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TITLE Alternative uses for co-products: Harnessing the potential of valuable compounds from meat processing chains

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1	Alternative uses for co-products: Harnessing the potential of valuable compounds from
2	meat processing chains
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26	Abstract
27	Opportunities for exploiting the inherent value of protein-rich meat processing co-products,
28	in the context of increased global demand for protein and for sustainable processing systems,
29	are discussed. While direct consumption maybe the most profitable route for some, this
30	approach is influenced greatly by local and cultural traditions. A more profitable and
31	sustainable approach may be found in recognizing this readily available and under-utilized
32	resource can provide high value components, such as proteins, with targeted high value

33 functionality of relevance to a variety of sectors. Applications in food & beverages, petfood

biomedical and nutrition arenas are discussed. Utilization of the raw material in its entirety is
a necessary underlying principle in this approach to help maintain minimum waste
generation. Understanding consumer attitudes to these products, in particular when used in
food or beverage systems, is critical in optimizing commercialization strategies.

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39 Highlights

40 - Opportunities for extracting additional value from meat processing chain

41 - Meat co-products excellent source of proteins, vitamins, minerals and bio-actives

42 -High added-value uses can be developed to revalorize co-products

- 43
- 44

45 Keywords

46 Meat co-products, bioactive, protein, consumers, legislation, offal, valorisation, by-products

47

48 **1. Introduction**

Global pressures for increased supply of foods with high protein content, coupled with the 49 ever increasing demand for more efficient processing and increased financial rewards, 50 51 provides the imperative for the food industry to examine all processing streams with a view to adding or recovering value. Based on its amino acid profile, animal protein can provide a 52 53 complete protein with a high biological value. In addition, meat, and in particular red meat, 54 provides a rich source of macro and micro-nutrients. Demand for this high quality product is 55 expected to continue to expand (FAO, 2009), which will lead to the increased production of the lower value 'non meat' products such as offal, blood, etc., which have been estimated to 56 57 account for approximately 54-56% of the bovine and 48% of the porcine live animal weight (Marti, Johnson, & Mathews Jr, 2012; Vernooij, 2012). In this review, such non-meat 58 59 products are referred to as co-products, similar to the view adopted in Australia. Classification as animal by-products (ABPs) is not always relevant as this classification under 60 European legislation would restrict the use of these parts in the human food chain. While 61 many countries have secured markets for them, in general these tend to be low value markets, 62 often necessitating substantial transport costs. Even more, some of the products will carry a 63 neutral (no disposal cost) or negative (costs money to dispose of) value. Making better use of 64 the many co-products arising from meat processing, is not only important from a 65 sustainability perspective, but also offers opportunities to the industry to increase their value. 66

As discussed by many authors (Lynch, Mullen, O'Neill, & Álvarez, 2017; Mullen, Álvarez, 67 Pojić, Hadnadev, & Papageorgiou, 2015; Toldrá, Mora, & Reig, 2016) many of the materials 68 are rich sources of valuable components such as protein, lipids, biomolecules etc. While some 69 products command a reasonable value, and provide a nutritious food (Jayathilakan, Sultana, 70 Radhakrishna, & Bawa, 2012) when consumed directly, for others it may be more valuable to 71 extract separate components for discreet downstream applications. A five-stage universal 72 73 recovery process was suggested by Galanakis (2012) to describe this process of transforming 74 raw material into a final higher value product. While some of these steps can be eliminated or 75 merged this gives a good overview of the process. Both traditional and emerging technologies of relevance are discussed in a later section. Optimal exploitation brings many challenges 76 such as product hygiene and stabilisation, economies of scale, product purity, consumer 77 attitudes and the requirement for efficient processing, which minimises the impact on the 78 79 environment. These challenges can be addressed both through good industry practice and an applied research approach. Important principles to be guided by include that of total 80 exploitation, avoiding the generation of additional waste (Waldron, 2007), and the cascading 81 principle within the bioencomy discourse which is about ensureing the highest value is 82 extracted first with lower value options only considred once the higher ones have been 83 84 exhausted.

Encouraging optimal utilisation of raw materials, early in the production chain, can help 85 86 ameliorate food waste losses before the product becomes more dispersed along the food chain. The search for alternative protein sources has advanced in recent years (Van Huis et 87 88 al., 2013), and provides a relevant approach to meeting global protein requirements. However, it is imperative to recognise that we have a readily available and under-utilised 89 90 resource ripe for exploitation in co-products arising from meat processing. As there is a link 91 between meat consumption and demand for protein-rich feed for livestock (Pimentel & 92 Pimentel, 2003; Trostle, 2008), it is imperative that all meat processing related proteins are optimally exploited. This review covers applications relating to use in food and beverage 93 systems, petfood, nutrition focused and bioactive & biomedical applications. Market and 94 legislative information is provided in additional to an overview of relevant technologies and 95 96 consumer attitudes.

97

98 2. Applications

99 **2.1 Food ingredients**

100 Among other attributes proteins can impart techno-functional properties within a food or beverage system. For example they may influence water holding capacity, viscosity, gelation, 101 emulsification and foaming. The ability of proteins to function in this manner is determined 102 by intrinsic e.g. amino acid profile, molecular structure and surface hydrophobicity, and by 103 extrinsic factors such as pH temperature, and ionic strength. Collagen, a fibrous, structural 104 105 protein, is one of the most abundant proteins. It is probably the animal protein most regularly employed in food production. Following extraction and hydrolysis it can be transformed into 106 gelatine whose strong gelling properties are relevant to a variety of products such as soups, 107 108 gravies, desserts or dairy products (Hettiarachchy, Sato, Marshall, & Kannan, 2012). Gelatine is mainly extracted from skin and bones. However, other offal such as lung, tongue, 109 trachea, large blood vessels, or tendons are, also, sources of collagen. Collagen has very 110 good film forming capacity which is exploited to generate casings and novel packaging 111 (Gómez-Guillén, Giménez, López-Caballero, & Montero, 2011). Other more specific 112 applications of collagen are described below in section 2.4, and have also been reviewed by 113 various authors (Ferraro, Anton, & Santé-Lhoutellier, 2016; Zeugolis, Paul, & Attenburrow, 114 115 2009). A variety of commercial collagen based products are available for use in food systems. Examples of companies supplying them include Rousselot, Collapro and Devro, who provide 116 117 both clean label binders, gelling and textural agents or collagen based sausage casing.

Blood plasma comes second in line in meat co-product derived protein most employed as a 118 119 food ingredient. Following hygienic collection, centrifugation leads to the recovery of plasma and cellular fractions (Lynch et al., 2017). Plasma can be used as it is, or dehydrated for 120 121 storage as a dry powder (Pares, Toldra, Saguer, & Carretero, 2014). Blood plasma is a versatile product, which presents good emulsifying, gelling, foaming and solubility 122 123 properties. Applications of plasma in food industry see it used as a binder in meat products, egg replacers in bakery, protein-rich pasta, fat replacers or even polyphosphate substitute 124 (Hsieh & Ofori, 2011; Hurtado et al., 2011). Some examples of plasma proteins currently in 125 the market are: Fibrimex® (purified fibrinogen used as binder), Immunolin® (concentrated 126 immunoglobulins, which improve the immune system)(Abu-Akkada & Awad, 2015), Myored 127 (natural red colorant), Vepro 95 HV (purified globin used as emulsifier) or Plasma Powder 128 129 FG (plasma with increased content of fibrinogen used as cold binder) (Ofori & Hsieh, 2012).

Apart from those two sources, the use of offal as sources of proteinaceous food ingredients has not received a lot of research funding. In the last decade research has been published describing the extraction and characterisation of functional proteins from various offal such as: liver (Steen et al., 2016; Zou et al., 2017), lung (Selmane, Christophe, & Gholamreza, 2008), heart (Dewitt, Gomez, & James, 2002), viscera (Bhaskar, Modi, Govindaraju, Radha,
& Lalitha, 2007) or bones. (Linder, Fanni, Parmentier, Sergent, & Phan-Tan-Luu, 1995).
Very few patents, using offal as an ingredient, have been filed in the last 20 years; for
instance a patent filed in 2001 relates the use of offal as an ingredient for cheaper sausages or
burger patties (Van, 2001). But, in spite of the advances generated in food processing, most
of the patents found in the literature are from the 1960's and 1970's.

At a very basic, low value level, protein extracts from offal can be used as extender in 140 processed meat products (Heinz & Hautzinger, 2007). Recently, the use of pork tongue and 141 142 pork head meat as sausage ingredients has been reported, with no significant differences in some quality characteristics compared to controls (Choi, Hwang, et al., 2016). Similar results 143 were observed when the final products were hamburger patties (Choi, Jeon, et al., 2016). 144 However, considering the functional properties of many of these proteins, there is a strong 145 argument to be made for capitalising on this high-value potential to more fully support a 146 sustainable use of natural resources. It is important to also remember that, as well as 147 providing proteins with nutritive value or good techno-functional characteristics, these co-148 products are well established as being rich in minerals and vitamins (Mullen & Álvarez, 149 150 2016).

151

152 **2.2 Nutritional quality of co-products**

153 Nutritional quality of proteins has traditionally been defined based on the amino acid profile, and the ability to provide specific patterns of amino acids for protein synthesis as measured 154 155 by animal growth or nitrogen balance in humans (Millward, Layman, Tomé, & Schaafsma, 2008). However, a more current definition describes this term as the ability of the protein to 156 157 achieve specific metabolic activities. Currently, the objective of protein quality evaluation is determining the ability of a protein to reach maintenance needs, along with special needs as 158 growth, pregnancy or lactation (WHO, 2007). The up-to-date method approved for protein 159 quality determination is the protein digestibility-corrected amino acid score (PDCAAS). In 160 this method the single most limiting amino acid is identified, when compared to an 161 appropriate reference pattern, which defines the amino acid score. According to FAO, 162 Lysine, Methionine and Tryptophan are the most limiting amino acids in poor protein 163 sources. Table 1 shows the amount of these three amino acids, the total protein content and 164 the percentage of essential amino acids in a variety of offal from pork, beef and lamb. It can 165 be seen that offal, generally, is a good source of essential (namely Ile, Leu, Lys, Met, Phe, 166 Thr, Trp, Vali) and limiting amino acids. However, offal rich in connective tissue as ears, 167

feet, or lips, present a lowest protein quality, since connective tissue is mainly composed by
glycine, proline and alanine, which are not essential amino acids (Mullen & Álvarez, 2016).
This lack of essential amino acids can be overcome by tailor blending ingredients to achieve

- a balanced amino acid profile in the final product.
- Besides proteins, meat co-products can be a good source of minerals (Ca, Fe, Mg, Cu or Se) 172 or vitamins (niacin, Vit. B12, folate or Vit. C). Heme iron, mainly provided by meat and fish, 173 is more absorbable that non-heme iron (<15%); and also, although the mechanism is not well 174 defined, heme iron is more absorbed in the presence of meat, what is known as the "meat 175 176 factor" (López & Martos, 2004). Thus, it is worthy to explore if protein derived from offal can also increase the bioavailability of iron, since they are a good iron source. Finally, the 177 offal fat content can vary from less than 1% in blood, up to 17% in tongue; with high 178 contents of saturated fatty acids. Among fats, it is important to draw attention to cholesterol 179 when referring to co-products, as some have high levels of this compound, as for example 180 kidneys, liver or brain (Mullen & Álvarez, 2016). 181
- 182 It is clear that meat co-products are rich sources of macro and micro-nutrients. Both these raw 183 materials and various extracts generated from them can be incorporated in products for 184 targeted end-users such as the elderly, sports-active or dairy intolerant consumers (Montero 185 Castillo, Ligardo, Alejandro, González, & Cristina, 2015).
- 186

187 **2.3 Bioactive compounds**

Bioactive compounds cover both peptides and other biomolecules which exert a 188 189 physiological benefit on consumption. Bioactive peptides or biopeptides are defined as short sequences of 2 to 30 amino acids that exert this effect. These sequences are encrypted in the 190 191 parent protein and must be released from the parent protein to have an effect (Di Bernardini, Harnedy, et al., 2011; Ryan, Ross, Bolton, Fitzgerald, & Stanton, 2011). In order to release 192 193 these encrypted amino acid sequences, three different techniques are employed: enzymatic hydrolysis by means of commercial proteases (Sarmadi & Ismail, 2010); proteolytic 194 microorganism; or after fermentation or aging processes (Escudero et al., 2013). After any of 195 these processes, hundreds of peptides are generated, but only a few of them show biological 196 197 activity. For this reason, a further purification step must be carried out. Techniques such as membrane filtration, chromatography or differential precipitation are commonly employed 198 (Power, Fernández, Norris, Riera, & FitzGerald, 2014). A different approach is to firstly 199 isolate the proteins, and then, carry out the hydrolysis step, which makes the further isolation 200 201 process easier. In order to optimise the screening process for bioactive peptide searching, in *silico* tools can be employed (Fu et al., 2016): these are useful to find the right protein/enzyme combination. As a complementary tool, the peptides of interest can be synthesized at lab scale, in such way further studies of toxicology, functionality, absorption, bioavailability or dose response can be performed (Korhonen & Pihlanto, 2006).

As discussed, the main component of meat co-products, on a dry basis, is protein; thus, these 206 by-products are a potential source of bioactive peptides which can be extracted, purified and 207 commercialised as a very high-added value product (Lemes et al., 2016). An ideal scenario is 208 one where proteins with key techno-functional characteristics are first extracted from the raw 209 210 material and the residual proteinaceous material subjected to biopeptide generation. An example of this is seen on the ReValueProtein project (Álvarez, Lynch, Drummond, & 211 Mullen, 2016) where techno-functional proteins are extracted from lung and the non-212 solubilised material used to generate bioactive peptides. Compared to other source materials 213 relatively few studies have looked at generating bioactive peptides from meat co-products. 214

Examples of bioactive peptides obtained from blood and meat are published or reviewed by 215 (Bah, Bekhit, Carne, & McConnell, 2013; Di Bernardini, Harnedy, et al., 2011; Toldra, 216 Aristoy, Mora, & Reig, 2012). However, a more limited number of studies can be found 217 using offal as a source (Di Bernardini, Rai, et al., 2011; Lee et al., 2010; O'Sullivan, Lafarga, 218 219 Hayes, & O'Brien, 2016). The types of activities reported have included antioxidant, antimicrobial, antihypertensive, anti-diabetic, anti-hypercholesterolemic or mineral binding 220 221 (Sharma, Singh, & Rana, 2011). A number of important points need to be addressed to fully capitalise on the value of these peptides. Ensuring that the peptides reach the target organ, are 222 223 non-toxic and both bioavailable and bioactive in the body is critical. The scale of processing and the economic feasibility of same are also key considerations: which can be somewhat 224 225 mitigated by readily available, low price raw materials combined with novel hydrolysis and purification steps (Agyei & Danquah, 2011). 226

227 Other molecules of interest, as health promoting compounds include creatine, carnosine, carnitine, anserine and taurine. These naturally occurring low molecular weight molecules 228 can be found mainly in skeletal muscles and have many positive effects as: antioxidant, 229 lowering cholesterol, improved calcium absorption, preventing muscle myopathy, or 230 preventing heart disease (Williams, 2007) hence exudate and cook-out may be good sources. 231 It has also been reported that heart or liver can be a good source of these compounds 232 (Hoffmann, Waszkiewicz-Robak, & Świderski, 2010). Though not the subject of this review, 233 many other active compounds can be found in meat and may also be present in co-products, 234 235 such as: conjugated linoleic acid (CLA), chondroitin sulphate, Coenzyme Q10, espermidine, choline, lipoic acid, or glutathione (Arihara & Ohata, 2008). The main biological activitiesare summarized in Table 2.

238

239 **2.4 Biomedical applications**

Natural biomaterials such as collagen present many opportunities outside of food sector and 240 in particular can serve as raw materials in biomedical applications. The term 'collagen' 241 encompasses a family of glycoproteins that are characterised by a repeating [Glycine-X-Y]n 242 amino acid sequence (X is usually proline and Y is frequently hydroxyproline). To-date, 243 244 twenty-nine genetically distinct collagen types have been described that can be classified in four broad categories [fibrous (e.g. types I, II, III), non-fibrous (e.g. types IV, VII, XXVIII), 245 filamentous (e.g. types VI, VIII, X) and fibril associated with interrupted triple helices (e.g. 246 types IX, XII, XIV) collagens], based on their primary structure, molecular weight, charge 247 profile along the helix, length of the triple helix, size and shape of the terminal globular 248 domains. In vertebrates, collagen is the major constituent of connective tissues, comprising 249 almost 25 % of total body proteins, 75 % of the dry weight of skin, 80 % of the organic 250 matter in bones and 90 % of tendon and corneal tissues. The abundance of this extracellular 251 matrix protein, in addition to its inherent bioactivity, molecular recognition signals, 252 253 controllable mechanical and degradation properties and its ability to be reconstructed in various three-dimensional architectures (e.g. fibres, sponges, films, spheres, hydrogels), make 254 255 it an ideal raw material for biomaterials, tissue engineering and drug / gene / cell delivery applications (Abbah et al., 2015; Kielty & Grant, 2002; Sorushanova et al., Submitted; 256 257 Thomas et al., 2016).

In biomedicine, porcine and bovine skin and tendon tissues are the main source of collagen 258 259 type I, whilst collagen type II is extracted from porcine and bovine articular cartilage. Fish industry by-products (e.g. skin, scales and bones) have also been used to extract collagen for 260 261 the fabrication of biomaterials, but to a smaller extent. Independently of the source (e.g. mammalian or fish), purified collagens are extracted using primarily dilute acids with or 262 without enzymes. Dilute acids disassociate mild intermolecular aldimine cross-links, whilst 263 proteolytic enzymes (e.g. pepsin) are effective even against stable ketoimine bonds, not only 264 increasing the yield for up to 10 times, but also lowering immune response in patients, 265 through the removal of the antigenic sequence P-determinant, located at the telo-peptide 266 regions (Bruckner & Prockop, 1981; Lynn, Yannas, & Bonfield, 2004; D. I. Zeugolis, Paul, 267 & Attenburrow, 2008). While collagen and collagen-like molecules can be produced using 268 269 various cell types, recombinant systems and peptide synthesis, these technologies are still

very expensive, of low yield, and produce unstable collagens. Compared to animal extracted
collagen they have increased susceptibility to proteolytic degradation, restricting their use in
biomedicine (Browne, Zeugolis, & Pandit, 2013; Fichard, Tillet, Delacoux, Garrone, &
Ruggiero, 1997; Frischholz et al., 1998; Myllyharju et al., 1997; Vuorela, Myllyharju, Nissi,
Pihlajaniemi, & Kivirikko, 1997). All these factors support continued expansion of the use of
collagen in the biomedicine, food and cosmetic industry.

276

277 2.5 Pet food and animal feed

278 As nutritious sources of protein, fat and micronutrients many products are highly suitable for use in companion animal diets (Cramer et al., 2007; Rivera, 1998). As rich sources of protein 279 and as a result of regular use in pet food formulations they have been considered a major 280 contributor to the growth of the pet food industry (Corbin, 1992). In canine diets they provide 281 good sources of digestible nutrients in particular proteins and fat (Murray et al 1997). 282 Palatability is an important consideration in pet foods formulation. Co-products can also 283 serve as palatability enhancers (Boskot, 2009) which will have a direct impact on product 284 acceptability. The pet food market has seen strong growth with pet humanisation and 285 premiumisation being considered key drivers. As well as being incorporated into petfoods 286 287 these products are also sold in pet stores as treat type products e.g. bone, hides, ears etc. Animal feeds play an important role in the global food industry as they influence greatly the 288 289 ability to produce animals in an economical way. In particular, following rendering, coproducts arising from meat processing form an important constituent in animal feeds 290 291 (Jayathilakan, Sultana, Radhakrishna, & Bawa, 2012).

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293 **3. Market overview**

The categorization and utilization of a secondary stream as a co-product is dependent on several factors such as local and export market demands, existing profitable applications and a favourable and feasible commercialization route. Broadly speaking, depending on local cultural habits, the market for 5th quarter products such as offal and blood is shared between exports, pet food, animal feed and local direct consumption.

299 Clear opportunities for exploring a variety of meat co-products have been identified over the 300 years in scientific research studies and market trend reports. However, an effective market 301 analysis of this area needs to take account not only of the latest R&D and scientific advances 302 in the recovery and reutilization of compounds from co-product materials, but also of 303 industry activities currently in place, as well as existing market and commercial scenario of the original products, since re-direction of a stream may have impacts on established supply
chains and ancillary commercialization routes. Several markets with existing or future
potential based on co-products are briefly presented below.

The protein market is generally divided by applications: food and beverage, sports nutrition, 307 animal feed, cosmetics and personal care, pharmaceuticals, etc. For the sports nutrition 308 category in particular, protein stands as the main driving force (Schmidt, 2014). In Ireland 309 and other markets, protein powders are still the most popular type of sports nutrition product, 310 but other more convenient formats such as protein ready-to-drink products and protein bars 311 312 are quickly becoming more popular (Hickey, 2014). As reported by Glanbia "the global performance nutrition market, at retail selling price, is approximately \$10.1 billion with the 313 USA accounting for 63%, and other international markets accounting for 37%" 314 (Glanbia.com, 2017). Although whey proteins still dominate this market, due to 315 diversification of products and applications, blends incorporating other sources of proteins 316 317 are increasingly gaining ground.

The market size for functional foods may be difficult to measure, as the exact definition of a functional food can vary between regions, countries or even companies. Nonetheless, some analysts value it at up to USD 190 Bn worldwide, showing a growing of 6–25% per year, and with the same perspective for the next 5 years (Calvo, Martorell, Genovés, & Gosálbez, 2016).

323 The global collagen market is forecast to expand at a 9.4 % compound annual growth rate from 2015 to 2023 and the market is projected to rise to US\$ 9.37 billion by 2023 ("Collagen 324 325 market - Global industry analysis, size, share, growth, trends, and forecast 2015 - 2023," Transparent Market Research Report 2016). Currently, around 60% is due to gelatine 326 327 production (Global Market Insights report, 2015). Other markets which go beyond food and beverage include the use of collagen for biomedical applications and the manufacture of 328 329 advanced biomaterials, for tissue regeneration for example, an area reported to be expanding rapidly (Global Market Insights report, 2015). Collagen has also attracted much interest as a 330 source of functional peptides, with applications in the health, cosmetics and personal care 331 332 areas.

The challenges in obtaining approval for peptides as ingredients with beneficial health properties has undoubtedly influenced the growth of this market, but for products which receive approval there are clear rewards. In 2012 while approved peptide drugs were estimated to command only about 2% of the global drug market the value represented approximately USD 20 billion (Sun, 2013).

Key markets for animal blood include the food industry, animal feed and pet food 338 applications. It is estimated that the food industry utilizes about 30% of blood produced from 339 slaughter in different forms, for example plasma used as an emulsifier or whole blood used in 340 traditional products (Bah et al., 2013). The value of the US dried animal blood plasma market 341 for animal feed in 2010 was estimated in USD 29 million per annum (Wilson, 2011). The 342 possibility of obtaining bioactive peptides from blood could lead to higher value products 343 being produced, giving access to the expanding functional and nutraceutical applications 344 market. Further efforts for the extraction and valorisation of protein from animal blood may 345 346 also increase the use of blood as a source of techno-functional compounds for food and beverage applications. 347

While the export market for some 5th quarter products offer the greatest opportunity to increase margins from edible offal, consumer perceptions in relation to food scares or disease outbreaks may have an unjustifiable impact of product values. Additionally, an understanding of specific market requirements is necessary, to satisfy demands in terms of customs and expectations.

353

354 **4. Consumer studies**

355 Humans seek novelty and variety in their diet for nutrition and health reasons and to satisfy pleasure seeking motives (Al-Shawaf, Lewis, Alley & Buss, 2015). However they are also 356 357 wary of foods with which they are unfamiliar, due to fears in relation to food safety and health. This led to the phrase "the omnivores dilemma" to reflect individual's competing 358 359 motives of neophobia and neophilia (Martins, Pelchat & Pliner, 1997). Reflecting differences 360 in cultures, meat co-products are regarded as delicacies in some countries, however most 361 consumers in developed countries are unfamiliar with meat co-products, such that many have never tasted offal and most have a very limited knowledge about how to prepare and cook it. 362 Some may even regard them as unhealthy or waste (Frewer & Gremmen, 2007). 363

Three important factors may lead to the rejection of unfamiliar food products: negative 364 sensory properties, harmful consequences and "ideational" factors (Rozin & Fallon, 1987). 365 Henchion et al (2016) found such factors at play in consumer evaluations of a range of 366 367 product concepts that incorporated ingredients derived from offal. On the negative side, consumers were concerned with the taste and texture of such products, did not see significant 368 benefits, were disgusted with the idea that the ingredients originated in live animals and even 369 questioned if such products were edible. However this research also found that processing 370 371 could have a positive influence on acceptance through its influence on the physical form of the ingredient. Changing the physical form of the ingredient resulted in "de-animalising" the ingredient and for some consumers shifted the focus away from emotive, ideational factors, and allowing them to consider the product's characteristics and benefits. The impact of processing is however complex with increased processing resulting in negative perceptions regarding healthiness and naturalness in some contexts.

Integrating unfamiliar foods into existing culinary and dietary practices was identified as supporting acceptance in the context of insect-based foods (Looy, Dunkel, & Wood, 2014). Research by Henchion et al (2016) reports similar findings with regards to meat co-products suggesting that this strategy could also work for meat co-products. They argue that meat coproducts should be presented to consumers as ingredients rather than finished products to give them flexibility in terms of how they are integrated into consumers' routines.

The role of experts, celebrity chefs and friends in providing evidence that consumption of unfamiliar products is safe and socially acceptable is highlighted in relation to meat coproducts in some niche markets. Boutique butchers with traditional "fancy meats" and restaurants featuring nose-to-tail eating are becoming more prevalent in the UK for example, resulting in changes in the consumer landscape, with a claim by Datamonitor (2014) that "offal is officially in", due to experimentation, consciousness of value and increased availability.

390

391 **5. Legislation**

To explore and commercialise the range of available co-products of animal origin it is 392 393 necessary to understand relevant legislation, rules and regulations governing access to source materials and markets. Information and guidance on how material is to be collected, handled, 394 395 stored, processed, etc. will determine not only the value of the final product, but crucially ensure compliance for downstream uses. It can be said that legislation was developed for two 396 397 distinct groups: unusable and usable by-products (Leoci, 2014). For unusable by-products, associated regulations lay down handling and disposal rules to safeguard the environment and 398 public health. On the other hand, for by-products posing no risk to health or the environment, 399 and for which a viable commercial application exists, regulations may relate to materials' 400 401 collection, transport and further processing.

In the European Union, the term animal by-product (ABP), is reserved for products of animal origin that are not intended for human consumption and use or disposal of ABP is strictly controlled. Depending on the risk they pose, ABPs are divided into categories 1, 2 and 3, category 1 being the highest in risk. The classification of some 5th quarter products as edible or inedible can be influenced by collection and handling conditions and on local or existing
export markets. If intended for human consumption, 5th quarter products are covered under
regulations for food of animal origin, and should not be classified as ABPs.

Since the bovine spongiform encephalopathy (BSE) crises, rules for the prevention, control 409 and eradication of certain transmissible spongiform encephalopathies (TSEs) set out in 410 Regulation 999/2001, have irreversibly excluded parts of the carcass from the food chain. 411 The regulation covers the production, placing on the market and, in some cases, the export of 412 animals and animal products. For feed and pet food, for example, only category 3 material is 413 414 allowed. Directive 2008/98/EC on waste includes a definition and the main conditions which must be met by a substance to be classified as a by-product. The regulation encourages the 415 reuse, reutilization and recycling of non-hazardous by-products. Also relevant, Directive 416 2008/1/EC prohibits discharging of blood into digesters and sets down an obligation of using 417 blood for other purposes. 418

Regulation (EC) No 1069/2009 includes health rules as regards animal by-products and
derived products not intended for human consumption (uses other than for food). However,
some products are not covered by this Regulation. These include cosmetic products
(Directive 76/768/EEC); medical implants (Directive 90/385/EEC); medical devices
(Directive 2003/32/EC); In vitro diagnostic medical devices (Directive 89/79/EC); veterinary
medical products (Directive 2001/82/EC); and medicinal products (Directive 2001/83/EC).

425 For parts of the carcass intended for human consumption, the "Hygiene Package": Regulation (EC) 852/2004, Regulation (EC) 853/2004 and Regulation (EC) 854/2004), must apply, as 426 427 well as the Regulation (EC) 1169/2011 for the provision of food information to consumers. From the perspective of food hygiene regulations, offal is defined as: "fresh meat other than 428 that of the carcass, including viscera and blood" (EC) 854/2004). However, from an 429 ingredient labelling perspective, Regulation (EC) 1169/2011 restricts the definition of "meat" 430 431 to skeletal attached muscles. Any other parts of the animal must be declared separately and the meat species must be identified on the label (e.g.: "beef heart protein" or "bovine heart 432 protein"), which can have an impact on consumer perception of these food ingredients. 433

Also relevant, assuming they are utilised in products for these target audiences, Regulation (EU) No 609/2013 on food intended for infants and young children, food for special medical purposes, and total diet replacement for weight control, lays down general rules on the composition and preparation of foods that are specially designed to meet the particular nutritional requirements of the persons for whom they are intended. 439 Regulation 1333/2008 (latest consolidated version published February 2016) covers food additives and lists substances that are not considered as additives, and hence not covered by 440 this regulation, which includes blood plasma, protein hydrolysates and their salts. These are 441 also not considered meat and must be declared separately on the label. Annex II to Regulation 442 1333/2008, amended by Regulation (EU) 601/2014, lays down a list of approved food 443 additives and their conditions of use. Food additives are listed on the basis of the categories 444 of food to which they may be added to, with Category 8 covering processed and unprocessed 445 meat. Food enzymes are covered separately by Regulation (EC) 1332/2008. 446

447 In the United States, Title 9 of the Code of Federal Regulations (CFR) provides the rules governing animals, and animal products. It includes two groups of protein contributing 448 ingredients: Group 1 includes ingredients of livestock or poultry origin from muscle tissue 449 which are skeletal or found in the edible organs, with or without the accompanying and 450 overlying fat, and the portions of bone, skin, sinew, nerve, and blood vessels which normally 451 accompany the muscle tissue, not separated in the process of dressing, as well as meat by 452 products; mechanically separated (species); and poultry products. Ingredients processed by 453 hydrolysis, extraction, concentrating or drying are part of Group 2 Protein-Contributing 454 Ingredients along with any other ingredient which contributes protein. An up-to-date list of 455 456 substances considered as safe and suitable ingredients for the production of meat, poultry and egg products is published in a regularly revised directive from the USDA-FSIS, currently 457 458 Revision 39, from January, 2017. The list includes beef protein as a food grade substance approved as generally recognized as safe (GRAS), for use in meat, poultry and egg products. 459 Federal regulation (USDA-FSIS, 1999) lays down the rules and regulations for collection and 460 processing of blood when intended for human consumption. 461

Meat protein ingredients are considered non-allergenic and are generally "minimally processed". Additionally, their nomenclature is more easily recognised. Although this can make them more consumer-friendly, it may also negatively affect consumer perception. Table summarizes the current legislative scenario in Europe, regarding the use of meat coproducts.

467 **6. Relevant technologies**

Traditional methods such as chemical precipitation, ultrafiltration, extrusion, lyophilisation,
isoelectric solubilization-precipitation or solvent extraction are widely employed in order to
extract and concentrate proteins and peptides from waste materials (Galanakis, 2015).
Depending on the raw material, and the final application of the extract, one of these methods,
or a combination of them, has to be selected. However, some of the methods above

mentioned, may have a negative impact on the activity of the desired compounds, since heat
treatment (for instance evaporation, pasteurization or extrusion) or denaturing chemicals
(ethanol, strong acids or alkalis) might irreversibly affect the structure and chemical
properties of the target compounds. In the particular case of proteins and peptides, negative
processing effects are (Korhonen, Pihlanto-Leppäla, Rantamäki, & Tupasela, 1998):

- 478 i) Heat treatment: destruction of heat sensitive peptides, cross linkages, protein479 denaturation
- 480 ii) pH: racemization, destruction of amino acids, risk of oxidation
- 481 iii) Membrane filtration: change in amino acid composition

482 iv) Storage: destruction of lysine and oxidation.

These changes can lead to a loss of protein functionality, nutritive value or bioactivity; reducing the final value of the product and narrowing their field of application. In order to minimise or prevent such modifications, novel technologies are being developed. Such technologies usually overcome the negative impacts of thermal treatments and additionally, minimise significantly the amount, and hence the effect, of solvents and reagents employed for protein/peptide extraction.

Most promising and emerging technologies for protein extraction are those based on 489 490 ultrasounds (Kadam, Tiwari, Álvarez, & O'Donnell, 2015); pulsed electric fields (PEF) (Soliva-Fortuny, Balasa, Knorr, & Martín-Belloso, 2009); high hydrostatic pressures (HHP) 491 492 (Li, Zhu, Zhou, & Peng, 2012); sub-critical water hydrolysis (SWH) (Marcet, Alvarez, Paredes, & Diaz, 2016); or laser ablation (Boutinguiza et al., 2007). Among these 493 494 techniques, ultrasounds and PEF have been reported to produce negligible changes in protein composition and structure (Chandrapala, Zisu, Palmer, Kentish, & Ashokkumar, 2011; 495 496 Garde-Cerdán, Arias-Gil, Marsellés-Fontanet, Ancín-Azpilicueta, & Martín-Belloso, 2007). On the other hand, HHP and SWH, can produce drastic changes in protein structure as non-497 498 reversible unfolding or hydrolysis (Smeller, 2002). Other technologies which hold potential for these products include aqueous two phase extraction, foam mat drying, and others 499 reported by Galanakis (2012). Table 4 presents how, through the application of currently 500 available technologies, is possible to obtain high added-value products from the various co-501 502 products generated by the meat industry.

There is little evidence of the application of many of these emerging technologies for extracting value from meat co-products. More effort has been seen in optimizing the use of these emerging technologies on vegetables, seaweeds and dairy products. There clearly is a pressing need to examine the relevance of transferring these developments into the meat sector in terms for example of both recovery yields and impact on functionality of finalproduct.

509

510 **7. Conclusions**

511 Opportunities clearly exist for extracting additional value from meat processing chains. As 512 well as ensuring optimal use of such a protein rich material this approach also supports 513 sustainable practices across the sector sector and will be increasingly necessary in a world of 514 growing demand and constrained resources. Science and technology driven approaches must 515 be coupled with market, legislative and consumer knowledge for successful 516 commercialisation of the downstream products.

517

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867 Tables

Table 1: Total protein content (% wet basis), individual essential amino acids (g/100 g protein) and

869	total essential amino acids	(%EAA) in selected offal across	a number of species.
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Product	Specie	Prot. (%)	Lys	Met	Trp	% EAA
Blood	Cattle	17-18	9.7	2.4		60.6
	Porcine	18.5	9.0	2.3	1.5	58
Brain	Cattle	10.5	6.0	2.1	4.7	42.8
	Porcine	10.3	7.8	2.0	4.2	46.8
	Sheep	12.3	6.4	2.0	1.1	42.8
Heart	Cattle	17	8.2	2.6	1.1	47.1
	Porcine	17	8.3	2.6	1.2	47.7
	Sheep	18	7.5	2.2	1.1	43.8
Kidney	Cattle	15.3	6.6	2.1	1.4	45.2
	Porcine	15.4	7.2	2.1	1.3	48.0
	Sheep	18.0	6.5	2.0	1.4	43.9
Liver	Cattle	21	6.9	2.5	1.4	49.1
	Porcine	19	7.7	2.5	1.4	48.9
	Sheep	20.3	5.4	2.1	1.2	42.7
Lung	Cattle	17	7.1	2.0	0.9	41.5
	Porcine	15	7.3	1.6	0.9	37.8
	Sheep	12.5	6.5	1.8	0.9	40.6
Spleen	Cattle	19	7.2	1.8	1.0	48.0
	Porcine	17.9	7.5	1.8	1.0	41.9
	Sheep	17.2	7.7	1.9	1.1	48.5
Tongue	Cattle	17.1	7.7	2.1	0.8	45.8
	Porcine	16.3	8.2	2.2	1.2	40.2
	Sheep	15.3	7.1	2.1	1.0	40.4
Connective tissue		85%	35	07	0.0	11 07
collagen		0.570	5.5	0.7	0.0	11.7/

870 Data from Venegas, 1996.

	Compound	Concentration	Biological function
		(mg/g)	
	Conjugated	3-8 (beef fat)	Reduced risk of colorectal cancer and diabetes,
	linoleic acid		antioxidant, immunomodulatory.
	Carnosine	2.7	Antioxidant
		(pork shoulder)	
	Anserine	nd	Antioxidant, metal chelating
	L-carnitine	1.3 (beef tight)	Lowing-cholesterol, calcium absorption, prevention
		0.10.0.00	of muscle myopathy
	Glutathione	0.12-0.26	Antioxidant, defence against toxicological and
		(beef muscle)	pathological processes
	Taurine	0.77 (beef muscle)	Eye health, preventing heart disease, essential in
	~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~ ~		some stages(lactation or immune challenge)
	Coenzyme Q10	0.02 (beef muscle)	Antioxidant
	Creatine	3.5 (beef muscle)	Enhancing muscle performance
879	*Adapted from (A	rihara and Ohata (200	8))
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878 Table 2: Summary of bioactive compounds from meat sources

894	Table 3: Main Regul	ations and Directives	regarding the use	of meat pro	cessing co-produc	cts
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895 in Europe

GeneralReg.(EC) 999/2001rules for the prevention, control and eradication of certain transmissible spongiform encephalopathies in animalsGeneralDirective 2008/98/ECconcepts and definitions related to waste managament, such as definitions of waste, recycling, recovery Directive 2008/1/ECDirective 2008/1/ECintegrated pollution prevention and control health rules as regards animal by-products and derived products not intended for human consumptionFor human consumption"Hygiene Package"•Reg.(EC) 852/2004on the hygiene of foodstuffs; specific rules for products of animal origin; specific rules for products of animal origin intended for human consumptionFor human consumptionReg.(EC) 1169/2011 Reg.(EC) 1169/2011 Reg.(EC) 1169/2013Reg.(EC) 1169/2011 Reg.(EC) 1333/2008provision of food information to consumers food intended for infants and young children, food for special medical purposes, and total diet replacement for weight controlReg.(EC) 1333/2008 Directive 2003/32/ECenzymes used in food Directive 2003/32/ECDirective 2003/32/EC Directive 2001/83/EECmedical implants Directive 2001/82/ECDirective 2001/82/EC Directive 2001/83/ECn vitro diagnostic medical devices Vet medical products	Scope	Reference	Short description
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		Directive 2001/83/EC	medicinal products

Product	Revalorizing techniques	High added-value products
Liver	Enzymatic hydrolysis	Antioxidant peptides
Heart	Isoelectric solubilization/precipitation	High value protein with low ash, fat
		and cholesterol
	Phosphate buffer washing	Myofibrillar concentrate as
		texturizing
Skin	Collagen recovery	Barrier membrane, drug delivery,
		fibroblast scaffolds, bioengineered
		tissues
	Enzymatic hydrolysis and	Antioxidant peptides and liver
	chromatographic purification	protectors
	Collagen hydrolysis	Antioxidant activity, antimicrobial
		properties antihypertensive,
		biomimetic tissue
Blood	Enzymatic hydrolysis	Antioxidant, antibacterial,
		antihypertensive or iron-binding
		peptides
	Chemical hydrolysis	Pre-digested peptides for animal and
		pet food
	Ethanol precipitation	Purified protein as food ingredient
		Peptides and biopreserved blood
	High pressure treatment	Amino acid and peptides production
	Subcritical water hydrolysis	
Bone	Subcritical water	Hydroxyapatite and collagen
	Alkaline extraction	New kind of sausages
Lung	Isoelectric solubilization -	Protein concentrates with good
U	precipitation and membrane filtration	functional properties
Feathers	Keratinolytic bacteria fermentation	Keratinolytic protease production,
and hair	and enzymatic hydrolysis	culture medium, soil assessment.
	5 5 5	separation membranes

Table 4: Examples of revalorising techniques for adding value to several meat co-products.

907 *Adapted from Mullen, Álvarez, Pojić, Hadnadev, & Papageorgiou, 2015.