants of goat milk caseins

478 The five principal proteins in goat milk (α_{s1} -casein, α_{s2} -casein, β -casein, β -lactoglobulin and 479 α -lactalbumin) were reported to closely resemble their homologs in cow milk (Jenness 1980). 480 Research at the time suggested that goat milk lacked the homolog of bovine α_{s1} -casein, the 481 most abundant protein in cow milk. Jenness (1980) attributed goat milk's reputed more easily 482 digested, softer curd, to the lack of α_{s1} -casein in goat milk; yet he acknowledged that no direct 483 experimental evidence was yet available on the subject. However, research in the late 1980s 484 would reveal interesting findings about goat α_{s1} -casein. The JDS was one of the first journals to 485 publish a manuscript on the topic in English, when Ambrosoli et al. (1988) reported that 486 coagulation properties (coagulation time, rate of curd formation and curd firmness) and 487 composition of goat milk with low and high α_{s1} -casein content differed. They reported that 488 goat milk with low α_{s1} -casein had faster coagulation time, while milk with high α_{s1} -casein had 489 higher levels of components and produced firmer curds. Later, Mora-Gutiérrez et al. (1991) 490 demonstrated, using isoelectric precipitation and reversed-phase HPLC, that milk from Alpine 491 and Nubian dairy goats could be divided into low, medium and high- α_{s1} -casein-producing 492 groups. The authors proposed the idea of genetic regulation of α_{s1} -casein production, stopping

493 short of suggesting it to be a breed-specific trait. In subsequent years, it was realized that at 494 least ten different genetic variants influenced the α_{s1} -casein phenotype expressed, and genetic 495 variants were associated with breeds, milk composition, and coagulation properties; those 496 works were not presented in JDS (Martin and Addeo 1996; Clark and Sherbon 2000). Later, it 497 was reported that at least 16 alleles are associated with different rates of α_{s1} -casein protein 498 synthesis in goats (Caroli et al. 2006).

499

500 Cebo et al. (2012) demonstrated that genetic polymorphisms at the α_{s1} -casein locus affect both 501 structure and composition of milk fat globules. At mid-lactation, goats displaying high-type α_{s1} -502 casein genotypes produced larger fat globules and had lower levels of polar lipids in the MFGM 503 than goats with null α_{s1} -casein genotype. More work in this area should be expected in the 504 coming years, since the authors suggest that genetic polymorphism in goats may be a tool to 505 provide clues into lipid secretion pathways in the mammary epithelial cell (Cebo et al. 2012). 506 Advances in metabolomics, using hyphenated gas chromatography-mass spectrometry and 507 multivariate data analysis techniques, enabled Caboni et al. (2016) to characterize low 508 molecular weight polar metabolites in milk of 28 goats with different α_{s1} -casein genotypes in 509 Italy. Upregulated compounds associated with weak genotypes included sugars and polyols, 510 while upregulated compounds associated with strong genotypes included citric and aconitic 511 acids (Caboni et al. 2016).

512

513 Characterizations of caprine κ -casein genotypes were reported in JDS by several authors (Coll et 514 al. 1993, 1995; Angiolillo et al. 2002; and Yahyaoui et al. 2003). Coll et al. (1995) characterized 515 the nucleotide sequence of the cDNA and the promoter region of the κ -casein gene. Angiolillo 516 et al. (2002) characterized three variants of goat κ -casein (designated A, B, and C) in Spanish, 517 French, German and Italian goat breeds. Yahyaoui and colleagues (2003) proposed a 518 nomenclature for the different alleles representing κ -casein variants. The full coding region of 519 the κ -case gene, including two new genetic variants were described, along with allele 520 distribution among 210 animals representing different European goat breeds and 23 Spanish 521 wild goats. The technique described by Yahyaoui et al. (2003) allowed the rapid and

simultaneous genotyping of all known κ-casein variants; use of such a system could enable
selection of milk for various industrial applications.

524

525 Growth in goat population and goat research

526 Goat milk research began to blossom in the 1970s, along with dairy goat populations in the US. 527 According to Leach (1980), the number of registered dairy goats in the US increased from 3,611 528 in 1955 to 32,459 in 1976. Additionally, herds enrolled in the National Cooperative Dairy Herd 529 Improvement Program (DHIP) increased from none in 1960 to 1,611 in 1978 (Leach 1980). By 530 1987, approximately 129,225 milk goats were counted (on 15,443 farms) in the USDA APHIS 531 Census, with approximately 17 million kg of milk produced; however, it was acknowledged that 532 the census does not always capture all animals (USDA, APHIS, CEI, 2003). Considering the 533 number of dairy goats not on test in the US, Haenlein (1978) estimated that closer to 350 534 million kg of milk were produced by US dairy goats annually in the 1970s and 1980s. Assuming 535 that the census captured only 60% of the true population, by 2002, the dairy goat population 536 had grown to 407,105 in the US (USDA, APHIS, CEI, 2003). By 2012, the US dairy goat 537 population was approximately 413,540 (USDA NASS).

538

539 It wasn't until Haenlein (1978) published "Dairy goat management" in JDS, that statistically 540 significant published research about nutritional and breeding management, behavior, and 541 economics of milk production of dairy goats was comprehensively reported. Around this time, 542 Larson (1978) suggested caution to animal scientists for using the dairy goat as a model in 543 lactation studies. Some of the most obvious differences, he pointed out, were the gross 544 structural differences between goats and cows, and differing milk constituents. He also 545 summarized the important differences in susceptibility to metabolic diseases associated with 546 lactation and differing rates of metabolism affecting transfer of dietary and administered 547 materials into milk.

548

549 A full issue of JDS was dedicated to dairy goats in 1980, resulting from the 1979 ADSA

550 International Symposium on Dairy Goats (Haenlein 1980). Issue 10 contained 14 manuscripts

551 related to dairy goats. Perhaps the most comprehensive summary of goat research at the time, 552 "Composition and characteristics of goat milk: Review 1968-1979" (Jenness 1980) was one of 553 them. Twenty-seven (11%) of the references cited were manuscripts published in the JDS. 554 Some of the key findings during the period from 1968 through 1979 included the observance 555 that although fat globules of goat milk resemble cow milk, goat milk lacks agglutinin, which 556 causes fat globules of cow milk to cluster when cooled (Jenness and Parkash, 1971). This, 557 coupled with the fact that goat milk contains a higher proportion of small fat globules than 558 large (Schultz and Chandler, 1921; Jenness 1980), explains why goat milk is called "naturally 559 homogenized". However, it was not until Cerbulis et al. (1982) that the lipid distribution of goat 560 milk was formally investigated and reported. Goat milk resembled cow milk fat with respect to 561 lipid fractions of whole milk and cream, containing 97 to 99% free lipid (97% of which was in the 562 form of triglycerides) and 1 to 3% bound lipid (containing neutral lipid, glycolipid and 563 phospholipid). However, goat skim milk contained more free lipid than cow milk, likely because 564 of the higher proportion of small globules (Cerbulis et al. 1982).

565

566 In the same issue as Jenness (1980), a review of research on goat milk products was published 567 (Lowenstein et al. 1980). Loewenstein and colleagues referenced 136 publications pertaining to 568 preparation of consumer products from goat milk; an additional 183 manuscripts were included 569 as "supplementary bibliography". Through their review, they concluded that, until that date, 570 cheese was the only extensively-studied goat milk product, and additional research of goat milk 571 products is needed. Perhaps partially in response, characterization of goat milk flavors surged 572 in JDS in the 1980s and 1990s (Chilliard et al. 1984; Iverson et al. 1989; Ha and Lindsay 1991; 573 Martín-Hernández et al. 1992; Jin and Park 1995; Attaie and Richter 1996a and b). The 574 characteristic "goaty" aroma of goat milk products results from the volatile FA that are found in 575 higher quantities in goat milk and milk products compared to cow milk. Branch-chain FA, 576 including 4-ethylocatanoic (goat-like or "goaty") and 4-methyloctanoic (mutton-like) acids, from 577 goat and sheep milks provide distinguishing flavors to varietal cheeses (Ha and Lindsay 1991). 578 They reported an absence of 4-ethylocatanoic acid in cow milk cheeses, and suggested that the 579 flavor compound, in particular, distinguished cow from goat and sheep cheeses. Additionally,

580 the presence of phenols, particularly p-cresol and 3- and 4-ethylphenols (sheep-like flavors) 581 were unique to sheep cheeses (Ha and Lindsay, 1991). Attaie and Richter (1996a) 582 demonstrated that ripening time significantly affected the concentrations of FFA in Cheddar-583 like hard goat cheeses up to 12 weeks, and that the percentage of NaCl or the ratio of salt to 584 moisture (S/M) did not affect FFA or lipolysis. In their companion paper (Attaie and Richter 585 1996b), it was shown that firmness of the Cheddar-like cheeses decreased up to 18 weeks, but 586 no significant change occurred between weeks 18 and 24. Cheeses with higher salt (highest 587 S/M) remained the most firm, explained by the lower hydration of the protein and less freedom 588 of movement for the protein molecules, larger amount of intact casein, and firmer casein 589 matrix (Attaie and Richter 1996b).

590

591 With the growing importance of dairy goats came the need for design of breeding programs. 592 Iloeje et al. (1981) were among the first. They evaluated 21,845 records of dairy goats on Dairy 593 Herd Improvement tests from 1965 to 1976. The relative importance of herd (22-31% of total 594 variation in milk and fat yields and 15 to 25% of variation in fat%), doe (16 to 25% or total 595 variation in milk yield, fat yield and fat%), sire (8 to 10% of the total variation), and year-season 596 effects (8 to 14% of total variation) were found to be similar to those for dairy cattle. Ali et al. 597 (1983) followed up, with a study of 42,618 records of goats with 125 days or more in milk, to 598 examine relationships among lactation and reproduction traits. Milk and fat yields were 599 affected by breed, parity, age after fitting parity, and month of conception. The authors 600 recommended a reduction in the number of days dry since it was found to be negatively 601 correlated with milk and fat yield in subsequent lactation (Ali et al. 1983).

602

603 Decades of adulteration detection

As the appreciation for and value of goat milk increased, methods to detect of goat milk with cow milk became necessary. Methods were published in Bulgaria (1929), Norway (1952), and France (1959) before the US. Aschaffenburg and Dance (1968) were among the first to publish methods to detect cow milk in goat milk by gel electrophoresis. Furtado (1983) utilized discontinuous polyacrylamide gel electrophoresis (PAGE) for detection of cow milk in

609 pasteurized goat milk. Because of the naturally-lower amount of α_{s1} -casein in goat milk than 610 cow milk, a frontal band, missing from the pattern of genuine goat milk and possessing the 611 same electrophoretic mobility as bovine α_{s1} -casein, could be directly related to the amount of 612 cow milk added to the goat milk. Iverson and Sheppard (1989) demonstrated that adulteration 613 of sheep and goat cheeses was occurring throughout the world by evaluating the fatty acid 614 profiles of 134 cheeses using programmed temperature gas-liquid chromatography of fatty acid 615 butyl esters. Goat and sheep milk cheeses exhibited a characteristically different lower chain 616 length fatty acid pattern than cow milk cheeses. The mean lauric:capric fatty acid (C_{12} : C_{10}) ratio 617 became proportionally larger with increased substitution of cow milk for goat or sheep milk in 618 cheese making. Later, Molina et al. (1996) reported on the use of Western blotting of native 619 and denatured bovine β -lactoglobulin to detect addition of bovine milk to non-bovine milk 620 cheeses. Native PAGE of whey or isoelectric focusing of β -lactoglobulin isolated from the casein 621 fractions was followed by immunodetection with anti-bovine β -lactoglobulin antiserum. 622 Immunoblotting of the native-PAGE plates of whey proteins from cheese allowed detection of 623 heat-denatured whey proteins or pasteurized cow milk added to goat cheese at less than 1% 624 adulteration. Even more recently, López-Calleja et al. (2004) utilized species-specific 625 polymerase chain reaction techniques to detect sheep and goat milk adulteration with cow 626 milk. The use of a forward primer complementary to a conserved DNA sequence, along with a 627 reverse primer specific for cow, yielded a 223-bp fragment from cow milk DNA, whereas no 628 amplification signal was obtained in sheep or goat milk DNA. When applied to raw, pasteurized, 629 or sterilized milk mixtures of cow-sheep and cow-goat, the specific detection of cow milk had a 630 good sensitivity threshold (0.1%). In follow-up work, López-Calleja et al. (2005) validated the 631 effectiveness of the technique to authenticate the purity of sheep milk, with similar sensitivity 632 threshold (0.1%). Adulteration continues to be of concern today. In Brazil, a study was 633 requested by the association of small-holder producers to investigate and to inhibit 634 adulteration practices (Rodrígues et al. 2012). A duplex PCR assay was developed, standardized 635 and validated on 160 fresh bulk goat milk samples. The detection limit was 0.5% bovine milk in 636 goat milk; 41.2% of the goat milk present in the market was adulterated with bovine milk at the 637 time (Rodrígues et al. 2012). Also using PCR, Golinelli et al. (2014) reported that all locally

638 produced goat cheeses tested (20 lots of 4 brands in Brazil) were adulterated with cow milk, 639 even though labels did not indicate addition of cow milk. Additionally, almost half of the 102 640 regular consumers invited to participate in triangle tests were able to perceive adulteration of 641 goat cheese with 10% (vol/vol) cow milk (Colinelli et al. 2014). Chen et al. (2016) used 642 proteomics to quantify the percentage of cow milk added to goat or sheep milks or dairy 643 products. Signature tryptic peptides in β -lactoglobulin were used as markers. The ultra-644 performance liquid chromatography triple quadrupole-mass spectrometry method was found 645 to have high accuracy, selectivity, linearity and precision. Similar to many previous studies, 646 adulteration was found in most of the commercial samples purchased (Chen et al. 2016).

647

648 Microbiology and safety

649 In the early part of the 20th century, livestock in the US were commonly infected with Brucella 650 species. While cows were often carriers of *Brucella abortis*, goats carried *Brucella melintensis*. 651 These were the early days of determining the current Federal Food and Drug Administration's 652 "60-day rule", stating that cheesemakers use pasteurized milk, or age raw milk cheese for at 653 least 60 days at not less than 35°F (1.5°C), which was established in 1950 (21 CFR 133; FDA 654 HHS, 2016). Gilman et al. (1946) evaluated the length of time that *B. abortis* (the cow-borne 655 source of Brucellosis) survived in Cheddar cheese. Uniquely, they also evaluated the survival of 656 B. melintensis (the goat-borne source of Brucellosis) in goat cheeses, as fresh goat cheeses 657 made from unpasteurized milk had been implicated in human undulant fever cases (Gilman et 658 al. 1946). However, no documentation of cases was provided in the manuscript. Gilman et al. 659 (1946) also reported, similar to other work of the day, that aging for about 60 days would 660 provide "reasonable assurance of the absence of viable B. abortus in commercial Cheddar 661 cheese." Surprisingly, although Gilman and colleagues did not conduct experiments with B. 662 melintensis, based upon previous research, they reported that *B. melintensis* may live longer 663 than 60 days in cheese, stating "goat milk cheese presents a special problem." Between 1965 664 and 1983, outbreaks of Brucellosis in Colorado and Texas in were linked to consumption of 665 cheeses made from unpasteurized goat milk sourced from the US or Mexico, or consumed 666 while US residents were visiting the Mediterranean basin, Far East, Middle East, and South

America (CDC, 1983; Eckman, 1975; Young and Suvannoparrat, 1975). Because of vaccination
programs and vigilance, since the 1980s, *Brucella* species have essentially been eradicated from
US livestock. There has only been one case of *B. melintensis* (not *B. abortis*) reported since, a
single cow in Texas, in 1999 (USDA APHIS CEI. 1999).

671

672 Milk-borne infections were more common before pasteurization was discovered in the late 19th century and commonly implemented in the 20th century, but outbreaks related to consumption 673 674 of unpasteurized milk remain a concern (Langer et al., 2012). In Scotland, milk pasteurization 675 was mandated in 1983, but not England or Wales, and sale of unpasteurized sheep and/or goat 676 milk was not prohibited anywhere in Great Britain at the time Sharp et al. (1985) wrote. 677 Nonetheless, more cases of foodborne illness were related to cow milk than goat milk during 678 that time (Sharp et al., 1985). In the US, Michigan was the first state to require milk 679 pasteurization, in 1948; in 1987, interstate shipment of raw milk was prohibited by the FDA 680 (Langer et al., 2012). In the period between 1993-2006, a disproportionate number (150-times 681 higher incidence) of outbreaks of foodborne illness were associated with non-pasteurized than 682 pasteurized dairy products, and in states that allow sales of raw milk (Langer et al. 2012). 683 Between 2007-2012, 4 outbreaks were associated with goat milk compared to 77 associated 684 with cow milk (Mungai et al. 2015). Goat milk and milk products have tended to stay out of the 685 food safety news spotlight, with a few exceptions, again, typically associated with 686 unpasteurized products, and mostly outside of the US (Bielaszewska et al. 1997; Hatchette et al. 687 2001; Hogerwerf et al. 2011; Lai et al. 2015; Méndez Martínez et al. 2003; McIntyre et al. 2002). 688

689 Because of their particularly high virulence and negative consequences in humans

690 contamination with shiga toxin-producing *Escherichia coli* (STEC) and *Listeria monocytogenes* 691 are of particular concern to dairy producers and processors. Until the 1990s, there was a lack 692 of information of *L. monocytogenes* in goat milk compared to the information available on cow 693 and sheep milk. Because of the high mortality rate associated (30%) with listeriosis, Gaya and 694 colleges (1996) evaluated incidence of *Listeria* species (spp.) in caprine milk in Spain. The 695 incidence of *Listeria* spp. in samples from bulk tanks of 405 farms was 4.15%. There was a peak

696 during autumn and winter months, compared to the reported spring peak for cows. The 697 findings confirmed the risk for *Listeria* contamination of cheese made of raw caprine milk. Of 698 796 raw milk cheeses obtained in 2006 and 2007 in Switzerland, 3.7% and 6.3% were positive 699 for pathogenic STEC, respectively (Stephan et al. 2008). Of the 63 goat cheeses evaluated, 4 700 goat milk soft cheeses and 1 goat milk semihard cheese (7.9%) were positive for STEC. The 701 various serotypes were not enumerated in samples, so it is unknown if levels were high enough 702 to cause foodborne illness. Nonetheless, authors concluded that raw milk cheeses continue to 703 be a potential vehicle for transmission of pathogenic STEC to humans (Stephan et al. 2008). 704

705 D'Amico et al. (2008) studied the overall pathogen presence in 138 samples of cow, sheep and 706 goat raw milk, from 11 farmstead cheese operations in Vermont, US. The study targeted 707 Staphylococcus aureus, Salmonella spp., L. monocytogenes, and E. coli O157:H7. Goat milk 708 samples had lower incidence (18.4%) of *S. aureus* than cow (27.4% or sheep (85.7%) milk 709 samples. S. aureus was present at levels that could not lead to a risk to produce heat-stable 710 enterotoxins (<100,000 CFU/mL); S. aureus is not considered a pathogen of high risk during 711 cheesemaking due to competition with active starter culture. E. coli O157:H7, Salmonella spp., 712 and L. monocytogenes were not found in goat milk at levels that could cause foodborne 713 infections but outbreaks from these microorganisms could still occur if conditions allow. 714 D'Amico and colleges (2008) concluded that since bacteria were found in very low levels in goat 715 milk, improper storage could still facilitate pathogen growth, and some properties of raw milk 716 cheeses could lead to the survival and growth of certain pathogens to risky levels.

717

Rahimit and Alian (2013) evaluated raw milk of buffaloes, cows, goats, sheep and camels, in
Iran, to test for the presence of *S. aureus* and its different enterotoxins. Twenty-two samples of
the total 200 tested positive for *S. aureus*, with the highest prevalence found in buffalo milk,
followed by cow, sheep, goat and camel, respectively. The enzyme-linked immunoabsorbent
assay (ELISA) technique was used to identify *S. aureus* enterotoxins; 45.6% of the samples
produced an enterotoxin. Cow milk produced enterotoxin types A, B, and D; buffalo milk
produced enterotoxin types A and D; sheep milk produced enterotoxin types A, and C; goat milk

produced enterotoxin type D. They concluded that there is a high potential risk of
staphylococcal food poisoning from drinking raw milk, especially if hygiene practices are not
followed.

728

729 Of additional importance to the dairy industry are the many spoilage microorganisms, which 730 compromise the organoleptic, nutritious, and biochemical characteristics. Because of their 731 widespread occurrence and capability to grow in pipelines, bulk tanks and milking machines in 732 dairy processing plants, Scatamburlo et al. (2015) characterized the proteolytic activity of 733 Pseudomonas spp. isolated from 61 Brazilian goat milk samples from 12 farms. Pseudomonas 734 spp. were confirmed by Polymerase Chain Reaction (PCR) by using a genus-specific region of the 735 16s DNA. Mean Pseudomonas spp. counts ranged from 3.0 to 4.8 log CFU/mL, a smaller range 736 than previously reported for cow milk (1.0 to 6.6 log CFU/mL) (De Oliviera et al. 2006). The 737 high proportion of proteolytic *Pseudomonas* spp. found in the analysis is indicative of a need for 738 greater attention to sanitation along the production and processing chain.

739

740 Goat milk cheeses (n = 75) were among 273 cheeses included in a landmark piece of work 741 presented by Trmčić et al. (2016), where it was proposed that coliform testing no longer be 742 used to assess the safety of cheese. For decades, coliform bacteria have been used as indicator 743 organisms for assessment of unsanitary conditions of manufacture. In fact, coliform testing in 744 pasteurized milk was recommended by the US Public Health Service in the earliest edition of 745 the Grade "A" Pasteurized Milk Ordinance (PMO) published in 1924 (Martin et al. 2016) and the 746 current tolerance limit for coliforms in milk is no more than 10 CFU/mL (HHS, PHS, FDA, 2015). 747 However, coliforms are a diverse group, and some research has shown that only a fraction have 748 been identified as factors of fecal contamination and a wider fraction is environmental (Martin 749 et al. 2016). Milk pasteurization, low pH, low water activity and other cheese characteristics 750 were found to significantly contribute to lower prevalence of coliforms in cheese (Trmčić et al. 751 2016). Water activity was the only factor identified as determining the concentration at which 752 coliforms are present in cheese. Although the prevalence of coliforms does not significantly 753 differ in milk of cows, goats and sheep (D'Amico et al. 2008; D'Amico and Donnelly 2010),

cheese manufactured from goat milk showed a higher risk of coliform detection (Trmčić et al.
2016). The authors proposed several reasons for higher coliform contamination or outgrowth:
1) intrinsic factors in goat milk and/or goat cheese (not including pH or water activity), 2)
procedures involved in making goat cheese (not including pasteurization and cheese rind
treatment), and/or 3) goat cheese producers may represent small facilities with reduced
resources related to food safety (Trmčić et al. 2016).

760

761 International advances in goat milk products research

762 The value of goats was realized in many countries well before the US recognized their 763 importance, and Haenlein (1980) documented some of the international symposia and 764 references in his 1980 manuscript published in JDS. At the time, few international works 765 related to dairy goats had been published in JDS. Since then, the international importance of 766 goat milk and milk products has not been overlooked in JDS publications. Fundamental goat 767 milk research, conducted by international authors, has been key to advancing dairy goat 768 science. Additionally, JDS has been a reservoir for international work of regional interest, 769 including works documenting chemical composition and nutritive value of the milk of native 770 goat breeds in Saudi Arabia (Sawaya et al. 1984) and regional goat cheeses in Italy, Turkey, 771 Spain and Portugal (Fontecha et al. 1990; Freitas and Malcata 2000; Martínez-Hernández and 772 Juárez 1989; Martínez-Hernández et al. 1992; Trani et al. 2016; Yuceer et al. 2009).

773

774 Goat milk has not traditionally been used extensively for production of mozzarella or other 775 pasta filata cheeses because the stretching process is not always successful. Italian 776 investigators, Imm and colleagues (2003), were the first to investigate functionality and physic-777 chemical characteristics of goat milk mozzarella. Batches of cow or goat milk were 778 standardized to 3.2% fat and pasteurized, then made into low moisture, part-skim mozzarella 779 cheese using standard procedures (Kosikowski and Mistry 1999). No difference was noted in 780 meltability between caprine and bovine mozzarella. The free oil was always lower in goat milk 781 mozzarella, which was attributed to intrinsic differences in goat cheese fat and protein matrix. 782 Although it improved by aging the cheese 3 to 4 weeks, it decreased in subsequent weeks of

783 storage. Authors confirmed that structural degradation by proteolysis weakened gel matrix and 784 improved melting characteristics; proteolysis occurred more quickly in bovine than caprine 785 cheeses. The authors recommended additional research to better understand the contribution 786 of fat globule size, polymorphic structure and casein micelle structure on melting properties. 787 Recently, Niro et al. (2014) partially substituted cow milk with goat or sheep milks to produce 788 acceptable Caciocavallo cheese. From a sensory standpoint, cow Caciocavallo cheeses were 789 characterized by higher scores for sweetness, elasticity, adhesiveness and humidity (moisture). 790 Mixed cow/sheep cheeses had higher scores for intensity of flavor, acidic, astringent, friability 791 and salty attributes. Mixed cow/goat cheeses solubility (fast melt in mouth), intensity of 792 aroma, and bitter attributes predominated (Niro et al. 2014). Faccia et al. (2015) reported 793 satisfactory production of Fior di latte cheeses from sheep and goat milk after methodological 794 modifications from the standard cow Fior di latte process.

795

796 In Ethiopia, a research project was initiated in 2007 to address the increasing demand for goat 797 milk cheese (Mestawet et al. 2013). The goat breeds, adapted to the climatic conditions of 798 Ethiopia, had high casein content and good milk coagulating properties. The new mutation in 799 the α_{s1} -casein gene that they identified, which yielded milk with very high α_{s1} -casein, may 800 provide opportunities for genetic improvements (Mestawet et al. 2013). Because of interest in 801 intensive dairy goat production and value-added goat milk products in Brazil, Fonseca et al. 802 (2013) evaluated the influence of lipolytic bacteria in raw goat milk upon goat milk powder 803 during storage. Although lipolytic psychrotrophs did not increase during 5 d of raw milk 804 storage, and psychrotrophs were killed by pasteurization, peroxide value, C_8 and C_{10} FA 805 concentrations and total FFA content of milk powder increased and rancid flavors increased 806 during 180 d of powder storage. Authors concluded that raw goat milk intended for powder 807 should be processed within 3 days of collection (Fonseca et al. 2013).

808

809 In Norway, as consumption of the brown whey cheese (Brunost) has decreased, the interest in

810 rennet- and acid-coagulated cheeses has increased (Inglingstad et al. 2014). As a result, the

811 need for higher quality goat milk has increased in recent years. Inglingstad et al. (2014)

812 examined the effect of two different types of pasture (cultivated and rangeland) and two 813 different hay qualities (high and low quality) on goat milk composition and renneting 814 properties. Milk from pastured goats was superior (higher casein and fat) to those on hay, and 815 goats on cultivated pasture had the highest yield; cultivated pasture yielded milk with higher α_{s1} -casein and κ -casein (better renneting properties) compared to other treatments 816 817 (Inglingstad et al. 2014). Providing additional support to the findings, Revilla et al. (2016) 818 analyzed the antioxidant capacity of 224 cheese samples in Spain, prepared using mixtures of 819 milk from cows, sheep and goats, in two manufacturing seasons (winter and summer) and over 820 6 months of ripening. Although animal species was not a significant factor correlated with total 821 antioxidant capacity, statistically significant correlations were found between total antioxidant 822 capacity and season of manufacturing (higher antioxidant activity in summer cheeses), time of 823 ripening, retinol, % fat, % protein, and some minerals (K, Mg, Na, and P) (Revilla et al. 2016).

- 824
- 825 SUMMARY AND FUTURE DIRECTIONS
- 826

827 One of the earliest domesticated animals in the world, goats will always be an important part of 828 human culture. Their compact size (compared to cows) makes them appealing from a herd 829 management and milking standpoint. Additionally, physiological differences render unique 830 physical characteristics to goat milk in terms of flavor profile, fat globule size, coagulation 831 properties and allergenicity, making goat milk the dairy product of choice for many consumers. 832 Economic demand for goat milk and milk products, based on the differences between goat and 833 cow milk and milk products has advanced methods to detect adulteration across the globe. It is 834 expected that such work will continue.

835

Although the track record of safety of goat milk and milk products is good, research must be continued to ensure the safety and quality of these products, particularly with the emergence of new foodborne pathogens and spoilage microorganisms. When Casper et al. (1998) first looked at seasonal changes in goat whey, and promoted food industry applications of goat whey, sustainability was not yet a buzz word. But now, and into the future, additional research

- 841 into methods to sustainably feed goats, responsibly improve productivity, ecologically manage
- 842 effluents, and creatively utilize goat whey will be essential to responsibly manage the global
- 843 dairy industry.
- 844

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1411	Goats' milk quackery. J. Paediatr. Child Health. 41: 569-571.

- 1412 Table 1. Analysis of 355 individual goat milk samples from 21 herds, collected across 16 months
- 1413 (adapted from Lythgoe (1940)).

Month	% Total solids	% Fat	% Solids nonfat	% Lactose	% Proteins	% Ash	Protein- fat ratio
December and January	14.5	5.08	9.42	4.78	3.99	0.84	0.78
February	14.56	5.13	9.43	4.87	3.97	0.85	0.78
March	14.08	4.80	9.28	5.03	3.74	0.76	0.80
May, June and July	12.24	3.79	8.45	4.66	3.34	0.77	0.86
August	11.44	3.37	8.07	4.32	2.99	0.78	0.89
September	12.29	3.98	8.31	4.49	3.16	0.79	0.82

1415

1416 Table 2. Average composition of milk of various mammals (adapted from Roadhouse and

1417 Henderson, 1950).

Species	% Total solids	% Fat	% Lactose	% Protein	% Ash
Human	12.57	3.70	6.98	1.63	0.21
Cow	13.10	4.00	4.90	3.50	0.70
Goat	12.86	4.09	4.20	3.71	0.78
Camel	12.39	5.40	3.30	3.00	0.70
Ewe	16.43	6.18	4.17	5.15	0.93

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1420 Table 3. Physicochemical properties of commingled goat milk (adapted from Guo et al. 2001).

	0		
	Ν	$X \pm SD$	Range
Fat (%)	50	$\textbf{3.61} \pm \textbf{0.47}$	3.00-4.40
Lactose (%)	50	4.47 ± 0.15	4.13 – 4.73
Crude protein (%)	50	$\textbf{3.47} \pm \textbf{0.21}$	3.19 – 3.86
Casein (%)	50	2.57 ± 0.15	2.34 – 2.86
Non-protein nitrogen (% of crude	49	5.04 ± 0.34	4.40 - 5.65
protein)			
Total solids (%)	50	12.38 ± 0.71	11.17 – 13.44
Ash (%)	50	0.82 ± 0.04	0.79 – 0.89
Calcium (%)	50	0.15 ± 0.01	0.12 - 0.17
Phosphorus (mg/kg)	50	$\textbf{0.13}\pm\textbf{0.02}$	0.10 - 0.16
Sodium (mg/kg)	49	672 ± 125	380 – 977
Magnesium (mg/kg)	49	160 ± 24	100 – 217
Zinc (mg/kg)	49	$\textbf{4.59} \pm \textbf{1.93}$	1.30 - 9.50
Specific gravity	50	1.0235 ± 0.0007	1.0224 - 1.0262

1422	Table 4.	Fatty acid composition (mole %) of milk fat triglycerides of five species, up to C _{20:0}	

		-		-	
1423	(adapted	from	Kuksis ((1967)).

Fatty acid	Human	Jersey cow	Holstein cow	Goat	Sheep
4:0	-	9.8	8.5	8.2	10.3
6:0	-	5.0	2.9	6.9	3.4
8:0	-	2.4	1.4	5.8	2.3
10:0	0.6	4.8	2.3	7.9	3.4
12:0	3.0	4.1	2.1	1.9	1.8
14:0	5.3	11.8	7.5	2.6	5.0
15:0	0.6	1.7	1.2	0.7	0.9
16:0	26.5	36.5	28.0	16.0	20.9
16:1	4.0	1.1	1.6	1.2	1.2
16:2	-	-	-	-	-
17:0	1.1	0.8	0.7	2.4	2.9
18:0	7.8	8.6	14.6	14.3	15.5
18:1	37.6	13.0	26.5	30.4	27.2
18:2	10.0	0.4	1.5	1.7	2.9
18:3	0.6	-	-	-	2.4
20:0	-	-	Trace	-	Trace

1425

1426Table 5. Concentration of total fatty acids in colostrum of goats and cows (adapted from Attaie1427et al. 1993).

/			
Fatty acid	Nubian goats ¹	Alpine goats ¹	Holstein cows ²
		Mean (µg/g of fat)⁺	
Butanoic acid	304.51 ^A	202.67 ^A	226.12 ^A
Hexanoic acid	385.66 ^A	239.44 ^B	235.45 ^B
Heptanoic acid	5.31 ^A	4.63 ^A	4.46 ^A
Octanoic acid	520.68 ^A	297.80 ^B	162.28 ^B
4-Ethyloctanoic	13.66 ^A	12.52 ^A	10.46 ^A
acid			
Decanoic acid	1513.70 ^A	766.99 ^B	256.10 ^c
9-Decenoic acid	36.22 ^A	18.34 ^B	19.66 ^в
Undecenoic acid	10.07 ^A	7.26 ^A	3.69 ^B
Dodecanoic acid	792.72 ^A	437.79 ^B	302.35 ^B
A D C			

1428 ^{A, B, C}Means in a row with the same superscript are not different (P > 0.05).

1429 ¹Means are average of seven samples with duplicate and triplicate determinations.

1430 ²Means are average of four samples with triplicate determinations.