

Elsevier Editorial System(tm) for Food

Research International

Manuscript Draft

Manuscript Number: FOODRES-D-19-02381R2

Title: Microencapsulation of Vitamins in Food Applications to Prevent Losses in Processing and Storage: A Review

Article Type: Review Article

Keywords: Microencapsulation; Vitamins; Food processing; Food Storage

Corresponding Author: Dr. Jibin He, PhD

Corresponding Author's Institution: Teesside University

First Author: SHABANA PRAVEEN DHAKAL, MSc

Order of Authors: SHABANA PRAVEEN DHAKAL, MSc; Jibin He, PhD

Abstract: The food consumption trends have long since shifted from demanding simple calories and essential nutrients in order to support the basic human body functions to demanding a balanced nutrition supply in order to achieve optimal health. Vitamins play a vital role in human health, yet are often lost or destroyed during food processing before they reach consumers, as they are highly prone to degradation by environmental factors. Microencapsulation technology is a technology aiming to protect sensitive compounds from environmental elements. It is widely used in pharmaceutical and cosmetic industries but its application in food production are few. This article reviews microencapsulation studies conducted in food with a specific focus on protecting vitamins from processing and stage losses. We found that although current technologies have the potential to create vitamin microcapsules, none could meet all the criteria for a successful product. To develop suitable vitamin microcapsules which are processing stable, digestible and safe to consume, we recommend further studies to focus on seeking and developing porous and thermal stable carbohydrate or protein based wall materials derived from natural food ingredients.

- Principles of microencapsulation technologies, selection of wall materials, and release mechanisms are reviewed.
- The impact of environmental factors on vitamin stability are complicated and not as established as commonly believed.
- Existing technologies cannot produce vitamin microcapsules that are processing stable, digestible and safe to consume, mainly due to the lack of thermal stable wall materials that are digestible.
- Further studies should focus on developing thermal stable carbohydrate or protein based wall materials derived from natural food ingredients.

1 Introduction

2 ‘Vitamins’ are defined as a group of micronutrients that cannot be synthesised by a human
3 body (Ottaway, 1993). Thirteen identified compounds are further classified into fat-soluble
4 (A, D, E and K) and water-soluble vitamins (B₁, B₂, B₃, B₅, B₆, B₇, B₉, B₁₂ and C). Vitamins
5 play vital roles in human life with their specific functions to regulate metabolic and cellular
6 functions, promote health, reproduction and growth, and prevent diseases (Lešková et al.,
7 2006). Deficiency of vitamins can lead to severe diseases, such as scurvy, beriberi and night
8 blindness. To ensure the optimal intake of vitamins for the total population, more than 60
9 countries around the world have implemented fortification plans to fortify staple foods with
10 vitamins (Teleki, Hitzfeld, & Eggersdorfer, 2013). In 2006, the World Health Organization
11 (WHO) and Food and Agriculture Organization (FAO) published micronutrient fortification
12 guidelines to ensure the best practice in fortification plans (Allen, de Benoist, Dary, &
13 Hurrell, 2006).

14 Vitamins are chemically reactive compounds. During food processing and storage, the
15 stability of many vitamins is affected by chemical and physical factors such as light,
16 temperature, pH and oxygen levels (Gregory, 2008), leading to significant nutrient losses in
17 the end products such as canned products (Rickman, Barrett, & Bruhn, 2007). To prevent
18 these losses, food industries aspire to seek methods that can protect vitamins in production
19 and storage. Microencapsulation is a modern technology which integrates bioactive
20 substances (vitamins, enzyme, phenols, molecules, cells) in the specific coating in order to
21 protect them from the environmental elements. The process of encapsulating can be carried
22 out in gases, liquid droplets or small solid particle form in micro sized (1-1000 µm) coatings
23 (Nazzaro, Orlando, Fratianni, & Coppola, 2012).

24 Microencapsulation is a widely explored subject in many fields. In food, most current studies
25 are focusing on bioactive compounds such as antioxidants, antimicrobial compounds and
26 bacteria. In this work, we aim to review technologies that can be scaled up easily and discuss
27 the possibility of utilising them to produce processing stable vitamins in order to create long
28 shelf life products with better nutritional values.

29 **Microencapsulation**

30 Microencapsulation is a process of encasing micron-sized materials in a polymeric shell. The
31 material to be encapsulated may be referred to as the internal phase, core material, fill,
32 payload phase or active agent, whilst the encapsulating material may be referred to as
33 membrane, carrier material, coating, shell, matrix, external phase or wall material (Zuidam &
34 Shimoni, 2010). The microcapsule implies core-wall structure and can be categorised as
35 reservoir and matrix systems. In the reservoir, the core is coated with the wall material, whilst
36 in a matrix system, the core material is embedded within a continuous network of the matrix
37 material lacking a distinctive external wall (Augustin & Hemar, 2009; Singh, Hemant, Ram,
38 & Shivakumar, 2010). The applied pressure can lead to the breakage of the reservoir capsules
39 and hence, the release of its contents. In a matrix type, the active agent is dispersed over the
40 carrier material either in the form of small droplets or more homogenously (Zuidam &
41 Shimoni, 2010). The capsules can be mononuclear where one core material is encapsulated
42 by a shell, or can be aggregated where a capsule consists of multiple-cores (Nazzaro et al.,
43 2012) or can be multi-layered (Augustin & Hemar, 2009). The capsules can be spherical,
44 cylindrical, oval, and irregular shaped (Zuidam & Shimoni, 2010).

45 The study and application of microencapsulation initially started in the 1930s to make
46 carbonless copy papers (Fanger, 1974). Since then it was extensively employed in many
47 industrial applications including pharmaceuticals, cosmetics and textiles, its application in the
48 food industry has been recognised relatively recently. Microencapsulation is considered a
49 costly process; hence the requirement must be established primarily (Poncelet, 2006).
50 Microencapsulation in food aims to preserve valuable and sensitive components by protecting
51 them from adverse environmental conditions, to prolong shelf-life and to ensure delivery of
52 the encapsulated material to the target location at the desired time (Augustin & Hemar, 2009;
53 Sobel, Versic, & Gaonkar, 2014). For example, protection of polyunsaturated fatty acids and
54 vitamins from oxidation, and protection of probiotics in gastric transit (Poncelet, 2006).
55 Moreover, microencapsulation has the potential to protect sensitive compounds, preserve
56 desirable flavours and aromas or mask unpleasant appearance (Huang, Yu, & Ru, 2010).

57 Microencapsulating technique can facilitate the convenient handling of materials by allowing
58 the conversion of a liquid food material into a solid or free-flowing powder (Sobel et al.,
59 2014) or vice versa (Huang et al., 2010). An additional benefit of microencapsulation is to

60 provide a suitable concentration and consistent dispersion of the core material. These
61 functionalities have been widely exploited in drug and vaccine delivery in the pharmaceutical
62 sectors and are increasingly being used to add value to novel food products in the food
63 industry (Olive Li, Dueik González, & Diosady, 2014).

64 **Wall materials**

65 A careful selection of wall material is vital as it imposes the encapsulation efficiency and
66 stability of the microcapsule. The most appropriate encapsulate material for food applications
67 should have the following properties: not reactive with the core, able to seal and retain the
68 core intact inside the capsule, able to effectively protect the core against adverse
69 environmental conditions, lack any unpleasant taste, edible and non-toxic, and economically
70 viable (McClements, Decker, & Weiss, 2007; Nazzaro et al., 2012).

71 Generally, the ideal wall material should have non-compatible physical properties to the core,
72 such as hydrophobic wall material and hydrophilic core or vice-a-versa. Table 1 summarises
73 the widely used wall materials in food. The properties of these wall materials were reviewed
74 in details by Wandrey, Bartkowiak, and Harding (2010) and Sobel et al. (2014), and updated
75 more recently by Javier D. Hoyos-Leyva, Bello-Pérez, Alvarez-Ramirez, and Garcia (2018)
76 on modified starch wall materials and by Reineccius (2019) on protein wall materials.
77 Attempts at using combinations of wall materials were carried out with various levels of
78 success (Santana, Cano-Higueta, de Oliveira, & Telis, 2016; Tontul & Topuz, 2013). There is
79 a noticeable gap in the area of utilising microbial protein and lipids as wall material.
80 Microbial proteins have been gradually recognised as a good supplement to the current
81 protein supply (Matassa, Boon, Pikaar, & Verstraete, 2016), which is contributed to by the
82 growth of the meat-free market (Parkes, 2018). Microbial lipids have been suggested as
83 alternative food sources (Bharathiraja, Sridharan, Sowmya, Yuvaraj, & Praveenkumar, 2017)
84 as well as a source for essential fatty acids (Béligon, Christophe, Fontanille, & Larroche,
85 2016). Both could be potential wall materials for microencapsulation in food.

86 **Microencapsulation Techniques**

87 Microencapsulation is a multidisciplinary area involving the knowledge and techniques of
88 physics, physical chemistry, polymer chemistry, colloid chemistry, biochemistry,
89 biotechnology, and material science. The methods to prepare microcapsules are broadly
90 divided into two major processes; physical and chemical. A liquid or a gas is normally used

91 as a suspending medium. Interfacial and in situ polymerisation, complex coacervation, and
92 solvent evaporation from emulsions technologies use liquids whereas gas is employed in the
93 fluidised-bed coating, spray-drying or spray-cooling and co-extrusion (Schrooyen, van der
94 Meer, & De Kruif, 2001). The physical methods comprise of spray chilling, spray drying,
95 fluid bed coating, pan coating, rotary disk atomization and coextrusion, whilst the chemical
96 process comprises of phase separation, simple and complex coacervation and interfacial
97 polymerisation. The majority of the processes involve the production of droplets of the active
98 core materials (in gas, liquid or powder form) and subsequently surrounding the droplets by
99 encapsulating materials in a gas or liquid phase with the exception of the preparation of
100 liposomes, melt extrudates, and the use of natural encapsulates like yeast cells (Zuidam &
101 Shimoni, 2010). Oxley (2014) categorised these techniques into four general categories:
102 atomisation, spray coating and extrusion are physical based processes, while emulsion-based
103 processes are chemical based. The following content focuses on techniques commonly used
104 for vitamin encapsulation processes.

105 **Spray drying**

106 Spray drying is the oldest microencapsulation technique, which is believed to have been
107 originally used in the 1930s to encapsulate flavours in gum acacia (Shahidi & Han, 1993). It
108 is the most commonly used microencapsulation method due to its ability to evaporate
109 moisture rapidly and maintain a low temperature in the particles. This method is widely
110 employed for the encapsulation of oils, flavours and fragrances, and considered suitable to
111 encapsulate a number of liquids and solids materials. Spray-dried lactose was introduced to
112 the pharmaceutical market in the 1960s and used as an excipient for direct compression
113 (Gunsel & Lachman, 1963), solid dispersions (Takahashi, Chen, Okamoto, & Danjo, 2005)
114 and more recently to manufacture dry powders (White et al., 2005).

115 The general steps for microencapsulation in the spray drying method involves
116 homogenisation of the core materials and wall materials to create an emulsion followed by
117 atomisation into the drying chamber (Bakry et al., 2016). Typically, wall materials used in the
118 spray-drying method are polysaccharides and proteins (S. J. Lee & Wong, 2014).
119 Incorporation with other techniques can increase the range of applications of spray drying.
120 Moraes et al. (2013) investigated the production of proliposomes incorporating β -carotene by
121 spray-drying, and Rodr  du  ez-Huezo, Pedroza-Islas, Prado-Barrag  n, Beristain, and Vernon-

122 Carter (2004) studied microencapsulation by the spray-drying of multiple emulsions
123 containing carotenoids.

124 **Spray chilling/cooling**

125 In spray chilling, a homogenous mixture of core and molten lipid coating materials are
126 atomised through a sprayer into cooled air below the melting point of the lipid. The cooled or
127 chilled air solidifies the lipid around the core particles. Typically coating materials in spray-
128 cooling would be a lipid or its hydrophobic derivatives with a reasonably high melting point,
129 where a core would be hydrophilic materials such as water-soluble vitamins and enzymes
130 (Kashappa Goud H. Desai & Jin Park, 2005). Some of the food applications are summarised
131 by Favaro-Trindade, Okuro, and Matos Jr (2015). Other studies since then utilised the
132 technique for probiotics (Arslan-Tontul & Erbas, 2017), flavouring agent 2-acetyl-1-pyrroline
133 zinc chloride (2AP-ZnCl₂) (Yin & Cadwallader, 2018, 2019), and antioxidant
134 proanthocyanidin-rich cinnamon extract (Tulini et al., 2017).

135 Yet it is possible to use this method to create microcapsules with hydrophobic cores and a
136 hydrophobic wall. A patent (Zoet, Grandia, & Sibeijn, 2012) utilises spray chilling for
137 vitamin D using fat as the wall material, claiming the process is a cheaper alternative to
138 complex coacervation and suitable to be processed at a temperature of 80-90°C for animal
139 feed products. Only two studies were found using the same approach. Paucar et al. (2016)
140 applied the same method on vitamin D₃ and found limited protection of the core at 25°C.
141 Gamboa, Gonçalves, and Grosso (2011) tested the technique on tocopherols and found good
142 retention rate at room temperature, however, the study did not include data for control
143 (unprotected tocopherols).

144 Another approach is double layered microencapsulation. The hydrophobic core is first coated
145 with a hydrophilic coat using spray drying then coated again with a hydrophobic coat using
146 spray cooling. This approach was tested on fish oil, sacha inchi oil (Fadini et al., 2018) and
147 probiotics (Arslan-Tontul & Erbas, 2017).

148 **Coacervation**

149 Coacervation is defined as the separation of a colloidal solution into two liquid phases
150 (Burwell, 1976). There are two types of coacervation processes: simple and complex. Simple
151 coacervation uses a single polymer to form coacervates through the addition of salt or
152 dissolving agent (Timilsena, Akanbi, Khalid, Adhikari, & Barrow, 2019), the dissolving

153 agents could be alcohol or acetone (Elzoghby, Elgohary, & Kamel, 2015). Simple
154 coacervation is relatively less studied in food. Kafirin, a sorghum prolamin storage protein,
155 has been used to encapsulate tannins (Links, Taylor, Kruger, & Taylor, 2015). Sutaphanit and
156 Chitprasert (2014) used gluten coacervates for microencapsulation of holy basil essential oil.
157 Hydroxypropyl methylcellulose (HPMC) simple coacervation is used to encapsulate fish oil
158 (Wu & Xiao, 2005).

159 In complex coacervation, two or more polymer solutions with opposite charges
160 electrostatically interact with each other in water resulting in two immiscible liquid phases:
161 the polymer-rich dense phase and the polymer-poor continuous phase. The former is
162 commonly called coacervate and can be used as a coating for a wide range of core materials.
163 The first investigation on the gelatin-gum arabic system was investigated by Bungenberg de
164 Jong and Kruyt in 1929 (de Kruif, Weinbreck, & de Vries, 2004), which became a primary
165 choice for the food industry (Thies, 2003). The complex coacervation generally involves
166 several steps shown in Figure 1. The core materials are firstly dispensed in a cationic polymer
167 water solution (normally a protein). Then an anionic polymer solution is added (normally a
168 carbohydrate). With the adjustment of pH and temperature, coacervate microdroplets start
169 separating from the continuous polymer phase. When water-insoluble core materials are
170 presented, the coacervate microdroplets form to interact with the core surface and gradually
171 form a continuous shell. A crosslinking agent such as transglutaminase and glutaraldehyde
172 are often used to strengthen the gel. The complex coacervation technique can be applied to
173 many water-insoluble compounds such as fatty acids, fat soluble vitamins and flavours.
174 More detailed applications using complex coacervation in food have been reviewed by
175 Eghbal and Choudhary (2018) and Timilsena et al. (2019).

176 Although the technique is considered as complex, time-consuming and costly compared to
177 spray drying, it can offer many advantages including: mild temperature changes, high shell
178 integrity, high core loading, maximum encapsulation efficiency, and great controlled-release
179 (Gouin, 2004). Many physicochemical parameters influence the complex coacervation
180 process. Some are environmental such as pH, temperature, pressure, and stirring; others are
181 based on properties of the polymers including molecular weight, ionic strength,
182 concentration, and the protein/polysaccharide ratio. Schmitt and Turgeon (2011) considered
183 pH, ionic strength, protein to polysaccharide ratio and total biopolymer concentration as the
184 most important factors.

185 **Extrusion**

186 The earliest pioneer work using extrusion for encapsulation appeared in the late 1950s in the
187 flavour industry, known as melt injection (Castro et al., 2016). Despite the early discovery,
188 commercial extrusion for the microencapsulation of active agents is a relatively less used
189 process in the food industry (Oxley, 2014). In the extrusion method, a core material is
190 incorporated into molten carbohydrate mass, and then the mixture is forced through a series
191 of dyes and collected in a bath of dehydrating solution (Reineccius, 2019). The coating
192 hardens in the bath, then is separated, dried, and sized. Common coating materials used
193 include glucose, glucose syrup, sucrose, maltodextrin, and glycerine, which can be used as a
194 single or a mixture of compounds. The primary advantage of this method is the protection of
195 the core compounds from oxygen as they are completely isolated from the air by the wall
196 material (Gibbs, Kermasha, Alli, & Mulligan, 1999).

197 Some considered coextrusion as a separate technology to extrusion (Gouin, 2004; Oxley,
198 2014). From the engineering point of view, coextrusion is a variation of the extrusion
199 technology allowing layered construction of a material (Mount, 2011). In extrusion, a wall
200 material matrix containing cores is formed before being extruded; where in coextrusion, core
201 materials and wall materials are injected through separate pathways to the inner and outer
202 parts of a double fluid nozzle. The combined materials then are extruded through a dye and
203 form round beads due to Raleigh instabilities (Gouin, 2004). One of the advantages of
204 coextrusion is the possibilities of producing multi-layered microcapsules.

205 **Liposome Entrapment**

206 Liposome was first discovered by Alec Bangham in 1965 (Bangham, Standish, & Watkins,
207 1965). Liposomes are vesicles made of bilayer mostly composed of phospholipids, which are
208 formed as a result of dispersion of phospholipids into aqueous solution and exposure to high
209 shear rates through microfluidisation or colloid mill. Phospholipids have amphipathic
210 characteristics, which means they contain both hydrophilic and hydrophobic head groups.
211 The polar end can bind to water-soluble particles, whilst the non-polar end binds to oil-
212 soluble particles. When suspended in an aqueous solution containing water soluble core
213 molecules, the phospholipids form a sheet first as their non-polar ends align with each other,
214 then form a sphere (liposome) with their hydrophilic ends inside the sphere, trapping part of
215 the aqueous solution with the core materials. One advantage of this method is that it allows
216 the core material to maintain high water activity, where the other methods discussed above

217 deprived water from the core in the process (Mirafzali, Thomas, & Tallua, 2014). This is
218 important for applications requiring high water activity for the core to be active, such as
219 enzymes and probiotics. Phospholipids are abundant membrane lipids which naturally
220 occurred in animal cells, this potentially gave liposome another advantage on
221 biocompatibility. Due to this advantage, liposome became a useful delivery system for drugs
222 (Li et al., 2019). Many studies use liposome to encapsulate nisin, a natural antibacterial
223 compound (Lopes, Barreto Pinilla, & Brandelli, 2019; Niaz et al., 2018). Also cheese is of
224 particular interest in liposome application, as entrapped enzymes can be used to control the
225 ripening process (Kheadr, Vuillemand, & El-Deeb, 2003; Laloy, Vuillemand, Dufour, &
226 Simard, 1998). Liposome could also be used reversely to entrap a fat-suitable material to
227 create more water suitable particles. These methods included thin film hydration, freeze-
228 thaw, rapid expansion of supercritical solution (RESS) and precipitation from gas saturated
229 solution (PGSS), which were reviewed in details by Sherry, Charcosset, Fessi, and Greige-
230 Gerges (2013).

231 Various factors limit the usage of liposome in the food industry however. The first is
232 stability. As a lipid, phospholipids are prone to oxidation and hydrolysis as well as sensitive
233 to low or high pH and heat, particularly when unsaturated phospholipids are used in order to
234 reduce cost or encapsulate fat-soluble cores by trapping them between the phospholipid
235 molecules (Mirafzali et al., 2014). Typically, this results with the liposomes being required to
236 be stored at chilled temperature (Sebaaly, Jraij, Fessi, Charcosset, & Greige-Gerges, 2015)
237 and it can be difficult for them to survive processing. This however can be improved by
238 incorporating additional coatings such as pectin (Lopes, Pinilla, & Brandelli, 2017).

239 **Release Mechanism**

240 Although the main purpose of microencapsulation is to protect the core materials from
241 environmental elements, it is meaningless if the cores cannot be released or are released at the
242 wrong time or place. Therefore, the release mechanism of the core at the appropriate place
243 and time is as important as the protection capacity for microencapsulation.

244 The main factor to the release rate is associated with interactions between the carrier and the
245 core materials. Additionally, other factors that influence the core release are the volatility of
246 the core, the ratio of the core and the wall material, particle size, and viscosity of the wall
247 material (Ganza-González, Anguiano-Igea, Otero-Espinar, & Blanco Méndez, 1999).

248 Controlled release is a strategy to release bioactive agent at the target site, at the target rate
1 249 and at the target time. The release profile of a microcapsule describes its release triggers and
2
3 250 release kinetics. The mechanisms of release are described below, in practice, a combination
4
5 251 of more than one mechanism is used. Singh et al. (2010) summarised the mechanisms of
6
7 252 microcapsules delivery in drugs into diffusion, dissolution, osmosis and erosion. We consider
8
9 253 some of the release triggers such as enzymes, pH and temperature to be individual
10
11 254 mechanisms as they can be controlled independently to each other in various delivery
12
13 255 situations.

14
15
16 256 **Diffusion:** occurs especially when the microcapsule shell is intact. Dissolution fluid dissolves
17
18 257 the core by penetrating the wall material leading to leakage through the pores or interstitial
19
20 258 channels (Gunder, Lippold, & Lippold, 1995). Early work suggested the final release relies
21
22 259 on the penetration rate of microcapsule by dissolution fluid, the dissolving rate of the core
23
24 260 into the dissolution fluid, and the leakage and dispersion rate of the core (Higuchi, 1963). The
25
26 261 core can be released with or without shell swelling (Shahidi & Han, 1993). In practice, the
27
28 262 rate of diffusion is also influenced by the type of core material and wall material, the
29
30 263 type/size/shape of the microcapsule (Siepmann & Siepmann, 2012).

31
32 264 **Dissolution:** when the coating gradually dissolves, the cores are released. Therefore the
33
34 265 release rate is determined by the dissolution rate (Gunder et al., 1995), which is influenced by
35
36 266 the thickness of the coating and its solubility in the dissolution fluid. An example of this
37
38 267 mechanism is the release of microencapsulated coffee flavours in contact with water, whilst
39
40 268 the flavour can be protected from heat, light and oxidation in the dry state (Frascareli, Silva,
41
42 269 Tonon, & Hubinger, 2012).

43
44 270 **Osmosis:** the polymer shell of the microcapsule resembles a semi permeable membrane. The
45
46 271 osmotic pressure difference between inside and outside of the microcapsule lead to the
47
48 272 movement of the core through small pores in the shell (Singh et al., 2010).

49
50 273 **Biodegradation:** in a biodegradation release mechanism, enzymes such as proteases and
51
52 274 lipases hydrolyse proteins or lipids, respectively, leading to release of the core. Hickey,
53
54 275 Kilcawley, Beresford, and Wilkinson (2007) discovered that enzymatic degradation enhances
55
56 276 the ripening of cheddar cheese by 50% faster than the conventional ripening process.

277 **pH:** the principle of the pH release profile is to alter the solubility of the wall material by
278 changing the pH of the environment. For instance, microencapsulation based on the pH
279 release mechanism enables the release of riboflavin in the intestine where the pH is alkaline,
280 skipping through the acidic pH of the stomach (de Farias et al., 2018).

281 **Temperature:** alteration in temperature enables core release. This mechanism is based on
282 two concepts: temperature-sensitive release and fusion-activated release. The former is based
283 on the expansion or collapse of the wall material at a critical temperature, whereas the latter
284 one is based on the melting of the wall material when temperature rises (da Silva et al., 2014).

285 **Pressure:** a release is triggered when pressure is applied to the core material. Releasing
286 flavours from chewing gum during mastication is an example of this mechanism (Wong, Yu,
287 Curran, & Zhou, 2009).

288 **Interests and challenges**

289 Microencapsulation is widely studied and applied in pharmaceuticals and has become a
290 significant interest in the food industry. The technology also found its applications in
291 cosmetic (Martins, Barreiro, Coelho, & Rodrigues, 2014), metal coating (Wazarkar et al.,
292 2016) and textile (Arantzazu, Marina, Ana, & Maria, 2018). For food applications, there are
293 many challenges in developing efficient and suitable microencapsulation systems. In our
294 opinion, the greatest challenge is to find a suitable material and method to create coatings
295 able to survive aggressive food processes. Although many novel processing technologies that
296 are less aggressive are emerging to the food industry, most commonly used technologies still
297 involve a large amount of mechanical operation and heat treatments at high temperature.
298 Most microencapsulation studies only considered the stability of the microcapsules in storage
299 temperatures, where few tested their stability under processing conditions. On the other side
300 of this challenge, a microcapsule is only useful if it can be released at the desired time and
301 location, which is in the human digestion system for most food applications. Therefore, the
302 release mechanism of the microcapsules must be effective. To add to the difficulty, the
303 selection of materials is restricted for food applications as well as they must be suitable for
304 human consumption, and the economic feasibility for large scale production must be
305 considered. Despite these challenges, the high potential of microencapsulation led to a
306 substantial rise of scientific interests in this field. A quick survey using “microencapsulation”

307 as the keyword for topics on Web of Science shows the number of publications rose from 147
308 in 1998 to 860 in 2018.

309 **Vitamins in food system**

310 Due to an increase in popularity of functional food, food industries have begun to add
311 vitamins to food and beverages. However, many of the micronutrients including vitamins
312 have limited stability and are highly reactive with many other food compounds. During food
313 processing and storage periods, the stability of many vitamins can be affected by various
314 chemical and physical parameters such as light, temperature, and pH. Moreover, the flavour
315 of vitamins can be strong and even unpleasant, making the food less appealing (Delompré,
316 Guichard, Briand, & Salles, 2019). Suitable microencapsulation methods can contribute to
317 the protection of these sensitive compounds by eliminating or limiting nutrient loss, masking
318 their flavour and appearance, and allowing controlled release of the compounds (Huang et al.,
319 2010)

320 **Vitamins and strategies to prevent their deficiency**

321 In 1912, Casimir Funk discovered ‘vital amine’ which was later known as vitamin (Teleki et
322 al., 2013). The isolation of all 13 major vitamins had been accomplished and their synthesis
323 had begun by the mid-20th century (Mozaffarian, Rosenberg, & Uauy, 2018). Vitamins take
324 part in various biochemical functions in the human body, hence they are classed as essential
325 micronutrients. For example, some B-group vitamins perform as coenzymes to provide
326 energy, vitamin C participates in numerous physiological processes, such as the immune
327 response and iron absorption (Bender, 2009). The majority of the vitamins have to be
328 supplied through diet as they cannot be synthesised in the human body with the exception of
329 some amounts of niacin and vitamin D (Drouin, Godin, & Pagé, 2011).

330 A diet deprived of vitamins can result in various deficiency diseases like pernicious anaemia,
331 pellagra, beriberi, scurvy, dermatitis, ariboflavinosis, enteritis, and many more (Gregory,
332 2008). Only an insignificant amount of water soluble vitamins, with the exception of vitamin
333 B₆ and B₁₂, are stored in the body (Bender, 2009). Fat-soluble vitamins are stored in lipid
334 fractions such as adipose tissues, however, the absorption rate can be poor such as in the case
335 of vitamin E (Barasi, 2003). The micronutrient deficiency is often unnoticed, therefore often
336 termed ‘hidden hunger’. Micronutrient deficiencies are most predominant in developing
337 countries due to their diet lacking variety (Ritchie & Roser, 2020). Many strategies have been

338 proposed to prevent vitamin deficiencies, broadly speaking, they can be categorised into
1 339 either supplementation or food-based strategies (Gibson, 2011). Dietary diversification is a
2 340 food-based strategy encouraging and helping the population to consume more diverse types
3 341 of food in order to achieve a more balanced diet. It is ideal in theory as there is no
4 342 modification of the food nor additional risks from taking supplements. However, it requires
5 343 social and even cultural changes in the community, which can be hindered by economic,
6 344 social and cultural barriers (Akhtar, Ismail, Atukorala, & Arlappa, 2013; Nyantakyi-
7 345 Frimpong, 2017).

14 346 Supplementation is a direct approach, which has the potential to rapidly correct the
15 347 micronutrients status in the human body. The supplement industry took this opportunity to
16 348 market single and multivitamins as a means to protect against their deficiency (Mozaffarian
17 349 et al., 2018). However, the effectiveness of the vitamin supplement is widely debatable
18 350 within the science community. Some find they might be effective in a particular case, such as
19 351 vitamin D on reducing the onset of preeclampsia (Fu, Ma, Liu, Wang, & Guo, 2018) and
20 352 multivitamin supplementation on reducing the risk of autism spectrum disorder in children
21 353 (Guo, Li, Zhai, & Ding, 2019). Others argued that they made no difference or even carried
22 354 higher risks than a balanced diet (Combet & Buckton, 2019; Hamishehkar, Ranjdoost,
23 355 Asgharian, Mahmoodpoor, & Sanaie, 2016; Jenkins et al., 2018; Tucker, Safadi, & Friedman,
24 356 2015). Fortification is a supplementation strategy of adding nutrients to food at levels higher
25 357 than originally found. This is to either restore micronutrients lost during processing, provide
26 358 a more nutritionally balanced food than their original form, or provide a special purpose food
27 359 designed to perform a specific function (Allen et al., 2006). Food fortification has been
28 360 widely used for many decades and has been considered as one of the most effective public
29 361 health measures to prevent micronutrient deficiency disorders in developed countries (Teleki
30 362 et al., 2013). However, the efficiency of vitamin fortification often is not maximised due to
31 363 the vitamin instability in processing and storage. To overcome this issue, microencapsulation
32 364 can be a good solution to enhance the effectiveness of the fortification.

365 **Vitamin loss in processing and storage**

366 Vitamins are heterogenous compounds with different chemical and physical properties. A
367 common attribute they share is their instability in food, whether they are occurring naturally
368 or synthetically added (Ottaway, 2010). Various factors can affect the stability of vitamins in

369 storage (Figure 2). Individual vitamin's response to these factors varies. Several textbooks
370 and reviews provided tables overviewing the stability or sensitivity of vitamins to main
371 environmental factors including light, oxygen, heat, moisture/humidity, and pH. The sources
372 of these tables are mostly from early publications such as Harris (1988) adopted by Gregory
373 (2008). As Gregory rightly advised, the conclusions of these early works are oversimplified.
374 Therefore caution is needed when using these tables. We conducted a mini review using
375 "stability" and "Vitamin A" as keywords, aiming to seek evidence to support the indications
376 in Harris's table. Strong evidence was found supporting light, temperature and oxygen as
377 primary causes of vitamin A loss in storage (Fávaro, Iha, Mazzi, Fávaro, & Bianchi, 2011;
378 Ferguson, Emery, Price-Davies, & Cosslett, 2014; Gupta, Arora, Sharma, & Sharma, 2019;
379 Hemery et al., 2015). Hemery et al. (2018) also found some evidence suggesting higher
380 humidity could also have a negative effect however only when samples were packed in paper
381 bags which has a limited oxygen barrier capacity, but not in PET/aluminium bags. We were
382 able to find only one study which directly addressed the effect of pH on vitamin A stability.
383 Carlotti, Rossatto, Gallarate, Trotta, and Debernardi (2004) tested the stability of retinyl
384 palmitate in hydroxy ethyl cellulose hydrogels under UV light and found natural pH (from
385 5.6 to 7.0) had no impact on retinyl palmitate's photostability, however, lower and higher pH
386 (4.0 and 8.0) decreased the photostability of retinyl palmitate, which is contrary to Harris'
387 indication that vitamin A is stable in an alkaline environment. There are also other factors not
388 presented on the table such as the presence of fat/lipid and the type of fat/lipid could impact
389 the stability of vitamin A. Lau, Kakuda, and Arnott (1986) and Moccand et al. (2016) both
390 found fat/lipid has a protective effect on vitamin A against degradation, and Moccand et al.
391 (2016) found a more saturated fat/lipid has a stronger protective effect. This mini exercise
392 demonstrated that the environmental influences on vitamin stability are complicated and
393 understudied despite vitamins having been discovered for over 100 years.

394 Loss of vitamins occurring during cooking processes have been reported in many early food
395 and nutrition studies. Ascorbic acid and thiamine in vegetables and fruits are the most
396 susceptible vitamins to degradation during cooking (Harris, 1988). Scott and Bishop (1986)
397 found pasteurisation resulted in a 25% loss of vitamin C in raw milk, which increased to 30%
398 and up to 60% during UHT and sterilisation respectively. Blanching is a common pre-process
399 step used to alter the properties of raw materials before main processes such as baking, frying
400 and canning. Sungpuag, Tangchitpianvit, Chittchang, and Wasantwisut (1999) reported

1 401 blanching of vegetables caused 7-11% loss of retinol, while boiling resulted in the highest
2 402 loss of 43% in cooked traditional Thai food.

3
4 403 Regardless of whether freezing is considered as a processing step or storage step, the loss of
5 404 nutrients in freezing and thawing is well established. Loss of thiamine in meat and ascorbic
6 405 acid in fruits and vegetables were found to be directly proportional to the frozen storage time
7 406 (Kramer, 1977). Adams and Erdman (1988) reported significant vitamin leaches during
8 407 thawing. Although most of the vitamins are chemically stable in frozen fruits and vegetables
9 408 for periods of up to a year, losses of vitamin C have been found to occur at temperatures as
10 409 low as -23°C (Ottaway, 2010). Refrigeration is regarded as a good storage method for
11 410 retaining nutrients in food because no ice crystallisation occurs and there are limited
12 411 chemical/biological changes due to the relatively low temperature. However the chilled
13 412 temperature combined with unsuitable humidity still can result in wilting of fruits and
14 413 vegetables, which lead to the loss of vitamins (Severi, Bedogni, Manzieri, Poli, & Battistini,
15 414 1997).

16 415 Several modern commercial nutrition databases such as “SELF Nutrition Data” includes
17 416 information on nutrition retention or loss in processes. Most of these are based on a dataset
18 417 from the NLG project carried out in Europe (Nutrient Losses and Gains in the preparation of
19 418 foods project 1983-1993) (Bergström, 1994) or USDA Table of Nutrient Retention Factors
20 419 produced by United States Department of Agriculture (USDA) (Nutrient Data Lab, 2017).
21 420 Both datasets are available for the public.

22 421 **Microencapsulation application for vitamins**

23 422 **Previous studies**

24 423 Microencapsulation is relatively new in the food industry, and much of the research is
25 424 focussed on flavours and colouring. A few reviews have been published on individual
26 425 vitamins including vitamin A (Gonçalves, Estevinho, & Rocha, 2016; Loveday & Singh,
27 426 2008) and vitamin C (Abbas, Da Wei, Hayat, & Xiaoming, 2012). Perhaps it is because
28 427 Vitamin C is known as one of the most perishable vitamins in processing and storage, whilst
29 428 vitamin A is widely used in food and beverage formulation as a colouring, an antioxidant or a
30 429 nutrient. Table 2 summarises encapsulation methods and wall material used, product particle
31 430 size and their release and retention behaviour from recent studies on vitamins for food
32 431 applications.

432 In the last decade, very few coating materials were known to be of food grade standard. With
433 the requirement in mind that, to protect the vitamins in food products, the delivery system
434 must be derived from permitted food ingredients via regulated food processing systems.
435 Recently, more studies in edible coating materials have been seen, particularly abundances in
436 the field of hydrocolloids. The potential of using taro starch to encapsulate ascorbic acid was
437 tested by J. D. Hoyos-Leyva, Chavez-Salazar, Castellanos-Galeano, Bello-Perez, and
438 Alvarez-Ramirez (2018). Although only limited encapsulation efficiency was achieved
439 (20.9%, whereas it is normally in the range of 50-60% for this technique), higher stability at
440 high humidity and temperature during the storage period was reported. Food protein-based
441 carriers for vitamin and other bioactive delivery have also been reported and reviewed (L.
442 Chen, Remondetto, & Subirade, 2006; Egan, Jacquier, Rosenberg, & Rosenberg, 2013;
443 Livney, 2010). Combinations of alginate and pectin (alg-pec) polymers were reported as a
444 suitable carrier for Folic Acid (FA) (Alborzi, Lim, & Kakuda, 2013; Madziva, Kailasapathy,
445 & Phillips, 2005, 2006; Shrestha, Arcot, & Yuliani, 2012).

446 Liposomes delivery system has been on the rise in the latest decades (Gouin, 2004), the
447 system has been used to deliver enzymes, proteins, flavours, antioxidants, and vitamins.
448 Several studies have been reported on the use of liposomes to stabilise vitamins including
449 ascorbic acid (Jiao, Wang, Yin, Xia, & Mei, 2018; Marsanasco, Márquez, Wagner, del V.
450 Alonso, & Chiaramoni, 2011; Wechtersbach, Poklar Ulrih, & Cigić, 2012), retinol (S. C. Lee,
451 Lee, Kim, & Lim, 2005; S. C. Lee et al., 2002) and α -tocopherol (Marsanasco et al., 2011).
452 Most liposome bilayers are formed by soy phosphatidylcholine (PC), but ascorbic acid and
453 pantothenic acid have been coated in dipalmitoyl-phosphatidylcholine (DPPC)
454 (Wechtersbach et al., 2012) and phospholipon 90 G (Ota et al., 2018). Retinol incorporated
455 into PC liposomes at 0.01:1- 0.05:1 retinol to PC ratios, showed almost 100% encapsulation
456 efficiency and improved stability under all conditions. In the alkaline buffer, 90% of retinol
457 was retained after 8 days of storage whereas only about 10% of free retinol was remaining
458 after 1 day (S. C. Lee et al., 2002). Effects of cholesterol (CH) addition in liposomes have
459 also been studied in vitamins. S. C. Lee et al. (2005) added CH to PC at various ratios into
460 the formation of liposomes and found CH increased the encapsulation efficiency and
461 stability. Wechtersbach et al. (2012) used DPPC with CH to form liposomes capsules of
462 ascorbic acid. They found half-lives of ascorbic acid were increased 300 fold along with
463 improved thermostability even during high temperature and low time pasteurisation (72°C).

1 464 Ota et al. (2018) stabilised pantothenic acid by capturing the vitamin into liposomes then
2 465 extruding with polysaccharide polymers (sodium alginate and pectin) and found improved
3 466 stability. After 14 days of storage, only 20% release occurred at pH 4 and improved
4 467 thermostability was also reported.
5
6

7 468 Solid lipid nanoparticles (SLN) as a drug and cosmetic delivery system have been a growing
8 469 trend. SLNs are submicron-sized lipid emulsions consisting of solid lipid instead of liquid
9 470 lipids (Mukherjee, Ray, & Thakur, 2009). This system has several advantages when
10 471 considering food production, including being easily scalable, can be made without using
11 472 solvents and great biocompatibility (Aditya & Ko, 2015). An obvious limitation of SLN is its
12 473 suitability for encapsulating hydrophilic cores. There are very few studies which encapsulate
13 474 vitamins in SLN for food application. Salminen, Gömmel, Leuenberger, and Weiss (2016)
14 475 encapsulated vitamin A and β -carotene in SLNs made from Tristearin using *Quillaja*
15 476 *saponaria* extract and lecithin as an emulsifier. They found the oxidative stability differed
16 477 significantly between the two cores. The oxidative stability of unprotected cores was not
17 478 tested, therefore, the protective effect of the SLN was not concluded in that study. Similarly,
18 479 Mehrad, Ravanfar, Licker, Regenstein, and Abbaspourrad (2018) also entrapped β -carotene,
19 480 but in SLN composed of palmitic acid and corn oil. They have found improved chemical
20 481 stability via measurement of the loss of colour intensity.
21
22
23
24
25
26
27
28
29
30
31
32

33 482 In recent years, some studies reported on the use of nanomaterial coating such as gated
34 483 mesoporous silica particle (MSP). Pérez-Esteve et al. (2015) proposed an MSP delivery
35 484 system to encapsulate FA and reported enhanced stability and *in vitro* target release of the
36 485 system. Ruiz-Rico et al. (2017) used the delivery system in fortification of fruit juice with
37 486 MSP-FA microcapsule. Only 10% of FA was released in the juice over the testing period at
38 487 pH 3.5, however, most of FA were released at pH 7.5 in less than two hours. This indicates
39 488 the significant potential of MSP as a smart delivery system which can protect vitamins from
40 489 an acidity environment in the stomach while allowing nutrient release in the small intestine
41 490 which has a near natural pH of 6 (Fallingborg, 1999). Many may consider MSP and other
42 491 nanosilica potentially harmful, most likely linked to silicosis, a lung disease caused by
43 492 exposure to crystalline silica dust (Leung, Yu, & Chen, 2012). However the adverse health
44 493 effects of nanosilica are still unclear and under further investigation (Murugadoss et al.,
45 494 2017).
46
47
48
49
50
51
52
53
54
55
56
57
58
59

495 A delivery system that has similar release behaviours to MSP but is based on natural edible
1 496 ingredients was found by Ahmad, Qureshi, Maqsood, Gani, and Masoodi (2017). They used
2
3 497 horse chestnut starch (HCS) and β - cyclodextrin as wall materials to entrap FA using spray
4
5 498 drying. Although the encapsulation yields were still less than 60%, it indicated the potential
6
7 499 of developing a smart delivery system using natural edible ingredients.

9
10 500 Nanoencapsulation is an emerging technology (Gouin, 2004) aiming to produce smaller
11
12 501 particle size compared to the traditional microencapsulation. Nanoemulsion has been
13
14 502 frequently used, particularly in the formation of liposomes (Jiao et al., 2018; Ota et al., 2018;
15
16 503 Wechtersbach et al., 2012). The nanoemulsion process was also employed in MSP delivery
17
18 504 systems (Pérez-Esteve et al., 2015; Ruiz-Rico et al., 2017) and electrospinning (Alborzi et al.,
19
20 505 2013) to encapsulate FA, in spray drying for vitamin D (Moeller, Martin, Schrader,
21
22 506 Hoffmann, & Lorenzen, 2018) and β -carotene (Liang, Huang, Ma, Shoemaker, & Zhong,
23
24 507 2013), and in emulsion encapsulation of vitamin E (C. C. Chen & Wagner, 2004). Higher
25
26 508 encapsulation efficiencies and vitamin stabilities were reported in all cases. The benefit of
27
28 509 nanoemulsion is increased stability of emulsion due to the smaller droplet size (Jafari,
29
30 510 Assadpoor, Bhandari, & He, 2008).

31
32 511 Lešková et al. (2006) stated that retinol, folate, thiamine and vitamin C are the most unstable
33
34 512 vitamins during processing, with retention below 40%. In contrast to the abundant attempt to
35
36 513 encapsulate the other three vitamins, we found only one study on encapsulation of thiamine
37
38 514 in food. Juveriya Fathima, Fathima, Abhishek, and Khanum (2016) used phosphatidylcholine
39
40 515 as the wall material to create nanocapsules of thiamine and found good encapsulation
41
42 516 efficiency, however the retention in storage was not tested, and chloroform was used in the
43
44 517 process therefore it is not suitable for human consumption. There is also a lack of similar
45
46 518 studies on biotin, pantothenic acid, B₁₂, riboflavin, vitamin D, and Vitamin K. This might be
47
48 519 led by considerations that these vitamins are relatively stable (Lešková et al., 2006) therefore
49
50 520 no further protections are needed. However, as we have demonstrated above using vitamin A
51
52 521 as an example as well as discussed by Lešková et al., the stability/sensitivity of many
53
54 522 vitamins are understudied and not fully understood. Therefore, further work in that area and
55
56 523 subsequently potential encapsulation studies are needed. Also, protection from environmental
57
58 524 factors is only one useful aspect of microencapsulation. There could be many other
59
60 525 applications focusing on processing functionality such as coating a water soluble vitamin
61
62 526 with a hydrophobic coating in order to incorporate it better in fat rich food.

527 **Select suitable wall materials**

528 In order to discuss more effectively in this section, we set the following criteria for a wall
529 material that would be suitable to produce processing stable vitamin microcapsules:

- 530 • Able to protect cores from environmental elements in food processing including high
531 temperature, pressure, mechanical stresses, oxygen, acidity and alkaline, moisture,
532 and other reactive compounds
- 533 • Digestible in the human digestive system
- 534 • Safe for human consumption, ideally derived from natural edible ingredients
- 535 • Ideally odourless and colourless
- 536 • Good solubility in water
- 537 • Inexpensive

538 Carbohydrates, or specifically, polysaccharides are good candidates for wall materials,
539 considering their capability of forming larger porous molecular structures (L. Liu, Shen,
540 Zhang, Li, & Zhu, 2018), they are mostly from natural food ingredients and widely used in
541 food industry, and many of them are digestible in the human digestion system. Fathi, Donsi,
542 and McClements (2018) reviewed carbohydrate based microencapsulation systems and found
543 a variety of potential applications including increased acid resistance, increased solubility in
544 natural pH, and increased heat resistance. However, none of the attempts reviewed could
545 satisfy the criteria discussed above. Proteins, similarly to carbohydrates, are good candidates.
546 Fathi et al. (2018) summarised the physicochemical properties of proteins that were used as
547 wall materials for food ingredients. Collagen, gelatin and casein were suggested as having
548 high-temperature stability, where gelatin and casein also have good solubility in water.
549 However, there is a lack of experiment data to prove these concepts. For lipid based
550 encapsulations, methods like SLN are useful for improving storage stability of fat soluble
551 vitamins, however, they are not suitable for products requiring further thermal processing as
552 the solid fat coats will melt and therefore lose their protective properties. On the other hand,
553 liposome entrapment could be used as an initial coating method to improve water solubility
554 of fat soluble vitamins (Sherry et al., 2013), subsequently enabling them to be encapsulated in
555 a hydrophilic coating.

556 **Suitable encapsulation techniques**

557 Similar to the selection of wall materials, general criteria should be set when selecting
558 techniques for vitamin microencapsulation: a) avoid heat, mechanical stress, and oxygen, b)
559 safe for food production, c) scalable, d) inexpensive to operate. Different applications would
560 also require individual criteria such as the physical format of the end product, particle
561 size/structure/morphology and solubility in water/oil. For an application to incorporate
562 vitamins in thermal processed food such as cans a selection of suitable techniques is critical
563 as they are susceptible to many elements commonly occur in processes, particularly heat
564 (Bergström, 1994). Yet low temperature techniques, including spray cooling and liposome
565 entrapment, were most used when lipids/fats are wall materials which will lose their
566 protective properties when heated. Therefore, none-thermal techniques would be most
567 suitable for this application. Vacuum spray drying limited the heating and oxygen in the
568 process, it has been successful utilised to produce highly viable dry probiotics (Semyonov,
569 Ramon, & Shimoni, 2011) and bovine serum albumin microcapsules (Freitas, Merkle, &
570 Gander, 2004). Super critical fluid, particularly super critical carbon dioxide, technologies
571 have been widely used in pharmaceutical microencapsulation application (Soh & Lee, 2019).
572 It has been used to encapsulate β -carotene (de Paz, Martín, & Cocero, 2012; de Paz, Martín,
573 Duarte, & Cocero, 2012), vitamin B2 (Couto, Alvarez, & Temelli, 2017) and vitamin E
574 (Prieto & Calvo, 2017). Although these studies were not designed to produce processing-
575 stable products, the many unique properties of super critical CO₂ allow it to act as a solvent,
576 anti-solvent, solute, drying medium, and foaming agent in the microencapsulation process
577 (Soh & Lee, 2019), which provides many possibilities for designing a suitable process.
578 Membrane emulsification is a relatively new technique for producing a range of products
579 from simple oil-in-water (O/W) and water-in-oil (W/O) emulsions to complex microcapsules
580 (Vladislavljević & Williams, 2005). The general principle of membrane emulsification is to
581 force the dispersed phase through a porous membrane into a moving continuous phase
582 (Laouini, Fessi, & Charcosset, 2012; Wang et al., 2020). A variation of this technique is to
583 pre-mix the dispersed phase and continuous phase, then focus the mixture through a porous
584 membrane (Eisinaite, Juraite, Schroën, & Leskauskaite, 2016). It is characterised by its low
585 energy consumption, good control of droplet size distribution, and overall mildness of the
586 process (Laouini et al., 2012). Membrane emulsification has been used for various food
587 applications including encapsulation of polyphenols (Wang et al., 2020), vitamin E (Laouini

588 et al., 2012) and vitamin B12 (Matos, Gutiérrez, Iglesias, Coca, & Pazos, 2015). Another
589 advantage of membrane emulsification is that it has good scalability (Piacentini, Giorno,
590 Dragosavac, Vladisavljević, & Holdich, 2013) and industrial solutions are already available
591 from companies such as Micropore, UK.

592 **Design of future study**

593 One of the difficulties we experienced in this review work is that many studies did not
594 provide a full set of information in order to determine the efficiency of the tested wall
595 material/delivery system. For the purpose of producing a processing stable vitamin
596 microcapsule for food applications, the system needs to provide information on: a)
597 encapsulation efficiency; b) vitamin retention under various food processing conditions, for
598 example, retention after a F6 sterilisation process; c) validation of retention in food
599 applications; d) release rate in an environment similar to the human gastrointestinal tract. If
600 the microcapsules were produced using food safe materials and procedures, organoleptic
601 properties should also be tested. The aim of an encapsulation efficiency test is to determine
602 the quantity of the vitamin molecules captured by the encapsulation process. Test methods
603 can be designed based on differences of the physiochemical properties between the
604 microcapsule and the vitamin. For example, this could be solubility differences in water (K.
605 G. H. Desai & Park, 2005) or solubility differences in solvents (Romo-Hualde, Yetano-
606 Cunchillos, González-Ferrero, Sáiz-Abajo, & González-Navarro, 2012). Many studies we
607 have seen did not include release rate data, and very few were tested in an environment
608 similar to the human digestion system. A standardised static *in vitro* digestion module has
609 been established by (Minekus et al., 2014) and been used in many food digestibility studies
610 (T. Liu et al., 2016; Smith et al., 2015). It is fairly easy to set up and does not require
611 specialised equipment such as a Dynamic Gastrointestinal Simulator (DGS). However, the
612 information it can provide is invaluable as only the released vitamins are available for
613 absorption and meaningful for health.

614 **Conclusion**

615 Vitamins are essential for human health, and their deficiency can lead to several diseases.
616 Fortification is an important strategy to battle vitamin deficiency, however, many of the
617 vitamins are susceptible to degradation in food processing. Microencapsulation can improve
618 their stability, help them survive food processes and storage and ensure they are released in a

619 suitable part of the human digestion system in order to provide maximum benefits. The
620 stability/sensitivity of many vitamins is understudied and not fully understood. This hinders
621 the research interests in encapsulation of some vitamins which are traditionally considered as
622 stable. To achieve the optimal encapsulation efficiency and retention of the vitamin, it is
623 essential to select appropriate wall materials and methods based on the chemical and physical
624 properties of the vitamins to be encapsulated. Wall material properties and processing
625 conditions dictate the morphology, particle size and final characteristics of the encapsulated
626 vitamins. Seeking digestible wall materials which can also protect vitamin cores from food
627 processing elements is critical for creating long shelf life food products with fortified
628 vitamins.

629 **References**

- 630 Abbas, S., Da Wei, C., Hayat, K., & Xiaoming, Z. (2012). Ascorbic acid: microencapsulation
631 techniques and trends—a review. *Food Reviews International*, 28(4), 343-374.
632 doi:<https://doi.org/10.1080/87559129.2011.635390>
- 633 Adams, C. E., & Erdman, J. W. (1988). Effects of home food preparation practices on
634 nutrient content of foods. In E. Karmas & R. S. Harris (Eds.), *Nutritional Evaluation*
635 *of Food Processing* (pp. 557-605). Dordrecht: Springer Netherlands.
- 636 Aditya, N. P., & Ko, S. (2015). Solid lipid nanoparticles (SLNs): delivery vehicles for food
637 bioactives. *RSC Advances*, 5(39), 30902-30911.
638 doi:<https://doi.org/10.1039/C4RA17127F>
- 639 Ahmad, M., Qureshi, S., Maqsood, S., Gani, A., & Masoodi, F. A. (2017). Micro-
640 encapsulation of folic acid using horse chestnut starch and β -cyclodextrin:
641 Microcapsule characterization, release behavior & antioxidant potential during GI
642 tract conditions. *Food Hydrocolloids*, 66, 154-160.
643 doi:<https://doi.org/10.1016/j.foodhyd.2016.11.012>
- 644 Akhtar, S., Ismail, T., Atukorala, S., & Arlappa, N. (2013). Micronutrient deficiencies in
645 South Asia – current status and strategies. *Trends in Food Science & Technology*,
646 31(1), 55-62. doi:<https://doi.org/10.1016/j.tifs.2013.02.005>
- 647 Alborzi, S., Lim, L.-T., & Kakuda, Y. (2013). Encapsulation of folic acid and its stability in
648 sodium alginate-pectin-poly (ethylene oxide) electrospun fibres. *Journal of*
649 *Microencapsulation*, 30(1), 64-71. doi:<https://doi.org/10.3109/02652048.2012.696153>
- 650 Allen, L., de Benoist, B., Dary, O., & Hurrell, R. (2006). *WHO Guidelines on food*
651 *fortification with micronutrients*. Retrieved from
652 https://apps.who.int/iris/bitstream/handle/10665/43412/9241594012_eng.pdf?ua=1
- 653 Arantzazu, V., Marina, R., Ana, B., & Maria, C. G. (2018). Recent trends in
654 microencapsulation for smart and active innovative textile products. *Current Organic*
655 *Chemistry*, 22(12), 1237-1248.
656 doi:<http://dx.doi.org/10.2174/1385272822666180430130528>

- 657 Arslan-Tontul, S., & Erbas, M. (2017). Single and double layered microencapsulation of
1 658 probiotics by spray drying and spray chilling. *LWT - Food Science and Technology*,
2 659 *81*, 160-169. doi:<https://doi.org/10.1016/j.lwt.2017.03.060>
3
- 4 660 Augustin, M. A., & Hemar, Y. (2009). Nano- and micro-structured assemblies for
5 661 encapsulation of food ingredients. *Chemical Society Reviews*, *38*(4), 902-912.
6 662 doi:<https://doi.org/10.1039/b801739p>
7
- 8 663 Bakry, A. M., Abbas, S., Ali, B., Majeed, H., Abouelwafa, M. Y., Mousa, A., & Liang, L.
9 664 (2016). Microencapsulation of oils: a comprehensive review of benefits, techniques,
10 665 and applications. *Comprehensive Reviews in Food Science and Food Safety*, *15*(1),
11 666 143-182. doi:<https://doi.org/10.1111/1541-4337.12179>
12
- 13 667 Bangham, A. D., Standish, M. M., & Watkins, J. C. (1965). Diffusion of univalent ions
14 668 across the lamellae of swollen phospholipids. *Journal of Molecular Biology*, *13*(1),
15 669 238-IN227. doi:[https://doi.org/10.1016/S0022-2836\(65\)80093-6](https://doi.org/10.1016/S0022-2836(65)80093-6)
16
- 17 670 Barasi, M. E. (2003). Vitamins. In *Human nutrition: a health perspective*. London; New
18 671 York: Arnold.
19
- 20 672 Béliçon, V., Christophe, G., Fontanille, P., & Larroche, C. (2016). Microbial lipids as
21 673 potential source to food supplements. *Current Opinion in Food Science*, *7*, 35-42.
22 674 doi:<https://doi.org/10.1016/j.cofs.2015.10.002>
23
- 24 675 Bender, D. A. (2009). The Vitamins. In M. J. Gibney, H. H. Vorster, F. Kok, & Nutrition
25 676 Society. (Eds.), *Introduction to human nutrition* (2nd ed.). Oxford: Blackwell Science.
26
- 27 677 Bergström, L. (1994). *Nutrient losses and gains in the preparation of foods* (Rapport 32/94).
28 678 Retrieved from
29 679 [http://www.fao.org/uploads/media/Bergstroem_1994_32_Livsmedelverket_nutrient](http://www.fao.org/uploads/media/Bergstroem_1994_32_Livsmedelverket_nutrient_losses_and_gains.pdf)
30 680 [losses_and_gains.pdf](http://www.fao.org/uploads/media/Bergstroem_1994_32_Livsmedelverket_nutrient_losses_and_gains.pdf)
31
- 32 681 Bharathiraja, B., Sridharan, S., Sowmya, V., Yuvaraj, D., & Praveenkumar, R. (2017).
33 682 Microbial oil – A plausible alternate resource for food and fuel application.
34 683 *Bioresource Technology*, *233*, 423-432.
35 684 doi:<https://doi.org/10.1016/j.biortech.2017.03.006>
36
- 37 685 Burwell, R. L. (1976). Definitions and terminology. In R. L. Burwell (Ed.), *Manual of*
38 686 *symbols and terminology for physicochemical quantities and units* (pp. 74-86):
39 687 Pergamon.
40
- 41 688 Carlotti, M. E., Rossatto, V., Gallarate, M., Trotta, M., & Debernardi, F. (2004). Vitamin A
42 689 palmitate photostability and stability over time. *International Journal of Cosmetic*
43 690 *Science*, *55*(3), 233-252. doi:https://doi.org/10.1111/j.1467-2494.2004.00233_1.x
44
- 45 691 Castro, N., Durrieu, V., Raynaud, C., Rouilly, A., Rigal, L., & Quellet, C. (2016). Melt
46 692 extrusion encapsulation of flavors: a review. *Polymer Reviews*, *56*(1), 137-186.
47 693 doi:<https://doi.org/10.1080/15583724.2015.1091776>
48
- 49 694 Chen, C. C., & Wagner, G. (2004). Vitamin E nanoparticle for beverage applications.
50 695 *Chemical Engineering Research and Design*, *82*(11), 1432-1437.
51 696 doi:<https://doi.org/10.1205/cerd.82.11.1432.52034>
52
- 53 697 Chen, L., Remondetto, G. E., & Subirade, M. (2006). Food protein-based materials as
54 698 nutraceutical delivery systems. *Trends in Food Science & Technology*, *17*(5), 272-
55 699 283. doi:<https://doi.org/10.1016/j.tifs.2005.12.011>
56

- 700 Combet, E., & Buckton, C. (2019). Micronutrient deficiencies, vitamin pills and nutritional
1 701 supplements. *Medicine*, 47(3), 145-151.
2 702 doi:<https://doi.org/10.1016/j.mpmed.2018.12.004>
3
- 4 703 Couto, R., Alvarez, V., & Temelli, F. (2017). Encapsulation of vitamin B2 in solid lipid
5 704 nanoparticles using supercritical CO₂. *The Journal of Supercritical Fluids*, 120, 432-
6 705 442. doi:<https://doi.org/10.1016/j.supflu.2016.05.036>
7
- 8 706 da Silva, P. T., Fries, L. L. M., de Menezes, C. R., Holkem, A. T., Schwan, C. L., Wigmann,
9 707 E. F., . . . da Silva, C. D. (2014). Microencapsulation: concepts, mechanisms, methods
10 708 and some applications in food technology. *Ciencia Rural*, 44(7), 1304-1311.
11 709 doi:<https://doi.org/10.1590/0103-8478cr20130971>
12
- 13 710 de Farias, S. S., Siqueira, S. M. C., Cunha, A. P., de Souza, C. A. G., dos Santos Fontenelle,
14 711 R. O., de Araújo, T. G., . . . Ricardo, N. M. P. S. (2018). Microencapsulation of
15 712 riboflavin with galactomannan biopolymer and F127: Physico-chemical
16 713 characterization, antifungal activity and controlled release. *Industrial Crops and*
17 714 *Products*, 118, 271-281. doi:<https://doi.org/10.1016/j.indcrop.2018.03.039>
18
- 19 715 de Kruif, C. G., Weinbreck, F., & de Vries, R. (2004). Complex coacervation of proteins and
20 716 anionic polysaccharides. *Current Opinion in Colloid & Interface Science*, 9(5), 340-
21 717 349. doi:<https://doi.org/10.1016/j.cocis.2004.09.006>
22
- 23 718 de Paz, E., Martín, Á., & Cocero, M. J. (2012). Formulation of β-carotene with soybean
24 719 lecithin by PGSS (particles from gas saturated solutions)-drying. *The Journal of*
25 720 *Supercritical Fluids*, 72, 125-133. doi:<https://doi.org/10.1016/j.supflu.2012.08.007>
26
- 27 721 de Paz, E., Martín, Á., Duarte, C. M. M., & Cocero, M. J. (2012). Formulation of β-carotene
28 722 with poly-(ε-caprolactones) by PGSS process. *Powder Technology*, 217, 77-83.
29 723 doi:<https://doi.org/10.1016/j.powtec.2011.10.011>
30
- 31 724 Delompré, T., Guichard, E., Briand, L., & Salles, C. (2019). Taste Perception of Nutrients
32 725 Found in Nutritional Supplements: A Review. *Nutrients*, 11(9), 2050.
33 726 doi:<https://doi.org/10.3390/nu11092050>
34
- 35 727 Desai, K. G. H., & Jin Park, H. (2005). Recent developments in microencapsulation of food
36 728 ingredients. *Drying Technology*, 23(7), 1361-1394. doi:<https://doi.org/10.1081/DRT-200063478>
37 729
38
- 39 730 Desai, K. G. H., & Park, H. J. (2005). Encapsulation of vitamin C in tripolyphosphate cross-
40 731 linked chitosan microspheres by spray drying. *Journal of Microencapsulation*, 22(2),
41 732 179-192. doi:10.1080/02652040400026533
42
- 43 733 Drouin, G., Godin, J.-R., & Pagé, B. (2011). The genetics of vitamin C loss in vertebrates.
44 734 *Current genomics*, 12(5), 371-378. doi:<https://doi.org/10.2174/138920211796429736>
45
- 46 735 Egan, T., Jacquier, J.-C., Rosenberg, Y., & Rosenberg, M. (2013). Cold-set whey protein
47 736 microgels for the stable immobilization of lipids. *Food Hydrocolloids*, 31(2), 317-
48 737 324. doi:<https://doi.org/10.1016/j.foodhyd.2012.11.008>
49
- 50 738 Eghbal, N., & Choudhary, R. (2018). Complex coacervation: encapsulation and controlled
51 739 release of active agents in food systems. *LWT - Food Science and Technology*, 90,
52 740 254-264. doi:<https://doi.org/10.1016/j.lwt.2017.12.036>
53
54
55
56
57
58
59
60
61
62
63
64
65

- 741 Eisinaite, V., Juraite, D., Schroën, K., & Leskauskaite, D. (2016). Preparation of stable food-
1 742 grade double emulsions with a hybrid premix membrane emulsification system. *Food*
2 743 *Chemistry*, 206, 59-66. doi:<https://doi.org/10.1016/j.foodchem.2016.03.046>
3
- 4 744 Elzoghby, A. O., Elgohary, M. M., & Kamel, N. M. (2015). Implications of protein- and
5 745 peptide-based nanoparticles as potential vehicles for anticancer drugs. In R. Donev
6 746 (Ed.), *Advances in Protein Chemistry and Structural Biology* (Vol. 98, pp. 169-221):
7 747 Elsevier BV.
- 9 748 Fadini, A. L., Alvim, I. D., Ribeiro, I. P., Ruzene, L. G., Silva, L. B. d., Queiroz, M. B., . . .
10 749 Rodrigues, R. A. F. (2018). Innovative strategy based on combined
11 750 microencapsulation technologies for food application and the influence of wall
12 751 material composition. *LWT - Food Science and Technology*, 91, 345-352.
13 752 doi:<https://doi.org/10.1016/j.lwt.2018.01.071>
14 753
- 16 753 Fallingborg, J. (1999). Intraluminal pH of the human gastrointestinal tract. *Danish medical*
17 754 *bulletin*, 46(3), 183-196.
- 19 755 Fanger, G. O. (1974). Microencapsulation: a brief history and introduction. In J. E.
20 756 Vandegaer (Ed.), *Microencapsulation: Processes and Applications* (pp. 1-20).
21 757 Boston, MA: Springer US.
- 23 758 Fathi, M., Donsi, F., & McClements, D. J. (2018). Protein-based delivery systems for the
24 759 nanoencapsulation of food ingredients. *Comprehensive Reviews in Food Science and*
25 760 *Food Safety*, 17(4), 920-936. doi:<https://doi.org/10.1111/1541-4337.12360>
- 27 761 Favaro-Trindade, C. S., Okuro, P. K., & Matos Jr, F. (2015). Encapsulation via spray
28 762 chilling/cooling/congealing. *Handbook of Encapsulation and Controlled Release*.
29 763 *CRC Press, Boca Raton*, 71-88.
- 31 764 Fávoro, R. M. D., Iha, M. H., Mazzi, T. C., Fávoro, R., & Bianchi, M. d. L. P. (2011).
32 765 Stability of vitamin A during storage of enteral feeding formulas. *Food Chemistry*,
33 766 126(3), 827-830. doi:<https://doi.org/10.1016/j.foodchem.2010.09.014>
34 766
- 36 767 Ferguson, T. I., Emery, S., Price-Davies, R., & Cosslett, A. G. (2014). A review of stability
37 768 issues associated with vitamins in parenteral nutrition. *e-SPEN Journal*, 9(2), e49-e53.
38 769 doi:<https://doi.org/10.1016/j.clnme.2014.01.001>
39
- 40 770 Frascareli, E. C., Silva, V. M., Tonon, R. V., & Hubinger, M. D. (2012). Effect of process
41 771 conditions on the microencapsulation of coffee oil by spray drying. *Food and*
42 772 *Bioproducts Processing*, 90(3), 413-424.
43 773 doi:<https://doi.org/10.1016/j.fbp.2011.12.002>
44
- 45 774 Freitas, S., Merkle, H. P., & Gander, B. (2004). Ultrasonic atomisation into reduced pressure
46 775 atmosphere—envisaging aseptic spray-drying for microencapsulation. *Journal of*
47 776 *Controlled Release*, 95(2), 185-195. doi:<https://doi.org/10.1016/j.jconrel.2003.11.005>
48
- 49 777 Fu, Z.-m., Ma, Z.-z., Liu, G.-j., Wang, L.-l., & Guo, Y. (2018). Vitamins supplementation
50 778 affects the onset of preeclampsia. *Journal of the Formosan Medical Association*,
51 779 117(1), 6-13. doi:<https://doi.org/10.1016/j.jfma.2017.08.005>
52
- 54 780 Gamboa, O. D., Gonçalves, L. G., & Grosso, C. F. (2011). Microencapsulation of tocopherols
55 781 in lipid matrix by spray chilling method. *Procedia Food Science*, 1, 1732-1739.
56 782 doi:<https://doi.org/10.1016/j.profoo.2011.09.255>
57
58
59

- 783 Ganza-González, A., Anguiano-Igea, S., Otero-Espinar, F. J., & Blanco Méndez, J. (1999).
1 784 Chitosan and chondroitin microspheres for oral-administration controlled release of
2 785 metoclopramide. *European Journal of Pharmaceutics and Biopharmaceutics*, 48(2),
3 786 149-155. doi:[https://doi.org/10.1016/S0939-6411\(99\)00040-5](https://doi.org/10.1016/S0939-6411(99)00040-5)
- 5 787 Gibbs, B. F., Kermasha, S., Alli, I., & Mulligan, C. N. (1999). Encapsulation in the food
6 788 industry: a review. *International Journal of Food Sciences and Nutrition*, 50(3), 213-
7 789 224. doi:<https://doi.org/10.1080/096374899101256>
- 9 790 Gibson, R. S. (2011). Strategies for preventing multi-micro nutrient deficiencies: a review of
10 791 experiences with food-based approaches in developing countries. In B. Thompson &
11 792 L. Amoroso (Eds.), *Combating micronutrient deficiencies : food-based approaches*:
12 793 Food and Agricultural Organisation of the United Nations, CABI.
- 14 794 Gonçalves, A., Estevinho, B. N., & Rocha, F. (2016). Microencapsulation of vitamin A: A
15 795 review. *Trends in Food Science & Technology*, 51, 76-87.
16 796 doi:<https://doi.org/10.1016/j.tifs.2016.03.001>
- 18 797 Gouin, S. (2004). Microencapsulation: industrial appraisal of existing technologies and
19 798 trends. *Trends in Food Science & Technology*, 15(7), 330-347.
20 799 doi:<https://doi.org/10.1016/j.tifs.2003.10.005>
- 22 800 Gregory, J. F. (2008). Vitamins. In S. Damodaran, K. L. Parkin, & O. R. Fennema (Eds.),
23 801 *Fennema's food chemistry* (4th ed.). Boca Raton ; London: CRC Press.
- 25 802 Gunder, W., Lippold, B. H., & Lippold, B. C. (1995). Release of drugs from ethyl cellulose
26 803 microcapsules (diffusion pellets) with pore formers and pore fusion. *European*
27 804 *Journal of Pharmaceutical Sciences*, 3(4), 203-214. doi:[https://doi.org/10.1016/0928-0987\(95\)00009-3](https://doi.org/10.1016/0928-0987(95)00009-3)
- 29 805
- 31 806 Gonsel, W. C., & Lachman, L. (1963). Comparative evaluation of tablet formulations
32 807 prepared from conventionally-processed and spray-dried lactose. *Journal of*
33 808 *Pharmaceutical Sciences*, 52(2), 178-182. doi:<https://doi.org/10.1002/jps.2600520219>
- 34 809 Guo, B.-Q., Li, H.-B., Zhai, D.-S., & Ding, S.-B. (2019). Maternal multivitamin
35 810 supplementation is associated with a reduced risk of autism spectrum disorder in
36 811 children: a systematic review and meta-analysis. *Nutrition Research*.
37 812 doi:<https://doi.org/10.1016/j.nutres.2019.02.003>
- 38 813 Gupta, C., Arora, S., Sharma, A., & Sharma, V. (2019). Evaluation of effective storage
39 814 conditions and in-vitro bioaccessibility of vitamin A from native and modified sodium
40 815 caseinate -vitamin A complexes. *LWT - Food Science and Technology*, 111, 284-290.
41 816 doi:<https://doi.org/10.1016/j.lwt.2019.05.048>
- 42 817 Hamishehkar, H., Ranjdoost, F., Asgharian, P., Mahmoodpoor, A., & Sanaie, S. (2016).
43 818 Vitamins, are they safe? *Advanced pharmaceutical bulletin*, 6(4), 467-477.
44 819 doi:<https://doi.org/10.15171/apb.2016.061>
- 45 820 Harris, R. S. (1988). General Discussion on the Stability of Nutrients. In E. Karmas & R. S.
46 821 Harris (Eds.), *Nutritional Evaluation of Food Processing* (pp. 3-5). Dordrecht:
47 822 Springer.
- 48 823 Hemery, Y. M., Fontan, L., Moench-Pfanner, R., Laillou, A., Berger, J., Renaud, C., &
49 824 Avallone, S. (2015). Influence of light exposure and oxidative status on the stability

- 825 of vitamins A and D3 during the storage of fortified soybean oil. *Food Chemistry*,
1 826 184, 90-98. doi:<https://doi.org/10.1016/j.foodchem.2015.03.096>
- 2
3 827 Hemery, Y. M., Laillou, A., Fontan, L., Jallier, V., Moench-Pfanner, R., Berger, J., &
4 828 Avallone, S. (2018). Storage conditions and packaging greatly affects the stability of
5 829 fortified wheat flour: Influence on vitamin A, iron, zinc, and oxidation. *Food*
6 830 *Chemistry*, 240, 43-50. doi:<https://doi.org/10.1016/j.foodchem.2017.07.084>
- 8 831 Hickey, D. K., Kilcawley, K. N., Beresford, T. P., & Wilkinson, M. G. (2007). Lipolysis in
9 832 cheddar cheese made from raw, thermized, and pasteurized milks. *Journal of Dairy*
10 833 *Science*, 90(1), 47-56. doi:[https://doi.org/10.3168/jds.S0022-0302\(07\)72607-3](https://doi.org/10.3168/jds.S0022-0302(07)72607-3)
- 12 834 Higuchi, T. (1963). Mechanism of sustained- action medication. Theoretical analysis of rate
13 835 of release of solid drugs dispersed in solid matrices. *Journal of Pharmaceutical*
14 836 *Sciences*, 52(12), 1145-1149. doi:<https://doi.org/10.1002/jps.2600521210>
- 16 837 Hoyos-Leyva, J. D., Bello-Pérez, L. A., Alvarez-Ramirez, J., & Garcia, H. S. (2018).
18 838 Microencapsulation using starch as wall material: a review. *Food Reviews*
19 839 *International*, 34(2), 148-161. doi:<https://doi.org/10.1080/87559129.2016.1261298>
- 21 840 Hoyos-Leyva, J. D., Chavez-Salazar, A., Castellanos-Galeano, F., Bello-Perez, L. A., &
22 841 Alvarez-Ramirez, J. (2018). Physical and chemical stability of l-ascorbic acid
23 842 microencapsulated into taro starch spherical aggregates by spray drying. *Food*
24 843 *Hydrocolloids*, 83, 143-152. doi:<https://doi.org/10.1016/j.foodhyd.2018.05.002>
- 26 844 Huang, Q., Yu, H., & Ru, Q. (2010). Bioavailability and delivery of nutraceuticals using
27 845 nanotechnology. *Journal of Food Science*, 75(1), R50-R57.
28 846 doi:<https://doi.org/10.1111/j.1750-3841.2009.01457.x>
- 30 847 Jafari, S. M., Assadpoor, E., Bhandari, B., & He, Y. (2008). Nano-particle encapsulation of
31 848 fish oil by spray drying. *Food Research International*, 41(2), 172-183.
32 849 doi:<https://doi.org/10.1016/j.foodres.2007.11.002>
- 34 850 Jenkins, D. J. A., Spence, J. D., Giovannucci, E. L., Kim, Y.-i., Josse, R., Vieth, R., . . .
35 851 Sievenpiper, J. L. (2018). Supplemental vitamins and minerals for CVD prevention
36 852 and treatment. *Journal of the American College of Cardiology*, 71(22), 2570-2584.
37 853 doi:<https://doi.org/10.1016/j.jacc.2018.04.020>
- 39
40 854 Jiao, Z., Wang, X., Yin, Y., Xia, J., & Mei, Y. (2018). Preparation and evaluation of a
41 855 chitosan-coated antioxidant liposome containing vitamin C and folic acid. *Journal of*
42 856 *Microencapsulation*, 35(3), 272-280.
43 857 doi:<https://doi.org/10.1080/02652048.2018.1467509>
- 45 858 Juveriya Fathima, S., Fathima, I., Abhishek, V., & Khanum, F. (2016). Phosphatidylcholine,
46 859 an edible carrier for nanoencapsulation of unstable thiamine. *Food Chemistry*, 197,
47 860 562-570. doi:<https://doi.org/10.1016/j.foodchem.2015.11.005>
- 49 861 Kheadr, E. E., Vuillemand, J. C., & El-Deeb, S. A. (2003). Impact of liposome-encapsulated
50 862 enzyme cocktails on cheddar cheese ripening. *Food Research International*, 36(3),
51 863 241-252. doi:[https://doi.org/10.1016/S0963-9969\(02\)00166-7](https://doi.org/10.1016/S0963-9969(02)00166-7)
- 53
54 864 Kramer, A. (1977). Effect of storage on nutritive value of food. *Journal of Food Quality*,
55 865 1(1), 23-55. doi: <https://doi.org/10.1111/j.1745-4557.1977.tb00998.x>

- 866 Laloy, E., Vuilleumard, J.-C., Dufour, P., & Simard, R. (1998). Release of enzymes from
1 867 liposomes during cheese ripening. *Journal of Controlled Release*, 54(2), 213-222.
2 868 doi:[https://doi.org/10.1016/S0168-3659\(97\)00265-4](https://doi.org/10.1016/S0168-3659(97)00265-4)
3
- 4 869 Laouini, A., Fessi, H., & Charcosset, C. (2012). Membrane emulsification: A promising
5 870 alternative for vitamin E encapsulation within nano-emulsion. *Journal of Membrane*
6 871 *Science*, 423-424, 85-96. doi:<https://doi.org/10.1016/j.memsci.2012.07.031>
- 8 872 Lau, B. L. T., Kakuda, Y., & Arnott, D. R. (1986). Effect of milk fat on the stability of
9 873 vitamin A in ultra-high temperature milk. *Journal of Dairy Science*, 69(8), 2052-
10 874 2059. doi:[https://doi.org/10.3168/jds.S0022-0302\(86\)80636-1](https://doi.org/10.3168/jds.S0022-0302(86)80636-1)
- 12 875 Lee, S. C., Lee, K. E., Kim, J. J., & Lim, S. H. (2005). The effect of cholesterol in the
13 876 liposome bilayer on the stabilization of incorporated Retinol. *Journal of Liposome*
14 877 *Research*, 15(3-4), 157-166. doi:<https://doi.org/10.1080/08982100500364131>
- 16 878 Lee, S. C., Yuk, H. G., Lee, D. H., Lee, K. E., Hwang, Y. I., & Ludescher, R. D. (2002).
17 879 Stabilization of retinol through incorporation into liposomes. *Journal of biochemistry*
18 880 *and molecular biology*, 35(4), 358-363.
20 881 doi:<https://doi.org/10.1080/08982100500364131>
- 22 882 Lee, S. J., & Wong, M. (2014). Nano-and microencapsulation of phytochemicals. In H. S.
23 883 Kwak (Ed.), *Nano- and Microencapsulation for Foods* (1st ed., pp. 119-165): John
24 884 Wiley & Sons.
- 26 885 Lešková, E., Kubíková, J., Kováčiková, E., Košická, M., Porubská, J., & Holčíková, K.
27 886 (2006). Vitamin losses: retention during heat treatment and continual changes
28 887 expressed by mathematical models. *Journal of Food Composition and Analysis*,
29 888 19(4), 252-276. doi:<https://doi.org/10.1016/j.jfca.2005.04.014>
- 31 889 Leung, C. C., Yu, I. T., & Chen, W. (2012). Silicosis. *Lancet*, 379(9830), 2008-2018.
32 890 doi:[https://doi.org/10.1016/s0140-6736\(12\)60235-9](https://doi.org/10.1016/s0140-6736(12)60235-9)
- 34 891 Li, M., Du, C., Guo, N., Teng, Y., Meng, X., Sun, H., . . . Galons, H. (2019). Composition
35 892 design and medical application of liposomes. *European Journal of Medicinal*
36 893 *Chemistry*, 164, 640-653. doi:<https://doi.org/10.1016/j.ejmech.2019.01.007>
- 38 894 Liang, R., Huang, Q., Ma, J., Shoemaker, C. F., & Zhong, F. (2013). Effect of relative
39 895 humidity on the store stability of spray-dried beta-carotene nanoemulsions. *Food*
40 896 *Hydrocolloids*, 33(2), 225-233. doi:<https://doi.org/10.1016/j.foodhyd.2013.03.015>
- 42 897 Links, M. R., Taylor, J., Kruger, M. C., & Taylor, J. R. N. (2015). Sorghum condensed
43 898 tannins encapsulated in kafirin microparticles as a nutraceutical for inhibition of
44 899 amylases during digestion to attenuate hyperglycaemia. *Journal of Functional Foods*,
45 900 12, 55-63. doi:<https://doi.org/10.1016/j.jff.2014.11.003>
- 48 901 Liu, L., Shen, W., Zhang, W., Li, F., & Zhu, Z. (2018). Porous Starch and Its Applications. In
49 902 Z. Jin (Ed.), *Functional Starch and Applications in Food* (pp. 91-117). Singapore:
50 903 Springer Singapore.
- 52 904 Liu, T., Hamid, N., Kantono, K., Pereira, L., Farouk, M. M., & Knowles, S. O. (2016).
53 905 Effects of meat addition on pasta structure, nutrition and in vitro digestibility. *Food*
54 906 *Chemistry*, 213, 108-114. doi:<https://doi.org/10.1016/j.foodchem.2016.06.058>
- 56 907 Livney, Y. D. (2010). Milk proteins as vehicles for bioactives. *Current Opinion in Colloid &*
57 908 *Interface Science*, 15(1), 73-83. doi:<https://doi.org/10.1016/j.cocis.2009.11.002>

- 909 Lopes, N. A., Barreto Pinilla, C. M., & Brandelli, A. (2019). Antimicrobial activity of
1 910 lysozyme-nisin co-encapsulated in liposomes coated with polysaccharides. *Food*
2 911 *Hydrocolloids*, 93, 1-9. doi:<https://doi.org/10.1016/j.foodhyd.2019.02.009>
3
- 4 912 Lopes, N. A., Pinilla, C. M. B., & Brandelli, A. (2017). Pectin and polygalacturonic acid-
5 913 coated liposomes as novel delivery system for nisin: Preparation, characterization and
6 914 release behavior. *Food Hydrocolloids*, 70, 1-7.
7 915 doi:<https://doi.org/10.1016/j.foodhyd.2017.03.016>
8
- 9 916 Loveday, S. M., & Singh, H. (2008). Recent advances in technologies for vitamin A
10 917 protection in foods. *Trends in Food Science & Technology*, 19(12), 657-668.
11 918 doi:<https://doi.org/10.1016/j.tifs.2008.08.002>
12
- 13 919 Madziva, H., Kailasapathy, K., & Phillips, M. (2005). Alginate–pectin microcapsules as a
14 920 potential for folic acid delivery in foods. *Journal of Microencapsulation*, 22(4), 343-
15 921 351. doi:<https://doi.org/10.1080/02652040500100931>
16 921
17
- 18 922 Madziva, H., Kailasapathy, K., & Phillips, M. (2006). Evaluation of alginate–pectin capsules
19 923 in Cheddar cheese as a food carrier for the delivery of folic acid. *LWT - Food Science*
20 924 *and Technology*, 39(2), 146-151. doi:<https://doi.org/10.1016/j.lwt.2004.12.015>
21
- 22 925 Marsanasco, M., Márquez, A. L., Wagner, J. R., del V. Alonso, S., & Chiaramoni, N. S.
23 926 (2011). Liposomes as vehicles for vitamins E and C: an alternative to fortify orange
24 927 juice and offer vitamin C protection after heat treatment. *Food Research*
25 928 *International*, 44(9), 3039-3046. doi:<https://doi.org/10.1016/j.foodres.2011.07.025>
26
- 27 929 Martins, I. M., Barreiro, M. F., Coelho, M., & Rodrigues, A. E. (2014). Microencapsulation
28 930 of essential oils with biodegradable polymeric carriers for cosmetic applications.
29 931 *Chemical Engineering Journal*, 245, 191-200.
30 932 doi:<https://doi.org/10.1016/j.cej.2014.02.024>
31 932
32
- 33 933 Matassa, S., Boon, N., Pikaar, I., & Verstraete, W. (2016). Microbial protein: future
34 934 sustainable food supply route with low environmental footprint. *Microbial*
35 935 *biotechnology*, 9(5), 568-575. doi:<https://doi.org/10.1111/1751-7915.12369>
36
- 37 936 Matos, M., Gutiérrez, G., Iglesias, O., Coca, J., & Pazos, C. (2015). Enhancing encapsulation
38 937 efficiency of food-grade double emulsions containing resveratrol or vitamin B12 by
39 938 membrane emulsification. *Journal of Food Engineering*, 166, 212-220.
40 939 doi:<https://doi.org/10.1016/j.jfoodeng.2015.06.002>
41
- 42 940 McClements, D. J., Decker, E. A., & Weiss, J. (2007). Emulsion-based delivery systems for
43 941 lipophilic bioactive components. *Journal of Food Science*, 72(8), R109-124.
44 942 doi:<https://doi.org/10.1111/j.1750-3841.2007.00507.x>
45 942
46
- 47 943 Mehrad, B., Ravanfar, R., Licker, J., Regenstein, J. M., & Abbaspourrad, A. (2018).
48 944 Enhancing the physicochemical stability of β -carotene solid lipid nanoparticle (SLNP)
49 945 using whey protein isolate. *Food Research International*, 105, 962-969.
50 946 doi:<https://doi.org/10.1016/j.foodres.2017.12.036>
51
- 52 947 Minekus, M., Alminger, M., Alvito, P., Ballance, S., Bohn, T., Bourlieu, C., . . . Brodkorb, A.
53 948 (2014). A standardised static in vitro digestion method suitable for food-an
54 949 international consensus. *Food & Function*, 5(6), 1113-1124.
55 950 doi:<https://doi.org/10.1039/c3fo60702j>
56 950
57
58
59
60
61
62
63
64
65

- 951 Mirafzali, Z., Thomas, C. S., & Tallua, K. (2014). Applicaiton of liposomes in the food
1 952 industry. In A. Gaonkar, N. Vasisht, A. R. Khare, & R. Sobel (Eds.),
2 953 *Microencapsulation in the food industry : a practical implementation guide* (pp. 139-
3 954 150). San Diego, US: Elsevier Science & Technology.
- 5 955 Moccand, C., Martin, F., Martiel, I., Gancel, C., Michel, M., Fries, L., & Sagalowicz, L.
6 956 (2016). Vitamin A degradation in triglycerides varying by their saturation levels.
7 957 *Food Research International*, 88, 3-9.
9 958 doi:<https://doi.org/10.1016/j.foodres.2016.06.001>
- 10 959 Moeller, H., Martin, D., Schrader, K., Hoffmann, W., & Lorenzen, P. C. (2018). Spray- or
11 960 freeze-drying of casein micelles loaded with Vitamin D2: Studies on storage stability
12 961 and in vitro digestibility. *LWT - Food Science and Technology*, 97, 87-93.
14 962 doi:<https://doi.org/10.1016/j.lwt.2018.04.003>
- 16 963 Moraes, M., Carvalho, J. M. P., Silva, C. R., Cho, S., Sola, M. R., & Pinho, S. C. (2013).
17 964 Liposomes encapsulating beta-carotene produced by the proliposomes method:
18 965 characterisation and shelf life of powders and phospholipid vesicles. *International*
19 966 *Journal of Food Science & Technology*, 48(2), 274-282.
21 967 doi:<https://doi.org/10.1111/j.1365-2621.2012.03184.x>
- 23 968 Mount, E. M. (2011). Extrusion Processes. In M. Kutz (Ed.), *Applied Plastics Engineering*
24 969 *Handbook* (pp. 227-266). Oxford: William Andrew Publishing.
- 26 970 Mozaffarian, D., Rosenberg, I., & Uauy, R. (2018). History of modern nutrition science—
27 971 implications for current research, dietary guidelines, and food policy. *BMJ*, 361,
28 972 k2392. doi:<https://doi.org/10.1136/bmj.k2392>
- 30 973 Mukherjee, S., Ray, S., & Thakur, R. S. (2009). Solid lipid nanoparticles: a modern
31 974 formulation approach in drug delivery system. *Indian journal of pharmaceutical*
32 975 *sciences*, 71(4), 349-358. doi:10.4103/0250-474X.57282
- 34 976 Murugadoss, S., Lison, D., Godderis, L., Van Den Brule, S., Mast, J., Brassinne, F., . . . Hoet,
35 977 P. H. (2017). Toxicology of silica nanoparticles: an update. *Archives of toxicology*,
36 978 91(9), 2967-3010. doi:<https://doi.org/10.1007/s00204-017-1993-y>
- 38 979 Nazzaro, F., Orlando, P., Fratianni, F., & Coppola, R. (2012). Microencapsulation in food
39 980 science and biotechnology. *Current Opinion in Biotechnology*, 23(2), 182-186.
41 981 doi:<https://doi.org/10.1016/j.copbio.2011.10.001>
- 42 982 Niaz, T., Shabbir, S., Noor, T., Rahman, A., Bokhari, H., & Imran, M. (2018). Potential of
43 983 polymer stabilized nano-liposomes to enhance antimicrobial activity of nisin Z against
44 984 foodborne pathogens. *LWT - Food Science and Technology*, 96, 98-110.
46 985 doi:<https://doi.org/10.1016/j.lwt.2018.05.029>
- 48 986 Nutrient Data Lab. (2017). *USDA Table of Nutrient Retention Factors*. Retrieved from:
49 987 <http://dx.doi.org/10.15482/USDA.ADC/1409034>
- 51 988 Nyantakyi-Frimpong, H. (2017). Agricultural diversification and dietary diversity: a feminist
52 989 political ecology of the everyday experiences of landless and smallholder households
53 990 in northern Ghana. *Geoforum*, 86, 63-75.
54 991 doi:<https://doi.org/10.1016/j.geoforum.2017.09.003>
- 56 992 Olive Li, Y., Dueik González, V. P., & Diosady, L. L. (2014). Microencapsulation of
57 993 vitamins, minerals, and nutraceuticals for food applications. In A. G. Gaonkar, N.

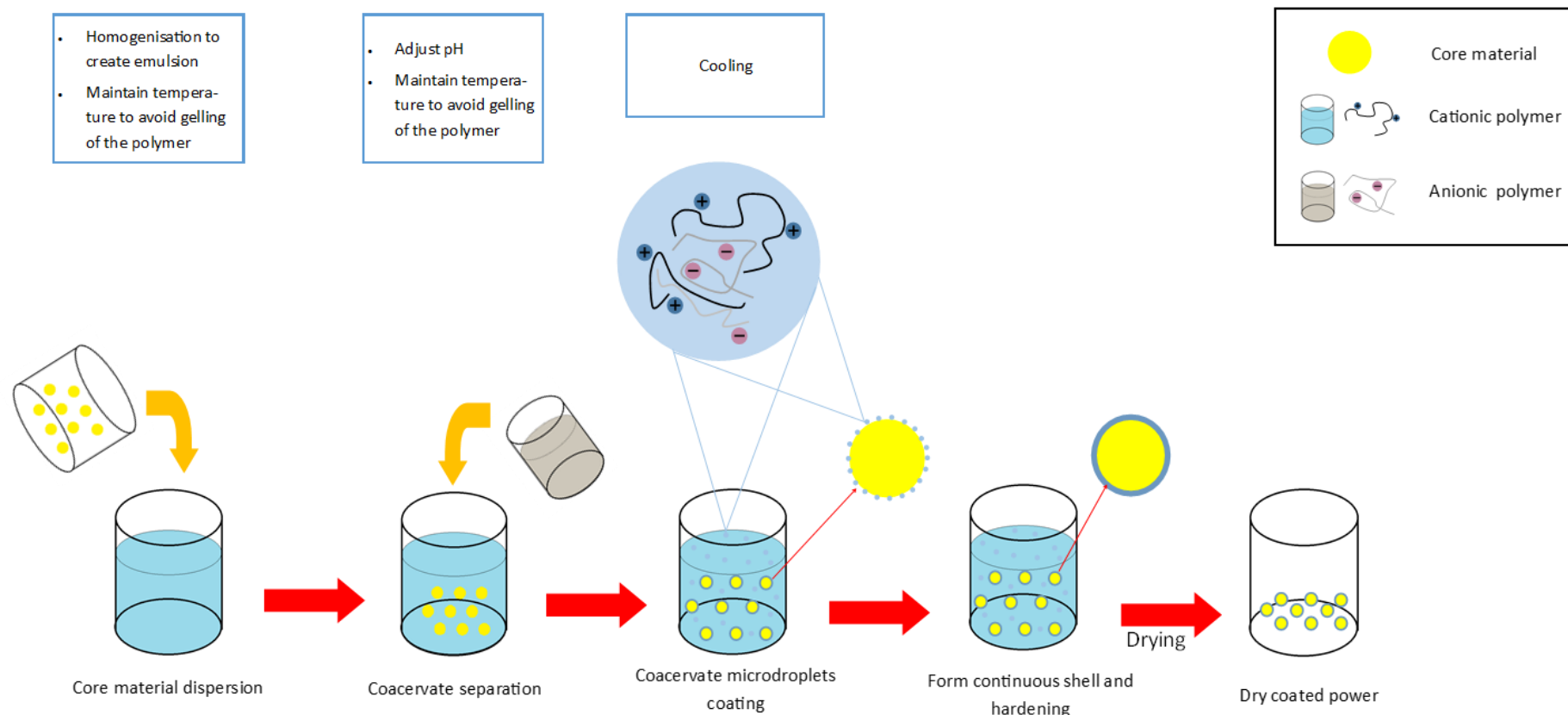
- 994 Vasisht, A. R. Khare, & R. Sobel (Eds.), *Microencapsulation in the Food Industry*
1 995 (pp. 501-522). San Diego: Academic Press.
2
- 3 996 Ota, A., Istenič, K., Skrt, M., Šegatin, N., Žnidaršič, N., Kogej, K., & Ulrih, N. P. (2018).
4 997 Encapsulation of pantothenic acid into liposomes and into alginate or alginate–pectin
5 998 microparticles loaded with liposomes. *Journal of Food Engineering*, 229, 21-31.
6 999 doi:<https://doi.org/10.1016/j.jfoodeng.2017.06.036>
- 8 1000 Ottaway, P. B. (1993). Stability of vitamins in food. In P. B. Ottaway (Ed.), *The Technology*
9 1001 *of Vitamins in Food* (pp. 90-113). Boston, MA: Springer US.
10
- 11 1002 Ottaway, P. B. (2010). Stability of vitamins during food processing and storage. In L. H.
12 1003 Skibsted, J. Risbo, & M. L. Andersen (Eds.), *Chemical deterioration and physical*
13 1004 *instability of food and beverages* (pp. 539-560): Woodhead Publishing.
- 15 1005 Oxley, J. (2014). Overview of microencapsulation process technologies. In A. Gaonkar, N.
16 1006 Vasisht, A. R. Khare, & R. Sobel (Eds.), *Microencapsulation in the food industry: a*
17 1007 *practical implementation guide* (pp. 37). San Diego, US: Elsevier.
- 19 1008 Parkes, A. (2018). *Meat-free Foods - UK - September 2018*. Retrieved from Mintel
20 1009 Academic: <http://academic.mintel.com/>
- 22 1010 Paucar, O. C., Tulini, F. L., Thomazini, M., Balieiro, J. C. C., Pallone, E. M. J. A., & Favaro-
23 1011 Trindade, C. S. (2016). Production by spray chilling and characterization of solid lipid
24 1012 microparticles loaded with vitamin D3. *Food and Bioprocess Processing*, 100, 344-
25 1013 350. doi:<https://doi.org/10.1016/j.fbp.2016.08.006>
26 1013
27
- 28 1014 Pérez-Esteve, É., Fuentes, A., Coll, C., Acosta, C., Bernardos, A., Amorós, P., . . . Barat, J.
29 1015 M. (2015). Modulation of folic acid bioaccessibility by encapsulation in pH-
30 1016 responsive gated mesoporous silica particles. *Microporous and Mesoporous*
31 1017 *Materials*, 202, 124-132. doi:<https://doi.org/10.1016/j.micromeso.2014.09.049>
- 33 1018 Piacentini, E., Giorno, L., Dragosavac, M. M., Vladisavljević, G. T., & Holdich, R. G.
34 1019 (2013). Microencapsulation of oil droplets using cold water fish gelatine/gum arabic
35 1020 complex coacervation by membrane emulsification. *Food Research International*,
36 1021 53(1), 362-372. doi:<https://doi.org/10.1016/j.foodres.2013.04.012>
37 1021
38
- 39 1022 Poncelet, D. (2006). *Microencapsulation: fundamentals, methods and applications*. Paper
40 1023 presented at the Surface Chemistry in Biomedical and Environmental Science,
41 1024 Dordrecht.
42
- 43 1025 Prieto, C., & Calvo, L. (2017). Supercritical fluid extraction of emulsions to nanoencapsulate
44 1026 vitamin E in polycaprolactone. *The Journal of Supercritical Fluids*, 119, 274-282.
45 1027 doi:<https://doi.org/10.1016/j.supflu.2016.10.004>
46
- 47 1028 Reineccius, G. (2019). Use of proteins for the delivery of flavours and other bioactive
48 1029 compounds. *Food Hydrocolloids*, 86, 62-69.
49 1030 doi:<https://doi.org/10.1016/j.foodhyd.2018.01.039>
- 51 1031 Rickman, J. C., Barrett, D. M., & Bruhn, C. M. (2007). Nutritional comparison of fresh,
52 1032 frozen and canned fruits and vegetables. Part 1. Vitamins C and B and phenolic
53 1033 compounds. *Journal of the Science of Food and Agriculture*, 87(6), 930-944.
54 1034 doi:<https://doi.org/10.1002/jsfa.2825>
- 56 1035 Ritchie, H., & Roser, M. (2020, 11/05/2020). Micronutrient Deficiency. *Our World in Data*.
57 1036 Retrieved from <https://ourworldindata.org/micronutrient-deficiency>

- 1037 Rodríguez-Huezo, M. E., Pedroza-Islas, R., Prado-Barragán, L. A., Beristain, C. I., &
11038 Vernon-Carter, E. J. (2004). Microencapsulation by spray drying of multiple
21039 emulsions containing carotenoids. *Journal of Food Science*, 69(7), 351-359.
31040 doi:<https://doi.org/10.1111/j.1365-2621.2004.tb13641.x>
- 51041 Romo-Hualde, A., Yetano-Cunchillos, A. I., González-Ferrero, C., Sáiz-Abajo, M. J., &
61042 González-Navarro, C. J. (2012). Supercritical fluid extraction and microencapsulation
71043 of bioactive compounds from red pepper (*Capsicum annum L.*) by-products. *Food*
81044 *Chemistry*, 133(3), 1045-1049. doi:<https://doi.org/10.1016/j.foodchem.2012.01.062>
- 101045 Ruiz-Rico, M., Pérez-Esteve, É., Lerma-García, M. J., Marcos, M. D., Martínez-Máñez, R.,
111046 & Barat, J. M. (2017). Protection of folic acid through encapsulation in mesoporous
121047 silica particles included in fruit juices. *Food Chemistry*, 218, 471-478.
131048 doi:<https://doi.org/10.1016/j.foodchem.2016.09.097>
- 151049 Salminen, H., Gömmel, C., Leuenberger, B. H., & Weiss, J. (2016). Influence of
161050 encapsulated functional lipids on crystal structure and chemical stability in solid lipid
171051 nanoparticles: Towards bioactive-based design of delivery systems. *Food Chemistry*,
181052 190, 928-937. doi:<https://doi.org/10.1016/j.foodchem.2015.06.054>
- 201053 Santana, A. A., Cano-Higuaita, D. M., de Oliveira, R. A., & Telis, V. R. N. (2016). Influence
211054 of different combinations of wall materials on the microencapsulation of jussara pulp
221055 (*Euterpe edulis*) by spray drying. *Food Chemistry*, 212, 1-9.
231056 doi:<https://doi.org/10.1016/j.foodchem.2016.05.148>
- 251057 Schmitt, C., & Turgeon, S. L. (2011). Protein/polysaccharide complexes and coacervates in
261058 food systems. *Advances in Colloid and Interface Science*, 167(1), 63-70.
271059 doi:<https://doi.org/10.1016/j.cis.2010.10.001>
- 291060 Schrooyen, P. M., van der Meer, R., & De Kruif, C. G. (2001). Microencapsulation: its
301061 application in nutrition. *Proceedings of the Nutrition Society*, 60(4), 475-479.
311062 doi:<https://doi.org/10.1079/PNS2001112>
- 321063 Scott, K. J., & Bishop, D. R. (1986). Nutrient content of milk and milk products: vitamins of
331064 the B complex and vitamin C in retail market milk and milk products. *International*
341065 *Journal of Dairy Technology*, 39(1), 32-35. doi:<https://doi.org/10.1111/j.1471-0307.1986.tb02356.x>
- 351066 Sebaaly, C., Jraj, A., Fessi, H., Charcosset, C., & Greige-Gerges, H. (2015). Preparation and
361067 characterization of clove essential oil-loaded liposomes. *Food Chemistry*, 178, 52-62.
371068 doi:<https://doi.org/10.1016/j.foodchem.2015.01.067>
- 381069 Semyonov, D., Ramon, O., & Shimoni, E. (2011). Using ultrasonic vacuum spray dryer to
391070 produce highly viable dry probiotics. *LWT - Food Science and Technology*, 44(9),
401071 1844-1852. doi:<https://doi.org/10.1016/j.lwt.2011.03.021>
- 411072 Severi, S., Bedogni, G., Manzi, A. M., Poli, M., & Battistini, N. (1997). Effects of cooking
421073 and storage methods on the micronutrient content of foods. *European Journal of*
431074 *Cancer Prevention*, 6 Suppl 1, S21-24. doi:<https://doi.org/10.1097/00008469-199703001-00005>
- 441075 Shahidi, F., & Han, X. Q. (1993). Encapsulation of food ingredients. *Critical Reviews in*
451076 *Food Science and Nutrition*, 33(6), 501-547.
461077 doi:<https://doi.org/10.1080/10408399309527645>

- 1080 Sherry, M., Charcosset, C., Fessi, H., & Greige-Gerges, H. (2013). Essential oils
1081 encapsulated in liposomes: a review. *Journal of Liposome Research*, 23(4), 268-275.
1082 doi:<https://doi.org/10.3109/08982104.2013.819888>
1083
1084 Shrestha, A. K., Arcot, J., & Yuliani, S. (2012). Susceptibility of 5-methyltetrahydrofolic acid
1085 to heat and microencapsulation to enhance its stability during extrusion processing.
1086 *Food Chemistry*, 130(2), 291-298.
1087 doi:<https://doi.org/10.1016/j.foodchem.2011.07.040>
1088
1089 Siepmann, J., & Siepmann, F. (2012). Modeling of diffusion controlled drug delivery.
1090 *Journal of Controlled Release*, 161(2), 351-362.
1091 doi:<https://doi.org/10.1016/j.jconrel.2011.10.006>
1092
1093 Singh, M. N., Hemant, K. S. Y., Ram, M., & Shivakumar, H. G. (2010). Microencapsulation:
1094 A promising technique for controlled drug delivery. *Research in pharmaceutical*
1095 *sciences*, 5(2), 65-77.
1096
1097 Smith, F., Pan, X. Y., Bellido, V., Toole, G. A., Gates, F. K., Wickham, M. S. J., . . . Mills, E.
1098 N. C. (2015). Digestibility of gluten proteins is reduced by baking and enhanced by
1099 starch digestion. *Molecular Nutrition & Food Research*, 59(10), 2034-2043.
1100 doi:<https://doi.org/10.1002/mnfr.201500262>
1101
1102 Sobel, R., Versic, R., & Gaonkar, A. G. (2014). Introduction to microencapsulation and
1103 controlled delivery in foods. In A. G. Gaonkar, N. Vasisht, A. R. Khare, & R. Sobel
1104 (Eds.), *Microencapsulation in the Food Industry* (pp. 3-12). San Diego: Elsevier
1105
1106 Soh, S. H., & Lee, L. Y. (2019). Microencapsulation and nanoencapsulation using
1107 supercritical fluid (SCF) techniques. *Pharmaceutics*, 11(1), 21.
1108 doi:<https://doi.org/10.3390/pharmaceutics11010021>
1109
1110 Sungpuag, P., Tangchitpianvit, S., Chittchang, U., & Wasantwisut, E. (1999). Retinol and
1111 beta carotene content of indigenous raw and home-prepared foods in Northeast
1112 Thailand. *Food Chemistry*, 64(2), 163-167. doi:[https://doi.org/10.1016/S0308-8146\(98\)00154-X](https://doi.org/10.1016/S0308-8146(98)00154-X)
1113
1114 Sutaphanit, P., & Chitprasert, P. (2014). Optimisation of microencapsulation of holy basil
1115 essential oil in gelatin by response surface methodology. *Food Chemistry*, 150, 313-
1116 320. doi:<https://doi.org/10.1016/j.foodchem.2013.10.159>
1117
1118 Takahashi, H., Chen, R., Okamoto, H., & Danjo, K. (2005). Acetaminophen particle design
1119 using chitosan and a spray-drying technique. *Chemical and Pharmaceutical Bulletin*,
1120 53(1), 37-41. doi:<https://doi.org/10.1248/cpb.53.37>
1121
1122 Teleki, A., Hitzfeld, A., & Eggersdorfer, M. (2013). 100 years of vitamins: the science of
1123 formulation is the key to functionality. *Kona Powder and Particle Journal*(30), 144-
1124 163. doi:<https://doi.org/10.14356/kona.2013015>
1125
1126 Thies, C. (2003). Microcapsules. In B. Caballero (Ed.), *Encyclopedia of food sciences and*
1127 *nutrition* (2 ed., pp. 3892-3903). Oxford: Elsevier.
1128
1129 Timilsena, Y. P., Akanbi, T. O., Khalid, N., Adhikari, B., & Barrow, C. J. (2019). Complex
1130 coacervation: Principles, mechanisms and applications in microencapsulation.
1131 *International Journal of Biological Macromolecules*, 121, 1276-1286.
1132 doi:<https://doi.org/10.1016/j.ijbiomac.2018.10.144>

- 1122 Tontul, I., & Topuz, A. (2013). Mixture design approach in wall material selection and
1123 evaluation of ultrasonic emulsification in flaxseed oil microencapsulation. *Drying*
1124 *Technology*, 31(12), 1362-1373. doi:<https://doi.org/10.1080/07373937.2013.795964>
- 1125 Tucker, B. M., Safadi, S., & Friedman, A. N. (2015). Is routine multivitamin supplementation
1126 necessary in US chronic adult hemodialysis patients?? A systematic review. *Journal*
1127 *of Renal Nutrition*, 25(3), 257-264. doi:<https://doi.org/10.1053/j.jrn.2014.09.003>
- 1128 Tulini, F. L., Souza, V. B., Thomazini, M., Silva, M. P., Massarioli, A. P., Alencar, S. M., . . .
1129 Favaro-Trindade, C. S. (2017). Evaluation of the release profile, stability and
1130 antioxidant activity of a proanthocyanidin-rich cinnamon (*cinnamomum zeylanicum*)
1131 extract co-encapsulated with α -tocopherol by spray chilling. *Food Research*
1132 *International*, 95, 117-124. doi:<https://doi.org/10.1016/j.foodres.2017.03.010>
- 1133 Vladislavljević, G. T., & Williams, R. A. (2005). Recent developments in manufacturing
1134 emulsions and particulate products using membranes. *Advances in Colloid and*
1135 *Interface Science*, 113(1), 1-20. doi:<https://doi.org/10.1016/j.cis.2004.10.002>
- 1136 Wandrey, C., Bartkowiak, A., & Harding, S. E. (2010). Materials for encapsulation. In N. J.
1137 Zuidam & V. Nedovic (Eds.), *Encapsulation technologies for active food ingredients*
1138 *and food processing* (pp. 31-100): Springer.
- 1139 Wang, J., Martínez-Hernández, A., de Lamo-Castellví, S., Romero, M.-P., Kaade, W.,
1140 Ferrando, M., & Güell, C. (2020). Low-energy membrane-based processes to
1141 concentrate and encapsulate polyphenols from carob pulp. *Journal of Food*
1142 *Engineering*, 281, 109996. doi:<https://doi.org/10.1016/j.jfoodeng.2020.109996>
- 1143 Wazarkar, K., Patil, D., Rane, A., Balgude, D., Kathalewar, M., & Sabnis, A. (2016).
1144 Microencapsulation: an emerging technique in the modern coating industry. *RSC*
1145 *Advances*, 6(108), 106964-106979. doi:<https://doi.org/10.1039/C6RA13237E>
- 1146 Wechtersbach, L., Poklar Ulrih, N., & Cigić, B. (2012). Liposomal stabilization of ascorbic
1147 acid in model systems and in food matrices. *LWT - Food Science and Technology*,
1148 45(1), 43-49. doi:<https://doi.org/10.1016/j.lwt.2011.07.025>
- 1149 White, S., Bennett, D. B., Cheu, S., Conley, P. W., Guzek, D. B., Gray, S., . . . Harper, N. J.
1150 (2005). EXUBERA: pharmaceutical development of a novel product for pulmonary
1151 delivery of insulin. *Diabetes Technology & Therapeutics*, 7(6), 896-906.
1152 doi:<https://doi.org/10.1089/dia.2005.7.896>
- 1153 Wong, S. W., Yu, B., Curran, P., & Zhou, W. (2009). Characterising the release of flavour
1154 compounds from chewing gum through HS-SPME analysis and mathematical
1155 modelling. *Food Chemistry*, 114(3), 852-858.
1156 doi:<https://doi.org/10.1016/j.foodchem.2008.10.030>
- 1157 Wu, K.-G., & Xiao, Q. (2005). Microencapsulation of fish oil by simple coacervation of
1158 hydroxypropyl methylcellulose. *Chinese Journal of Chemistry*, 23(11), 1569-1572.
1159 doi:<https://doi.org/10.1002/cjoc.200591569>
- 1160 Yin, Y., & Cadwallader, K. R. (2018). Spray-chilling encapsulation of 2-acetyl-1-pyrroline
1161 zinc chloride complex using hydrophobic materials: Feasibility and characterization
1162 of microcapsules. *Food Chemistry*, 265, 173-181.
1163 doi:<https://doi.org/10.1016/j.foodchem.2018.05.079>

1164 Yin, Y., & Cadwallader, K. R. (2019). Spray-chilling encapsulation of 2-acetyl-1-pyrroline
11165 zinc chloride using hydrophobic materials: Storage stability and flavor application in
21166 food. *Food Chemistry*, 278, 738-743.
31167 doi:<https://doi.org/10.1016/j.foodchem.2018.11.122>
4
51168 Zoet, F. D., Grandia, J., & Sibeijn, M. (2012). International patent WO/2012/047098.
6
71169 Zuidam, N. J., & Shimoni, E. (2010). Overview of microencapsulates for use in food
81170 products or processes and methods to make them. In N. J. Zuidam & V. Nedovic
91171 (Eds.), *Encapsulation technologies for active food ingredients and food processing*
101172 (pp. 3-29). New York, NY: Springer New York.
11
121173
13
14
15
16
17
18
19
20
21
22
23
24
25
26
27
28
29
30
31
32
33
34
35
36
37
38
39
40
41
42
43
44
45
46
47
48
49
50
51
52
53
54
55
56
57
58
59
60
61
62
63
64
65



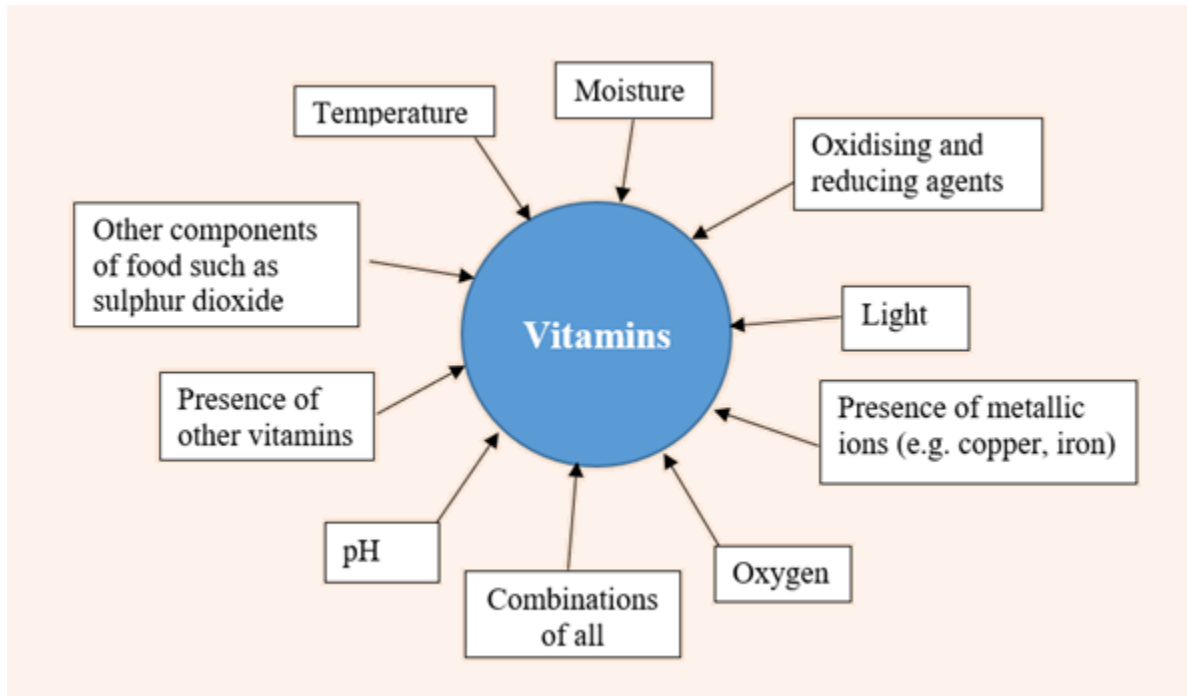
1

2 Figure 1. Schematic diagram of a complex coacervation encapsulation process. Recreated based on Ghosh (2006), Thies (2003) and Kim, Yang,
3 Park, Lim, and Cha (2017)

4

5

1



6

7 Figure 2. Main factors affecting vitamins' stability

1 Table 1. Wall materials commonly used for microencapsulation in the food industry

	Physical Property	Animal	Plant	Marine	Microbial	Synthetic/mineral
Carbohydrate	hydrophobic	Chitosan	Starch	Carrageenan	Xanthan gum	Cyclodextrin
			Glucose syrup	Agar	Gellan	Octenyl succinate starch
			Cellulose extracts	Alginate	Dextran	
			Guar gum		Curdlan	
			Pectin		Pullulan	
			Galactomannans			
			Soluble soy polysaccharides			
			Maltodextrins			
			Arabic gum			
			Karaya gum			
		Tragacanth gum				
Protein	hydrophobic	Gelatin	Gluten			
			Casein	Soy		
			Whey	Pea		
			Egg White	Rice		
			Lactoglobulin	Sorghum		
			Caseinate	Lupine		
				Zein		

Lipid	hydrophilic	Fat	Fat/oil	Ethyl cellulose
		Fatty acids	Fatty acids	Sorbitan esters
		Glycerides	Glycerides	Hydrogenated fat
		Phospholipids	Phospholipids	Paraffin Wax
		Beeswax	Plant sterols	Microcrystalline wax
		Shellac	Carnauba wax	
Polymers	hydrophobic			Polyethylene glycol
				Polyvinyl acetate
				Polyvinyl pyrrolidone

2 Modified based on Wandrey et al. (2010), Sobel et al. (2014), Javier D. Hoyos-Leyva et al. (2018), and Reineccius (2019)

3 Table 2. A summary of vitamin microencapsulation methods, wall material, encapsulation efficiency (EE), particle size, and release and
 4 retention from recent studies in food science

Core particle	Method	Wall materials	EE (%)	Particle size	Release and Retention notes	Source
Ascorbic acid	Spray drying	Taro starch	20.9	14.5 µm	Degradation rate was 0.006 at 13% RH and 0.029 at 72% RH at 55°C	(J. D. Hoyos-Leyva et al., 2018)
Ascorbic acid	W/O/W double emulsion	Sugar-free rebaudioside-sweetened model beverage	86.5-95	0.46- 0.80 µm	Higher entrapment efficiency, storage stability and improved chemical stability than single emulsion. 70.02-79.97% retention efficiency after 30 days Heat treatment at 80°C reduced the retention efficiency	(Kheynoor, Hosseini, Yousefi, Hashemi Gahruie, & Mesbahi, 2018)
Ascorbic acid and Folic acid	Liposome entrapment	Soybean phosphatidylcholine, cholesterol, chitosan, Tween 80	AA: 80.96 FA: 87.41	34.63- 138.58 - 249 nm	Hydroxyl radical scavenging activity-44.83- 57.15% Improved retention after 40 days storage- AA: 20.43-25.59% FA: 28.17-33.42% Control- 0.13-0.32%	(Jiao et al., 2018)

					Chitosan enhanced stability, encapsulation efficiency and antioxidant activity	
Ascorbic acid	Spray drying	Chitosan, modified chitosan and sodium alginate	Not tested	3 µm	43.6%- 45.4% yield Chitosan coating resulted in the highest yield and protection with release time 120 min.	(Jiao et al., 2018)
Ascorbic acid	Spray drying	Chitosan, tripolyphosphate cross-linking agent	45.05-58.30	6.1-9.0 µm	Crosslinked chitosan improved the efficiency of encapsulation High stability of ascorbic acid observed The increased concentration of crosslinking agent decreased encapsulation efficiency	(K. G. H. Desai & Park, 2005)
Ascorbic acid	Liposome	Soy phosphatidylcholine, stearic acid, calcium stearate	85.31-86.33	0.5-400 µm	Protective effect of vitamin C in orange juice from pasteurisation (65°C for 30 min) was found.	(Marsanasco et al., 2011)
Ascorbic acid	Thermal phase separation (TPS)	Ethyl cellulose, coacervation-inducing agent	TPS-NA	TPS 16.9-61.9 µm	Protection from oxidation, no colour change after one month storage	(M. S. Uddin, 2001)
	Melt dispersion (MED)	Carnauba wax	MED- NA	MED ~50 mm	Carnauba wax and β-cyclodextrin were effective coating material	

	Solvent evaporation (SE)	Ethyl cellulose, plasticizer (triethyl citrate)	SE- NA	SE NA		with higher encapsulation efficiency and stabilisation of acid
	Spray drying (SD)	gel, starch, ethyl cellulose, cyclodextrin,	-	SD- below 50	SD 90-280 µm	
β-Carotene	Spray drying	N-octenyl succinate anhydride modified starches (HI-CAP, CAPSUL and CAPSUL TA)		Not tested	118-159 nm	Retention after 30 days: 71.87% at RH 11% and 58.04% at RH 97% The relation between oxygen permeability of the matrixes and vitamin degradation was found.
β-Carotene (Piquillo pepper extract)	Emulsion-spray drying	Gum arabic		77.1	5.46 µm	β-Carotene were stable when entrapped in 35 days storage at room temperature. No other storage conditions were given. (Romo-Hualde, Yetano-Cunchillos, González-Ferrero, Sáiz-Abajo, & González-Navarro,

Retinol	Liposome entrapment	Combination of phosphatidylcholine and cholesterol	94.52- 99.31	31.7-41.42 μm	After 10 days of storage- 90% at pH 7, 87% at pH 9, and 39.46% at pH 5	(S. C. Lee et al., 2005)
Folic acid	Impregnation method (Pérez-Esteve et al., 2015)	Inorganic gated mesoporous silica particles (calcinated MCM-41)	75	862 \pm 59 nm	Higher stability facilitating target release, protection from photodegradation, thermostability, high retention (84–94.5%) after 28 days of storage.	(Ruiz-Rico et al., 2017)
Folic acid	Spray drying	Horse chestnut starch and β -cyclodextrin	Starch (SF)- 57.29 β -cyclodextrin (BF)- 76.10	SF 28.26-227.3 μm BF 30.09-145.9 μm	Protection from environmental factors ensuring release in the GI tract. Potential of horse chestnut starch in stabilising FA.	(Ahmad et al., 2017)
Folic acid	Spray drying	Rice starch with gum arabia or guar gum or xanthan gum or κ -carrageenan	65-85	15-45 μm	Thermostability, higher retention with κ -carrageenan and alg-pec, 97% retention after boiling the noodles for 3min 20s - 3min 40s	(Hau, 2008)

			combination of alginate and Low methoxyl pectin						
Folic acid	Electrospinning		Sodium alginate, pectin, polyethylene oxide	Not tested		140-131 nm		Almost 100% retention at controlled light and pH Higher stability when crosslinked with ethanol	(Alborzi et al., 2013)
5-methyl tetrahydrofolic acid	Combination of spray drying and extrusion		Combination of pectin and alginate	spray drying- 60 Extrusion- not tested				84-94.5% retention after thermal treatment (boiling/autoclaving)	(Shrestha et al., 2012)
Folic acid	Extrusion		alginate, pectin, xanthan gum, gelatine, iota-carrageenan in a single form or combination	55-89		Not tested		Alg-pec coating significantly enhanced the stability and pressure tolerance properties	(Madziva et al., 2005, 2006)
Pantothenic acid	Liposome and Extrusion		Phospholipon 90 G, alginate, pectin	Liposome- 75 Extrusion- 60		Liposome- 100-240 nm Extrusion- 240-300 μm		After 14 days of storage, only 20% release occurred at pH 4 Thermostable capsules formed No further effect of alginate-pectin was noted	(Ota et al., 2018)
Thiamine	Liposome		Phosphatidylcholine	97		108-1351 nm		Stability/retention was not tested. Chloroform was used to dissolve phosphatidylcholine, therefore not suitable for human	(Juveriya Fathima et al., 2016)

						consumption	
Riboflavin	Cold-set gelation and Immobilisation	Whey protein isolate	24.5-57	Not tested	Not tested	Fast release of riboflavin, no or little protection Controlling diffusional losses is required to make use of this carrier Release rate increased with the decreased hydrophobicity of the active	(O'Neill, Egan, Jacquier, O'Sullivan, & Dolores O'Riordan, 2014, 2015)
Riboflavin	Spray drying	Gum Arabic, β -Carotene	Not tested	Not tested	Not tested	20% photo protection Gum arabia alone or with β -carotene can protect the self-photoinduced degradation of RF, but with β -carotene is more efficient	(Boiero et al., 2014)
Vitamin B ₁₂	Coextrusion	Rice flour, guar gum, xanthan gum, carboxymethyl cellulose	Not tested	Not tested	Not tested	Best retention (77%) of vitamin B12 achieved at 140°C/150 rpm/4 kg/h extrusion parameters Carboxymethyl cellulose enhanced physico-functional properties	(Bajaj & Singhal, 2019)

Vitamin B ₁₂	Spray drying	chitosan, modified chitosan and sodium alginate	Not tested	Not tested	41.8%- 55.6% yield Chitosan coating resulted in the highest yield and protection with release time 120 min but rough surface	(Estevinho, Carlan, Blaga, & Rocha, 2016)
-------------------------	--------------	---	------------	------------	---	---

