- Systematic review on n-3 and n-6 PUFA intake in European countries in light of the 1
- current recommendations focus on specific population groups 2
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- 38

40	List of abbre	eviations
41	AI	adequate intake
42	ARA	arachidonic acid
43	ALA	α-linolenic acid
44	DHA	docosahexaenoic acid
45	DPA	docosapentaenoic acid
46	DRI	dietary reference intake
47	DRV	dietary reference value
48	EFSA	European Food Safety Authority
49	EPA	eicosapentaenoic acid
50	FA	fatty acid
51	FAO	Food and Agriculture Organization
52	LA	linoleic acid
53	PUFA	polyunsaturated fatty acid
54	WHO	World Health Organization

57 Abstract

Background: Earlier reviews indicated that in many countries adults, children and adolescents
consume on average less polyunsaturated fatty acids (PUFAs) than recommended by the
FAO/WHO.

Summary: We now systematically reviewed the intake of total and individual n-3 and n-6
PUFAs in European infants, children, adolescents, elderly, and pregnant/lactating women.
Results were evaluated against recommendations of the European Food Safety Authority (EFSA).

Key messages: Fifty-three studies from 17 different European countries reported intake of 65 total n-3 and n-6 PUFAs and/or individual n-3 or n-6 PUFAs in at least one of the specific 66 population groups: 10 in pregnant women, 4 in lactating women, 3 in infants 6-12 mo, 6 in 67 children 1-3 y, 11 in children 4-9 y, 8 in adolescents 10-18 y and 11 in elderly >65 y. Mean 68 linoleic acid intake was within the recommendation (4E%) in 52% of the countries, with 69 inadequate intakes more likely in lactating women, adolescents and elderly. Mean α-linolenic 70 acid intake was within the recommendation (0.5E%) in 77% of the countries. In 26% of the 71 countries, mean eicosapentaenoic acid and/or docosahexaenoic acid intake was as 72 recommended. These results indicate that intake of n-3 and n-6 PUFAs may be suboptimal in 73 74 specific population groups in Europe.

76 Introduction

N-3 and n-6 polyunsaturated fatty acids (PUFAs) — particularly the longer chain, more 77 unsaturated members of these families, but also the plant-derived essential members — play a 78 vital role in human health from conception onwards: through every stage of human 79 development, maturation and aging, including roles in cell membrane composition, 80 metabolism, signal transduction and amplification, as well as in gene expression [1]. An 81 adequate intake of PUFAs is of critical importance during early life and plays an essential 82 role in supporting growth and development. In addition, for the general population an 83 adequate intake of n-3 and n-6 PUFAs is recommended for prevention of cardiovascular 84 diseases [2]. However, specific population groups may have relatively higher requirements of 85 these fatty acids and thus more at risk of inadequate intake. These specific population groups 86 include pregnant and lactating women because of increased PUFA requirements for growth, 87 neurological and immune function of their infants [3,4]. Similarly, infants, children and 88 adolescents have relatively high nutrient requirements compared to adults to support rapid 89 growth and development, but complementary foods, school meals and dietary habits during 90 childhood and adolescence may not provide sufficient amounts of nutrients, notably PUFAs 91 [5-8]. Finally, elderly have altered nutrient requirements, because of changes in body 92 93 composition and physical activity and the presence of disease, and altered nutrient intakes 94 because of low variety of food they eat, reduced appetite, loss of sensory appreciation of food, dentition and swallowing problems, the presence of disease, and social issues and thus 95 may be at risk of inadequate PUFA intake, besides other important micronutrients [9]. 96

Recent reviews on dietary fat and fatty acid intake in different countries around the world 97 [10-14] are mainly based on data from national dietary surveys that do not include a 98 representative subsample of these specific subgroups of the population. Moreover, in many of 99 100 these dietary surveys, intake data of individual PUFAs are not reported. Therefore, the aim of this systematic review is to evaluate the available data on n-3 and n-6 PUFA intake in seven 101 102 specific population groups in Europe: (1) pregnant women, (2) lactating women, (3) infants (6-12 mo), (4) young children (1-3 y), (5) older children (4-10 y), (6) adolescents (10-18 y) 103 104 and (7) elderly (>65 y). First, an overview is given of the current available recommendations for PUFA intake in European countries. Second, the available intake data are summarised and 105

106 reported for each group, followed by a discussion of the gaps between reported intake and the

- 107 EFSA recommendations. Third, an overview of studies describing the main food sources of 108 the various PUFAs in these population groups is presented.
- 109

110 Methods

111 A more detailed methods description is given in Supplementary text 1.

112 Evaluation of current recommendations for PUFA intake in European countries

For the evaluation of current recommendations for PUFA intake in European countries, we 113 updated the systematic review on dietary reference intake, nutritional goals and dietary 114 guidelines for fats and fatty acids by Aranceta et al. that was published in 2012 [15] with 115 PUFA recommendations for specific population groups in Europe, using the search strategy 116 by Aranceta et al. [15], excluding trans fatty acids. The search was conducted using PubMed 117 and Scopus from January 2011 (date of Aranceta search) to April 20th 2015. In addition, a 118 manual search on individual European country recommendations was performed via Google 119 with latest update done in October 2016. 120

121 Evaluation of current intake of total and specific n-3 and n-6 PUFAs in the European

- 122 diet for the specific population groups
- 123
- 124 Criteria for considering studies in this review

Type of study: Observational studies and national dietary surveys were the primary focus of this review. Randomised control trials or other experimental studies were included only if they reported baseline data and/or data for the control group.

128 *Type of exposure:* Studies were included if they reported data on intake of at least **one** of the 129 following: α -linolenic acid (ALA), eicosapentaenoic acid (EPA), docosapentaenoic acid 130 (DPA), docosahexaenoic acid (DHA), total n-3 PUFAs, linoleic acid (LA), arachidonic acid 131 (ARA) or total n-6 PUFAs. We excluded studies reporting only the total PUFA intake 132 without any distinction on individual classes as mentioned above. *Type of population:* The following seven subgroups of the population were the focus of interest in this review: pregnant women, lactating women, infants (6-12 mo), young children (1-3 y), older children (4-10 y), adolescents (10-18 y), elderly (>65 y). The age ranges are defined in accordance with the EFSA Comprehensive European Consumption Database [16]. Studies in specific disease populations were excluded. Only studies conducted in European countries were included. Inclusion and exclusion criteria can be found in the abstract review form at PROSPERO with registration number CRD42014014717.

Period of time for exposure measurement: Studies conducted after the 1st of January 2000
were eligible for inclusion. Studies conducted before 2000, but reported in a publication
published after 2000, were excluded.

143 Search methods for identification of studies

PubMed, Scopus and Cochrane Central Register of Controlled Trials (CENTRAL) databases 144 were searched for papers published from January 2000 until November 2015, using text 145 words with appropriate truncation and relevant indexing terms. The search was in the form 146 [n-3 and n-6 PUFA terms] and [terms for intake] and [terms for the specific subgroup 147 considered] and [limit to humans] and [limit to 2000 - current]. The full search strategy for 148 the Ovid MEDLINE database can be accessed via PROSPERO file CRD42014014717; the 149 searches of the other databases were based on this strategy. Reference lists of all eligible 150 papers and relevant systematic reviews were searched for additional studies. 151

152 Data selection

The titles and abstracts of studies identified by the search were screened by a single reviewer and clearly irrelevant studies were excluded. The full text reports of all potentially relevant studies were obtained and assessed independently for eligibility by two independent reviewers. The systematic review software Covidence (<u>www.covidence.org</u>) was used to facilitate screening of the literature. Any disagreement was resolved by discussion.

158 **Data extraction**

Standardised forms were used for data extraction and management. For each included study,the following data were extracted and brought together into one large database: name of first

161 author, year of publication, year(s) of data collection, participant characteristics (age, sex, n, 162 the subgroup they belong to), method used for dietary assessment and intake data on total energy (MJ/day), total fat (g/day and E%), total PUFAs (g/day and E%), total n-6 PUFAs 163 (g/day and E%), LA (g/day and E%), ARA (mg/day), total n-3 PUFAs (g/day and E%), ALA 164 (g/day and E%), EPA (mg/day), DPA (mg/day) and DHA (mg/day). When necessary, units of 165 measurement were converted to a standard form (g/day, mg/day or energy percentage (E%))166 expressed in mean and SD to facilitate comparison across studies. When only median and 167 quartiles were given, the mean was calculated as the average of the median, 25th (P25) and 168 75th (P75) percentiles. The standard deviation was calculated as P75-P25/1.35. Unpublished 169 data for countries that participated in the HELENA study in adolescents were included based 170 171 on personal communication with the researchers involved, as the original publication only reported data of the overall international sample [17]. 172

173 **Data assessment**

174 All fatty acid intake data were evaluated against the EFSA recommendations [18], which are the most recent recommendations that are set by a recognised Europe-wide health authority. 175 Because of the absence of total n-3 and total n-6 PUFA intake recommendations, ALA and 176 LA recommendations were used to evaluate total n-3 or n-6 PUFA intake. ALA and LA are 177 the biggest contributors to total n-3 and total n-6 PUFA intake, respectively. Per country and 178 population group, intake data were extracted to evaluate whether the intake of the different 179 PUFAs was in line with the EFSA recommendations. Where multiple datasets were available 180 for one country and specific population group, the average of these datasets was calculated 181 (weighted for n), and this average was used to define whether the intake data were in line or 182 183 below the relevant EFSA recommendation.

184 Evaluation of major dietary contributors to PUFA intake

Another literature search was undertaken in PubMed to find data on dietary sources of PUFAs in the considered population subgroups as well as data on the PUFA composition of human milk. The search was conducted with terms describing fatty acids, diet and habits. The full search strategy can be found on PROSPERO with registration number CRD42014014717. In addition, all identified publications with intake data were screened for relevant data on food sources. For the data on dietary sources excluding human milk, studies 191 were excluded, if the data were collected before 2000. For data on human milk publications

- 192 from 1990 onwards were included because of limited available data published after 2000.
- 193

194 **Results**

195 Evaluation of recommendations for PUFA intake in European countries

In addition to the EFSA and FAO/WHO recommendations, we identified six individual country and four multi-country recommendations for intake of PUFAs. In general, recommendations were developed by scientific experts and health authorities after commissioning expert systematic evidence based reviews of mainly human observational and intervention studies in order to guide the authority's decision. Table 1 gives an overview of the recommendations identified.

The criteria used to set the specific recommendations (Table 1) are not always clearly described, but include provision for adequate growth for infants and children, prevention of clinical deficiency and provision for good health, encompassing prevention of cardiovascular diseases after the age of 2 years.

Mostly, two kinds of Dietary Reference Values (DRVs) were derived for PUFAs, the Adequate Intake (AI), e.g. set by EFSA and many European countries, and the Acceptable Macronutrient Distribution Range (AMDR), set by FAO/WHO [19]. For LA, the FAO/WHO additionally set an Estimated Average Requirement (EAR).

In general, the DRVs for the different n-3 and n-6 PUFAs for Europe and European countries 210 differ with regard to age group, type of fatty acid and unit of expression, which made it 211 difficult to compare. In general, DRVs were reported for LA, ALA and EPA+DHA or DHA, 212 while no recommendations were reported for DPA, and for ARA recommendations were only 213 214 reported for infants 0-6 mo of age. If guidelines were formulated for pregnant and breastfeeding women, they were focused on a specific increase in the intake of DHA. 215 Recommendations for children >2 y of age were not age-specific and were often similar to 216 those for adults. For elderly, no recommendations were derived, as no specific needs for any 217 of the PUFAs were deemed evident. 218

Based on these outcomes, and as explained in the method section, the EFSA recommendations [18] have been selected to evaluate adequacy of fatty acid intake data in the current review. In absence of EFSA recommendations for elderly, adult recommendations were used instead.

223 Evaluation of intake of total and specific n-3 and n-6 PUFAs

Altogether 5404 titles and abstracts were identified via the electronic, bibliographic and 224 additional expert searches, and 267 of them appeared to be potentially relevant. Finally, 49 225 studies fulfilled the inclusion criteria. For pregnant women, 10 different publications were 226 included reporting PUFA intake data in 11 different European countries. For lactating 227 women, only four studies from four different countries were included, all with a limited 228 sample size (14 to 63 women). For infants aged 6-12 mo, only three studies from three 229 different countries were found, of which the study in German infants reported intake data, 230 when children were 6 and 9 mo old [20]. For young children aged 1-3 y, six studies from six 231 different countries were included. For older children aged 4-9 y, 11 studies from 10 countries 232 were identified, of which some studies reported intake in multiple countries. For adolescents 233 aged 10-19 y, eight studies from 11 different countries reported data on individual PUFA 234 intake. For elderly aged ≥ 65 y, 11 studies from nine countries were included of which one 235 study from Hungary [21] reported data in elderly aged ≥ 60 y. Details of intake data can be 236 found in supplementary tables 1-7. 237

In general, intake data were reported for ALA, LA, total n-3 PUFAs, total n-6 PUFAs, EPA
and DHA, whereas for DPA and ARA, very little intake data were found.

Table 2 gives an overview of available intake data and summarizes the proportions of 240 countries where current intake data were in line with the EFSA recommendations, for each 241 population. Detailed information about the countries included in each population groups can 242 be found in the supplementary tables 1-7. Across all population groups, mean LA intake was 243 below the recommendation of \geq 4 E% in 48% of the countries, with low intake more likely in 244 lactating women, adolescents and elderly, whereas mean ALA intake was below the 245 recommendation of ≥ 0.5 E% in 23% of the countries (Table 2). Across all population groups, 246 247 mean EPA and/or DHA intake was lower than the EFSA recommendation in 76% of the

countries, and low intake was in particular of concern in pregnant and lactating women and in

249 infants, children and adolescents.

250 Evaluation of major dietary contributors to PUFA intake

251 Dietary sources of individual PUFAs

Data on the contribution of different food groups and supplements to the intake of the various 252 253 PUFAs were limited to three studies, including a Belgian study in children aged 2.5-6.5 y [22], the HELENA study in adolescents aged 12.5-17.5 y [17] and a study in Dutch elderly 254 aged ≥ 70 y [23]. In Belgian children, fats and oils were the major contributors to intakes of 255 LA (23.6%) and ALA (33.1%), followed by cereal products with 17.6% and 13.5%, 256 respectively [22]. Meat, poultry and eggs were the main contributors to ARA intake (72.0%), 257 and fish and seafood were the main contributors to EPA (83.5%), DPA (57.8%) and DHA 258 (75.7%) intake [22]. In adolescents in the multiple country HELENA study [17], the food 259 group "meat, fish, eggs and meat alternatives" was the largest contributor to the intake of LA 260 (31.7%), ALA (21.5), ARA (54.2%), EPA (92.3%), DPA (94.9%) and DHA (85.8%). In 261 Dutch elderly, fats and oils were the main contributor to LA (39%) and ALA (36%) intake, 262 whereas fish and shellfish (29%) and meat and meat products (28%) were the main 263 contributors to EPA and DHA intake. 264

265 Individual PUFA content of human milk

LA and ALA are the major PUFAs present in human milk. According to a review of 14 266 studies from nine European countries [24,25], the median (range) content of human milk is 267 11.0% wt/wt (6.9-16.4) for LA and 0.9% wt/wt (0.7-1.3) for ALA. A descriptive meta-268 analysis including 65 studies of 2474 women worldwide indicated a mean (± SD) 269 concentration of DHA and ARA in human milk of $0.32 \pm 0.22\%$ and $0.47 \pm 0.13\%$ of total 270 fatty acids, respectively [26]. European countries tended to report higher DHA levels [27], 271 with the highest DHA concentrations in coastal regions such as Greece, Italy and Spain 272 (Supplementary table 8) [26]. DHA levels in human milk vary considerably among women 273 and are strongly influenced by maternal diet, e.g. fish and seafood intake [27-30], whereas 274 275 ARA concentrations in human milk are less sensitive to maternal dietary ARA intake [26, 30]. 276

278 **Discussion**

We provided an overview of the available data on recommendations, dietary intake and sources of total and individual n-3 and n-6 PUFAs in European pregnant women, lactating women, infants, young children, older children, adolescents and elderly.

282 **Recommendations**

Since the review of Aranceta et al. [15] published in 2012, we found that some 283 recommendations had not been updated (FAO/WHO, EFSA, The Netherlands, Spain), while 284 others had (France, Nordic countries, Belgium). In addition, we identified new 285 recommendations that were not published at the time of the earlier review (DACH countries -286 Germany, Austria, Switzerland –, Poland; France and Spain for infants and children). Similar 287 to FAO/WHO and EFSA, no specific dietary recommendations were formulated for elderly 288 in Europe, despite the fact that some countries, (e.g. France, 2001) discussed the specific 289 290 dietary needs of the elderly [31].

For ARA, recommendations are limited to infants aged 0-6 mo and based on human milk 291 content. However, the functional effect of increased ARA intake in this age group, as well as 292 in other age groups, is still much discussed [32-33]. The role of preformed DHA in visual 293 development and brain growth and functioning in foetal life and early infancy has been 294 demonstrated and translated into specific recommendations for pregnant and lactating women 295 and infants. After the age of 2 y, based largely on their preventive effect on CVD and 296 beneficial effects for neurodevelopment, recommendations have been formulated for EPA + 297 DHA, since these PUFAs are often consumed in combination, for instance in seafood and 298 supplements. In contrast to recommendations in European countries and EFSA, some 299 authorities, such as those of Australia and New Zealand [34] have set an AI based on the 300 concept of essentiality, i.e. on the median intake for children and adults in a population 301 without apparent essential fatty acid deficiency, resulting in generally lower recommended 302 intakes. Whereas no recommendations for DPA were formulated in Europe, the health 303 authority from Australia and New Zealand included DPA in its recommendations, i.e. the 304 recommended intake for long chain n-3 PUFAs including EPA + DPA + DHA. 305

Germany and the Netherlands recently reviewed their intake recommendations, but did not 306 307 set any reference value for fatty acids, and rather formulated food-based dietary guidelines [35,36]. Moreover, two important authorities, the US Institute of Medicine and the WHO 308 Nutrition Guidance Expert Advisory Group (NUGAG) intend to update their PUFA intake 309 recommendations [37]. In 2014, the US and Canadian governments collaboratively indicated 310 that nutrient reference values for n-3 PUFAs needed to be updated with priority, based on 311 public health and/or policy importance. The committee is considering to incorporate chronic 312 disease endpoints into the setting of DRI values [38]. 313

314 Intake data

In summary, we found that current information on intake of PUFAs, with especially individual long chain PUFAs in specific age groups being at risk for an inadequate PUFA intake across Europe is limited, and we identified many gaps in the current knowledge. Our findings show that EFSA recommendations for intake of LA, ALA and EPA+DHA were not met in almost half, a quarter and three-quarters of the countries, respectively. This is in line with findings of earlier studies that have evaluated intakes of these fatty acids in children, adolescents and adults worldwide [12,13].

The lowest number of available studies was found for lactating women and infants. However, 322 these population groups are of particular interest as PUFA intake is very important for rapid 323 brain growth and development during infancy. In addition, our findings indicate that there is a 324 lack of data on ARA and DPA intake. The scarcity of data on ARA and DPA intakes limits 325 the assessment of intake adequacy especially for at-risk population groups. A potential 326 explanation is that information on individual PUFAs in the European food composition tables 327 is often missing. Therefore, it may be necessary to combine information from different food 328 composition tables in order to make a complete estimation of the individual PUFA intake. In 329 the future, it would be relevant that ARA and DPA are also included in food composition 330 analyses. 331

Furthermore, large heterogeneity between studies with regard to methodologies and data presentation, made it difficult to compare the available intake data across countries. First, studies used different methods for dietary assessments (e.g. 24 hr recall, food frequency questionnaires, dietary record). Some of these methods assess intake over a short term (1 to 3

d), which may not capture the intake of foods that are not consumed on a daily basis (e.g. fish 336 and seafood) and consequently underestimate intake of EPA and DHA. However, when 337 dietary assessments of the study population cover all weekdays and seasons, the estimation of 338 mean intakes on population level will be reasonable [39]. In a limited number of the 339 identified studies, statistical methods were applied to correct for this between-day variability. 340 However, even when diets were evaluated over a longer period, foods may have been 341 grouped together (e.g. n-3 PUFA rich fish and fish less rich in n-3 PUFAs), which may 342 reduce the specificity of intake data. Secondly, some studies took the use of food supplements 343 into consideration [40,41,42] or even selected participants based on very high seafood 344 consumption (a survey in French coastal populations) [43], while others specifically excluded 345 346 fish oil supplement users [44-46] and other studies just did not report whether or not supplements were taken into account. Also, in some countries the use of food supplements in 347 general (not specifically n-3 PUFA supplements) by pregnant and lactating women is quite 348 common (e.g. in the study of Rodriguez-Bernal, 55.8% of the women report the use of 349 multivitamin supplements) [42], which may explain the large variation in PUFA intake. 350 Unfortunately, no data could be found on the average use of n-3 PUFA supplements by 351 European pregnant women, which indicated the need to collect data on supplement use in 352 addition to dietary intake in future studies. Thirdly, different metrics were used to represent 353 the intake data: some studies reported the median, while others used the arithmetic or 354 geometric mean, sometimes in combination with the standard deviation or with a percentile 355 range. This shows a need to harmonise the way data on PUFA intake are presented; e.g. in 356 case of skewed distributions (e.g. EPA and DHA intake) medians should be reported. The 357 studies included in our review mainly reported data in means and only a few reported 358 medians. Moreover, comparing the mean PUFA intake of a population with the PUFA 359 recommendation does not allow to determine the percentage of the population not meeting 360 the PUFA recommendation. To do so, the distribution of the intake in the population has to 361 be known. Fourthly, different units to express the intake data were used: intake of fatty acids 362 can be expressed in (m)g/day, E%, or % of total fat. As it was not always possible to convert 363 intake to a common unit based on the available information in the publication, comparisons 364 365 between different studies and evaluation to EFSA recommendations were not possible for some data. 366

This review also showed that information on the foods and food groups contributing to the intake of different PUFAs, and in which amount they contribute, is largely lacking and if available, food groups did often not report data for individual PUFAs and were not comparable between studies.

371 Strengths and limitations

This study has some important strengths. First, the overview provided is based on the 372 application of a systematic and standardised approach to screen the literature and identify the 373 studies to be included. Secondly, data extraction from the selected studies was conducted in a 374 harmonised way in order to obtain useful data for comparison. However, some limitations 375 also need to be mentioned. First, most of the available studies were not based on a random 376 sampling procedure or on a national representative sample. Secondly, the EFSA 377 recommendations for LA and ALA intake were used to evaluate if intake of total n-6 PUFAs 378 and total n-3 PUFAs was adequate, which may have led to an overestimation of countries 379 with sufficient total n-6 and n-3 PUFA intake. While the approximation is small for n-6 380 PUFAs (LA contributes on average to ~99% of total n-6 PUFA intake), it is higher for n-3 381 PUFAs (ALA contributes on average to ~80% of n-3 PUFA intake). 382

383 Recommendations for future research

Given the limited data available on individual n-3 and n-6 PUFA intake in specific population 384 385 groups in Europe, our key recommendation is that the EU should develop harmonised data collection systems that will provide a robust and reliable database on the intake of individual 386 PUFAs. This would be needed to establish evidence-based guidelines for public health 387 programs aiming to improve fatty acid intake and for monitoring and evaluation of the 388 effectiveness of these programs. Moreover, greater national commitment to and consistency 389 in the provision of intake data is required in order to allow reasonable comparative analyses 390 between different countries. Future more detailed intake data should be used to re-evaluate 391 current recommendations for the general population and specific population groups, 392 393 including those with different pathologies and genetic polymorphisms. This would particularly be relevant for ARA and DPA for which extremely limited data are available. 394 While studies in infants have suggested the role of ARA in combination with DHA on 395 physical growth and cognitive development [47-50], more recent data indicate that ARA and 396

its derivatives may also play key roles in cardiovascular, inflammatory and immune function,
which have not yet been adequately investigated [34]. Similarly, the potential role of DPA in
cardiovascular disease, immune function, and psychiatric and cognitive health needs further
investigation [51].

401 **Conclusion**

The available data indicate that mean intake of EPA and DHA and to a lesser extent of LA 402 403 and ALA may be suboptimal compared to EFSA recommendations for a significant part of specific population groups in Europe. More nationally representative surveys including 404 subsamples of specific population groups and data on relevant individual PUFAs are required 405 to clarify the need for specific public health measures to optimise PUFA intake in Europe. 406 Also, recommendations for nutrient requirements should be developed for the elderly 407 population as well as recommendations for intake of total n-6 and n-3 PUFAs in all 408 409 population groups.

410 **Potential conflict of interest**

411 Dr Petisca is an employee of Bunge Europe, Dr Fleith is an employee of Nestlé Research 412 Centre, Dr Eilander is an employee of Unilever Research and Development, Dr Eussen is an 413 employee of Danone Nutricia Research, Prof. Forsyth is a consultant for DSM Nutritional 414 Products Ltd. and Prof. Calder is an advisor to Pronova BioPharma (part of BASF), Danone 415 Research Centre for Specialized Nutrition, DSM, Cargill, and Smartfish.

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Nestlé, SQM Europe, Ülker Bisküvi and Unilever (Nutrient Intake Optimisation Task Force) and Abbott Nutrition, Arla Foods, Danone, DSM, Mead Johnson Pediatric Nutrition Institute and Nestlé (Early Nutrition and Long-Term Health Task Force). For further information about ILSI Europe, please email <u>info@ilsieurope.be</u> or call +32 2 771 00 14. The opinions expressed herein and the conclusions of this publication are those of the authors and do not necessarily represent the views of ILSI Europe nor those of its member companies.

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intake data per survey centre.

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632 Table 1. Current recommendations for PUFA intake in specific population groups according to different national and international

- 633 authorities^a
- 634

Region/Country (Authority)	Age / Age group	Year	LA	ARA	Total n-3 PUFAs	ALA	EPA	DHA	EPA+DHA
Europe (EFSA) [18,52]	Pregnant and lactating women	2010	4.0 E%	-	-	0.5 E%	-	100.0-200.0 mg/d ^b	250.0 mg/d
	0-6 mo	2013	4.0 E%	140.0 mg/d	-	0.5 E%	-	100.0 mg/d	-
	7-24 mo	2010	4.0 E%	-	-	0.5 E%	-	100.0 mg/d	-
	> 2y	2010	4.0 E%	-	-	0.5 E%	-	-	250.0 mg/d
	Elderly (=Adults)	2010	4.0 E%	-	-	0.5 E%	-	-	250.0 mg/d
Worldwide FAO / WHO) [19]	Pregnant and lactating women	2008	2.5-9.0 E% (ADMR) 2.0 E% (EAR) 2.0-3.0 E%	-	0.5-2.0 E% (AMDR)	>0.5 E% (L-AMDR)	-	200.0 mg/d (ANR)	300.0 mg/d (ANR)
	0-6 mo	2008	Human milk	0.2-0.3 E% Human milk (U-AMDR)	-	0.2-0.3 E%	-	0.1-0.2 E% <0.8 E% (U-AMDR)	-
	6-24 mo	2008	3.0-4.5 E% < 10.0 E% (U-ADMR)	-	-	0.4-0.6 E% <3.0 E% (U-AMDR)	-	-	10.0-12.0 mg/kg BW
	2-4 y	2008	(U-AMDR)	-	0.5-2.0 E% (AMDR)	>0.5 E% (L-AMDR)	-	-	100.0-150.0 mg/d
	4-6 y	2008	2.5-9.0 E% (AMDR) 2.0 E% (EAR) 2.0-3.0 E%	-	0.5-2.0 E% (AMDR)	>0.5 E% (L-AMDR)	-	-	150.0-200.0 mg/d
	6-10 y	2008	2.5-9.0 E% (AMDR) 2 E% (EAR) 2.0-3.0 E%	-	0.5-2.0 E% (AMDR)	>0.5 E% (L-AMDR)	-	-	200.0-250.0 mg/d
	10-18 у	2008	2.5-9.0 E% (AMDR) 2.0 E% (EAR) 2.0-3.0 E%	-	0.5-2.0 E% (AMDR)	>0.5 E% (L-AMDR)	-	-	250.0-2000.0 mg/d (AMDR)
	Elderly (=Adults)	2008	2.5-9.0 E% (AMDR) 2.0 E% (EAR) 2.0-3.0 E%	-	0.5-2.0 E% (AMDR)	>0.5 E% (L-AMDR)	-	-	250.0-2000.0 mg/d (AMDR)
Belgium (Superior Health	0-6 mo	2009	4.4 g/d	-	-	0.5 g/d	-	-	-
Council of Belgium) ^e [53]	7-12 mo	2009	4.6 g/d	-	-	0.5 g/d	-	-	-
	1-18 y	2009	2.0-5.0 E% ^d	0.1-0.3 E%	-	0.5-1.5 E%	0.1-0.2 E%	0.1-0.4 E%	-
	Elderly (=Adults)	2009	>2.0 E%	-	1.3-2.0 E%	>1.0 E%	-	-	0.3 E%
Belgium (Superior Health	0-6 mo	2016	4.0 E%	-	-	1.0 E%	-	-	-
Council of Belgium) [53]	7-12 mo	2016	4.0 E%	-	-	1.0 E%	-	100.0 mg/d	-
	1-3 yrs	2016	4.0 E%	-	-	1.0 E%	-	100.0 mg/d	-
	3-18 yrs	2016	4.0 E%	-	-	1.0 E%	-	-	250.0-500.0 mg/d
	Elderly (=Adults)	2016	4.0 E%	-	-	1.0 E%	-	-	250.0-500.0 mg/d
France (ANSES) [4]	Pregnant women (2050 kcal)	2011	4.0 E%	-	-	1.0 E%	-	250.0 mg/d	500.0 mg/d
	Lactating women (2250 kcal)	2011	4.0 E%	-	-	1.0 E%	-	250.0 mg/d	500.0 mg/d
	0-6 mo	2011	2.7 E%	0.5% of total FA	-	0.5 E%	EPA < DHA	0.3% of total FA	-
	6-36 mo	2011	2.7 E%	-	-	0.5 E%	-	70.0 mg/d	-

	3-9 у	2011	4.0 E%	-	-	1.0 E%	-	125.0 mg/d	250.0 mg/d
	10-18 y	2011	4.0 E%	-	-	1.0 E%	-	250.0 mg/d	500.0 mg/d
	Elderly	2011	4.0 E%	-	-	1.0 E%	250.0 mg/d	250.0 mg/d	500.0 mg/d
Germany – Austria –	Pregnant and lactating women	2013	2.5 E%	-	-	0.5 E%	-	-	-
Switzerland (D-A-CH) ^c	0-4 mo	2013	4.0 E%	-	-	0.5 E%	-	> 200.0 mg/d	-
[54]									
	4-12 mo	2013	3.5 E%	-	-	0.5 E%	-	-	-
	1-4 y	2013	3.0 E%	-	-	0.5 E%	-	-	-
	4-19 y	2013	2.5 E%	-	-	0.5 E%	-	-	-
	Elderly	2013	2.5 E%	-	-	0.5 E%	-	-	-
Switzerland	Pregnant and lactating women	2013	-	-	-	-	-	> 200.0 mg/d	-
(Eidgenössische	Elderly (=adults)	2013	-	-	0.5-2.0 E%	-	-		500.0 mg/d
Ernährungskommission) [55]									
The Netherlands (Health	0-5 mo	2001	0.6 g/kg BW	0.0 g/kg BW	0.1 g/kg BW	-	-	0.0 g/kg BW	
Council of the	6 mo-18 y	2001	-	-	-	-	-	-	150.0-200.0 mg/d
Netherlands) [56]	Elderly (=adults)	2001	2.0 E%	1.0 E%		-	-		450.0 mg/d ^e
Nordic Countries ^c [57]	Pregnant and lactating women	2001	2.0 E/0	1.0 E/0	>1.0 E%	-	-	200.0 mg/d	450.0 mg/u
Norule Countries [57]	6-23 mo	2012	-	-	>1.0 E%-	-	-	200.0 mg/u	-
	> 2 y	2012	2.5 E% ^f	-	- 1.0 1.70-	> 0.5 E%	_	-	-
Poland [58]	Pregnant and lactating women	2012	4 E%	-	-	0.5 E%	250.0 mg/d	100.0-200.0 mg/d	-
i olanu [50]	1-2 y	2012	5 E%	_	_	1.0 E%	250.0 mg/u	100.0 mg/d (<2 y)	-
	2-18 y	2012	4 E%	_	-	0.5 E%	-		250.0 mg/d
	Elderly (=adults)	2012	4 E%			0.5 E%	-	-	250.0 mg/d
Spain (AEP) [1]	6 mo-2 y	2012	3.0-4.5 E%	-	-	> 0.5 E%	-	-	
Spain (AECOSAN) [59]	> 2 y	2000	5.0-4.5 E/0			2.0 g/d	_	<200.0 mg/d	
Span (ALCOSAN) [57]	Elderly (=Adults)	2014	-			2.0 g/d 2.0 g/d	-	<200.0 mg/d	-
	Enderry (nauno)	2017		-	-	2.0 g/u	_	-200.0 mg/u	_

635 Values are Adequate Intake unless otherwise noted

636 y, year. g/d, grams per day. mg/d, miligrams per day. E%, energy percentage. PUFAs, polyunsaturated fatty acids. - , no data available. LA, linoleic acid. ARA, Arachidonic acid. n-3, omega-3 PUFA. ALA, α-linolenic acid. EPA,

637 eicosapentaenoic acid. DHA, docosahexaenoic acid. ADMR, AccepSupplemental table Macronutrient Distribution Range . L-AMDR, Lower AccepSupplemental table Macronutrient Distribution Range. U-AMDR, Upper AccepSupplemental

638 table Macronutrient Distribution Range. EAR, estimated average requirement. ANR: average nutrient requirement, BW, body weight.

639

640 *Total n-6 PUFAs is not included in this Supplemental table as there are only recommendations set for the adult population and this is not a target group in our review.

641 ^bIn addition to the 250.0 mg of (EPA+DHA).

642 °Total n-6 PUFA has been set for adults: 4.0-8.0 E% (Belgium); 2.5-9.0E% (Switzerland).

643 ^dDRI unit not mentioned.

644 ^eAs 2 serving fish per week (Guidelines for healthy diet, 2006).

 $645 \qquad {}^{\rm f} {\rm Obtained \ by \ calculation \ (official \ recommendation \ is \ for \ total \ LA + ALA \ and \ for \ ALA).}$

647 Table 2. Overview of European countries meeting n-3 and n-6 fatty acid intake recommendations per population group if compared

48	with the EFSA recommendations ((% of countries with adequate intake).
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					Evaluation a	gainst recommendat	tions (% of countries wi	th adequate intake)	
	N studies	N countries	N subjects included	Total n-6 PUFAs	LA	ARA ^a	Total n-3 PUFAs	ALA	EPA+DHA
Recommendations				4.0 E% ^b	4.0 E%	-	0.5 E% ^b	0.5 E%	RI: 250 mg/d + 100-200 mg/d DHA
Pregnant women	10	11	6033	66 (2 out of 3)	75 (3 out of 4)	3 reported	100 (3 out of 3)	100 (4 out of 4)	33 (3 out of 9)
Lactating women	4	4	293	0 (0 out of 1)	0 (0 out of 3)	1 reported	100 (4 out of 4)	100 (2 out of 2)	50 (1 out of 2)
Recommendations				4 E% ^b	4 E%	-	0.5 E% ^b	0.5 E%	100 mg/d DHA ^c
Infants 6-12 mo	3	3	606	100 (2 out of 2)	100 (2 out of 2)	1 reported	100 (2 out of 2)	100 (2 out of 2)	0 (0 out of 2)
Children 1-3 y	6	6	1797	66 (2 out of 3)	75 (3 out of 4)	1 reported	100 (4 out of 4)	75 (3 out of 4)	0 (0 out of 2) ^c
Recommendations				4 E% ^b	4 E%	-	0.5 E% ^b	0.5 E%	≥250 mg/d
Children 4-9 y	11	10	10102	55 (5 out of 9)	66 (2 out of 3)	2 reported	78 (7 out of 9)	100 (3 out of 3)	0 (0 out of 3)
Adolescents 10-18 y	8	11	4988	50 (1 out of 2)	44 (4 out of 9)	8 reported	100 (3 out of 3)	70 (7 out of 10)	20 (2 out of 10)
Elderly >65 y	11	9	9091	66 (2 out of 3)	33 (2 out of 6)	4 reported	66 (2 out of 3)	50 (3 out of 6)	50 (3 out of 6)
Total of all population groups				61 (14 out of 23)	52 (16 out of 31)	20 reported	89 (25 out of 28)	77 (24 out of 31)	26 (9 out of 34)

649 y, year. mg/d, miligrams per day. E%, energy percentage. PUFAs, polyunsaturated fatty acids. LA, linoleic acid. ARA, Arachidonic acid. n-6, omega-6 PUFA. n-3, omega-3 PUFA. ALA, α-linolenic acid. EPA,

650 eicosapentaenoic acid. DHA, docosahexaenoic acid. RI, reference intake.

651

652 "No recommendation, only number of studies with intake data.

654 °For children >2 y recommendation 250 mg EPA+DHA per day.

655

^bBased on EFSA recommendations for LA and ALA.