Cheese whey recycling in the perspective of the circular economy: Modeling processes and the supply chain to design the involvement of the small and medium enterprises

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- 1 Cheese whey recycling in the perspective of the circular economy: modeling processes and the
- 2 supply chain to design the involvement of the small and medium enterprises
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6 Abstract

Rationale: Cheese whey has relevant nutritional, health and functional properties that could address
the dietary needs of a growing world population. Nevertheless, around 50% of whey produced
globally is not reused in the food system. A significant role in cheese production is played by small
and medium enterprises (SMEs) that process typical products labelled with a protected designation
of origin (PDO), representing both the cultural heritage and the major economical resource of specific
geographical regions.

Scope and Approach: This study is aimed at identifying the potential involvement of SMEs in the recycling of whey for food uses. Techno-economic analyses of whey recycling processes were reviewed to define efficient configurations and plant size. Safety and quality specifications for whey were then examined as well as traceability systems designed for improving transparency among the actors and stakeholders of the dairy supply chain.

Key Findings and Conclusion: The study led to conclude that a whey value chain could be built by various cheese SMEs and a stand-alone industrial unit for the downstream processing of whey into value-added foods. In this scenario, advantages could derive from the involvement of SMEs that belong to the same PDO cheese consortium, in terms of standardization of whey, third party certification and possible use of smart traceability tools. Hence, the SMEs producing PDO products could advance the application of the bioeconomy principles in the food system. Research challenges to promote the implementation of the conceived whey value chain are finally outlined. Keywords: cheese whey, traceability, Protected Designation of Origin, Internet of Things,
Blockchain technology, industrial symbiosis

27

28 1. Introduction

The concept of the bioeconomy has promoted the rethinking and redesign of food supply chains. In this context, food waste management has been identified as one potential solution for the transition towards the implementation of circular supply chains in the food system (Jurgilevich et al., 2016). Among various waste streams generated by the food systems, the materials that are diverted from the food chain but are still edible (or can be processed to become edible) are considered as the most relevant resources as they can be converted into new food ingredients and foods, thus addressing the dietary needs of a growing world population (Lavelli, 2021).

The cheese industry produces about 145 million tons of whey annually and only 79 million tons (i.e., 54%) are re-used in the food system, while 6 million tons are destined to non-food uses and 60 million tons are used as feed, fertilizer or waste (Ganju & Gogate, 2017).

Whey composition differs between sweet whey (pH 6.2 - 6.4) and acid whey (pH 4.6 - 5.0) and 39 typically comprises: 93.0 - 93-7% of water, 4.4 - 5.2% of lactose, traces or up to 0.8% of lactic acid, 40 41 0.6 - 1% of protein, 0.5% - 0.8% of minerals (Table 1, Jelen, Roginski, Fuquay, & Fox, 2003). Whey is highly pollutant, since its chemical oxygen demand (COD) can vary from 50,000 to 80,000 mg/L, 42 whereas its biochemical oxygen demand (BOD) is in the range from 40,000 to 60,000 mg/L 43 (Chatzipaschali, & Stamatis, 2012). A number of recycling processes have been proposed to obtain 44 value added products from whey for the food and non-food sectors (Ganju & Gogate, 2017; Lappa et 45 46 al., 2019; Rocha, & Guerra., 2020; Zandona, Blažić, & Jambrak 2021; Barba, 2021). Focusing on food uses, recycling processes comprise re-use of whey in cheese factories to produce other dairy 47 products, recycling to produce non-conventional products such as unfermented and fermented whey-48 49 derived beverages (Mehra, R.; Kumar, H.; Kumar, N.; Ranvir S., Jana, A. et al., 2021) or separation

of the lactose and protein fractions and distinct valorization. Lactose upgrading technologies include 50 enzymatic or microbial production of target molecules such as prebiotic compounds (galacto-51 oligosaccharides, lactulose, lactitol and lactobionic acid) and carotenoids or bioconversion in 52 biomasses of mushrooms or microalgae. Protein upgrading technologies include use as source of 53 54 essential amino acids, use as a functional ingredient (gelling agent, foaming agent and emulsifying agent) and enzymatic hydrolysis to obtain bioactive peptides (Lappa et al., 2019; Rocha, & Guerra., 55 2020; Zandona et al., 2021). Alternatively, lactose can also be reintroduced in the food system upon 56 conversion via fermentation in bacterial cellulose and fatty acid esters, which are used as structural 57 agents, such as components of edible film. Whey proteins can also be reintroduced in the food system 58 59 after chemical and physical modification as structural agents or delivery systems to protect bioactive 60 compounds (Lappa et al., 2019; Lavelli, & Sereikaite, 2022a; Lavelli, & Sereikaite, 2022b).

lactose prices 61 Whey, whey protein and are constantly monitored (https://www.clal.it/en/index.php?section=grafici_siero#wo) indicating a general attention of the 62 food companies towards these multifunctional ingredients. However, despite multiple options to 63 convert whey into value added foods, the implementation of recycling processes remains a challenge 64 as proven by low percentage of whey recovered in the food system. It is worth noticing that, within 65 the European Community approximately 200 cheese products have a Protected Designation of Origin 66 67 (PDO), i.e., they originate in a specific geographical area and all their production steps take place in the same geographical area. The PDO foods are processed at SMEs according to a specific and well-68 defined process and their conformity to the procedural document of the PDO is assessed by both a 69 70 consortium of producers and a third party certification body (Reg. CE 1151/2012). The PDO foods represent an important part of the culture, history, identity, heritage and local economy of a region or 71 72 country (Albuquerque, Oliveira, & Costa, 2018; Mattas et al., 2022).

The aim of this review was to identify the potential roles of SMEs in the dairy sector in supplying whey for further processing in the food system. First, a literature study was performed to define the unit operations involved in liquid whey processing and to draw the process configurations that have

been modeled and validated at various industrial scales (from 100 to 3000 t of whey inflow/d).
Subsequently, the literature study was directed to identify the safety and quality specifications of
whey needed to address quality requirements for value-added food applications. Then, the traceability
systems implemented in a cheesemaking process and in the entire dairy supply chain were reviewed,
which served as a basis to propose a framework for a transparent and trustable whey value chain and
to suggest future research challenges.

82 2. Unit operations and model processes for whey recycling in the food system

83 2.1. Overview of the main unit operations for whey recycling in the food system

One relevant feature of whey is its susceptibility to microbial growth due to the presence of viable 84 85 starter microorganisms added during fermentation and possible microbial contaminants. Therefore, at industrial level pasteurization of whey is applied in the first steps of recycling (USDEC, 2006). 86 Pasteurization is commonly performed at 75 °C for 15 s and can be combined with heat regeneration 87 of about 90% to 95%, which makes an average energy consumption of around 0.789 MJ/kg product 88 89 (as primary energy) (Ramirez et al., 2006). To further decrease the energy input, non-thermal 90 technologies for pasteurization have been proposed and studied at laboratory scale. There has been 91 considerable interest in the potential use of high-power/high-intensity ultrasounds, often in conjunction with mild heating (thermosonication), for the inactivation of microorganisms and 92 93 enzymes in milk (D'Incecco, Limbo, Hogenboom, Pellegrino, 2021). An application was also designed for whey treatment, including heating at 55 °C combined with ultrasound treatment at 480 94 95 W with a 20 kHz ultrasonic probe for 10 min, which led to approximately 3 log10 cycles reduction in microbial load (Barukcic, Jakopovic, Herceg, Karlovic, & Bozanic, 2015). Pulsed electric field 96 97 (PEF) treatments, generally combined with mild heating have also been explored for milk 98 pasteurization and found to achieve about 3 log10 cycles of microbial reduction (Alirezalu, 99 Munekata, Parniakov, Barba, Witt, et al. 2020). PEF treatment was specifically studied for microbial inactivation in a whey product, using a chamber with E set at 32 kV/cm, pulse width set at 3 µs, 100

residence time at 1.8 s., inlet temperature in the range 20 - 40 °C and outlet temperature 58 °C. As a 101 102 result, inactivation of up to 6.51 log10 cycles of the target microorganism Lysteria innocua was achieved (Schottroff, Gratz, Krottenthaler, Johnson, Bedard, & Jaeger, 2019). High hydrostatic 103 pressure (HPP) treatments were investigated for inactivation of microorganisms in milk and it has 104 been suggested that application of 600 MPa for 5 min is adequate to reduce the number of log10 105 cycles by 5-7, resulting in products comparable to traditionally pasteurized ones (Serna-Hernandez, 106 107 Escobedo-Avellaneda, García-García, Rostro-Alanis, & Welti-Chanes, 2021). However, specific information on HPP application for whey pasteurization is lacking. In general, research is still 108 necessary to efficiently replace heat pasteurization oh whey with non-thermal technologies. 109

110 Another relevant feature of whey composition is its high water content, hence evaporation and drying technologies are applied at industrial level. Drying is performed by spray-driers (USDEC, 2006). The 111 impact of the spray drier system on energy consumption is in the range 4.9 - 3.4 MJ/kg water 112 evaporated (as primary energy), where the highest consumption is for one stage system and the lowest 113 for a tree stages system. Commonly, evaporation is applied prior to spray-drying to remove water 114 with less energy input, which decreases in the range 0.9 - 0.2 MJ/kg water evaporated (as primary 115 energy), with increasing the number of stages from one to seven for evaporators with a thermal vapor 116 compressor, and decreases to 0.050 MJ/kg water evaporated (as primary energy) for a single stage 117 118 evaporator with a mechanical vapor compressor (Ramirez et al., 2006). Pre-concentration of whey is performed by membrane processes such as reverse osmosis (RO), in which only water can permeate 119 through the semi-permeable membrane, upon application of pressure to overcome the osmotic 120 121 pressure. Membrane processes also include forward osmosis (FO) where a concentrated draw solution (2 M NaCl) is placed on the other side of the RO membrane to drive the transport of water molecules 122 from whey and membrane distillation (MD), which is performed at 50 °C to selectively transport 123 vapor molecules through the semi-permeable membrane due to surface tension forces. Overall, 124 membrane filtration is efficient for water removal, with energy demand of 0.014 - 0.036 MJ/kg water 125 126 removed (as primary energy). Nonetheless, membrane concentration can only reach a maximum dry

weight of 12–20% due to the increasing viscosity of the retentate with increasing solid content(Ramirez et al., 2006).

Lactose, fat and protein that are the main solid components of whey are sometimes separated in 129 industrial plants by membrane technologies to implement distinct applications. Microfiltration (MF) 130 is applied because it allows the passage of small soluble proteins, peptides, lactose, minerals, non-131 protein nitrogen components and water, while fat globules are retained and can be reused in cream 132 133 (USDEC, 2006). Moreover, ultrafiltration membranes (UF) are applied to separate protein in the retentate from lactose in the permeate. To further improve the separation of protein and lactose, 134 diafiltration (DF) is performed, in which UF membranes are used and water is added to the retentate 135 136 (USDEC, 2006). In the UF process, fouling limits the efficacy of the separation. Approaches were studied to increase the efficacy of UF, including the application of electric fields or ultrasounds for 137 membrane cleaning. In one study using model solutions, electric fields were applied and the best 138 operating conditions found included the use of 5 mM NaCl and an electric potential of 30 V 139 (Corbaton-Baguena, Alvarez-Blanco, Vincent-Vela, Ortega-Navarro, Perez-Herranz, 2016). Use of 140 ultrasounds was studied and positive effects were found upon using a power of 30 W/L cleaning 141 solution, a frequency of 20 kHz for 30 min (Lujan-Facundo, Mendoza-Roca, Cuartas-Uribe, Alvarez-142 Blanco, 2017). However, these approaches were applied at pilot scale only. Upon UF, whey protein 143 144 fraction is spray-dried while lactose is generally produced by concentrating the UF permeate at 65 °C by evaporation followed by a slow cooling process to 20 - 25 °C to achieve supersaturation, where 145 lactose is crystallized. As for the other evaporation steps, lactose concentration is highly energy 146 147 demanding (Ramirez et al., 2006). To increase the efficiency of the process alternative approaches have been proposed at laboratory scale. In one study, the application of ultrasounds was investigated 148 and improved efficiency was observed using a power of 1.33×10^{-2} W/mL of whey at temperature 149 of 7 °C for 20 min, followed by immediate addition of the ethanol (95%, v/v) as antisolvent (Bund, 150 & Pandit, 2007). An alternative process was proposed in which subzero temperatures were reached, 151

where eutectic freeze crystallization occurred (Halfwerk, Yntema, Van Spronsen, & Van der Padt,2021).

Whey composition is also characterized by high amount of minerals, especially sodium, that is added 154 to the cheese curd to lower the water activity and remains in significant amounts in whey, thus limiting 155 its applications as food ingredient. Beside demineralization with membrane-based processes, ion 156 exchange resins are used, with an average consumption of 0.128 MJ/kg of NF whey retentate (electric 157 power) (Greiter et al., 2002). Electrodialysis was found to be more efficient than ion exchange resins, 158 with an energy demand of 0.0459 MJ/kg of NF whey retentate (electric power) (Greiter et al., 2002). 159 Similarly, an energy input of 0.058 MJ/kg of UF whey permeate (electric power) was also found to 160 161 be required for demineralization (Chen et al., 2020). Eutectic freeze concentration is an emerging technology that operates at the eutectic temperature of the UF whey permeate (-24 °C) to enable 162 simultaneous separation of ice and salt (Chen et al., 2020). The energy input for eutectic desalination 163 is 0.288 MJ/kg feed for cooling at the eutectic point and 0.450 - 0.400 MJ/kg feed (electric power) 164 for freeze crystallization, but the process was not studied at industrial scale (Chen et al., 2020). 165

In general, a critical aspect of the major unit operations involved in whey processing is the elevate 166 energy demand. Moreover, the high hygienic standards require cleaning-in-place operations, which 167 need an average energy input of 0.1 - 28 MJ/cleaning cycle (primary energy) (Ramırez et al., 2006). 168 Finally, energy input for cold storage is on average 0.59 MJ/m³ × d⁻¹ (Evans et al., 2015). High energy 169 demand is characteristic for the dairy sector as compared to the other food sectors (Ladha-Saburet al., 170 2019). For various dairy products, the energy demand was mapped from farm to fork (Malliaroudaki, 171 172 Watson, Ferrari, Nchari, & Gomes, 2022), while among various alternatives for whey processing drying to powder was only considered. This led to identify an average energy input of 6.5 MJ/kg of 173 whey, which is higher than the average input needed by its generating process, i.e., 3.3 MJ/kg of 174 cheese (Malliaroudaki et al., 2022). 175

176 2.2 Model processes for whey recycling in the food system

Considering the technological equipment and high energy input necessary for whey recycling, a 177 techno-economic analysis is necessary to define the plant size for the implementation of a sustainable 178 process. The simplest process for whey recycling, i.e., drying to obtain whey powder (WP) was 179 modeled for two cheese plants that operate with a milk inflow of 2000 - 3000 and 100 t/d, respectively 180 (Peters, 2005). As a result, the relevant impact of the plant size on process efficiency was evidenced, 181 since the process was validated for the highest plant while the feasibility of whey recycling in small 182 183 cheese-making companies remained uncertain (Peters, 2005). Accordingly, other studies have pointed that the relatively low amount of cheese whey produced by the SMEs is one of the main 184 barrier to the reutilization of this valuable byproduct (Rocha et al., 2020; Chalermthai et al., 2020; 185 186 Lindsay et al., 2020). One interesting approach to increase process efficiency is the integrated recovery of both WP and water (Aydiner et al., 2014). Two technological configurations were 187 validated with a whey inflow of $100 \text{ m}^3/\text{d}$. In the first configuration, whey was submitted to FO, using 188 189 a concentrated draw solution made with 2 M NaCl. The resulting retentate was concentrated and spray-dried to obtain WP. The permeate was submitted to MD at 50 °C. In the second configuration, 190 MD was used as a first stage and the permeate was submitted to RO. In both cases, 66% of water was 191 recovered (Aydiner et al., 2014) (Figure 1). Another configuration was validated to produce WP with 192 water recovery, but the plant size was 1500 t/d. In this latter process, whey was filtered to remove 193 194 fines (vacuum filter), pasteurized (plate heat exchanger, 72 °C for 15 s), centrifuged to remove 90% of fats and then submitted to RO to recover water and the retentate, which was then evaporated (six-195 stage evaporator, final temperature 45 °C) and spray-dried to obtain WP (Chamberland, Scott Benoit, 196 197 Doyen, & Pouliot, 2020). The WPs so far obtained contained lactose as the main component, at concentration of 53 - 75% and proteins in the range 11 - 15% (Table 1). 198

UF membranes represents the standard tool for production of whey protein concentrates (WPCs)
(USDEC, 2006). Integrated processes that recover both WPC and lactose are emerging, with respect
to the recovery of protein alone. A techno-economic analysis was performed for processing 700 m³/d
of whey, by comparing: a) the production of whey protein concentrates at 34 or 80 %, respectively

(WPC 34 and WPC 80); b) the production of WPC 34 or WPC 80 in combination with lactose; and 203 204 c) the production of WPC 34 or WPC 80 with ethanol obtained via lactose fermentation. The integrated production of WPC 80 and lactose was found to be the optimal solution (da Silva et al., 205 2015). According to the best proposed process, whey was filtered to remove fines (vacuum filter), 206 pasteurized (plate heat exchanger, 72 °C for 15 s), centrifuged to remove 90% of fats and then 207 submitted to UF and DF. The retentate rich in protein, with 21% solids was spray-dried (final 208 209 temperature of the powder 70 °C) to obtain WPC 80 with 96% solids (Table 1). The permeate was firstly concentrated by RO to 18% solids, evaporated to 50% solids (triple effect evaporator, final 210 temperature 55 °C) and then lactose was crystallized (at 15 °C) and spray-dried to a moisture content 211 212 of 4% (final temperature of the powder 70 °C) (da Silva et al., 2015) (Figure 2). A similar configuration was further analyzed in combination with enzymatic conversion of lactose to galacto-213 oligosaccharides by β -galactosidase and the process was found to be feasible for whey inflow of > 80 214 215 m³/h (Scott et al., 2016). An integrated whey recovery process to obtain both WPC and glucose and fructose syrup was also found to be economically feasible with a whey inflow of 500 m³/d. In this 216 latter process, the technological configuration for WPC obtainment was similar to that reported 217 previously (da Silva et al., 2015) but an evaporation step for the retentate (single stage, 80 °C) to 65% 218 of solids (single effect, 80 °C) was included before spray drying (Gomes et al., 2020) and a final 219 220 protein concentration of 50% was obtained (Table 1). The permeate stream was filtered using sodium bentonite in the filtration bed and submitted to the ion exchange to remove > 95% of cations in order 221 to avoid inhibition of the enzymatic process with immobilized lactase and glucose isomerase. Then, 222 223 the resulting liquid stream was bleached with activated carbon packed in columns and concentrated through vacuum evaporation up to a minimum of 70% total solids to obtain the sugar syrup (Gomes 224 et al., 2020) (Figure 2). Alternatively, a plant was designed to treat 100 m³/d of whey permeate from 225 UF whey, for further production of food grade lactic acid via fermentation by Lactobacillus 226 helveticus, followed by UF, ionic exchange, RO and evaporation (Gonzalez, Ivarez, Riera, & Ivarez, 227 2007). 228

To obtain a finished product with more than 90% of protein, i.e., whey protein isolate (WPI), two 229 230 main processes are used. In one case pasteurized whey is submitted to MF to remove fat and then to extensive DF to remove lactose and salts. Alternatively, pasteurized whey is demineralized with ion 231 exchange resins and then submitted to UF. In both processes, the resulting final retentate is 232 concentrated and spray-dried to obtain WPI (USDEC, 2006). Most of non-protein components of 233 whey are not present in WPI (Table 1). These processes are applied to large size plants, which operate 234 235 with 2000 – 3000 t whey/d (Peters, 2005). An alternative process operating with a whey inflow of $500 \text{ m}^3/\text{d}$ was proposed to recover whey protein without lactose and salts, including: heat treatment 236 at 90°C for 30 min to achieve protein denaturation, followed by precipitation with 237 238 carboxymethylcellulose (CMP) at 50°C, filtration and drying in a fluidized bed dryer to obtain whey protein precipitate (WPP). The permeate was concentrated in a multiple effect evaporator, then 239 lactose was crystallized and dried in a fluidized bed drier (Raout et al., 2022) (Figure 3). According 240 241 to this latter process, a complex of the denatured whey proteins and CMC was obtained, but the exact composition was not reported (Raout et al., 2022). 242

Value-added applications could be applied to WPC, WPI and lactose, such as the separation of whey
proteins, production of bioactive peptides from whey protein (Brandelli, Daroit, & Folmer Corrêa,
2015), improvement of whey protein functionality by physical and chemical treatments (Foegeding,
Davis, Doucet, & McGuffey, 2002; Doost, Nasrabadi, Wu, Ayun, & Van der Meeren, 2019) and
fermentation of lactose to produce value–added compounds or biomasses (Lappa et al., 2019; Rocha,
& Guerra., 2020; Zandona et al., 2021; Barba et al., 2021). However, these upgrading processes were
not modeled to define the optimal plant size.

250 **3.** Safety and quality specifications for whey recycling in the food system

251 *3.1 Safety specification for whey*

The development of food applications from whey recovery imposes to meet safety requirements. Five hazard categories have been identified in the literature, namely mycotoxins, antibiotics, pathogenic microorganisms, heavy metals and organic environmental pollutants.

Considering mycotoxins, aflatoxin M1 exhibits a high genotoxic activity and it has been classified by 255 256 IARC as a class 2B human carcinogen (IARC, 2002). Aflatoxin M1 derives from aflatoxin B1 that can be present in feed. Upon ingestion of contaminated feed by ruminants, aflatoxin B1 is converted 257 in secondary metabolites, among which aflatoxin M1 is the major oxidation metabolite, primarily 258 excreted in the urine and secondarily in the milk (Prandini et al., 2009). In one study, the carry-over 259 of aflatoxin M1 in cheese and whey was investigated, indicating that on average 55% of aflatoxin M1 260 261 is recovered in whey (Costamanaga et al., 2019). The carry-over of aflatoxin M1 from milk to whey deserves attention. In fact, even in case that this contaminant is below the limits fixed by the EU 262 regulation (Reg EC 1881/2006), whey drying causes an increase in its concentration. 263

264 Antibiotic residues in foods can lead to allergic reactions and to or transient disturbances in the microbiota (Oliver, Murinda, & Jarayao, 2011). Antibiotics used for the treatment of infectious 265 disease of livestock are found in milk (Lupton, Shappell, Shelver, & Hakk, 2018). In one study, goat 266 267 milk spiked with 18 antibiotics and then coagulated with rennet, the whey was separated and the transfer of antibiotics was investigated. The percent of antibiotic recovered in whey was found to 268 depend on their hydrophilicity and ability to interact with Ca²⁺ ions, varying from 100% for 269 amoxicillin, ampillicin, sulphadiazine to 90 - 60% for benzilpenicillin, cloxacillin, cefacetrile, 270 271 cefquinome, ceftiofur, erythromycin, while the other cephalosporins, aminoglycosides, macrolides, quinolones, sulphonamides and tetracyclines were not recovered in whey (Giraldo, Althaus, Beltran, 272 & Molina, 2017). A following study confirmed that some β -lactames (cephalexin, benzylpenicillin, 273 ampicillin and cloxacillin) were transferred less into the cheese curd than into the whey (Lányi et al., 274 275 2022). As for aflatoxin M1, the carry-over of antibiotic compounds from milk to whey deserves attention, even if they are present at limits below the threshold fixed by the Regulation (Reg EU 276 277 37/2010), due to their concentration by drying.

Pathogenic microorganisms that can be present in milk, are also of concerns for their possible
presence in whey, most likely due to inadequate hygienic practices (Wedel, Atamer, Dettling,
Wenning, Scherer, et al., 2022). *Salmonella* spp. and *Cronobacter* spp. do not survive after thermal

treatment. Afterward, post-heating handling can result in cross-contamination. Salmonella spp. was 281 282 found to be able to survive on dairy plant surface for at least 30 d (Margas, Meneses, Conde-Petit, & Dodd, Holaha, 2014). Similarly, Cronobacter spp. was isolated from dairy plants (Kent, Fitzgerald, 283 Hill, Stanton, & Ross, 2015). Listeria monocytogenes is a food-borne pathogen of great concern for 284 the dairy industry. It is also inactivated during heating at the high temperatures (> 80 - 95 °C). Then, 285 the psychrotrophic pathogen L. monocytogenes may outgrow in the absence or low presence of a 286 protective background microbiota (Sameli, & Samelis, 2021). Bacillus cereus spores demonstrated 287 the ability to survive to pasteurization. Evaporation, which is another thermal process applied in whey 288 processing, is also unable to inactivate the total load of mesophilic and thermophilic spore formers; 289 290 instead this treatment provides an environment where the spores can undergo outgrowth (Wedel et 291 al., 2022). Spray-drying effect on B. cereus depends on the strain, since Log-reductions in the range 1 – 4.7 were observed (Alvarenga, Brancini, Silva, da Pia, Campagnollo et al., 2018). Moreover, B. 292 cereus is able to form biofilm that facilitates spore dispersion, subsequent germination, multiplication 293 and enterotoxins production (Tirloni et al., 2017). Staphylococcus aureus is one of the major 294 etiological agents of mastitis in dairy cattle. These infections are a source of contamination of milk 295 and dairy products since up to 50 - 70% of S. aureus strains are able to produce under suitable 296 conditions extracellular heat stable staphylococcal enterotoxins (SEs) (Bogdanovicova, Necidov, 297 298 Harustiakov, & Janstov 2017). S. aureus is ubiquitous: detected in animals, cowsheds, farm and dairy 299 workers, processing environments and ultimately the dairy products including whey (Mehli, Hoel, Thomassen, Jakobsen, & Karlsen, 2017). 300

Heavy metals such as lead, cadmium, arsenic, chromium and mercury can accumulate in the body after intake for long years in high concentrations and lead to disorders in the heart, liver, kidney, nervous, blood, and lungs (Elgammal, Khorsheda, & Ismail, 2019; Bandara, Towle, & Monnot, 2021). Heavy metals can be found in milk due to intake of contaminated feed. Moreover, ruminant animals are exposed during grazing to intake a small portion from surface layers of soil and hence environmental pollution of the farming area due to industrial activities is also a vehicle for toxic metal

intake (Campos de Aquino et al., 2017; Castro-Gonzales et al., 2018; Elgammal et al., 2019). Casein
have a high affinity for metals, especially lead (Delavar et al., 2012). However, the transfer of heavy
metals from milk to whey can occur. In fact, presence of toxic metals in whey was estimated to be a
risk in case of elevated consumption (Ring, Sheehan, Lehane, & Furey, 2021).

311 Different polyhalogenated-compounds are present in the environment of industrialized areas and show high resistance to metabolism in humans. As a result, their concentrations can increase in tissues 312 thus causing toxic effects such as immunosuppression, decrease in reproductive potential and tumor 313 promotion (Van den Berget al., 2006). Compounds used as flame retardants tetrabromobisphenol A 314 (TBBPA), 2,4',4,5,5'- pentabromodiphenyl ether (BDE-99) and β-hexchlorocyclododecane (β-315 316 HBCD) and the coolant/ plasticizer/ hydraulic fluid/pesticide/flame retardant 2, 3', 4, 4', 5pentachlorobiphenyl (PCB-118), were found to possess high affinity for whey protein and, if present 317 in whey, upon UF are found in the retentate (Shelver, Lupton, Shappell, Smith, & Hakk, 2018). 318

The studies so far reported attest that for cheese-making companies that provide whey for further usein the food system, there is a need to guarantee that safety standards are met.

321 *3.2 Quality specifications for whey*

Multiple factors related to bovine and milk processing conditions, could affect whey composition and hence its performance in the production of value-added compounds, but information in literature is scattered.

One route to produce added value compounds from cheese whey is via lactose fermentation. In this context, bacteriophages infecting lactic acid bacteria are one concern related to the whey quality. In fact, bacteriophages can be present in raw milk, contaminate plant surfaces and generate aerosols, which remain in air for long periods (Pujato, Quiberoni, & Mercanti, 2018). Hence, when whey has to be recycled by lactic bacteria fermentation (Sipola et al., 2002; Pihlanto, 2001; FitzGerald, Murray, & Walsh, 2004), the presence of bacteriophages can cause failure or reduced process yield.

Antibiotics present in whey are another feature related to whey quality since they can decrease the yield of microbial-based processes. *Kluyveromyces marxianus* sensitivity towards antibiotics was

considered since it is an interesting yeast for whey valorization due to its ability to assimilate lactose
directly, without hydrolysis pretreatment (Chandra, Castillo-Zacarias, Delgado, & Parra-Saldívar,
2018). The fermentative capacity of *K. marxianus* was found to be reduced in whey by the presence
of antibiotics closed to the maximum residue limits fixed by the EU regulation (Eluk, Ceruti, Nagel,
& Althaus, 2019).

The yield of enzymatic processes to convert lactose into valuable compounds, such as galactooligosaccharides, lactulose as prebiotics and lactose esters as emulsifiers is affected by the nature of whey (acid, sweet) and by lactose concentration (Guerrero, Vera, Acevedo, & Illanes, 2015; Staron, Dabrowski, Cichon, & Guzik, 2018; Fischer & Kleinschmidt, 2021). Moreover, high sodium content inhibits the enzymatic conversion of lactose to glucose and fructose syrup (Gómez et al., 2020).

The second route to produce added value compounds from cheese whey is the use of protein fraction. 343 Hence, the quality of whey depends on total protein content, amino acid profile, protein digestibility 344 345 and bioavailability of essential amino acids (Brandão da Costa, Rocha Roiffé, da Silva de la Cruz, 2021). Moreover, the use of enzyme technology for the hydrolysis of whey proteins is applied to 346 generate bioactive peptides, which can exert physiological effects in vivo, such as antioxidant, 347 antihypertensive, antidiabetic, immunomodulatory, anticancer, 348 antimicrobial, opioid and hypocholesterolemic activities (Brandelli et al., 2015). 349

350 However, heating intensity applied during processing can affect nutritional quality and bioactivity of whey. In fact, during whey production, storage and processing, changes are induced in protein 351 structure, including unfolding, cross-linking and aggregation (Zhang, Zhou, Zhang, & Zhou, 2021). 352 353 Heating promotes the Maillard reaction between reducing sugars and amino acids, peptides, or proteins, with the main consequence to form the so-called blocked lysine, affecting bioavailability of 354 this essential amino acid. Denaturation temperatures for α -lactalbumin, bovine serum albumin, 355 immunoglobulin and β-lactoglobulin are 62, 64, 72 and 78 °C, while for lactoferrin denaturation 356 temperatures are 71 and 91 °C for the apo- and holo-protein, respectively (Zang et al., 2021). Milk 357 pasteurization at 72 °C for 15 s, does not affect the major whey protein, i.e., β-lactoglobulin. The 358

second major whey protein, i.e., α -lactalbumin, is denatured upon pasteurization but it can refold 359 360 upon cooling (deWit, & Klarenbeek, 1984). Interestingly, milk pasteurization at 72 °C for 15 s does not affect the immunoreactivity of lactoferrin (Navarro, Harouna, Calvo, Perez, & Sanchez, 2015) 361 and immunoglobulin (Riera et al., 2014). However, significant heat damage was observed in 362 commercial whey products. In fact, furosine that is an indicator of blocked lysine was found to be 363 present at levels up to 1125.7 mg/100 g protein (150 mg average) in whey (Rufián-Henares, Delgado-364 365 Andrade, Jiménez-Pérez, & Morales, 2007). Accordingly, in one study, fifty-two commercial whey samples were analyzed and 9% of them presented blocked lysine values exceeding 20%, while 50% 366 of them had over 6% of blocked lysine (Sánchez-Oliver, Contreras-Calderon, Puya-Braza, & Guerra-367 368 Hernández, 2018). Regarding the effect of heat treatments on the digestibility of whey protein, there are many reports but no consensus emerges, since either increased protein digestibility and decreased 369 protein digestibility was observed after processing, which probably depends on the intensity of 370 371 heating conditions (Zhang et al., 2021). Indeed, in fourteen commercial whey products in vitro protein digestibility ranged from 50.4 to 79.6% and different amino acid profiles were observed 372 (Pehlivanoglu, Bardakci, & Yaman, 2022). The changes of whey proteins allergenicity during heat 373 treatment also depend on heating temperature and processing time, resulting in either a decrease or 374 375 an increase (Zhang et al., 2021). As a result, more investigations are necessary to unravel the effect 376 of thermal processes on whey nutritional and health properties. On the other hand, another relevant 377 feature for whey valorization relies on its techno-functional properties, such as foaming, gelling and emulsifying ability, which are of interest in the development and stabilization of particular food 378 379 structures or for encapsulation of bioactive compounds (Lavelli & Serenikate, 2022a; Lavelli & Serenikate, 2022b). Whey techno-functional properties generally improve with increasing the 380 intensity of the thermal treatment (Foegeding et al., 2002). To control and enhance protein 381 functionality, emerging processes are also considered, such as ohmic heating (Ferreira, Machado, 382 Pereira, Vicente, Rodrigues, 2021), high hydrostatic pressure (Serna-Hernandezet al., 2021), and 383 ultrasound treatments (Carrillo-Lopez, Garcia-Galicia, Tirado-Gallegos, Sanchez-Vega, Huerta-384

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Jimenez, et al., 2021). Moreover, the covalent interaction between an amino group of a protein and a 385 carbonyl group of a polysaccharide through Maillard reaction is applied to produce conjugates with 386 improved characteristics such as heat, ionic strength and pH sensitivity (Doost et al., 2019). The 387 conjugation of whey proteins with small molecular weight reducing sugars is another potential 388 strategy to overcome the heat instability, aggregation and fouling (Wu et al., 2021a). Additionally, 389 whey protein-phenolics interactions are also intensively studied to generate phenolic protection, 390 improved delivery and bioavailability (Wu et al., 2021b). The route of protein conjugation will open 391 a new area for whey applications. It can be assumed that protein conjugation processes also pose 392 some quality requirements for the whey protein source, i.e. for instance protein purity, while a detailed 393 394 information on this aspect is lacking.

In general, a deeper insight into the relationship between whey composition and functionality would support the development of target applications. To this aim, the application of foodomics tools is a promising approach to pave the road for the exploitation of whey streams (Tsermoula, Khakimov, Nielsen, & Engelsen, 2012). Nevertheless, even if advancing knowledge is advisable, specification of at least some basic quality parameters of whey would increase the range of possible applications in the food system.

401 **4.** Traceability systems for whey and for a smart whey supply chain

402 *4.1 Whey traceability*

As discussed above, efficient processing of whey requires an inflow of 100 - 700 t/d, which is higher 403 than the amount produced at a SME. However, the SMEs could access the fresh whey market by 404 405 providing whey with guaranteed safety and quality specifications for further collection in large batches and processing. In this scenario, the application of an efficient traceability system for whey 406 would be necessary for food applications. Traceability systems enable the identification of the 407 companies that are involved in its production and are responsible for its safety/quality, thus providing 408 transparency within the value-added supply chain. The procedural document of a model PDO cheese 409 410 (https://www.politicheagricole.it/flex/cm/pages/ServeBLOB.php/L/IT/IDPagina/3340) was used as

an example to draw a traceability scheme at a SME, considering the analogies in traditional 411 412 cheesemaking processes (Fox, Guinee, Cogan, & McSweeney, 2017). The flow of materials and information for the model PDO cheese was described according to the basic requirements of the 413 international standard ISO 22005:2007 (Table 2). The ISO 22005:2007 standard includes the 414 identification of the relevant information capture points, "traceable materials" and information 415 captured. Information capture points are the processing steps in which recording information is 416 417 essential to maintain identification of raw materials, partially-processed materials and products. Traceable units are items upon which there is a need to retrieve predefined information and include 418 products that can be exchanged between two parties in the supply chain as well as partially processed 419 420 materials. In general, information to be captured includes product information, process information and quality information and comprised the answers to the following questions: a) who is responsible 421 at that stage? (supplier, transporter, or organization); what are the product definition and 422 423 safety/quality specifications? c) what quantity of product is being considered? d) when does the stage occur? (date); e) where does the stage occur? (within an organization: in which plant is the product 424 processed, if more than one plant is present?) (Lavelli, 2013). The traceability plan for the model 425 PDO cheese consists of 17 capture points (1 - 11 and 13 - 18 in Table 2) that altogether collect 426 information on the materials involved in the production of a single whey lot, including all preceding 427 428 materials, such as: milk, rennet, starters, brine. Two additional capture points (12 and 19) are 429 necessary in case of whey recovery for food applications. The link among whey, curd and cheese is also important because in case of detection of curd or cheese contamination, whey safety has also to 430 431 be assessed. Hence, the traceability procedure allows to retrieve the safety and quality control of every lot of cheese (and thus whey) produced, such as: cow identity and veterinary controls, analytical 432 characterization of milk (content of fat, protein and lactose, pH, cryogenic index, absence of 433 bacteriophages and antibiotics); processing parameters, including: temperature and time of 434 pasteurization, scalding and ripening; quality control on the final cheese lots, including pathogenic 435 436 microorganisms, hygiene indicators, content of fat, protein and lactose and furosine level. In this

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context, it is worth noticing that the food safety management systems of SMEs dedicated to PDO 437 production are continuously controlled by self-assessment. Moreover, these companies are 438 periodically submitted to audits performed by the producers' consortium and by the third party 439 certification body. For a PDO cheese, processing conditions and final product specifications are 440 strictly standardized by the PDO procedural document. As a result, whey lots obtained from a PDO 441 cheese are characterized by standardized safety parameters, nutritional and functional properties, 442 443 which could be further modulated by physical treatments, enzymatic hydrolysis or chemical conjugation for valuable target applications in the food system. 444

The traceability procedure was then exemplified with the indication of the history of a single cheese 445 446 (and whey) lot recorded at a SME (Figure 4), which showed all the actors involved, i.e., seven farms, one milk transporter, three suppliers for rennet, starters, brine and packaging materials, respectively. 447 The exemplary mass flow revealed that the amount of milk collected was 64047 L, while only 25231.7 448 449 kg was processed to obtain the selected cheese lot (Figure 4). The amount of whey obtained was 21469.7 kg, i.e., 85% of milk, which is similar to the percentage reported previously (Rocha et al., 450 2020). The total amount of cheese obtained was 13.9% of milk, i.e., 3517.02 kg (1620 packages, that 451 is lower than the difference between processed milk and whey recovered, because of moisture loss 452 during ripening). Based on the analysis of the exemplary PDO cheese, it can be calculated that a 453 454 single whey lot of about 100 t, which is the minimum daily amount necessary to process whey into valuable products according to the techno-economic analysis previously proposed, a total of 35 farms 455 and 5 cheese-making companies need to be involved as well as transporters and ingredients' suppliers. 456 457 A scenario was also drawn to analyze the capability for SMEs to continuously supply whey to a potential stand-alone whey processor. To this aim, 36500 t/year would be necessary, corresponding 458 to about 6000 t of cheese/year. Interestingly, according to the producers' association of the model 459 PDO, 8400 2020 460 t/year of cheese were produced in (https://www.politicheagricole.it/flex/cm/pages/ServeBLOB.php/L/IT/IDPagina/3340). Hence, 461

462 almost all the companies of the consortium, which are approximately 80, should be involved in the463 whey value chain.

464 *4.2. Models of traceable whey supply chains*

A whey value chain could be built with the participation of a large number of SMEs and would require 465 tracing extensive information related to both product safety and quality, as well as responsible actors 466 at each stage of processing and distribution. While studies on a sought after "whey value chain" are 467 lacking, solutions to improve the milk and dairy supply chain transparency, involving large number 468 of actors have been proposed in the literature and could easily be extended to guarantee whey 469 traceability. One of the first traceability frameworks was developed to trace the whole supply chain 470 471 of Parmigiano Reggiano PDO cheese (Regattieri, Gamberi, & Manzini, 2007). This latter study was based on collection of a part of the relevant data with sensors and another part with manual data entry 472 in a centralized database, which was accessible by the stakeholders. After production, information 473 was also traced by a radio-frequency identification (RFID) tag applied to a whole cheese (Regattieri 474 et al., 2007). The number of sensors and devices developed to perform automatic food controls are 475 continuously increasing and hence Internet of Thing (IoT) and the Distributed Ledger Technologies 476 (DLT) such as the blockchain technology have become a new paradigm of how data can be collected, 477 securely stored, integrated, and communicated among different stakeholders. The blockchain is a 478 479 shared (distributed) and decentralized ledger, i.e., there are thousands of secure copies of data on different devices (nodes) worldwide. A unique blockchain account is created for every participant 480 and authentication and authorization through blockchain guarantees that the data added to blockchain 481 482 actually come from a specific participant. It is possible to insert blocks of information in the chain, but it is not possible to modify and remove the blocks previously added to the chain. The security and 483 immutability of information within the blocks are guaranteed by a consensus protocol and encryption 484 (Di Pierro, 2017). The blockchain technology can improve the transparency of the agri-food chain by 485 creating a flow of immutable data that any stakeholder can consult. An application of the blockchain 486 technology was proposed to manage data generated in a "smart" dairy supply chain in order to 487

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increase transaction efficiency, and create a flexible pricing mechanism (Zhang et al., 2019). 488 489 According to this model supply chain, the actors involved (farms, factories, government, service providers) are intended to collect various data related to location, safety and quality of the products, 490 mainly based on automatic systems such as drones, sensors (for temperature, humidity), timers, RFID 491 tags as well as analytical procedures (for microbial counts, protein and fat determinations, etc.). 492 Hence the blockchain is used as a data storage. Moreover, the blockchain provides a computational 493 infrastructure to run smart contracts, which can be used to automate some procedures of quality 494 control since they can directly execute the agreement once the specified conditions are met. A smart 495 contract can be designed by the government to verify to what extent the process meets the regulatory 496 497 standards: once all the requirements are met, the regulatory approval will be automatically granted. Moreover, smart contracts could introduce a more transparent and flexible pricing mechanism. For 498 example, a range of values (target value and tolerance range) can be determined for each category of 499 500 data related to product safety and quality, based on which, the finished products are categorized into different quality grades, with the first grade being the highest quality and last grade failing to meet 501 the standards and therefore being illegal to sell in the market (Zhang et al., 2019). 502

An application study was performed to calculate the economic impact of implementing a blockchain 503 technology to a milk supply chain including 80 affiliated dairy farms, a single dairy plant, seven 504 505 transportation companies, five distribution centers and approximately three thousand retailers (Longo, Nicoletti, & Padovano, 2019). Based on the necessary information to track the history of the 506 final food product, the transactions' frequency was calculated for each actor. For instance, at the farm 507 508 level, considering fifty cows, two hundred and forty six transactions are recorded every day, namely fifty for rearing (one for each cow), a hundred for milking (two for each cow) and ninety six for milk 509 temperature during storage (one every 15 min). The presented results were particularly interesting 510 since it was demonstrated that the investments in hardware and software was not relevant for any of 511 the actors involved. Moreover, the total cost due to the transactions' fee did not increase the 512 513 product/service price (Longo et al., 2019).

Indeed, the blockchain technology is reputed as a useful tool for SMEs. Implementation of 514 frameworks based on blockchain technology have been designed in the context of the PDO cheese 515 production, to record relevant information bringing together the milk producers, the dairy companies, 516 transporters, distributors, PDO consortium, third-party auditors, and the final consumer (Giacalone 517 et al., 2021; Varavallo et al., 2022). Implementation of blockchain technology was also studied in 518 emerging economy contexts, i.e., the biggest cooperative of dairy farmers in Turkey (Mangla et al., 519 520 2021), the domestic dairy supply chain in Vietnam, which involves all parties in the supply chain and the Vietnamese government (Tan and Ngan, 2020), as well as the smallholder farmers and local milk 521 collection centers in Kenya (Rambim, & Awuor, 2020). 522

523 The availability of big data by IoT and DLT poses additional challenges, one being the real-time analysis and response, pattern recognition and forecasting and the other being the bottleneck and 524 congestion that may occur when dealing with millions of connected data sources. Hence, Edge 525 Computing (EC) has emerged as a new paradigm to solve those problem. An EC architecture was 526 proposed to manage the dairy supply chain, which consists of three principal layers: IoT, Edge and 527 Business Solution layers. At the IoT level, sensors are located in livestock farms, crops used to feed 528 livestock, cattle, dairy factories and means for transportation to collect data on ambient conditions 529 (temperature and relative humidity, hazardous gas sensors), power consumption, cattle real-time 530 531 location and health conditions, product real time location. At the edge level, the data are pre-processed and filtered by Data Analytics techniques, generating knowledge and reducing data traffic and 532 transmission costs to the cloud. At the business solution layer, a virtual organization of agents works 533 534 as a social machine, which provides the Decision Support System (DSS) and an alert management system which sends warning messages and corrective actions when the values obtained by 535 heterogeneous IoT networks indicate a hazardous situation. This architecture was tested in a single 536 dairy company and planned to be implemented on several dairy farms in the same region (Alonso et 537 al., 2020). 538

Overall, these model traceability systems have been designed to implement smart system for data 539 collection and sharing among stakeholders and producers. For instance, a network could be 540 implemented among SMEs located in a confined geographical area, including those involved in PDO 541 cheese production, and various stakeholders such as the PDO producer consortium, the third party 542 certification body and the competent authority (Figure 5). As a result, these model traceability 543 systems could also support collection of information on whey safety and quality and hence promote 544 whey recovery processes. Moreover, this latter network would provide a comprehensive data 545 assembly and analysis, which would result in a powerful tool that will increase dairy SMEs 546 competitiveness. 547

548 **5.** Conclusions

The techno-economic analyses applied to whey recovery processes led to conclude that the downstream processing into valuable food products are associated to high energy input, which could be mitigated in large size plants operating 100 or more ton of whey daily. Risk assessment approaches evidenced that for the recycling of whey in the food system, the prevention of various chemical and microbial contaminants is needed. Moreover, the control and improvement of whey nutritional properties and functionality by physical and chemical treatments can potentially open up a new area of value-added applications.

556 A single SME could contribute to the circular economy by supplying small whey lots to whey processors, for collection in large batches and further transformation into value-added products. 557 Interestingly, SMEs that operate according to the same PDO quality scheme could constitute a supply 558 559 chain that provides whey processors with traceable whey lots with standardized safety and quality specifications through a shared smart traceability architecture based on IoT, DLT and EC solutions, 560 within the context of industrial symbiosis. Hence, the PDO products, which have been conceived as 561 a strategy for rural development, fostering local values such as environmental stewardship, culture 562 and tradition could also support the strategies for by-product recovery, thus advancing the application 563 564 of the bioeconomy principles in the food system.

To implement a whey value-chain, future research should aim at the advancement of cost-effective technological solutions for whey processing. Moreover, deepening knowledge on the required quality specification of whey as correlated to target applications would support the application of the best technology available as related to the end-use. Finally, the definition of standardized audit protocols with specific focus on whey processing conditions and quality parameters, would increase transparency and trust in the whey supply chain.

571

572 **Declaration of competing interest**

573 None.

574

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578

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583 **References**

Albuquerque, T. G., Oliveira, B. M. P. P, & Costa, H. S. (2018). 25 years of European Union (EU)

- quality schemes for agricultural products and foodstuffs across EU Member States. *Journal of the Science of Food and Agriculture*, 98, 2475–2489. https://doi.org/10.1002/jsfa.8811
- 587 Alirezalu, K., Munekata, P. E. S., Parniakov, O., Barba, F. J., Witt, J et al. (2020). Pulsed electric
- field and mild heating for milk processing: A review on recent advances. *Journal of the Science*
- 589 *of Food and Agriculture*, *100*, 16–24. https://doi.org/0.1002/jsfa.9942

Alonso, R. S., Sittón-Candanedo, I., García, Ó., Prieto, J., & Rodríguez-González, S. (2020). An
intelligent Edge-IoT platform for monitoring livestock and crops in a dairy farming scenario.

 592
 Ad Hoc Networks, 98, 102047. https://doi.org/10.1016/j.adhoc.2019.102047

Alvarenga, V. O., Brancini, G. T. P., Silva, E. K., da Pia, A. K. R., Campagnollo et al. (2018). Survival
 variability of 12 strains of *Bacillus cereus* yielded to spray drying of whole milk. *International*

Journal of Food Microbiology, 286, 80–89. https://doi.org/10.1016/j.ijfoodmicro.2018.07.020

Aydiner, C., Sen, U., Topcu, S., Ekinci, D., Altinay, A. D., et al. (2014). Techno-economic viability
of innovative membrane systems in water and mass recovery from dairy wastewater. *Journal*

598 *of Membrane Science*, 458, 66–75. http://doi.org/10.1016/j.memsci.2014.01.058

Bandara, S. B., Towle, K. M., & Monnot, A. D. (2021). A human health risk assessment of heavy
metal ingestion among consumers of protein powder supplements. *Toxicology Reports*, 7,

601 1255–1262. https://doi.org/10.1016/j.toxrep.2020.08.001

Barba, F. J. (2021). An integrated approach for the valorization of cheese whey. *Foods*, *10*, 564.
https://doi.org/10.3390/foods10030564

Barukcic, I., Jakopovic, K. L., Herceg, Z., Karlovic, S., & Bozanic, R. (2015). Influence of high

605 intensity ultrasound on microbial reduction, physico-chemical characteristics and fermentation

of sweet whey. Innovative Food Science and Emerging Technologies, 27, 94–101.

607 https://doi.org/10.1016/j.ifset.2014.10.013

- Bogdanovicova, K., Necidov, L., Harustiakov, D., & Janstov, B. (2017). Milk powder risk assessment
 with *Staphylococcus aureus* toxigenic strains. *Food Control*, 73, 2–7.
 http://doi.org/10.1016/j.foodcont.2016.07.007
- 611 Brandão da Costa, B. R., Rocha Roiffé, R., da Silva de la Cruz, M. N. (2021). Quality control of
- 612 protein supplements: A review. International Journal of Sport Nutrition and Exercise
- 613 *Metabolism*, 31, 369–379. https://doi.org/10.1123/ijsnem.2020-0287

- Brandelli, A., Daroit, D. J., Folmer Corrêa, A. P. (2015). Whey as a source of peptides with
 remarkable biological activities. *Food Research International*, 73, 149–161
 http://doi.org/10.1016/j.foodres.2015.01.016
- Bund, R. K., & Pandit, A. B. (2007). Rapid lactose recovery from paneer whey using
 sonocrystallization: A process optimization. *Chemical Engineering and Processing*, *46*, 846–
 850. http://doi.org/10.1016/j.cep.2007.05.015
- 620 Campos de Aquino, L. F. M., de Oliveira Resende Ribeiro, R., Siqueira Simoes, J., Borges Mano, S.,
 621 Teixeira Mársico, E. et al. (2017). Mercury content in whey protein and potential risk for human
- health. Journal of Food Composition and Analysis, 59, 141–144. http://
 doi.org/10.1016/j.jfca.2017.02.014
- Carrillo-Lopez, L. M., Garcia-Galicia, I. A., Tirado-Gallegos, J. M., Sanchez-Vega, R., HuertaJimenez, M. et al. (2021). Recent advances in the application of ultrasound in dairy products:
 Effect on functional, physical, chemical, microbiological and sensory properties. *Ultrasonics*

627 *Sonochemistry*, 73, 105467. https://doi.org/10.1016/j.ultsonch.2021.105467

- 628 Castro-González, N. P., Calderón-Sánchez, F., Castro de Jesús, J., Moreno-Rojas, R., Tamariz-Flores,
- J. V., et al. (2018). Heavy metals in cow's milk and cheese produced in areas irrigated with
- 630 waste water in Puebla, Mexic. *Food Additives and Contaminants: Part B Surveillance*, 11, 33–
- 631 36. https://doi.org/10.1080/19393210.2017.1397060
- Chalermthai, B., Ashraf, M. T., Bastidas-Oyanedel, J-R., Olsen, B. D., Schmidt, J. E., et al. (2020).
 Techno-economic assessment of whey protein-based plastic production from a copolymerization process. *Polymers*, *12*, 847. https://doi:10.3390/polym12040847
- Chamberland, J., Benoit, S., Doyen, A., Pouliot, Y. (2020). Integrating reverse osmosis to reduce 635 water and energy consumption in dairy processing: a predictive analysis for Cheddar cheese 636 manufacturing plants. Journal of Water Process Engineering, 38, 101606. 637 https://doi.org/10.1016/j.jwpe.2020.101606 638

- Chandra, R., Castillo-Zacarias, C., Delgado, P., & Parra-Saldívar, R. (2018). A biorefinery approach
 for dairy wastewater treatment and product recovery towards establishing a biorefinery
 complexity index. *Journal of Cleaner Production*, *183*, 1184–1196.
 https://doi.org/10.1016/j.jclepro.2018.02.124
- Chatzipaschali, A. A., & Stamatis, A. G. (2012). Biotechnological utilization with a focus on
 anaerobic treatment of cheese whey: current status and prospects. *Energies*, *5*, 3492–3525.
 https://doi.org/10.3390/en5093492
- 646 Corbaton-Baguena, M. J., Alvarez-Blanco, S., Vincent-Vela, M.-C., Ortega-Navarro, E., Perez-
- Herranz, V. (2016). Application of electric fields to clean ultrafiltration membranes fouled with
 whey model solutions. *Separation and Purification Technology*, *159*, 92–99.
 http://dx.doi.org/10.1016/j.seppur.2015.12.039
- Costamagna, D., Gaggiotti, M., Chiericatti, C. A., Costabel, L., Audero, G. M. L., et al. (2019).
 Quantification of aflatoxin M1 carry-over rate from feed to soft cheese. *Toxicology Reports*, *6*,
 782-787 https://doi.org/10.1016/j.toxrep.2019.07.004
- da Silva, A. N., Perez, R., Minim, V. P. R., Sant'Anna Martins, D. D., & Minim, L. A. (2015).
 Integrated production of whey protein concentrate and lactose derivatives: What is the best
 combination? *Food Research International*, *73*, 62–74
 https://dx.doi.org/10.1016/j.foodres.2015.03.009
- Delavar, M., Abdollahi, M., Navabi, A., Sadeghi, M., Hadavand, S., et al. (2012). Evaluation and
 determination of toxic metals, Lead and Cadmium, in incoming raw milk from traditional and
 industrial farms to milk production factories in Arak, Iran. *Iranian Journal of Toxicology*. 6,
 630–634.
- DeWit, J. N., & Klarenbeek, G. (1984). Effects of various heat treatments on structure and solubility
 of whey proteins. *Journal of Dairy Science*, 67, 2701–2710. https://doi.org/10.3168/jds.S00220302(84)81628-8

- D'Incecco, P., Limbo, S., Hogenboom, J. A., Pellegrino, L. (2021). Novel technologies for extending
 the shelf life of drinking milk: Concepts, research trends and current applications. *LWT Food Science and Technology*, *148*, 111746. https://doi.org/10.1016/j.lwt.2021.111746
- Di Pierro, M. (2017). What is the blockchain? *Computing in Science and Engineering*, *19*, 92–95.
 https://doi.org/10.1109/MCSE.2017.3421554
- Doost, A. S., Nasrabadi, M. N., Wu, J., Ayun, O., & Van der Meeren, P. (2019). Maillard conjugation 669 670 as an approach to improve whey proteins functionality: A review of conventional and novel Food preparation techniques. **Trends** in Science & Technology, 91, 1–11. 671 https://doi.org/10.1016/j.tifs.2019.06.011 672
- Elgammal, S. M., Khorsheda, M. A., & Ismail, E. H. (2019). Determination of heavy metal content
 in whey protein samples from markets in Giza, Egypt, using inductively coupled plasma optical
 emission spectrometry and graphite furnace atomic absorption spectrometry: A probabilistic
 risk assessment study. *Journal of Food Composition and Analysis, 84*, 103300.
 https://doi.org/10.1016/j.jfca.2019.103300
- Eluk, D., Ceruti, R., Nagel, O., & Althaus, R. (2019). Effect of thermal treatment of whey
 contaminated with antibiotics on the growth of *Kluyveromyces marxianus*. *Journal of Dairy Research*, 86, 102–107. https://doi.org/10.1017/S0022029919000098
- European Commission (2006). Regulation (EC) No 1881/2006 of 19 December 2006 setting
 maximum levels for certain contaminants in foodstuffs. *Official Journal of European Communities*, L 364, 5-24.
- European Commission (2010). Regulation (EU) No 37/2010 of 22 December 2009 on
 pharmacologically active substances and their classification regarding maximum residue limits
 in foodstuffs of animal origin. *Official Journal of European Communities*, L 15, 1-72.
- European Parliament and Council (2012). Regulation (EU) No 1151/2012 of 21 November 2012 on
 quality schemes for agricultural products and foodstuffs. *Official Journal of European Communities*, L 343, 1-29.

- Evans, J., Foster, A., Huet, J-M., Reinholdt, L., Fikiin, K., et al. (2015). Specific energy consumption
 values for various refrigerated food cold stores. In: *Proceedings of the 24th International Congress of Refrigeration*, Yokohama, Japan. https://doi.org/10.13140/RG.2.1.2977.8400
- Ferreira, S., Machado, L., Pereira, R. N., Vicente, A. A., & Rodrigues, R. M. (2021). Unraveling the
- and electrochemical effects. *Innovative Food Science and Emerging Technologies*, 74,102831.

nature of ohmic heating effects in structural aspects of whey proteins – The impact of electrical

696 https://doi.org/10.1016/j.ifset.2021.102831

694

- Fischer, C., & Kleinschmidt, T. (2021). Valorisation of sweet whey by fermentation with mixed
 yoghurt starter cultures with focus on galacto-oligosaccharide synthesis. *International Dairy Journal*, *119*, 105068. https://doi.org/10.1016/j.idairyj.2021.105068
- FitzGerald, R. J., Murray, B. A., & Walsh, D. J. (2004). Hypotensive peptides from milk proteins.
 Journal of Nutrition, *134*, 980S–988S. https://doi.org/10.1093/jn/134.4.980S
- Foegeding, E. A., Davis, J. P., Doucet, D., & McGuffey, M. K. (2002). Advances in modifying and
 understanding whey protein functionality. *Trends in Food Science & Technology*, *13*, 151–159.
 https://doi.org/10.1016/S0924-2244(02)00111-5
- Fox, P. F., Guinee, T. P., Cogan, T. M., & McSweeney, P.L.H. (2017). Fundamentals of Cheese
 Science. (2nd ed.), Springer, New York, NY
- Ganju, S., & Gogate, P. R. (2017). A review on approaches for efficient recovery of whey proteins
 from dairy industry effluents. *Journal of Food Engineering*, 215, 84–96.
 http://doi.org/10.1016/j.jfoodeng.2017.07.021
- Giacalone, M., Santarcangelo, V., Donvito, V., Schiavone, O., & Massa, E. (2021). Big data for
- corporate social responsibility: blockchain use in Gioia del Colle DOP. Quality & Quantity, 55,
- 712 1945–1971. https://doi.org/10.1007/s11135-021-01095-w
- Giraldo, J., Althaus, R. L., Beltran, M. C., & Molina, M. P. (2017). Antimicrobial activity in cheese
- whey as an indicator of antibiotic drug transfer from goat milk. *International Dairy Journal*,
- 715 *69*, 40–44. https://doi.org/10.1016/j.idairyj.2017.02.003

- Gómez, J. A., Sánchez, Ó. J., & Correa, L. F. (2020). Techno-economic and environmental evaluation
 of cheesemaking waste valorization through process simulation using SuperPro designer. *Waste*
- 718 *and Biomass Valorization*, *11*, 6025–6045. https://doi.org/10.1007/s12649-019-00833-4
- Gonzalez, M. I., Ivarez, S. A, Riera, F., & Ivarez., R. A. (2007). Economic evaluation of an integrated
- process for lactic acid production from ultrafiltered whey. *Journal of Food Engineering*, 80,
- 721 553–561. https://doi.org/10.1016/j.jfoodeng.2006.06.021
- Guerrero, C., Vera, C., Acevedo, F., & Illanes, A. (2015). Simultaneous synthesis of mixtures of
 lactulose and galacto-oligosaccharides and their selective fermentation. *Journal of Biotechnology*, 209, 31–40. https://doi.org/10.1016/j.jbiotec.2015.06.394
- Halfwerk, R., Yntema, D., Van Spronsen, J. & Van der Padt, A. (2021). A sub-zero crystallization
 process for the recovery of lactose. *Journal of Food Engineering*, *308*, 110677.
 https://doi.org/10.1016/j.jfoodeng.2021.110677
- IARC- International Agency for Research on Cancer (2002). Aflatoxins, IARC Monographs on the
 Evaluation of Carcinogenic Risks to Humans 62 IARC Press, Lyon, France.
- ISO (2007). Traceability in the feed and food chain General principles and basic requirements for
 system design and implementation ISO 22005:2007. International Standard Organization,
 Geneve, CH
- Jelen, P., Roginski, H., Fuquay, J., & Fox, P. (2003). Whey processing: utilization and products.
 Encyclopedia of Dairy Science, 2739–2745
- Jurgilevich, A., Birge, T., Kentala-Lehtonen, J., Korhonen-Kurki, k., Pietikainen, K., Saikku, L., et
 al. (2016). Transition towards circular economy in the food system. *Sustainability*, *8*, 1–14.
 https://doi.org/10.3390/su8010069
- Kent, R. M., Fitzgerald, G. F., Hill, C., Stanton, C., & Ross, R. P. (2015). Novel approaches to
 improve the intrinsic microbiological safety of powdered infant milk formula. *Nutrients*, *7*,
- 740 1217-1244. https//doi.org/10.3390/nu7021217

- Ladha-Sabur, A., Bakalis, S., Fryer, P. J., & Lopez-Quiroga, E. (2019). Mapping energy consumption
- in food manufacturing. *Trends in Food Science & Technology*, 86, 270–280.
 https://doi.org/10.1016/j.tifs.2019.02.034
- Lányi, K., Darnay, L., László, N., Lehel, J., Friedrich, L. et al. (2022). Transfer of certain beta-lactam
 antibiotics from cow's milk to fresh cheese and whey. *Food Additives and Contaminants Part*
- A Chemistry, Analysis, Control, Exposure and Risk Assessment, 39, 52–60.
 https://doi.org/10.1080/19440049.2021.1973114
- Lappa, I. K., Papadaki, A., Kachrimanidou, V., Terpou, A., Koulougliotis, D., Eriotou, E., et al.
 (2019). Cheese whey processing: integrated biorefinery concepts and emerging food
 applications. *Foods*, *8*, 347. https://doi:10.3390/foods8080347
- Lavelli, V. (2013). High-warranty traceability system in the poultry meat supply chain: a mediumsized enterprise case study. *Food Control*, *33*, 148–156. https://
 doi.org/10.1016/j.foodcont.2013.02.022.
- Lavelli, V. (2021). Circular food supply chains Impact on value addition and safety. *Trends in Food Science &Technology*, *114*, 323–332. https://doi.org/10.1016/j.tifs.2021.06.008.
- Lavelli, V.; Sereikaite, J. (2022). Kinetic study of encapsulated β-carotene degradation in aqueous
 environments: A review. *Foods*, *11*, 317. https://doi.org/10.3390/foods11030317.
- Lavelli, V.; Sereikaite, J. (2022). Kinetic study of encapsulated β-carotene degradation in dried
 systems: A review. *Foods*, *11*, 437. https://doi.org/10.3390/foods11030437.
- Lindsay, M. J., Huang, K., Buchinger, B. A., Maravelias, C. T., Dumesic, J. A., et al. (2020). Catalytic
 production of glucose–galactose syrup from Greek yogurt acid whey in a continuous-flow
 reactor. *ChemSusChem*, *13*, 791–802. https://doi.org/10.1002/cssc.201902847.
- Congo, F., Nicoletti, L., & Padovano, A. (2019). Estimating the impact of blockchain adoption in the
- food processing industry and supply chain. *International Journal of Food Engineering*, *16*,
- 765 20190109. https://doi.org/10.1515/ijfe-2019-0109

- Lujan-Facundo, M. J., Mendoza-Roca, J. A., Cuartas-Uribe, B., & Alvarez-Blanco, S. (2017)
 Membrane fouling in whey processing and subsequent cleaning with ultrasounds for a more
 sustainable process. *Journal of Cleaner Production*, 143, 804–813.
 http://dx.doi.org/10.1016/j.jclepro.2016.12.043
- Lupton, S. J., Shappell, N. W., Shelver, W. L., & Hakk, H. (2018). Distribution of spiked drugs
 between milk fat, skim milk, whey, curd, and milk protein fractions: Expansion of partitioning
 models. *Journal of Agricultural and Food Chemistry*, 66, 306–314. https://doi.org/
 10.1021/acs.jafc.7b04463
- Malliaroudaki, M. I., Watson, N., Ferrari, R., Nchari L. N. (2022). Energy management for a net
 zero dairy supply chain under climate change. *Trends in Food Science & Technology*, xxx
 (xxxx) xxx. https://doi.org/10.1016/j.tifs.2022.01.015
- Mangla, S. K., Kazancoglu, Y., Ekinci, E., Liu, M., Ozbiltekin, M., & Sezer, M. D. (2021). Using
 system dynamics to analyze the societal impacts of blockchain technology in milk supply
 chains. *Transportation Research Part E: Logistics and Transportation Review*, *149*, 102289.
 https://doi.org/10.1016/j.tre.2021.102289
- 781 Margas, E., Meneses, N., Conde-Petit, B., & Dodd, C. E. R., Holaha, J. (2014). Survival and death
- kinetics of *Salmonella* strains at low relative humidity, attached to stainless steel surfaces.
 International Journal of Food Microbiology, 187, 33–40.
 http://doi.org/10.1016/j.ijfoodmicro.2014.06.027
- Mattas, K., Tsakiridou, E., Karelakis, C., Lazaridou, D., Gorton, M. et al. (2022). Strengthening the
 sustainability of European food chains through quality and procurement policies. *Trends in Food Science & Technology*, *120*, 248–253. https://doi.org/10.1016/j.tifs.2021.11.021
- Mehli, L., Hoel, S., Thomassen, B. G. M., Jakobsen, A. N., & Karlsen, H. (2017). The prevalence, 788 genetic diversity and antibiotic resistance of *Staphylococcus aureus* in milk, whey, and cheese 789 International 65. from artisan dairies. Dairy Journal. 20-27.790 farm 791 http://doi.org/10.1016/j.idairyj.2016.10.006

792	Mehra, R.; Kumar, H.; Kumar, N.; Ranvir S., Jana, A. et al. (2021). Whey proteins processing and					
793	emergent derivatives: An insight perspective from constituents, bioactivities, functionalities to					
794	therapeutic applications. Journal of Functional Foods, 87, 104760.					
795	https://doi.org/10.1016/j.jff.2021.104760					
796	Navarro, F., Harouna, S., Calvo, M., Perez, M. D., & Sanchez, L. (2015). Kinetic and thermodynamic					
797	parameters for thermal denaturation of ovine milk lactoferrin determined by its loss of					
798	immunoreactivity. Journal of Dairy Science, 98, 4328–4337. https://doi.org/10.3168/jds.2015-					
799	9403.					
800	Oliver, S. P., Murinda, S. E., & Jarayao, B. M. (2011). Impact of antibiotic use in adult dairy cows					
801	on antimicrobial resistance of veterinary and human pathogens: A comprehensive review.					
802	Foodborne Pathogens and Disease, 8, 337-355. https://doi:10.1089/fpd.2010.0730					
803	Pehlivanoglu, H., Feyza, H., Bardakci, M., Yaman, M. (2022). Protein quality assessment of					
804	commercial whey protein supplements commonly consumed in Turkey by in vitro protein					
805	digestibility-corrected amino acid score (PDCAAS). Food Science and Technology Campinas,					
806	42, 64720. https://doi.org/10.1590/fst.64720					
807	Peters, R. H. (2005). Economic aspects of cheese making as influenced by whey processing options.					
808	International Dairy Journal, 15, 537-545. https://doi:10.1016/j.idairyj.2004.11.009					
809	Pihlanto, A. (2001). Bioactive peptides derived from bovine whey proteins: opioid and ace-inhibitory					

810 peptides. *Trends in Food Science & Technology*, *11*, 347–356. https://doi.org/10.1016/S0924811 2244(01)00003-6.

- Prandini, A., Tansini, G., Sigolo, S., Filippi, L., Laporta, M. et al. (2009). On the occurrence of
 aflatoxin M1 in milk and dairy products. *Food Chemistry & Toxicology*, *47*, 984–991.
 http://doi.org/10.1016/j.fct.2007.10.005
- Pujato, S. A., Quiberoni, A., & Mercanti, D. J. (2018). Review article. Bacteriophages on dairy foods. *Journal of Applied Microbiology*, *126*, 14—30. https://doi.org/10.1111/jam.14062

- Rambim, D., & Awuor, F. M. (2020). Blockchain based milk delivery platform for stallholder dairy
 farmers in Kenya: Enforcing transparency and fair payment. 2020 IST-Africa Conference,
 9144071. www.IST-Africa.org/Conference2020
- Raut, S., Jain, S., Dhamole, P., & Agrawal, S. (2022). WPC manufacturing using thermal polyelectrolyte precipitation: A product quality and techno-economic assessment. *Journal of Food Engineering*, *315*, 110796. https://doi.org/10.1016/j.jfoodeng.2021.110796
- Regattieri, A., Gamberi, M., & Manzini, R. (2007). Traceability of food products: General framework
 and experimental evidence. *Journal of Food Engineering*, 81, 347–356.
 https://doi.org/10.1016/j.jfoodeng.2006.10.032
- Riera, F., Alvarez, A., Espi, A., Prieto, M., de la Roza, B., et al. (2014). Cow's milk with active 826 immunoglobulins against Campylobacter jejuni: Effects of temperature on immunoglobulin 827 Agriculture, Science of Food and 1205-1211. 828 activity. Journal of the 94. https://doi.org/10.1002/jsfa.6398 829
- Ring, G., Sheehan, A., Lehane, M., & Furey, A. (2021). Development, validation and application of 830 an ICP-SFMS method for the determination of metals in protein powder samples, sourced in 831 with risk assessment Irish consumers. Molecules, 832 Ireland, for 26. 4347. https://doi.org/10.3390/molecules26144347 833
- Rocha, J. M., & Guerra, A. (2020). On the valorization of lactose and its derivatives from cheese
 whey as a dairy industry by-product: an overview. *European Food Research and Technology*,
 246, 2161–2174. https://doi.org/10.1007/s00217-020-03580-2
- Rufián-Henares, J. A., Delgado-Andrade, C., Jiménez-Pérez, S., & Morales, F. J. (2007). Assessing
 nutritional quality of milk-based sport supplements as determined by furosine. *Food Chemistry*,
- 839 *101*, 573–578. https://doi.org/10.1016/j.foodchem.2006.02.016
- 840 Sameli, N., & Samelis, J. (2022). Growth and biocontrol of *Listeria monocytogenes* in Greek
 841 anthotyros whey cheese without or with a crude enterocin A-B-P extract: Interactive effects of

the native spoilage microbiota during vacuum-packed storage at 4 °C. *Foods*, 11, 334. https://doi.org/10.3390/foods11030334

- Sánchez-Oliver, A. J., Contreras-Calderon, J., Puya-Braza, J. M., & Guerra-Hernández, E. (2018).
 Quality analysis of commercial protein powder supplements and relation to characteristics
 declared by manufacturer. *LWT Food Science and Technology*, *97*, 100–108.
 https://doi.org/10.1016/j.lwt.2018.06.04
- Schottroff, F., Gratz, M., Krottenthaler, A., Johnson, N. B., Bedard, M. F. & Jaeger, H. (2019). Pulsed 848 electric field preservation of liquid whey protein formulations - influence of process 849 parameters, pH, and protein content on the inactivation of *Listeria innocua* and the retention of 850 851 bioactive ingredients. Journal of Food Engineering, 243, 142-152. https://doi.org/10.1016/j.jfoodeng.2018.09.003 852
- Scott, F., Vera, C., & Conejeros, R. (2016). Technical and economic analysis of industrial production
 of lactose-derived prebiotics with focus on galacto-oligosaccharides. In A., Guerrero, C., Vera,
- 855 C., Wilson, L., Conejeros, R., Scott, F. (Eds), Lactose-derived prebiotics; Illanes (pp. 261–284).
 856 Academic Press: San Diego, CA, USA
- Serna-Hernandez, S. O., Escobedo-Avellaneda, Z., García-García, R., Rostro-Alanis, M. J., & WeltiChanes, J. (2021). High hydrostatic pressure induced changes in the physicochemical and
 functional properties of milk and dairy products: A review. *Foods*, 10, 1867.
- 860 https://doi.org/10.3390/foods10081867
- Shelver, W. L., Lupton, S. J., Shappell, N. W., Smith, D. J., & Hakk, H. (2018). Distribution of
 chemical residues among fat, skim, curd, whey and protein fractions in fortified, pasteurized
 milk. *ACS Omega*, *3*, 8697-8708. https://doi.org/10.1021/acsomega.8b00762
- 864 Sipola, M., Finckenberg, P., Vapaatalo, H., Pihlanto-Leppala, A. et al. (2002). Alpha-lactorphin and
- beta-lactorphin improve arterial function in spontaneously hypertensive rats. *Life Science*, *71*,
- 866 1245–1253. https://doi.org/ 10.1016/s0024-3205(02)01793-9

- Staron, J., Dabrowski, J. M., Cichon, E., & Guzik, M. (2018). Lactose esters: Synthesis and
 biotechnological applications. *Critical Review in Biotechnology*, 38, 245–258.
 https://doi.org.10.1080/07388551.2017.1332571
- 870 Tirloni, E., Ghelardi, E., Celandroni, F., Bernardi, C., Casati, et al. (2017). Bacillus cereus in fresh
- 871 ricotta: Comparison of growth and haemolysin BL production after artificial contamination
- 872 during production or post processing. *Food Control*, 79, 272–278.
 873 http://doi.org/10.1016/j.foodcont.2017.04.008
- Tan, A., & Ngan, P. T. (2020). A proposed framework model for dairy supply chain traceability. *Sustainable Futures*, 2, 100034. https://doi.org/10.1016/j.sftr.2020.100034
- Tsermoula, P., Khakimov, B., Nielsen, J. H., Engelsen, S. B. (2021). Whey The waste-stream that
 became more valuable than the food product. *Trends in Food Science & Technology*, *118*, 230–
 241. https://doi.org/10.1016/j.tifs.2021.08.025
- USDEC, 2006. Reference Manual for US Whey and Lactose Products. V. Lagrange, Ed. U.S. Dairy
 Export Council Arlington, VA, US
- Van den Berg, M., Birnbaum, L. S., Denison, M., De Vito, M., & Farland, W. (2006). The 2005 world
 health organization reevaluation of human and mammalian toxic equivalency factors for
 dioxins and dioxin-like compounds. *Toxicological Sciences*, 93, 223–241. https://doi.org/
 10.1093/toxsci/kfl055
- Varavallo, G., Caragnano, G., Bertone, F., Vernetti-Prot, L., & Terzo, O. (2022). Traceability
 platform based on green blockchain: an application case study in dairy supply chain. *Sustainability*, *14*, 3321. https://doi.org/10.3390/su14063321
- Wedel, C., Atamer, Z., Dettling, A., Wenning, M., Scherer, S., et al. (2022). Towards low-spore milk
 powders: A review on microbiological challenges of dairy powder production with focus on
 aerobic mesophilic and thermophilic spores. *International Dairy Journal*, *126*, 105252.
- 891 https://doi.org/10.1016/j.idairyj.2021.105252

- Wu, G., Hui, X., Gong, X., Tran, K. N., Stipkovits, L., et al. (2021a). Functionalization of bovine 892 893 whey proteins by dietary phenolics from molecular-level fabrications and mixture-level combinations. Trends Food Science & Technology, 110, 107–119. 894 in https://doi.org/10.1016/j.tifs.2021.01.072 895
- Wu, J., Li, H., Ayun, Q., Doost, A. S., De Meulenaer, B., et al. (2021b). Conjugation of milk proteins
 and reducing sugars and its potential application in the improvement of the heat stability of
 (recombined) evaporated milk. *Trends in Food Science & Technology, 108,* 287–296.
 https://doi.org/10.1016/j.tifs.2021.01.019
- Zandona, E., Blažić, M., & Jambrak A. R. (2021). Whey utilisation: sustainable uses and
 environmental approach. *Food Technology and Biotechnology*, 59, 147–161.
 https://doi.org/10.17113/ftb.59.02.21.6968
- Zhang, Y., Xu, X., Liu, A., Lu, Q., Xu, L., & Tao, F. (2019). Blockchain-based trust mechanism for
 IoT-based smart manufacturing system. *IEEE Transactions on Computational Social Systems*,

905 6, 1386 – 1394. https://doi.org/10.1109/TCSS.2019.2918467

Zhang, L., Zhou, R., Zhang, J., & Zhou P. (2021). Heat-induced denaturation and bioactivity changes
of whey proteins. *International Dairy Journal*, *123*, 105175.
https://doi.org/10.1016/j.idairyj.2021.105175

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910 Captions to the Figures

911

Figure 1. Processing schemes for whey recycling to recover whey powder (WP) and water. Routes [1a] and [1b] were modelled by techno-economic analysis for an inflow of 100 t whey/d (Aydiner et al., 2014); route [2] for 1500 t whey/d (Chamberland et al., 2022). Operations that need high energy input, possibly mitigated by increasing plant size, are indicated in orange. WPs compositions are indicated in Table 1.

Figure 2. Processing schemes for whey recycling to recover whey protein concentrates (WPC) and lactose-derived products. Route [3] was modelled by techno-economic analysis for 700 t whey/d (da Silva et al., 2015); route [4] for 80 t whey/h (Scott et al., 2016); route [5] for 500 t whey/d (Gomes et al., 2020); route [6] for 100 t whey/d (Gonzales et al., 2007). Operations that need high energy input, possibly mitigated by increasing plant size are indicated in orange. WPCs compositions are indicated in Table 1.

924

Figure 3. Processing schemes for whey recycling to recover whey protein isolate (WPI) and whey
protein precipitate (WPP). Routes [7a] and [7b] were modelled by techno-economic analysis for 2000
- 3000 t whey/d (Peters, 2005); route [8] was modelled for 100 t whey/d (Raout et al., 2022).
Operations that need high energy input, possibly mitigated by increasing plant size are indicated in
orange. WPI and WPP compositions are indicated in Table 1.

930

Figure 4. Exemplary traceability scheme for a cheese lot. The mass flow, process timing andmaterials' lot numbers are indicated in the red, light blue and dark blue boxes, respectively.

933

Figure 5. A model smart whey supply chain, indicating the actors involved and the communicationtools.

Product	Moisture	Protein	Fat	Lactose	Ash	Reference
Sweet whey	93 - 93.7	0.6 - 1	0.5	4.6 - 5.2	0.5	Jelen et al., 2003
Acid whey	93 - 93.7	0.6 - 0.8	0.04	4.4 - 4.6	0.8	Jelen et al., 2003
WP (FO/MD)	4.89	14.02	1.86	53.36	13.91	Aydiner et al., 2014
WP (MD/RO)	4.51	11.99	5.36	57.85	9.12	Aydiner et al., 2014
WP (RO)	n.d.	15	0.57	75.4	n.d.	Chamberland et al., 2021
WPC 80	4.0	81.86	8.28	4.88	0.98	daSilva et al., 2016
WPC 50	4.99	50.24	n.d.	35.39	n.d.	Gómez et al., 2020
WPI	4.5	92	1	0.5	2	USDEC, 2006

Table 1. Composition of sweet and acid whey and whey products (g in 100 g).

WP, whey powder; WP (FO/MD), WP concentrated by forward osmosis and membrane distillation; WP (MD/RO), WP concentrated by membrane distillation and reverse osmosis; WP (RO), WP concentrated by reverse osmosis; WPC 80, whey protein concentrates at 80%, WPC 50, whey protein concentrates at 50%, WPI, whey powder isolates. n.d., not determined.

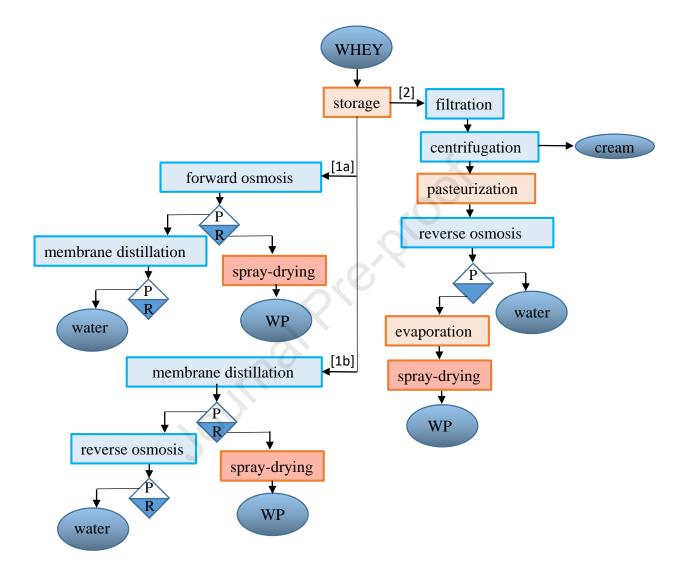
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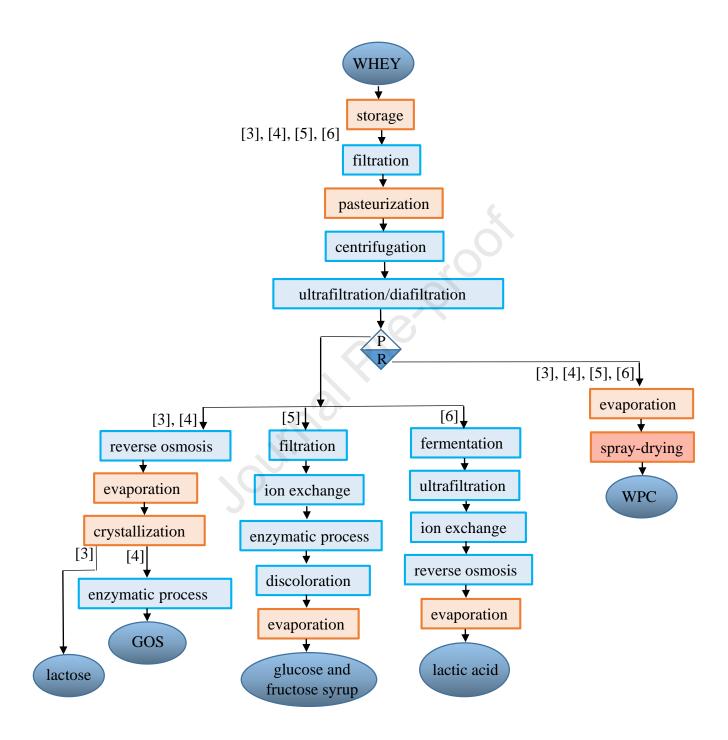
CP. Processing step	Traceable unit(s)	Information tracked	Note
At the farm			
1. Rearing	Cow	 Animal code Sanitary controls and eventual drug treatments 	Geographical location of the farms, breeding practices and admitted feed are indicated in the PDO procedural document
2. Milking	Raw milk in the cistern	 Cow Milk lot number and amount Date 	
3. Milk storage	As for point 2	- Temperature - Date and duration	
At the transporter			
4. Milk transport	As for point 2	 Transporter name Milk supplier name Milk lot number and amount Temperature Date and duration 	
At the cheesemaking S.			
5. Receiving of milk	As for point 2	 Milk supplier name Milk lot number and amount Transporter's name Temperature Safety/quality specification Date 	Milk characteristics are indicated in the procedural document of the PDO. Limits are fixed for fat: 4.01 -4.25 %; protein: 3.25 - 3.35 %; lactose: 5.00 - 5.15 %; cryoscopic index: -0.520 - 0.525 °C; pH 6.74
6. Milk storage	Raw milk mixture stored in the tank	 Milk lot numbers and amount Number of the tank Temperature Date and duration 	Maximum storage duration indicated in the PDO procedural document is 24 h at 4 °C
7. Milk pasteurization	Pasteurized milk mixture intended for the scheduled cheese production	 Mixed milk lot number and amount Storage tank number Time and temperature Date 	Pasteurization conditions: 72°C 15 s The lot number of the mixed milk is the same as that of the scheduled cheese
8. Receiving of rennet, starters, salt and packaging materials	Rennet/starters, salt and packaging materials in the suppliers' packages	 Supplier name Rennet/starters/salt/packaging materials' lot numbers and amounts Transporter's name Temperature (for rennet and starters) Date 	Rennet form young calves and cultures of <i>L. bulgaricus</i> and <i>S. thermophilus</i> are only admitted by procedural document of the PDO. Rennet performance and fermentation ability are checked to standardize cheese yield.
9. Rennet and starters storage	As for point 7	TemperatureDate and duration	Storage is performed at 4 °C.

Table 2. Traceability plan for a model PDO cheese according to the ISO 22005:2007 standard.

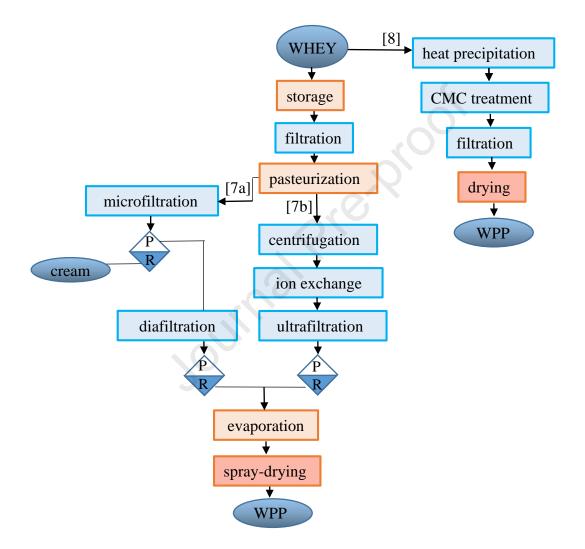
10. Filling of the cheesemaking vats	Mixture in the cheese-making vats	 Pasteurized milk mixture lot number and amount Rennet/starters lot numbers and amounts Date 	
11. Empting of the cheesemaking vats and filling of the mould	Curd in the moulds Whey in the tank	 Curd lot number, amount and number pieces Whey lot number and amount Date 	The lot number of the curd and whey obtained is the same as that of the scheduled cheese
12. Whey storage	Whey in the tank	Whey lot number and amountTemperatureQuality specificationsDate and duration	This step is needed to implement whey recovery
13. Curd scalding	Curds in the scalding room	 Curd lot number, amount and number pieces Temperature and relative humidity Date and duration 	Scalding conditions are indicated in the PDO procedural document. Temperature is 22-25 °C and duration is 8-16 h. Relative humidity is 90%.
14. Brine preparation	Brine in the can	Salt lot number and amountBrine lot number and amountDate	
15. Curd ripening	Curds in the ripening room	 Curd lot number, amount and number of pieces Brine lot number and amount Temperature and relative humidity Date and duration 	Ripening conditions are indicated in the PDO procedural document. Temperature is 2-6 °C and minimum ripening duration is 35 d. Relative humidity is 85-90%.
16. Pressing of the brand, primary and secondary packaging of cheese	Cheese in single packages assembled in boxes	 Cheese lot number, amount and number of final portions obtained Quality specifications Date 	Quality specifications for cheese are indicated in the procedural document of the PDO. Limits are fixed for dry matter: 46%; fat: 23%; protein: 17%; furosine <14 g/100g protein; pH 5.5; pathogenic microorganisms: absent; <i>Enterobacteriaceae</i> < 100 CFU/g; Staphylococci coagulase+ < 20 CFU/g.
17. Cheese storage	As for point 16	- Time and temperature	Shelf-life is 30 d at 4 °C.
At the transporter			
18. Cheese transport	As for point 16	 Cheese lot number, amount and number of portions delivered Customer's name Temperature Date and duration 	
19. Whey transport	Whey in the tanker	Whey lot number and amountWhey processor nameTemperature	This step is needed to implement whey recover processes at a plant size

CP, capture point

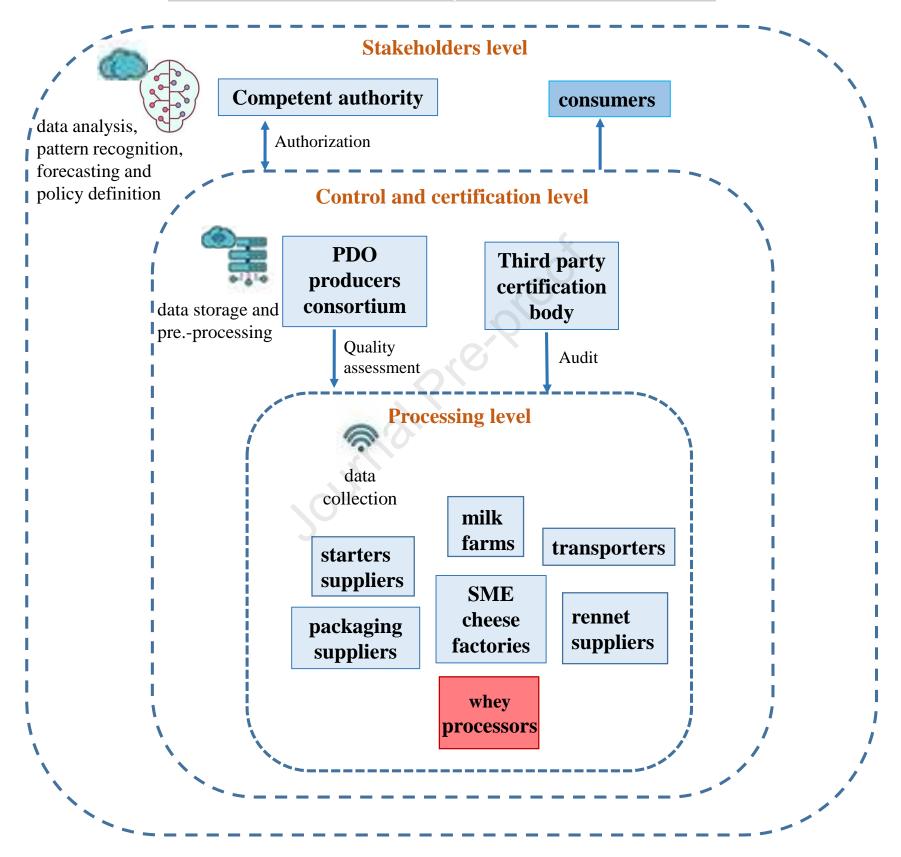








collected 64047 L milk RL 019028 RL 019029 storage 6/11 RL 019030 RL 019031 RL 019035 pasteurization 7/11 RL 019036 RL 019037 pasteurized 25231.7 kg milk receiving and Ŧ A007903A starters storage filling of the cheese-making vats GT144503 receiving and rennet storage empting of the cheese-making 7/11 vats and filling of the mould whey 25231.7 kg 14247 receiving and curd 3762.00 kg (1620 pieces) P1438 salt storage 7/11 brine addition and scalding receiving and ripening packaging 2820010 storage materials packaging 23/12 cheese 3517.02 kg (1620 pieces)



Highlights

- Efficient whey recycling processes can be performed with large whey inflow (> 100 t/d)
- High-quality Protected Designation of Origin (PDO) cheeses are made at small/medium enterprises (SMEs)
- SMEs can produce whey that meets safety quality requirements for recycling
- SMEs could contribute to whey recycling by suppling traceable whey lots to industrial processors
- Smart traceability schemes support a trustable whey value chain involving a multitude of actors
- SMEs processing PDO cheeses could advance the application of the bioeconomy in the food system