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Dietary polyphenol intake in Europe: the European Prospective Investigation into Cancer and Nutrition (EPIC) study

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Abstract

Background/Objectives Polyphenols are plant secondary metabolites with a large variability in their chemical structure and dietary occurrence that have been associated with some protective effects against several chronic diseases. To date, limited data exist on intake of polyphenols

in populations. The current cross-sectional analysis aimed at estimating dietary intakes of all currently known individual polyphenols and total intake per class and subclass, and to identify their main food sources in the European Prospective Investigation into Cancer and Nutrition cohort. **Methods** Dietary data at baseline were collected using a standardized 24-h dietary recall software administered to

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36,037 adult subjects. Dietary data were linked with Phenol-Explorer, a database with data on 502 individual polyphenols in 452 foods and data on polyphenol losses due to cooking and food processing.

Results Mean total polyphenol intake was the highest in Aarhus—Denmark (1786 mg/day in men and 1626 mg/day in women) and the lowest in Greece (744 mg/day in men and 584 mg/day in women). When dividing the subjects into three regions, the highest intake of total polyphenols was observed in the UK health-conscious group, followed by non-Mediterranean (non-MED) and MED countries. The main polyphenol contributors were phenolic acids (52.5–56.9 %), except in men from MED countries and in the UK health-conscious group where they were flavonoids (49.1–61.7 %). Coffee, tea, and fruits were the most important food sources of total polyphenols. A total of 437 different individual polyphenols were consumed, including 94 consumed at a level >1 mg/day. The most abundant ones were the caffeoylquinic acids and the proanthocyanidin oligomers and polymers.

Conclusion This study describes the large number of dietary individual polyphenols consumed and the high variability of their intakes between European populations, particularly between MED and non-MED countries.

Keywords Polyphenols · Dietary intake · Food sources · EPIC

Abbreviations

24-HDR	24-h dietary recall
EPIC	European Prospective Investigation into Cancer and Nutrition
MED	Mediterranean
s.e.	Standard error
SU.VI.MAX	SUplémentation en VItamines et Minéraux AntioXydants
USDA	US Department of Agriculture

Introduction

Polyphenols are plant secondary metabolites widely distributed in plant-based foods, such as tea, coffee, wine, fruit, vegetables, whole-grain cereals, and cocoa [1]. Dietary polyphenols constitute a large family of >500 different compounds with highly diverse structures from simple molecules, such as phenolic acids, to large ones, such as proanthocyanidin polymers [1]. According to their chemical structure, polyphenols are divided into four main classes: flavonoids, phenolic acids, lignans, and stilbenes [2]. In foods, flavonoids, lignans, and stilbenes are usually found as glycosides, whereas phenolic acids are most often present as esters with various polyols; some polyphenols such as flavanols are mainly present as aglycones (free

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forms) [1]. It is important to consider these structural variations as glycosylation/esterification greatly influences polyphenol absorption in the gut and bioavailability [3]. Over the last two decades, the literature on polyphenols has grown exponentially following the recognition of their antioxidant, anti-inflammatory, and anti-carcinogenic properties and more evidence for their potential beneficial effects upon health has been accumulated [4]. However, epidemiological data on the relationship between polyphenol intake and the risk of chronic diseases and mortality are still limited, especially in prospective studies, and largely concern flavonoids and lignans [2, 5–7]. To calculate polyphenol intake, most of these studies have used the US Department of Agriculture (USDA) databases on polyphenol contents in foods [8–10], which only contain data on flavonoids expressed as aglycones. The more recently published Phenol-Explorer database (www.phenol-explorer.eu) [11] is more comprehensive and gathers food composition data on all known polyphenols, either aglycones, glycosides or esters depending on how they are found in foods. Moreover, a new module of the Phenol-Explorer database contains data on the effects of cooking and food processing on polyphenol contents [12]. These retention factors permit to take into account the effects of food cooking and processing measurements when calculating polyphenol intake and to improve the reliability of such measurements [13].

Several descriptive papers on intakes of flavonoids and other polyphenols have been published using either the USDA databases [14–16], the Phenol-Explorer database [17–19] or custom databases [20]. The purpose of the present work is to describe intake of all currently known dietary polyphenols in several European countries using the more comprehensive Phenol-Explorer database and especially applying polyphenol-specific retention factors. The European Prospective Investigation into Cancer and Nutrition (EPIC) study offers a unique opportunity to estimate the intake of individual polyphenols, to identify their main food sources, and to compare these intakes between different European countries showing large variations in diets.

Materials and methods

Study population

The EPIC study is a large cohort study conducted in 10 European countries (Denmark, France, Germany, Greece, Italy, Norway, Spain, Sweden, the Netherlands, and the UK) and aims at investigating the role of diet and environmental factors on the etiology of cancer and other chronic diseases [21, 22]. Over half million participants were recruited mostly from the general population residing

within defined geographical areas, with some exceptions: women of a health insurance company for teachers and school workers (France), women attending breast cancer screening (Utrecht—the Netherlands and Florence—Italy), mainly blood donors (most centers in Italy and Spain) and a cohort consisting predominantly of vegetarians (the “health-conscious” group in Oxford, UK) [22]. The initial 23 EPIC administrative centers were redefined into 27 geographical regions relevant to the analysis of dietary consumption patterns [23].

Data used in the present work were derived from the EPIC calibration study, in which a 24-h dietary recall (24-HDR) was administered. The cohort comprises an 8 % ($n = 36,994$) random sample stratified by age, sex, and center, and weighted for expected cancer cases in each stratum of the whole EPIC cohort [23]. After exclusion of 941 subjects who were aged younger than 35 years or older than 74 years because of low participation in these age categories, and 16 individuals due to incomplete dietary information, 36,037 participants were included in the present analysis. Approval for the study was obtained from ethical review boards of the International Agency for Research on Cancer (IARC) and from all local participating institutions. All participants provided written informed consent.

Dietary and lifestyle information

Dietary assessment was performed with a single 24-HDR using specialized software (EPIC-Soft) [24], and administered in a face-to-face interview, except in Norway, where it was obtained by telephone interview [25]. Data on age, as well as on body weight and height, were self-reported by the participants during the 24-HDR interview. Data on other lifestyle factors, including educational level, physical activity, and smoking history, were collected at baseline through standardized questionnaires [23, 26]. The mean time interval between completion of the baseline questionnaire measures and the 24-HDR interview varied by country, and ranged from the same day to 3 years later [23].

Food composition database on polyphenols

The Phenol-Explorer database provides data on 502 polyphenol compounds in 452 plant-based foods collected from 638 scientific peer-reviewed articles [11]. All animal foods that contain none or only traces of plant polyphenols were excluded.

Phenol-Explorer contains data on four main polyphenol classes: flavonoids, phenolic acids, stilbenes, and lignans, as well as on a number of miscellaneous minor polyphenols that include tyrosols, alkylphenols (mainly alkylresorcinols), and alkylmethoxyphenols among others, described in the Phenol-Explorer database and in this manuscript as “other

polyphenols” [11]. Total polyphenol content was calculated as the sum of the contents of individual compounds expressed in mg/100 g food fresh weight. The data used in the present study were mainly acquired by chromatography without previous hydrolysis of the food extracts except for a few polyphenols bound to the food matrix. Lignans in all foods, ellagic acid in walnuts, and hydroxycinnamic acids in cereals, legumes, and olives were quantified by chromatography after previous basic or acid hydrolysis. Proanthocyanidin dimers were measured as individual compounds by chromatography without prior hydrolysis, whereas other proanthocyanidin oligomers (trimers, 4–6 and 7–10 oligomers) and proanthocyanidin polymers were measured as mixtures by normal-phase HPLC. Overall data on 463 individual polyphenols were used.

Some missing polyphenol content values for orange fruit and breakfast cereals were extrapolated from orange juice and wheat flour, respectively. The polyphenol content of different types of coffee, “American” or filtered diluted coffee and espresso, reported in the 24-HDR, was estimated by multiplying the polyphenol contents of “normal” filtered coffee from the Phenol-Explorer database by 0.4 and 2, respectively [27, 28]. When the type of herbal tea was unspecified in the 24-HDRs, the polyphenolic composition of a mix of herbal tea was applied on a country-by-country basis. The effect of food cooking and processing was accounted for by applying polyphenol-specific retention factors from Phenol-Explorer [12] to 24-HDR foods wherever relevant. Phenol-Explorer contains data on 1253 aggregated retention factors, including data on 161 polyphenols and 35 processes, such as domestic cooking, storage, and industrial processing.

Statistical analyses

Dietary polyphenol intakes were estimated using general linear models and presented as means and standard errors (s.e.) stratified by sex and center, adjusted for age, and weighted by season and weekday of the 24-HDR to control for different distributions of participants across seasons and 24-HDR days. The contribution of each polyphenol class, subclass, and family to the total polyphenol intake was calculated as a percentage according to three *ad hoc* European regions: (1) Mediterranean (MED) countries: all centers in Greece, Spain, Italy, and the south of France; (2) non-MED countries: all centers in the north-east and north-west of France, Germany, the Netherlands, UK general population, Denmark, Sweden, and Norway; (3) UK health-conscious group. The contribution of each food group to the intake of total and polyphenol classes by European region was also calculated as a percentage. Differences in polyphenol intakes stratified by European region were also compared using general linear models

according to the categories of sex, age (35–44, 45–54, 55–64, and 64–74), BMI (<25, 25–30, and >30 kg/m²), educational level (no formal education, primary school, technical/professional school, secondary school, university, or not specified), smoking status (never, former, current smoker, and not specified), and Cambridge physical activity index (inactive, moderately inactive, moderately active, and not specified). All these models were adjusted for sex, age (years), center, BMI (kg/m²), and energy intake, and weighted by season and day of 24-HDR. *P* values <0.05 (two-tailed) were considered significant. All analyses were conducted using the SPSS Statistics software (version 19.0; SPSS Inc.).

Results

A south-to-north gradient in the daily mean intake of total polyphenols was observed among EPIC centers, in both men and women (Fig. 1). The highest total polyphenol intake in both sexes was in Aarhus—Denmark (in men 1786 mg/day and in women 1626 mg/day), whereas the lowest intake was in Greece in both men (744 mg/day) and women (584 mg/day). Table 1 shows the mean intakes of total polyphenols and polyphenol classes adjusted for sex, age, BMI (except where stratified for these variables), center, and energy intake, and weighted by season and weekday of the 24-HDR. The intake of phenolic acids, stilbenes, and other polyphenols was higher in men than in women, whereas for flavonoids and lignans the opposite was observed. No differences were observed for total polyphenol intake between men and women. The intake of flavonoids, stilbenes, lignans, and other polyphenols increased with age, whereas the opposite was seen for total polyphenols and phenolic acids for which a maximum intake was observed at the age of 45–54 years. When comparing the intake by European region, the UK health-conscious group (1521 mg/day) had the highest intake of total polyphenols due to the higher intake of flavonoids. Non-MED countries (1284 mg/day) showed a higher total polyphenol intake when compared to MED countries (1011 mg/day), with phenolic acids as the main contributors. MED countries presented the highest intake of stilbenes compared to the other regions. Individuals with BMI < 25, and/or a university degree, and/or current smokers showed the highest intakes of total polyphenols. No differences in total polyphenol intake were observed between groups showing different physical activity levels (Table 1).

Phenolic acids were the main contributors to the total polyphenol intake in non-MED countries (57 and 53 % in men and women, respectively) and in women from MED countries (54 %) (Table 2). Flavonoids were the second most abundant contributors in these groups (38–44 %). In contrast, in the UK health-conscious group and in men

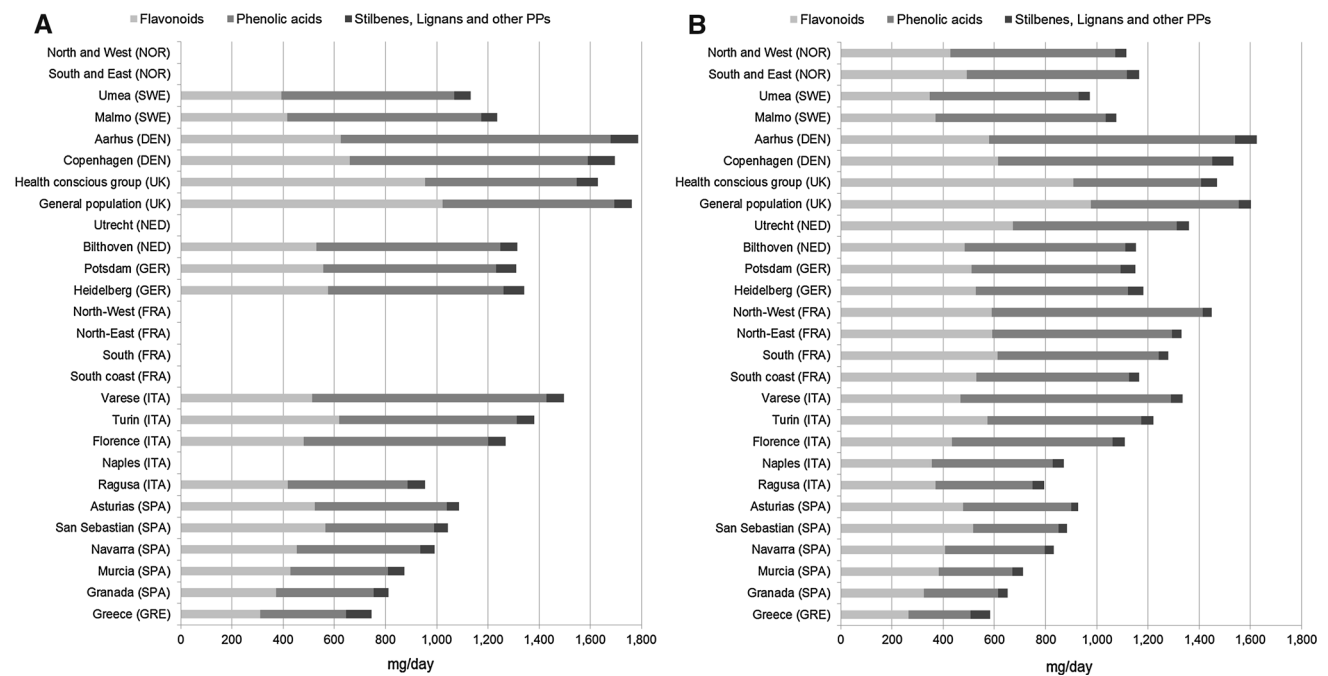


Fig. 1 Adjusted daily polyphenol mean intake (mg/day), stratified by sex (**a** men; **b** women) and center ordered from north to south, adjusted for age and weighted by season and weekday of dietary

recalls. *GRE* Greece, *SPA* Spain, *ITA* Italy, *FRA* France, *GER* Germany, *NED* the Netherlands, *DEN* Denmark, *SWE* Sweden, *NOR* Norway, *UK* United Kingdom

from MED countries, flavonoids were the largest contributor to total polyphenol intake ranging from 49 to 62 %, followed by phenolic acids (34–44 %). Stilbenes and lignans accounted for <0.7 % of total polyphenol intake with levels of intake not exceeding 3.1 and 9.1 mg/day, respectively, in any of the EPIC regions.

Regarding polyphenol subclasses, the two most important contributors to total polyphenol intake were hydroxycinnamic acids (ranging from 27 % in women from the UK health-conscious group to 53 % in men from non-MED countries) and flavanols (ranging from 28 % in men from non-MED countries to 48 % in women from the UK health-conscious group). They were followed by various subclasses: anthocyanidins, flavanones, flavonols, hydroxybenzoic acids, alkylphenols, and tyrosols which each accounted for 2–6 % of total polyphenol intake. Other subclasses were less remarkable, each one contributing less than 1 % to total polyphenol intake in most European regions.

The main food sources of polyphenols according to European regions are presented in Table 3. In MED countries, coffee (36 %), fruits (25 %), and wine (10 %) were the major dietary sources of polyphenols. In non-MED countries, coffee (41 %), tea (17 %), and fruits (13 %) were the most important food sources. In the UK health-conscious group, tea ranked first (41 %), followed by coffee (21 %) and fruits (9 %).

The main food sources of flavonoids were fruits in MED countries (45 %), and tea in both non-MED countries (34 %) and the UK health-conscious group (64 %). Coffee

was the most important dietary source of phenolic acids in all European regions (58–75 %). The major food sources of lignans were region-related: vegetable oils (26 %) in MED countries, cereal products (24 %) in non-MED countries, and seeds (49 %) in the UK health-conscious group. Wine was clearly the most prominent food source of stilbenes accounting for >92 % of intake in all regions.

A total of 437 individual polyphenols were consumed by the EPIC population (Table 2). Ninety-four polyphenols were consumed in a mean quantity of at least 1 mg/day, 118 polyphenols between 0.1 and 1 mg/day, 112 polyphenols between 0.01 and 0.1 mg/day, and 113 polyphenols between >0 and 0.01 mg/day (Supplemental Table 1). Finally, 26 additional polyphenols documented in the food composition table were either not consumed or their intake could not be calculated since the consumption of their food sources was not reported in dietary recalls (Supplemental Table 2). Mean intakes of the top 25 individual polyphenols are shown in Table 4 together with their three main food sources. Mean intakes of all individual polyphenols are presented in Supplemental Table 1.

In men from MED countries and participants from the UK health-conscious group, the top three most consumed polyphenols were 5-caffeoylquinic acid, proanthocyanidin polymers, and 4-caffeoylquinic acid, whereas for the other groups, they were 5-caffeoylquinic acid, 4-caffeoylquinic acid, and 3-caffeoylquinic acid.

A large variability in both the mean intake and the ranking of the top 10 most consumed polyphenols was

Table 1 Adjusted^a mean daily intakes of total and polyphenol classes by socio-demographic and lifestyle factors in the EPIC study

Stratification variable	N	Polyphenols (mg/day)			Flavonoids (mg/day)			Phenolic acids (mg/day)		
		Mean	s.e.	<i>P</i> value	Mean	s.e.	<i>P</i> value	Mean	s.e.	<i>P</i> value
Sex				0.16			<0.001			<0.001
Men	13,028	1177	8		492	5		625	6	
Women	23,009	1192	6		546	4		593	5	
Age				<0.001			<0.001			<0.001
35–44	3335	1167	14		490	8		623	11	
45–54	12,595	1200	7		504	5		641	6	
55–64	14,940	1187	7		531	4		598	6	
65–74	5167	1148	12		541	7		548	9	
European region				<0.001			<0.001			<0.001
Mediterranean countries	11,285	1011	7		449	5		503	6	
Non-Mediterranean countries	24,443	1284	5		522	3		700	4	
UK health-conscious group	309	1521	44		909	27		538	34	
Body mass index (kg/m ²)				0.002			<0.001			<0.001
<25	16,854	1194	7		542	4		594	6	
25–30	13,766	1186	7		510	4		620	6	
>30	5417	1150	11		471	7		625	8	
Educational level				<0.001			<0.001			0.029
None	1709	1124	21		488	13		578	17	
Primary completed	10,469	1134	8		457	5		621	7	
Technical/professional	8038	1168	10		511	6		602	8	
Secondary school	7152	1205	10		543	6		607	8	
University degree	8155	1275	9		598	6		618	7	
Smoking status				<0.001			<0.001			<0.001
Never smoker	17,483	1137	7		532	4		549	5	
Former smoker	10,288	1200	8		537	5		605	6	
Current smoker	7726	1265	9		470	6		739	7	
Physical activity				0.32			<0.001			0.14
Inactive	7463	1173	9		506	6		611	7	
Moderately inactive	11,969	1187	8		522	5		608	6	
Moderately active	8400	1187	9		534	6		596	7	
Active	6380	1198	10		549	6		591	8	
Stratification variable	N	Lignans (mg/day)			Stilbenes (mg/day)			Other polyphenols (mg/day)		
		Mean	s.e.	<i>P</i> value	Mean	s.e.	<i>P</i> value	Mean	s.e.	<i>P</i> value
Sex				<0.001			<0.001			<0.001
Men	13,028	2.5	0.2		3.0	0.1		55	0.5	
Women	23,009	3.6	0.1		2.4	0.0		46	0.3	
Age				0.002			0.009			<0.001
35–44	3335	2.8	0.3		2.4	0.1		48	0.8	
45–54	12,595	2.7	0.2		2.7	0.0		50	0.4	
55–64	14,940	3.3	0.1		2.8	0.0		52	0.4	
65–74	5167	3.5	0.2		2.8	0.1		52	0.7	
European region				<0.001			<0.001			<0.001
Mediterranean countries	11,285	3.6	0.1		3.1	0.0		53	0.4	
Non-Mediterranean countries	24,443	2.3	0.1		2.1	0.0		57	0.3	
UK health-conscious group	309	9.1	0.9		1.4	0.3		63	2.6	

Table 1 continued

Stratification variable	N	Lignans (mg/day)			Stilbenes (mg/day)			Other polyphenols (mg/day)		
		Mean	s.e.	<i>P</i> value	Mean	s.e.	<i>P</i> value	Mean	s.e.	<i>P</i> value
Body mass index (kg/m ²)				0.043			0.036			<0.001
<25	16,854	3.3	0.1		2.7	0.0		52	0.4	
25–30	13,766	2.8	0.1		2.7	0.0		50	0.4	
>30	5417	3.1	0.2		2.5	0.1		49	0.6	
Educational level				0.036			<0.001			0.011
None	1709	3.3	0.4		2.7	0.1		52	1.2	
Primary completed	10,469	2.7	0.2		2.3	0.1		50	0.5	
Technical/professional	8038	2.9	0.2		2.7	0.1		50	0.5	
Secondary school	7152	3.2	0.2		2.8	0.1		50	0.5	
University degree	8155	3.4	0.2		3.2	0.1		52	0.5	
Smoking status				0.08			<0.001			<0.001
Never smoker	17,483	3.2	0.1		2.4	0.0		50	0.4	
Former smoker	10,288	3.0	0.2		3.0	0.1		52	0.4	
Current smoker	7726	2.8	0.2		3.0	0.1		50	0.5	
Physical activity				0.05			0.002			0.001
Inactive	7463	3.5	0.2		2.6	0.1		49	0.5	
Moderately inactive	11,969	3.4	0.2		2.8	0.0		51	0.4	
Moderately active	8400	2.8	0.2		2.9	0.1		51	0.5	
Active	6380	3.0	0.2		2.7	0.1		53	0.6	

^a Means and standard error (s.e.) were computed using general linear models adjusted for sex, age, center, energy intake, and body mass index (as appropriate) and weighted by season and weekday of recall

P value is for differences in means

observed between European regions (Fig. 2). For example, 5-caffeoylquinic acid intake in men ranged from 149 mg/day from MED countries to 231 mg/day from non-MED countries (Table 4). However, the main food sources for each individual polyphenol by European region were very similar and therefore are listed for the EPIC study as a whole (Table 4 and Supplemental Table 1). Coffee was the major food source (>78 %) of caffeoyl- and feruloylquinic acids, whereas tea was the main food source (>99.5 %) of (+)-gallic acid, (-)-epigallocatechin, (-)-epigallocatechin 3-O-gallate, and 5-O-galloylquinic acid. Apples, pears, and chocolate products were the main food sources of proanthocyanidin oligomers and polymers. Hesperidin was provided exclusively from citrus products (citrus fruit and juices). Bread and cereal products were the main sources of ferulic acid and alkylphenols (5-heneicosylresorcinol and 5-nonadecylresorcinol).

Discussion

In the current study, dietary intake of total polyphenols, polyphenol classes, and individual polyphenols was estimated across 10 European countries, using a standardized 24-HDR and the Phenol-Explorer database, allowing easy

comparisons between groups of individuals and countries. Main food sources were also identified, and the influence of some socio-demographic characteristics was assessed. Most previous descriptive studies on polyphenols used the USDA food composition data to document intake of approximately 50 polyphenol aglycones [14–16, 20, 29–31]. In the current study, intakes of 437 polyphenols, either glycosides, esters or aglycones are described. This study follows one of our earlier studies in which intakes of 337 polyphenols in the French SUPplémentation en Vitamines et Minéraux AntioXydants (SU.VI.MAX) cohort were reported [17].

In addition to the large number of polyphenols and diversity of countries considered, the present study is also the first one to take into account the effects of cooking and processing on polyphenol contents in foods by applying retention factors recently collected in the Phenol-Explorer database [12]. In our previous studies in EPIC, a common retention factor for each cooking method (boiled, fried, and microwaved) was eventually applied irrespective of the polyphenol compound and food considered [15, 31]. This simple approach could only lead to a crude estimation of the effects of cooking on polyphenol intake. Even after applying retention factors, mean polyphenol intakes in our French EPIC centers (1166–1449 mg/day) were slightly

Table 2 Number of individual polyphenols and contribution of classes and subclasses of total polyphenols, and the top three most consumed polyphenols for each polyphenol class and subclass by sex and European region in the EPIC study

Polyphenol classes and subclasses	MED countries (%)		non-MED countries (%)		UK Health-conscious group (%)		Number of individual polyphenols	Top three most consumed individual polyphenols
	Men	Women	Men	Women	Men	Women		
Total polyphenols	100	100	100	100	100	100	437	5-Caffeoylquinic acid, 4-caffeoylquinic acid, 3-caffeoylquinic acid
Flavonoids	49.1	40.9	37.5	43.5	58.2	61.7	234	Proanthocyanidin polymers, proanthocyanidin 4-6 oligomers, proanthocyanidin 7-10 oligomers
Anthocyanins	5.4	4.3	3.2	3.7	2.2	2.9	65	Cyanidin 3-O-rutinoside, malvidin 3-O-glucoside, delphinidin 3-O-rutinoside
Chalcones	0.0	0.0	0.0	0.0	0.0	0.0	2	Xanthohumol, butein
Dihydrochalcones	0.3	0.2	0.2	0.2	0.2	0.2	4	Phloridzin, phloretin 2'-O-xylosylglucoside, 3-hydroxyphloretin 2'-O-glucoside
Dihydroflavonols	0.8	0.2	0.2	0.2	0.1	0.1	2	Dihydromyricetin 3-O-rhamnoside, dihydroquercetin 3-O-rhamnoside
Flavanols	34.1	28.7	27.7	32.0	43.5	47.8	25	Proanthocyanidin polymers, proanthocyanidin 4-6 oligomers, proanthocyanidin 7-10 oligomers
Flavan-3-ol monomers	29.1	21.5	16.6	18.1	21.7	22.3	10	(+)-Gallocatechin, (–)-epicatechin, (–)-epigallocatechin 3-O-gallate
Proanthocyanidins	9.1	5.7	4.7	5.3	5.6	6.4	11	Proanthocyanidin polymers, proanthocyanidin 4-6 oligomers, proanthocyanidin 7-10 oligomers
Theaflavins	24.8	19.2	14.4	15.6	18.3	18.8	4	Theaflavin 3'-O-gallate, theaflavin 3,3'-O-digallate, theaflavin
Flavanones	3.3	3.0	2.6	3.3	4.0	3.0	17	Hesperidin, narirutin, didymin
Flavones	1.5	1.5	0.7	0.9	1.2	1.0	38	Apigenin 6,8-di-C-glucoside, apigenin 6,8-C-galactoside-C-arabinoside, apigenin 6,8-C-arabinoside-C-glucoside
Flavonols	3.7	2.9	2.8	3.2	4.8	5.1	68	Quercetin 3-O-rutinoside, quercetin 3-O-glucoside, quercetin 3,4'-O-diglucoside
Isoflavonoids	0.0	0.1	0.1	0.1	2.2	1.6	13	Genistin, 6'-O-malonylgenistin, daidzin
Phenolic acids	43.6	54.4	56.8	52.5	36.4	34.4	100	5-Caffeoylquinic acid, 4-caffeoylquinic acid, 3-caffeoylquinic acid
Hydroxybenzoic acids	3.3	3.6	3.5	4.1	6.0	7.4	27	5-O-Galloylquinic acid, gallic acid, ellagic acid
Hydroxycinnamic acids	40.2	50.7	53.3	48.4	30.4	27.0	68	5-Caffeoylquinic acid, 4-caffeoylquinic acid, 3-caffeoylquinic acid
Hydroxyphenylacetic acids	0.1	0.0	0.0	0.0	0.0	0.0	4	4-Hydroxyphenylacetic acid, homovanillic acid, 3,4-dihydroxyphenylacetic acid
Hydroxyphenylpropanoic acids	0.0	0.0	0.0	0.0	0.0	0.0	1	Dihydro-p-coumaric acid
Stilbenes	0.5	0.1	0.2	0.1	0.1	0.1	7	Piceatannol 3-O-glucoside, resveratrol 3-O-glucoside, d-viniferin
Lignans	0.3	0.4	0.1	0.2	0.7	0.5	27	Secoisolariciresinol, lariciresinol, sesamol
Other polyphenols class	6.4	4.2	5.3	3.6	4.7	3.4	69	5-Heneicosylresorcinol, 5-nonadecylresorcinol, 5-heptadecylresorcinol
Alkylmethoxyphenols	0.2	0.2	0.3	0.2	0.2	0.1	3	4-Ethylguaiaicol, 4-vinylguaiaicol, 4-vinylsyringol

Table 2 continued

Polyphenol classes and subclasses	MED countries (%)		non-MED countries (%)		UK Health-conscious group (%)		Number of individual polyphenols	Top three most consumed individual polyphenols
	Men	Women	Men	Women	Men	Women		
Alkylphenols	2.1	1.6	4.0	2.5	3.7	2.6	14	5-Heneicosylresorcinol, 5-nonadecylresorcinol, 5-heptadecylresorcinol
Tyrosols	3.6	1.9	0.4	0.4	0.5	0.4	14	Tyrosol, 3,4-DHPEA-EDA, oleuropein-aglycone
Other polyphenols subclass	0.6	0.5	0.7	0.5	0.3	0.2	38	Pyrogallol, catechol, eugenol
Cucurminoids	0.0	0.0	0.0	0.0	0.0	0.0	3	Curcumin, demethoxycurcumin, bisdemethoxycurcumin
Furanocoumarins	0.0	0.0	0.0	0.0	0.0	0.0	4	Bergapten, isopimpinellin, xanthotoxin
Hydroxybenzaldehydes	0.1	0.0	0.0	0.0	0.0	0.0	5	Syringaldehyde, protocatechuic aldehyde, p-anisaldehyde
Hydroxybenzoketones	0.0	0.0	0.0	0.0	0.0	0.0	2	2,3-Dihydroxy-1-guaiacylpropanone, 3-methoxyacetophenone
Hydroxycinnamaldehydes	0.0	0.0	0.0	0.0	0.0	0.0	2	Ferulaldehyde, sinapaldehyde
Hydroxycoumarins	0.0	0.0	0.0	0.0	0.0	0.0	6	4-Hydroxycoumarin, esculin, esculetin
Hydroxyphenylpropenes	0.1	0.0	0.3	0.2	0.0	0.0	5	Eugenol, acetyl eugenol, anethole
Methoxyphenols	0.0	0.0	0.0	0.0	0.0	0.0	1	Guaiacol
Naphthoquinones	0.0	0.0	0.0	0.0	0.0	0.0	1	Juglone
Phenolic terpenes	0.0	0.0	0.0	0.0	0.0	0.0	3	Carnosic acid, rosmanol, carnosol
Other polyphenols family	0.4	0.4	0.3	0.3	0.2	0.1	6	Pyrogallol, catechol, phlorin

Values are percentages derived from general lineal models adjusted for sex, age, and center and weighted by season and weekday of dietary recalls

higher than in the French SU.VI.MAX cohort (1193 mg/day) [17], indicating that the use of retention factors does not seem to have high impact on the estimated total polyphenol intake. This can be explained by the fact that these retention factors mainly concern vegetables and legumes [12] which were not major contributors to polyphenol intake in the present or previous studies [14, 15, 17, 20]. It is still important to highlight that polyphenol losses due to cooking and food processing can have a great impact on the intake of some individual polyphenols mainly present in vegetables, such as quercetin 3,4'-O-diglucoside that is only present in onions [1].

Using a single 24-HDR for each of the 36,037 subjects across 10 European countries in EPIC, 437 different individual polyphenols have been documented. Intakes of 347, 337, and 290 polyphenols have been reported utilizing Food Frequency Questionnaires and the Phenol-Explorer database in the Polish arm of the HAPIEE (Health, Alcohol and Psychosocial factors In Eastern Europe) study [19], in the SU.VI.MAX study [17], and in the Spanish PREDIMED (PREvención con DIeta MEDiterránea) study [18], respectively. However, a similar number of individual polyphenols (around 100) were consumed at a level higher than 1 mg/day on average in these three studies [17, 18].

Interestingly, although the number of most abundant polyphenols (>1 mg/day) is similar by region/cohort, there are considerable differences in the individual polyphenols included in that group. For example, oleuropein (tyrosol subclass) was highly consumed in men from MED countries (3.3 mg/day, rank 49), but not in men from non-MED countries (0.3 mg/day, rank 147). This is due to the fact that some polyphenols are exclusively present in a few characteristic foods (e.g., oleuropein is only found in olives and olive oil) and that their consumption largely varies according to regions. Another example is delphinidin 3-O-rutinoside (anthocyanidin subclass) only found in blackcurrants; its intake varied from 7.0 mg/day (rank 31) in men from non-MED countries to 0.04 mg/day (rank 245) in men from MED countries. This illustrates the need for further detailed analyses of polyphenols in foods not commonly consumed in Western countries and poorly described in the Phenol-Explorer or USDA databases, to document their intake across different geographical areas of the world.

Our results show a wide range of total polyphenol intake and a south-to-north geographical gradient. Mean intake was as much as three times higher in men from Aarhus—Denmark (1786 mg/day) than in women from Greece (584 mg/day). When stratified by regions, total polyphenol

Table 3 Contributions of food groups and some specific foods to the intake of total polyphenols and polyphenol classes by European region in the EPIC study

Food items	Polyphenols (%)			Flavonoids (%)			Phenolic acids (%)		
	MED countries	non-MED countries	UK Health	MED countries	non-MED countries	UK Health	MED countries	non-MED countries	UK Health
Potatoes and other tubers	1.0	1.3	0.9	0.0	0.0	0.0	1.9	2.4	2.6
Vegetables	4.6	1.7	1.5	4.7	1.5	0.6	5.0	1.9	2.5
Leafy vegetables	1.2	0.6	0.1	0.8	0.4	0.1	1.7	0.8	0.1
Fruiting vegetables	1.8	0.3	0.2	1.1	0.2	0.0	2.6	0.4	0.3
Root vegetables	0.1	0.1	0.1	0.0	0.0	0.0	0.2	0.2	0.4
Cabbages	0.1	0.1	0.2	0.0	0.1	0.1	0.1	0.1	0.4
Grain and pod vegetables	0.4	0.1	0.2	0.7	0.0	0.0	0.1	0.2	0.6
Onion, garlic	0.8	0.3	0.3	1.7	0.7	0.4	0.0	0.0	0.0
Stalk vegetables, sprouts	0.1	0.0	0.0	0.1	0.1	0.0	0.0	0.0	0.0
Mixed salad, mixed vegetables	0.3	0.1	0.3	0.2	0.1	0.0	0.3	0.2	0.7
Legumes	3.4	0.8	4.0	6.9	1.9	4.4	0.8	0.1	0.5
Fruits	25.2	12.5	8.6	44.9	24.6	10.0	9.0	4.0	4.2
Citrus fruit	2.3	0.8	0.6	5.0	1.8	0.7	0.0	0.0	0.0
Apple and pear	11.7	6.4	3.9	22.2	13.0	4.9	3.8	1.8	1.7
Grape	2.1	0.8	0.9	4.6	1.8	1.1	0.1	0.0	0.1
Stone fruits	5.5	2.3	1.1	8.9	3.5	1.4	3.3	1.5	1.1
Berries	1.2	1.6	1.4	2.5	3.3	0.9	0.2	0.4	0.6
Other fruits	0.5	0.5	0.6	0.9	1.1	0.9	0.1	0.1	0.3
Olives	1.9	0.2	0.2	0.7	0.1	0.0	1.6	0.2	0.2
Nuts	1.5	0.3	0.6	0.6	0.3	0.1	2.5	0.3	0.7
Seeds	0.1	0.1	0.3	0.0	0.0	0.0	0.2	0.0	0.4
Dairy products	0.3	0.6	0.7	0.6	1.3	1.8	0.2	0.1	0.1
Cereal products	3.7	7.0	4.4	1.2	0.6	0.5	2.4	7.0	4.0
Flour, flakes, starches, semolina and pastry	0.2	0.1	0.2	0.2	0.1	0.1	0.1	0.1	0.2
Pasta, rice, other grains	1.4	0.4	0.4	0.5	0.2	0.0	1.5	0.4	0.8
Bread, crispbread, rusks and crackers	1.8	6.0	1.4	0.4	0.2	0.0	0.6	6.1	1.1
Breakfast cereals	0.2	0.5	2.4	0.0	0.1	0.3	0.2	0.5	2.0
Cakes, biscuits and sweets	5.0	7.0	7.7	9.9	15.7	10.3	0.9	0.7	1.4
Cakes and biscuits	1.7	1.9	2.6	2.8	3.8	0.9	0.6	0.4	0.9
Chocolate, candy bars, paste, confetti	2.6	4.2	4.6	5.9	10.1	0.0	0.1	0.1	0.2
Confectionary non-chocolate, candied fruits	0.1	0.1	0.1	0.1	0.1	0.8	0.0	0.0	0.2
Ice cream, water ice	0.5	0.6	0.4	1.1	1.4	2.7	0.0	0.0	0.0
Sugar, honey, jam and syrup	0.1	0.2	0.0	0.0	0.3	5.8	0.1	0.1	0.1
Meat, fish and eggs	0.0	0.1	0.0	0.1	0.1	0.0	0.0	0.0	0.0
Fats and oils	1.6	0.2	0.3	0.1	0.0	0.0	0.5	0.1	0.3
Vegetable oils	1.6	0.1	0.2	0.1	0.0	0.0	0.5	0.1	0.2
Margarine and animal fats	0.0	0.1	0.1	0.0	0.0	0.0	0.0	0.0	0.0
Non-alcoholic beverages	42.3	62.6	65.7	12.1	43.9	67.8	72.8	81.1	80.9
Fruit and vegetable juices	1.4	3.3	2.7	3.0	7.4	3.5	0.1	0.3	0.4
Carbonated/soft/isotonic drinks	0.1	0.7	0.4	0.2	1.7	0.4	0.0	0.0	0.0

Table 3 continued

Food items	Polyphenols (%)			Flavonoids (%)			Phenolic acids (%)		
	MED countries	non-MED countries	UK Health	MED countries	non-MED countries	UK Health	MED countries	non-MED countries	UK Health
Coffee	35.9	40.9	20.9	0.0	0.1	0.0	70.6	74.6	57.5
Tea	4.6	17.4	40.8	8.6	34.2	63.8	1.7	5.9	21.2
Herbal tea	0.3	0.3	0.9	0.3	0.5	0.1	0.4	0.2	1.7
Alcoholic beverages	10.4	5.1	3.7	18.2	9.0	3.9	3.3	1.9	2.0
Wine	10.2	4.6	3.5	18.1	8.6	3.8	3.2	1.4	1.7
Beer, cider	0.1	0.4	0.2	0.1	0.2	0.1	0.1	0.5	0.4
Spirits and cocktails	0.0	0.1	0.0	0.1	0.2	0.0	0.0	0.0	0.0
Condiments, spices, and sauces	0.6	0.3	0.5	0.4	0.4	0.4	0.3	0.1	0.2
Other products	0.3	0.5	1.0	0.4	0.7	0.2	0.1	0.2	0.3
Food items	Lignans (%)			Stilbenes (%)			Other polyphenols (%)		
	MED countries	non-MED countries	UK Health	MED countries	non-MED countries	UK Health	MED countries	non-MED countries	UK Health
Potatoes and other tubers	0.5	1.0	0.5	0.0	0.0	0.0	0.0	0.0	0.0
Vegetables	11.2	9.8	9.0	0.0	0.0	0.0	0.1	0.0	0.2
Leafy vegetables	0.3	0.2	0.1	0.0	0.0	0.0	0.0	0.0	0.0
Fruiting vegetables	7.4	3.2	1.9	0.0	0.0	0.0	0.0	0.0	0.0
Root vegetables	0.4	0.9	0.7	0.0	0.0	0.0	0.0	0.0	0.0
Cabbages	2.0	4.4	5.3	0.0	0.0	0.0	0.0	0.0	0.0
Grain and pod vegetables	0.3	0.3	0.6	0.0	0.0	0.0	0.0	0.0	0.0
Onion, garlic	0.1	0.0	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Stalk vegetables, sprouts	0.5	0.5	0.2	0.0	0.0	0.0	0.1	0.0	0.1
Mixed salad, mixed vegetables	0.2	0.4	0.3	0.0	0.0	0.0	0.0	0.0	0.0
Legumes	2.8	0.6	1.2	0.2	0.0	0.1	0.0	0.0	0.0
Fruits	10.7	7.1	4.9	1.9	2.9	3.0	17.8	3.1	3.4
Citrus fruit	4.2	2.6	1.4	0.0	0.0	0.0	2.0	0.9	0.8
Apple and pear	1.9	1.2	0.8	0.0	0.0	0.0	0.6	0.2	0.1
Grape	0.3	0.2	0.2	1.3	1.0	1.3	0.0	0.0	0.0
Stone fruits	2.4	1.1	0.7	0.0	0.0	0.0	0.0	0.0	0.0
Berries	0.6	1.1	0.7	0.6	1.6	1.3	0.0	0.0	0.0
Other fruits	1.2	0.9	1.1	0.1	0.3	0.4	0.0	0.0	0.0
Olives	0.1	0.0	0.0	0.0	0.0	0.0	15.2	2.0	2.5
Nuts	0.2	0.2	0.1	0.0	0.1	0.1	0.0	0.0	0.0
Seeds	4.2	15.6	48.8	0.0	0.0	0.0	0.0	0.0	0.0
Dairy products	0.0	0.1	0.1	0.0	0.1	0.3	0.0	0.0	0.0
Cereal products	13.7	24.3	6.6	0.0	0.0	0.0	34.6	68.5	73.9
Flour, flakes, starches, semolina and pastry	0.4	0.3	0.2	0.0	0.0	0.0	1.0	0.7	1.4
Pasta, rice, other grains	0.7	0.6	0.1	0.0	0.0	0.0	8.0	3.4	2.8
Bread, crispbread, rusks and crackers	12.5	21.9	4.8	0.0	0.0	0.0	23.5	60.4	26.7
Breakfast cereals	0.1	1.4	1.4	0.0	0.0	0.0	2.1	4.0	43.1
Cakes, biscuits and sweets	19.8	2.1	2.6	0.1	0.6	0.5	2.8	2.6	2.5
Cakes and biscuits	1.3	1.8	1.2	0.1	0.3	0.1	2.8	2.5	2.3

Table 3 continued

Food items	Lignans (%)			Stilbenes (%)			Other polyphenols (%)		
	MED countries	non-MED countries	UK Health	MED countries	non-MED countries	UK Health	MED countries	non-MED countries	UK Health
Chocolate, candy bars, paste, confetti	0.0	0.1	0.1	0.1	0.2	0.3	0.0	0.0	0.0
Confectionary non-chocolate, candied fruits	18.5	0.0	1.3	0.0	0.0	0.0	0.0	0.0	0.2
Ice cream, water ice	0.0	0.1	0.0	0.0	0.1	0.1	0.0	0.0	0.0
Sugar, honey, jam and syrup	0.0	0.1	0.0	0.0	0.0	0.0	0.0	0.0	0.0
Meat, fish and eggs	0.0	0.1	0.1	0.0	0.0	0.0	0.1	1.0	0.3
Fats and oils	26.1	11.7	10.0	0.0	0.0	0.0	23.2	2.1	4.0
Vegetable oils	26.0	11.5	9.7	0.0	0.0	0.0	22.8	1.1	2.6
Margarine and animal fats	0.1	0.1	0.3	0.0	0.0	0.0	0.5	1.0	1.5
Non-alcoholic beverages	0.9	5.3	4.1	0.1	1.4	2.5	9.0	12.9	7.4
Fruit and vegetable juices	0.1	0.2	0.1	0.0	0.2	2.5	0.9	0.5	0.3
Carbonated/soft/isotonic drinks	0.0	0.2	0.0	0.0	1.2	0.1	0.0	0.0	0.0
Coffee	0.5	3.0	1.2	0.0	0.0	0.0	8.1	11.7	7.1
Tea	0.3	1.6	2.6	0.0	0.0	0.0	0.0	0.0	0.0
Herbal tea	0.1	0.3	0.2	0.0	0.0	0.0	0.0	0.7	0.0
Alcoholic beverages	3.9	4.3	2.3	97.2	94.4	93.2	7.6	6.2	5.1
Wine	3.7	2.9	1.7	96.9	92.3	92.3	7.3	4.6	4.3
Beer, cider	0.3	1.3	0.5	0.0	0.1	0.6	0.2	1.5	0.8
Spirits and cocktails	0.0	0.1	0.0	0.3	2.0	0.4	0.0	0.1	0.0
Condiments, spices, and sauces	5.5	16.5	8.7	0.4	0.2	0.2	3.9	1.5	2.1
Other products	0.4	1.3	0.9	0.0	0.2	0.1	0.8	2.0	1.0

Values are percentages derived from general linear models adjusted for sex, age, and center and weighted by season and weekday of dietary recalls

intake in the non-MED countries was higher than that in the MED countries. This small gradient in total polyphenol intake was mainly due to a 1.5-fold higher intake of phenolic acids in northern EPIC cohorts. Hydroxycinnamic acids, and more specifically the 5-, 4-, and 3-caffeoylquinic acids, are by far the highest contributors to total polyphenol in non-MED countries due to the high coffee consumption in this region, which explains almost 90 % of the phenolic acid intake as similarly observed in previous studies [17, 20, 30]. Intakes of phenolic acids in the current study are much lower than those reported in our previous study [30]. This is explained by the fact that in the present work, we took into account differences in polyphenol concentrations according to coffee types. People from non-MED countries usually drink a diluted type of filtered coffee, whereas in MED countries coffee is often consumed as espresso, a coffee fivefold more concentrated than the diluted filtered one [28].

Flavonoid intake was slightly higher in non-MED countries compared to MED countries. In the UK health-

conscious group, flavonoid intake was almost twice that of the other two regions. This is mainly due to the high intake of flavan-3-ol monomers, with tea being the main source as described previously [32]. It is worth bearing in mind that, thearubigins were not included in the present study, since only crude analytical data are available [33]. However, if these data were considered, the contribution of flavonoids to total polyphenol intake in countries with a traditional tea culture would be significantly higher [33]. Conversely, in MED countries, the relatively high intake of flavonoids is due to proanthocyanidins, mainly coming from fruits [31, 32]. Intakes of flavonoid sub-classes, that are particularly abundant in fruits and fruit products (anthocyanidins, flavanones, dihydrochalcones, and dihydroflavonols), were higher in MED countries as observed earlier [34, 35]. Isoflavonoid intake is very low in European [17, 20, 36] and other Western populations [14, 16] in comparison with Asian populations [37], but the intake in the UK health-conscious group was quite large, due to the high consumption of soya products in this mainly vegetarian cohort

Table 4 Adjusted^a mean intakes (mg/day) of top 25 most consumed individual polyphenols and their main food sources by European region in the EPIC study

Polyphenol (mg/day)	Polyphenol subclass	ALL		MED countries				non-MED countries			
				Men		Women		Men		Women	
		Mean	se	Mean	se	Mean	se	Mean	se	Mean	se
5-Caffeoylquinic acid	Hydroxycinnamic acids	195	1.1	149	2.8	178	4.1	231	1.7	201	1.2
4-Caffeoylquinic acid	Hydroxycinnamic acids	133	0.9	83	2.1	113	3.4	166	1.4	142	1.0
3-Caffeoylquinic acid	Hydroxycinnamic acids	122	0.8	83	2.0	109	3.0	147	1.3	129	0.9
Proanthocyanidin polymers (>10 mers)	Flavanols	78	0.6	103	2.6	74	1.4	75	1.4	74	0.8
Proanthocyanidin 4–6 oligomers	Flavanols	49	0.4	60	1.2	46	0.8	48	0.8	48	0.5
Ferulic acid	Hydroxycinnamic acids	38	0.3	17	0.7	15	0.5	77	0.9	35	0.4
Proanthocyanidin 7–10 oligomers	Flavanols	34	0.2	46	1.0	33	0.5	33	0.5	32	0.3
Hesperidin	Flavanones	27	0.3	22	0.7	19	0.4	27	0.7	30	0.5
5-Feruloylquinic acid	Hydroxycinnamic acids	26	0.2	16	0.4	22	0.7	32	0.3	27	0.2
(+)-Gallic acid	Flavanols	24	0.3	3.4	0.2	10	0.3	27	0.6	32	0.4
(-)-Epicatechin	Flavanols	21	0.1	19	0.3	16	0.2	21	0.3	22	0.2
5-O-Galloylquinic acid	Hydroxybenzoic acids	20	0.2	2.8	0.2	8.8	0.3	23	0.5	27	0.4
4-Feruloylquinic acid	Hydroxycinnamic acids	19	0.1	12	0.3	16	0.5	23	0.2	20	0.1
Proanthocyanidin trimers	Flavanols	19	0.2	20	0.5	17	0.4	19	0.4	19	0.2
(-)-Epigallocatechin 3-O-gallate	Flavanols	18	0.2	2.8	0.2	7.8	0.3	21	0.4	24	0.3
Procyanidin dimer B2	Flavanols	18	0.1	21	0.3	15	0.2	18	0.2	18	0.1
(-)-Epicatechin 3-O-gallate	Flavanols	15	0.1	3.4	0.1	6.8	0.2	17	0.3	20	0.2
Procyanidin dimer B1	Flavanols	14	0.1	17	0.4	14	0.3	13	0.2	15	0.1
(-)-Epigallocatechin	Flavanols	14	0.1	2.4	0.1	6.2	0.2	17	0.3	19	0.2
(+)-Catechin	Flavanols	14	0.1	17	0.3	10	0.1	14	0.2	14	0.1
Gallic acid	Hydroxybenzoic acids	13	0.2	11	0.9	11	0.7	12	0.2	14	0.3
Cyanidin 3-O-rutinoside	Anthocyanins	11	0.4	12	0.8	17	1.1	7.4	0.6	12	0.5
5-Heneicosylresorcinol	Alkylphenols	10	0.1	8.4	0.2	5.6	0.1	15	0.1	10	0.1
5-Nonadecylresorcinol	Alkylphenols	10	0.1	6.8	0.2	4.4	0.1	17	0.2	10	0.1
3-Feruloylquinic acid	Hydroxycinnamic acids	9.3	0.1	6.0	0.2	8.0	0.2	11	0.1	10	0.1

Polyphenol (mg/day)	Polyphenol subclass	UK health-conscious group				Top 3 main food sources
		Men		Women		
		Mean	se	Mean	se	
5-Caffeoylquinic acid	Hydroxycinnamic acids	159	11.5	134	9.2	Coffee (78.2 %), potatoes (7.3 %), apples and pears (6.0 %)
4-Caffeoylquinic acid	Hydroxycinnamic acids	102	9.9	84	7.5	Coffee (97.5 %), tea (1.5 %), stone fruits (0.4 %)
3-Caffeoylquinic acid	Hydroxycinnamic acids	88	8.6	75	6.8	Coffee (92.4 %), stone fruits (6.0 %), tea (0.4 %)
Proanthocyanidin polymers (>10 mers)	Flavanols	133	16.6	118	11.4	Apples and pears (24.6 %), chocolate products (19.8 %), legumes (11.4 %)
Proanthocyanidin 4-6 oligomers	Flavanols	64	6.5	66	7.0	Apples and pears (33.1 %), chocolate products (24.9 %), wine (7.5 %)
Ferulic acid	Hydroxycinnamic acids	41	6.6	31	3.5	Bread (66.2 %), pasta and rice (9.7 %), breakfast cereals (6.3 %)
Proanthocyanidin 7-10 oligomers	Flavanols	50	5.5	44	4.1	Apples and pears (39.8 %), chocolate products (17.5 %), legumes (8.4 %)
Hesperidin	Flavanones	46	6.9	35	5.0	Fruit juices (66.2 %), citrus fruit (28.6 %), carbonated/soft drinks (3.9 %)

Table 4 continued

Polyphenol (mg/day)	Polyphenol subclass	UK health-conscious group				Top 3 main food sources
		Men		Women		
		Mean	se	Mean	se	
5-Feruloylquinic acid	Hydroxycinnamic acids	19	2.0	15	1.5	Coffee (99.7 %), root vegetables (0.2 %), cream desserts (0.1 %)
(+)-Gallic catechin	Flavanols	73	9.0	81	6.0	Tea (99.6 %), wine (0.3 %), fruiting vegetables (0.1 %)
(-)-Epicatechin	Flavanols	33	2.8	37	2.1	Tea (35.9 %), apples and pears (25.7 %), wine (11.4 %)
5-O-Galloylquinic acid	Hydroxybenzoic acids	60	7.5	67	5.0	Tea (100 %)
4-Feruloylquinic acid	Hydroxycinnamic acids	14	1.4	11	1.1	Coffee (99.8 %), root vegetables (0.1 %), cream desserts (0.1 %)
Proanthocyanidin trimers	Flavanols	22	2.7	28	4.3	Chocolate products (34.5 %), apples and pears (25.3 %), wine (7.9 %)
(-)-Epigallocatechin 3-O-gallate	Flavanols	48	5.9	53	3.9	Tea (99.5 %), herbal tea (0.5 %)
Procyanidin dimer B2	Flavanols	26	2.1	27	1.5	Apples and pears (45.3 %), tea (24.1 %), wine (15.3 %)
(-)-Epicatechin 3-O-gallate	Flavanols	43	4.6	46	3.1	Tea (87.4 %), herbal tea (8.8 %), wine (2.8 %)
Procyanidin dimer B1	Flavanols	23	2.4	26	1.7	Tea (43.6 %), wine (15.8 %), stone fruits (14.4 %)
(-)-Epigallocatechin	Flavanols	38	4.6	42	3.1	Tea (99.1 %), wine (0.3 %), herbal tea (0.2 %)
(+)-Catechin	Flavanols	23	2.0	24	1.3	Tea (30.7 %), wine (29.6 %), apples and pears (7.7 %)
Gallic acid	Hydroxybenzoic acids	26	3.0	29	2.0	Tea (60.4 %), wine (16.3 %), mixed fruits (12.8 %)
Cyanidin 3-O-rutinoside	Anthocyanins	5.2	1.9	8.8	2.1	Stone fruits (64.5 %), fruit juices (12.8 %), carbonated/soft drinks (7.2 %)
5-Heneicosylresorcinol	Alkylphenols	22	2.1	14	1.0	Bread (81.0 %), pasta and rice (8.7 %), breakfast cereals (7.5 %)
5-Nonadecylresorcinol	Alkylphenols	17	1.6	10	0.6	Bread (88.6 %), pasta and rice (5.1 %), breakfast cereals (4.4 %)
3-Feruloylquinic acid	Hydroxycinnamic acids	6.7	0.7	5.4	0.5	Coffee (97.9 %), stone fruits (1.4 %), root vegetables (0.3 %)

MED Mediterranean

^a Means and standard error (s.e.) were computed using general linear models adjusted for sex, age, and center and weighted by season and weekday of recall

[36] and the use of flour fortified with soya protein in the United Kingdom [38].

Lignans are minor contributors to total polyphenol intake as reported earlier (<0.7 %) [17, 18, 20]. We estimated the intake of 29 lignans, including those usually assessed in other cohort studies, such as lariciresinol, matairesinol, pinoresinol, and secoisolariciresinol [36, 39]. One of the three most widely consumed lignans was sesamol, a lignan not considered in previous studies on polyphenol intake. This highlights the importance of using the most comprehensive databases when evaluating the relationships between lignan/polyphenol intake and health outcomes.

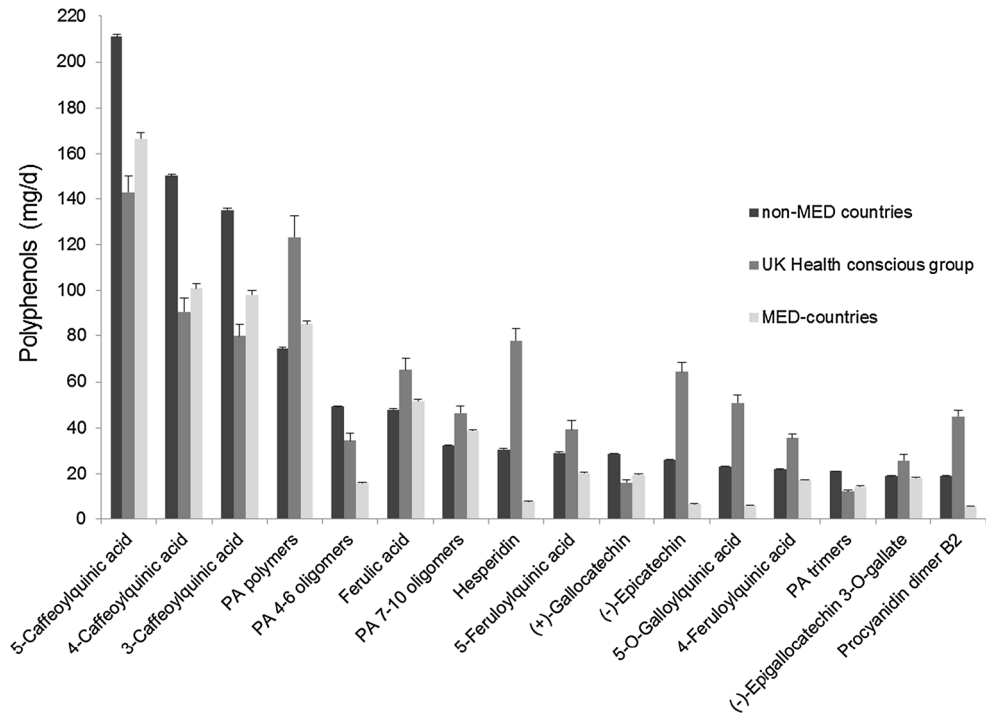
Resveratrol is the most well-known stilbene, and in the last decade, it has received much attention due to its ability to modulate the expression of sirtuin deacetylases, and to

extend life span of model organisms [40]. However, the intake of stilbenes in the European populations is very low, and their intake does not exceed 3.1 mg/day in agreement with values previously published [17, 18, 41].

Other polyphenols beside the four classes described above, include alkylphenols, contributing from 1.6 % of total polyphenols in women from MED countries to 4.0 % in men from non-MED countries, where whole-grain cereals are more widely consumed. They also include tyrosols (0.4–3.6 % of total polyphenols) characteristic of foods more abundant in the diet of MED countries, such as olive oil, olives, and wine [11]. Intakes of other polyphenols were almost negligible (<0.7 % of total polyphenol).

The present study shows a number of strengths, in particular the large number of EPIC participants, the use of the Phenol-Explorer database as the most comprehensive food

Fig. 2 Adjusted daily intake (mg/day), mean and standard error, of the top 10 polyphenol of each European region adjusted for age, sex, and center and weighted by season and weekday of dietary recall in the EPIC study. *MED* Mediterranean, *PA* proanthocyanidins



composition database for polyphenols, the uniform application of retention factors to take into account the effects of cooking and food processing on polyphenol contents in foods, and the use of a standardized 24-HDR in all participants, allowing to easily compare results across countries. However, our study has some limitations. Our results cannot be totally generalizable, since not all the EPIC cohorts are population-based [21, 22]. Another limitation of this study is that each person contributed only one 24-HDR, hence variation in intakes cannot be evaluated at the individual level and it is less likely to reflect true usual individual diet. However, analyses were weighted by season and weekday of dietary recall, and 24-HDR is a useful method to describe the average dietary intake of a group, particularly when estimated from a large number of subjects [42]. A relevant weakness is also the likely underestimation of total polyphenol intake, due to some missing or insufficient food composition data for polyphenols such as thearubigins [33] or proanthocyanidins which are not easily measured in foods as well as the lack of data on the content of polyphenol-rich additives in processed foods [2], and the omission of herb/plant supplements in the present analysis [43]. Another limitation that may result in inaccuracies in intake estimates for the major coffee polyphenols is linked to the lack of detailed data on polyphenol composition for various types of coffee drinks, according to brewing methods and cultivars (Arabica vs. Robusta) [27, 44]. A similar limitation exists for herbal teas, which were insufficiently documented in the 24-HDR and vary widely in polyphenol composition. The lack of food composition

data in these polyphenol-rich foods, such as coffee and herbal tea, was minimized estimating their composition as described in the methodology section.

In conclusion, this study provides the most detailed description of polyphenol intake in a large European multi-center study. It shows a large heterogeneity in both the nature of polyphenols and levels of intake across countries, particularly between MED and non-MED countries. The main food sources for individual polyphenols were similar between regions. However, the major food sources for total polyphenols and specific polyphenol subclasses may vary widely according to regions or countries and food preferences since many compounds show a discrete distribution in foods and can be specific of a food or group of foods consumed in particular regions and not in others. This can be particularly important for polyphenols which would show highly potent and specific biological properties. Lastly, we also showed that socio-demographic, anthropometric, and lifestyle factors were associated with differential intake of polyphenols. These descriptive data provide a unique platform to further investigate the role of polyphenols in the prevention of diseases and in healthy aging in one of the largest European cohorts with detailed dietary data.

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