Comparison of 2 methods for estimating the prevalences of inadequate and excessive iodine intakes $1-3$

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ABSTRACT

Background: Prevalences of iodine inadequacy and excess are usually evaluated by comparing the population distribution of urinary iodine concentration (UIC) in spot samples with established UIC cutoffs. To our knowledge, until now, dietary intake data have not been assessed for this purpose.

Objective: Our objective was to compare 2 methods for evaluating the prevalence of iodine inadequacy and excess in sex- and life stage–specific subgroups of the US population: one that uses UIC cutoffs, and one that uses iodine intake cutoffs.

Design: By using the iodine concentrations of foods measured in the US Food and Drug Administration's Total Diet Study (TDS), dietary intake data from the NHANES 2003–2010, and a file that maps each NHANES food to a TDS food with similar ingredients, we estimated each NHANES participant's iodine intake from each NHANES food as the mean iodine concentration of the corresponding TDS food in samples gathered over the same 2-y period. We calculated prevalences of iodine inadequacy and excess in each sexand life stage–specific subgroup by both the UIC cutoff method and the iodine intake cutoff method—using the UIC values and dietary intakes reported for NHANES participants who provided both types of data—and compared the prevalences across methods.

Results: We found lower prevalences of iodine inadequacy across all sex- and life stage–specific subgroups with the iodine intake cutoff method than with the UIC cutoff method; for pregnant females, the respective prevalences were 5.0% and 37.9%. For children aged \leq 8 y, the prevalence of excessive iodine intake was high by either method. Conclusions: The consideration of dietary iodine intake from all sources may provide a more complete understanding of population prevalences of iodine inadequacy and excess and thus better inform dietary guidance than consideration of UIC alone. Methods of adjusting UIC for within-person variation are needed to improve the accuracy of prevalence assessments based on UIC. Am J Clin Nutr 2016;104(Suppl):888S–97S.

Keywords: dietary surveys, food content, iodine, Total Diet Study, urinary iodine, variability

INTRODUCTION

Prevalences of iodine inadequacy and excess are usually evaluated by comparing the population distribution of urinary iodine concentrations $(UICs)^9$ in spot samples with established cutoffs for median UIC concentrations in sex- and life stage– specific population subgroups, such as those developed by the WHO (1). To our knowledge, until now, the use of dietary intake data for this purpose has not been assessed.

Because UIC mostly reflects recent iodine intake, large day-today variability in iodine intake is reflected in large day-to-day variability in UIC (2). For that reason, as discussed elsewhere in this supplement issue, UIC measured in spot samples or single 24-h collections is not a reliable measure of the iodine status of individuals (3). In addition, UIC is not informative as to specific dietary sources of iodine or the amounts obtained from each source; thus, it is not helpful for formulating dietary guidance.

In contrast, a reliable means of assessing dietary intake could be used to monitor the iodine status of individuals over time and to inform the interpretation of UIC in individuals. However, there are also limitations to the use of dietary survey data for assessing iodine status. One issue is that food intakes may be under- or overreported (4). In addition, the iodine concentration of many foods is poorly characterized and its variability is often high. A discussion of the causes of variability in the iodine concentration of foods (5) and a statistical assessment of that variability (6) are presented elsewhere in this supplement issue.

Three sets of Dietary Reference Intakes for iodine have been developed for children and adults by the Food and Nutrition Board of the Institute of Medicine: Estimated Average Requirements (EARs), Recommended Dietary Allowances (RDAs), and Tolerable Upper Intake Levels (ULs) (7). The RDA is intended for use

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³ Supplemental Tables 1–5 are available from the "Online Supporting Material" link in the online posting of the article and from the same link in the online table of contents at http://ajcn.nutrition.org.

⁹ Abbreviations used: EAR, Estimated Average Requirement; FNDDS, Food and Nutrient Database for Dietary Studies; RDA, Recommended Dietary Allowance; TDS, Total Diet Study; UIC, urinary iodine concentration; UL, Tolerable Upper Intake Level.

primarily as a goal for an individual's usual intake, whereas the EAR is used to assess the population prevalence of inadequate intakes.

In the present study, our first aim was to develop a method for using dietary intake data from NHANES and data on the iodine concentration of foods from the US Food and Drug Administration's Total Diet Study (TDS) to estimate total usual daily iodine intakes. Our second aim was to compute the prevalences of iodine inadequacy and excess across sex- and life stage– specific subgroups of NHANES participants by comparing the estimated total usual daily iodine intakes for each subgroup with subgroup-specific EARs and ULs. Our third aim was to identify and use EAR-like and UL-like UIC cutoffs to calculate subgroup-specific prevalences of iodine inadequacy and excess among NHANES participants who provided both UIC data and dietary intake data. Our final aim was to compare prevalences of iodine inadequacy and excess calculated by using iodine intake cutoffs with those estimated by using UIC cutoffs.

METHODS

We used SAS version 9.3 (SAS Institute) for all statistical analyses unless otherwise noted. A flowchart summarizing the study protocol is shown in Figure 1.

NHANES data

NHANES, an annual survey of the civilian, noninstitutionalized US population conducted by the CDC's National Center for Health Statistics, collects dietary intake data for all participants \geq 1 y old on 2 survey days and measures UIC in nontimed spot urine samples from participants ≥ 6 y old on the first survey day (8). In the present study, we examined the dietary intake and UIC data of NHANES participants for the years $2003-2004$, $2005-2006$, $2007-2008$, and $2009-2010$. During this time period, the CDC measured UIC by using variations on an inductively coupled plasma mass spectrometry method (9, 10).

Dietary intake data

NHANES collects data on the foods, plain drinking water, and beverages consumed from midnight to midnight of the previous day and the types and amounts of dietary supplements consumed in the previous 30 d. We included in our study all NHANES participants aged ≥ 1 y with complete 24-h dietary data for both survey days ($n = 31,352$) and analyzed the dietary iodine intake data for each day separately.

UIC data

The CDC measured UIC in spot samples from approximately one-third of the $2003-2004$, $2005-2006$, and $2009-2010$ participants aged \geq 6 y and all of the 2007-2008 participants aged \geq 6 y. All of the NHANES participants with urinary iodine data $(n = 15,545)$ were included in our study.

Dietary iodine evaluation

We estimated iodine intake from food, drinking water, dietary supplements, salt used in cooking, and salt used at the table.

Iodine from drinking water

On the basis of the reported amount of total plain drinking water consumed by NHANES participants on each dietary survey day, we calculated iodine intake from drinking water by assuming an iodine concentration of 9.2 μ g/L, the median of values (ranging from 0.1 to 25 μ g/L) measured at various US locations in available studies (11, 12).

Iodine from dietary supplements

We calculated the daily iodine intake from dietary supplements on the basis of the fractional number of days they were used, the number of servings per day, and the iodine content per serving as listed on the product label. Dietary supplements containing kelp are very high in iodine. We excluded kelp-containing supplements when calculating the iodine intakes of participants who reported the use of such supplements $(n = 39)$ to avoid biasing the summary statistics toward high values inconsistent with the overall distributions of the iodine intake from supplements and the total iodine intake.

Iodine from salt used at the table

Table salt contains 387.6 mg Na/g (13); if the salt is iodized, the iodine concentration is nominally 45 μ g/g (14). First, we estimated the daily intake of sodium from salt used at the table, as described below. Next, we estimated the daily intake of table salt (in g) as the sodium intake (in mg) divided by 387.6. Under the assumption that \sim 70% of table salt sold in the United States is iodized (14), we estimated the daily intake of iodine from table salt (in μ g) as the table salt intake (in g) multiplied by 45×0.7 . For NHANES participants who indicated the use of "ordinary salt" (i.e., table salt), we assigned a default intake of 580 mg sodium from salt used at the table to all ages on the basis of the mean value for adults reported in a 1991 study (15). We are not aware of a later study in adults or any similar study in children. For participants who indicated the use of "lite salt," we assigned a default table salt sodium intake of one-half that value (290 mg). For participants whose response was "salt substitute," "other," or "don't use," we assigned a default table salt sodium intake of zero. If the frequency of salt use was "very often," "occasionally," or "rarely," we multiplied the default table salt sodium intake by 1, 0.5, or 0.25, respectively. A similar approach has been used for estimating total sodium consumption (16).

Iodine from foods

The Food and Drug Administration's TDS measures the concentrations of iodine and other nutrients in a "market basket" of 286 foods collected 4 times/y, once from each of 4 geographic regions (17, 18). The foods recorded by NHANES participants are drawn from the ~ 6200 foods in the USDA's Food and Nutrient Database for Dietary Studies (FNDDS). The FNDDS provides data on the concentrations of 65 nutrients and food components, but iodine is not among them (19). To account for the iodine content of foods recorded by NHANES participants, we used a file that maps each FNDDS food to a TDS food on the basis of the similarity of their ingredients (JH Spungen, unpublished data, 2014). Because there are far fewer TDS foods than FNDDS foods, multiple FNDDS foods are mapped to a single TDS food. For example, 15 natural cheeses listed in the FNDDS are mapped to natural cheddar cheese in the TDS. With the use of the TDS foods as surrogates for the FNDDS foods, we

890S JUAN ET AL.

in cooking and at the table on the correlation between UIC and iodine intake.

FIGURE 1 Flowchart of the study protocol. EAR, Estimated Average Requirement; FNDDS, Food and Nutrient Database for Dietary Studies; TDS, Total Diet Study; UIC, urinary iodine concentration; UL, Tolerable Upper Intake Level.

estimated the iodine concentration of each food recorded by NHANES participants as the mean iodine concentration of the corresponding TDS food in samples collected over the same 2-y period as the NHANES data.

Iodine from salt in prepared foods

The TDS kitchens use only noniodized salt. To account for the use of iodized salt in home food preparation, first we estimated the amount of salt used in preparing 47 TDS foods on the basis of recipes for similar foods in the FNDDS (20). We then calculated iodine intake from salt used in cooking as described above for table salt.

Total usual daily iodine intakes

We estimated total usual daily iodine intakes using the "usual intake" method developed at Iowa State University (21). By taking the inter- and intraindividual variability in food consumption into account, the Iowa State University method diminishes the impact of over- and underreporting.

Correlation of UIC with sources of dietary iodine

We examined the relation between UIC and various combinations of dietary iodine sources by calculating the Pearson correlation coefficient at a significance criterion of $P < 0.05$.

Prevalence of inadequate and excessive intakes

We calculated prevalences of iodine inadequacy and excess from urinary data using SAS version 9.3 and from iodine intake data using Personal Computer Software for Intake Distribution Estimation (PC-SIDE version 1.1, 2003; Department of Statistics and Center for Agricultural and Rural Development, Iowa State University).

Iodine intake cutoff method

Table 1 shows subgroup-specific EARs, RDAs, and ULs developed by the Food and Nutrition Board of the Institute of Medicine. To avoid overestimating the prevalence of iodine inadequacy in a population, it is crucial to use the EAR and not the RDA as the cutoff (22, 23). We estimated the prevalence of iodine inadequacy as the prevalence of total usual iodine intakes below the subgroup-specific EAR. Similarly, we estimated the prevalence of excessive iodine intake as the prevalence of total usual iodine intakes above the subgroup-specific UL.

UIC cutoff method

Because UIC is typically used to assess the iodine adequacy of populations, we set the RDA-like UIC cutoffs to the lowest WHO guideline median UIC cutoffs for adequate intake: $150 \mu g/L$ for pregnant females and 100 μ g/L for other population subgroups, including lactating females and children ≥ 6 y of age (1). Similarly, we set the UL-like UIC cutoffs to the WHO guideline median UIC cutoffs for excessive intake: 500 μ g/L for pregnant females and 300 μ g/L for other population subgroups, excluding lactating females and including children ≥ 6 y of age (1). There is no established WHO UIC cutoff guideline for excessive iodine intake in lactating females.

We calculated EAR-like UIC cutoffs as follows (24):

$$
EAR(\mu g/d) / RDA(\mu g/d) \times RDA-like \text{ UIC cutoff}(\mu g/L) \quad (1)
$$

We estimated the subgroup-specific prevalence of iodine inadequacy as the prevalence of UIC values below the relevant EARlike UIC cutoff. Similarly, we estimated the subgroup-specific prevalence of excessive iodine intake as the prevalence of UIC values above the relevant UL-like UIC cutoff. The UIC cutoffs for each subgroup are shown in Table 1.

RESULTS

Distributions of iodine intake and UIC

Table 2 reports mean and median values of total usual iodine intake and UIC across sex- and life stage–specific subgroups for NHANES participants who reported data for both measures on survey day 1 (the "matched" population, $n = 13,043$). Percentiles of the distributions of total usual iodine intake and UIC in these subgroups are shown in Supplemental Tables 1 and 2, respectively. Percentiles of the distribution of total usual iodine intake in subgroups of the larger, intersecting population of NHANES participants who provided dietary intake data on both survey days (the "intake" population, $n = 31,352$) are shown in Supplemental Table 3. Unlike the matched population, the intake population includes children as young as 1 y old.

Prevalence of iodine inadequacy

Table 3 presents, for the matched population, a comparison of the prevalence of iodine inadequacy calculated on the basis of total usual iodine intake with that calculated on the basis of UIC. In every sex- and life stage–specific subgroup, the prevalence of iodine inadequacy on the basis of total usual iodine intake was markedly lower than that based on UIC. Based on iodine intake, both boys and girls aged 6–8 y and 9–13 y had a 0.0% prevalence of iodine inadequacy; the corresponding prevalences based on UIC were 5.8% and 7.9% in boys and 11.2% and 12.5% in girls, respectively. The highest calculated prevalence of inadequacy based on iodine intake, 5.0%, was for pregnant females; the corresponding prevalence based on UIC was 37.9%.

Calculated prevalences of iodine inadequacy in the intake population across 4 iodine intake scenarios of varying completeness are shown in Table 4 for females and in Supplemental Table 4 for males. The scenarios included the following: I) foods and dietary supplements, 2) the latter plus drinking water, 3) the latter plus salt used in cooking, and 4) the latter plus salt used at the table (identified as total usual iodine intake). When salt used in cooking and at the table was included in the calculation (scenario 4 compared with scenario 2), the calculated

TABLE 1	
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Calculated UIC cutoffs for sex- and life stage–specific subgroups based on Dietary Reference Intakes for iodine¹

¹Urinary data were available only for survey participants ≥ 6 y of age; for that reason, we did not develop UIC cutoffs for children <6 y of age. The EARs, RDAs, and ULs are Dietary Reference Intakes developed by the Food and Nutrition Board of the Institute of Medicine (7). EAR, Estimated Average Requirement; RDA, Recommended Dietary Allowance; NC, not calculated; UIC, urinary iodine concentration; UL, Tolerable Upper Intake Level. ²

²EAR-like UIC cutoffs were calculated as follows: RDA-like UIC cutoff (μ g/L) \times EAR (μ g/d)/RDA (μ g/d).

³RDA-like UIC cutoffs were set at the lowest WHO guideline median UIC cutoffs for adequate iodine intake: 150 μ g/L for pregnant females and 100 μ g/L for other population subgroups, including lactating females and children \geq 6 y of age (1).

⁴UL-like UIC cutoffs were set at the WHO guideline median UIC cutoffs for excessive iodine intake: 500 µg/L for pregnant females and 300 μ g/L for other population subgroups, excluding lactating females and including children ≥ 6 y of age (1).

TABLE 2

Mean and median estimated usual iodine intakes and UICs in sex- and life stage-specific subgroups of the "matched" population¹

¹The "matched" population consisted of the participants in NHANES 2003-2010 who provided both UIC data and dietary intake data. The estimated usual iodine intake is based on iodine from foods, plain drinking water, dietary supplements, salt used in cooking, and salt used at the table. The iodine concentration of drinking water was assumed to be 9.2 μ g/L. UIC, urinary iodine concentration.

prevalence of iodine inadequacy decreased from 11.9% to 3.7% in pregnant females and from 6.3% to 1.2%, from 7.9% to 1.2%, and from 6.3% to 0.8% in the 3 subgroups of nonpregnant females aged 14–50 y.

Prevalence of iodine excess

Table 3 also presents, for the matched population, a comparison of the prevalence of iodine excess calculated on the basis of total usual iodine intake with that calculated on the basis of UIC. The prevalences of iodine excess based on iodine intake in boys and girls aged 6–8 y were 59.8% and 53.0%, respectively; the corresponding prevalences based on UIC were somewhat lower: 41.7% and 35.1%, respectively. For all other subgroups, the calculated prevalences of iodine excess based on iodine intake were in the range of 0.0–0.9%, values that are markedly lower than those based on UIC (5.9–32.6%).

The calculated prevalences of iodine excess in the intake population across the same 4 iodine intake scenarios defined above are shown in Table 5 for females and in Supplemental Table 5 for males. The largest prevalences of iodine excess were found in children <6 y of age. On the basis of total usual iodine intake (scenario 4), prevalences of $\geq 82\%$ and $\geq 49\%$ were calculated for children of either sex aged 1–3 y and 4–8 y, respectively.

The relative contributions of foods, drinking water, dietary supplements, salt used in cooking, and salt used at the table to the calculated iodine intake of the intake population are shown in Figure 2 for males and Figure 3 for females. Consistent with the data presented in Supplemental Table 4, the major sources of iodine intake were foods, dietary supplements, and salt used in cooking.

Correlations between UIC and various combinations of dietary iodine sources are shown in Table 6. Participants with UICs >1000 μ g/L (n = 106) were excluded from the correlation analyses to avoid biasing the results toward nonrepresentative data pairs at the extreme high end of the UIC distribution. All combinations of dietary iodine sources tested were significantly correlated with UIC ($P < 0.0001$). However, the correlations were not strong and there was little or no change across the various combinations of dietary sources. In nonpregnant females aged 14–50 y, r ranged from 0.12 to 0.15, and in pregnant females aged $14-50$ y, r ranged from 0.36 to 0.40.

DISCUSSION

According to our analysis of NHANES UIC data for 2003– 2010, the median UICs of all sex- and life stage–specific subgroups of the US population aged ≥ 6 y other than pregnant females exceeded the WHO's cutoff of $100 \mu g/L$ for iodine adequacy (1). Our findings are consistent with other analyses of NHANES data on the adequacy and stability of iodine status in the general US population since 2000 (25–27). In addition, according to our analysis, the median UIC of pregnant females (\sim 140 μ g/L) was slightly below the WHO criterion for adequacy, 150 μ g/L. Other studies reported median UICs of 125– 145 μ g/L for pregnant females surveyed by NHANES between 2001 and 2010 (28, 29).

A previous study found that salt used in cooking was not significantly associated with an increase in median UIC among females aged 15–44 y (28). Consistent with that result, we found that the correlation between UIC and iodine intake did not improve by accounting for iodized salt used in cooking and at the table. The observed lack of improvement in the correlation might be explained in part by our inability to account for variability in the iodine content of iodized salt, which depends on temperature, humidity, and length of storage; concentrations ranging from 15 to

TABLE 3

Prevalence rates of iodine inadequacy and iodine excess in sex- and life stage–specific subgroups of the "matched" population based on UIC or estimated total usual iodine intake

¹The "matched" population consisted of the 13,043 participants in NHANES 2003-2010 who provided both UIC data and dietary intake data. EAR, Estimated Average Requirement; NA, not available (the SE cannot be computed); UIC, urinary iodine concentration; UL, Tolerable Upper Intake Level. ²

 T The rate of iodine inadequacy is calculated as the percentage of the subgroup with estimated iodine intake below the EAR; the rate of iodine excess is calculated as the percentage of the subgroup with estimated iodine intake above the UL. The estimated total usual iodine intake is based on iodine from foods, plain drinking water, dietary supplements, salt used in cooking, and salt used at the table. The iodine concentration of drinking water was assumed to be 9.2μ g/L.

 $3³$ The rate of iodine inadequacy is calculated as the percentage of the subgroup with UIC below the EAR-like cutoff; the rate of iodine excess is calculated as the percentage of the subgroup with UIC above the UL-like cutoff.

80 mg/kg have been reported (2, 14). In addition, our method of calculating the iodine intake attributable to table salt had the effect of inflating the amount consumed by NHANES participants who used noniodized salt at the table and, at the same time, depressing (by \sim 30%) the amount consumed by NHANES participants who used iodized salt. In addition, because salt intake (measured as urinary excretion of sodium) increased slightly in US adults from the years 1988–1991 to 2010 (30), our reliance on 1991 data are

TABLE 4

Prevalence of iodine inadequacy by age and pregnancy status in female members of the "intake" population, based on estimated usual iodine intake from various sources¹

	Prevalence of iodine inadequacy, 2%					
	Iodine from foods and DS	Iodine from foods, DS, and DW	Iodine from foods, DS, DW, and S_C	Iodine from foods, DS, DW, S_C , and S_T		
Nonpregnant						
$1-3$ y	0.5 ± 0.2	0.5 ± 0.2	0.3 ± 0.1	0.2 ± 0.1		
$4-8y$	0.1 ± 0.1	0.1 ± 0.1	$0.0 \pm NA$	$0.0 \pm NA$		
$9 - 13$ y	0.6 ± 0.3	0.5 ± 0.3	0.2 ± 0.1	$0.0 \pm NA$		
$14 - 18$ y	7.7 ± 1.6	6.3 ± 1.4	3.4 ± 1.0	1.2 ± 0.5		
$19 - 30$ y	10.5 ± 1.7	7.9 ± 1.5	3.2 ± 1.0	1.2 ± 0.5		
$31 - 50y$	8.6 ± 1.1	6.3 ± 1.0	2.3 ± 0.6	0.8 ± 0.3		
$51 - 70$ y	9.8 ± 1.0	7.8 ± 0.9	3.4 ± 0.6	1.7 ± 0.4		
≥ 71 y	9.0 ± 1.2	7.6 ± 1.1	3.7 ± 0.8	2.3 ± 0.6		
Pregnant						
$14 - 50$ y	14.6 ± 2.6	11.9 ± 2.4	5.9 ± 1.8	3.7 ± 1.4		

¹The "intake" population consisted of the 31,352 participants in NHANES 2003-2010 who provided dietary intake data on both survey days. The iodine concentration of DW was assumed to be 9.2 µg/L. DS, dietary supplements; DW, drinking water; NA, not available (the SE cannot be computed); S_C , salt used in cooking; S_T , salt used at the table.

²Values are percentages \pm SEs of individuals in each age group with iodine intakes below the Estimated Average Requirement for each iodine intake scenario shown.

TABLE 5

Prevalence of excessive iodine intake by age and pregnancy status in female members of the "intake" population, based on estimated usual iodine intake from various sources

	Prevalence of excessive iodine intake, 2%					
	Iodine from foods and DS	Iodine from foods, DS, and DW	Iodine from foods, DS, DW, and S_C	Iodine from foods, DS, DW, S_C , and S_T		
Nonpregnant						
$1-3$ y	78.3 ± 1.8	79.2 ± 1.8	82.9 ± 1.8	84.2 ± 1.7		
$4-8$ y	36.3 ± 1.9	38.0 ± 1.9	43.7 ± 1.8	49.2 ± 1.7		
$9 - 13$ y	0.5 ± 0.3	0.5 ± 0.3	0.6 ± 0.3	0.9 ± 0.4		
$14 - 18$ y	$0.0 \pm NA$	$0.0 \pm NA$	$0.0 \pm NA$	$0.0 \pm NA$		
$19 - 30y$	$0.0 \pm NA$	$0.0 \pm NA$	$0.0 \pm NA$	$0.0 \pm NA$		
$31 - 50$ y	$0.0 \pm NA$	$0.0 \pm NA$	$0.0 \pm NA$	$0.0 \pm NA$		
$51 - 70$ y	0.4 ± 0.1	0.5 ± 0.1	0.4 ± 0.1	0.5 ± 0.2		
≥ 71 y	$0.0 \pm NA$	$0.0 \pm NA$	$0.0 \pm NA$	$0.0 \pm NA$		
Pregnant						
$14 - 50y$	$0.0 \pm NA$	$0.0 \pm NA$	0.1 ± 0.1	0.2 ± 0.2		

¹The "intake" population consisted of the 31,352 participants in NHANES 2003-2010 who provided dietary intake data on both survey days. The iodine concentration of DW was assumed to be 9.2 μ g/L. DS, dietary supplements; DW, drinking water; NA, not available (the SE cannot be computed); S_c , salt used in cooking; S_T , salt used at the table.

²Values are percentages \pm SEs of individuals in each age group with iodine intake above the Tolerable Upper Intake Level for each iodine intake scenario shown.

more likely to have underestimated adult use of salt at the table than to have overestimated such use. Finally, our reliance on a value obtained in adults may have overestimated or underestimated children's use of salt at the table.

Our estimates of iodine intake have several limitations. First, the thousands of foods potentially consumed by NHANES participants do not necessarily provide the same iodine content as the 286 TDS foods to which they were matched. Thus, our mapping strategy may have over- or underestimated iodine intake from the foods consumed by NHANES participants. In addition, for foods with highly variable iodine concentrations, our use of the mean iodine concentration rather than the median may have overestimated iodine content, thereby inflating the estimated

iodine intake of individuals who reported consuming those foods (31). Another potential limitation in assessing iodine intake by using 24-h dietary recalls is that saltwater fish and seafood, both important sources of dietary iodine, tend to be consumed episodically in the Western diet (32). Episodic intake might be missed by two 24-h recalls, resulting in underestimation of some participants' intakes. Less often, intake might be overestimated for a participant who rarely consumes fish or seafood but does so coincident with the survey. Another concern is that by excluding kelp-containing supplements when calculating the iodine intakes of the 39 NHANES participants who reported using such supplements, we may have greatly underestimated their iodine intake from supplements as well as their total iodine intake.

FIGURE 2 Proportions of mean iodine intake contributed by 5 dietary sources, calculated for male participants in NHANES 2003–2010 with dietary intake data for both survey days.

FIGURE 3 Proportions of mean iodine intake contributed by 5 dietary sources, calculated for female participants in NHANES 2003–2010 with dietary intake data for both survey days. The right-most column (labeled "p") presents data for pregnant females; the other columns exclude data from pregnant females.

In general, food intake data are subject to both under- and overreporting of the consumption of foods with high iodine content, which can result in both under- and overestimation of the prevalence of iodine inadequacy and excess. However, the impact of individual reporting errors was reduced by our adjustment of individual intake distributions for intra- and interindividual variation.

In all of the subgroups, the estimated prevalence of iodine inadequacy based on total usual dietary iodine intake was substantially lower than that based on UIC. One explanation for this

discrepancy concerns the timing of urine collection. In NHANES, UIC was measured in single spot urine samples that were nontimed (i.e., collection times varied). If sampling times tended to precede the meal with the highest iodine intake, this would have biased the UIC toward values that underestimate 24-h intake. In one study, UIC measured in timed spot urine samples showed a good correlation with median 24-h urinary iodine excretion (33). In addition to the uniformity produced by timing the spot urines, the participants' iodine intakes in that study may have been more homogeneous than those of the general US population because

TABLE 6

Correlation between UIC and various measures of dietary iodine intake in sex- and life stage–specific subgroups of the "matched" population¹

		Correlation between UIC and iodine intake ² from				
	n	Foods and DW	Foods and DS	Foods, DW, and DS	Foods, DW, DS, and S_C	Foods, DW, DS, S_C , and S_T
Age						
$6 - 18y$	3875	0.23	0.25	0.25	0.24	0.24
\geq 19 y	8726	0.16	0.21	0.20	0.19	0.19
Sex						
Male	6057	0.20	0.23	0.22	0.21	0.21
Female ³	6294	0.15	0.19	0.18	0.17	0.17
Pregnancy status						
Nonpregnant, 14–50 y	3073	0.12	0.15	0.14	0.13	0.13
Pregnant, $14-50y$	250	0.36	0.40	0.39	0.40	0.40

¹The "matched" population consisted of the 13,043 participants in NHANES 2003-2010 who provided both UIC data and dietary intake data. Analysis was based on dietary intake data gathered at the in-person interview on survey day 1, the same day that urine was collected for analysis. Kelp-containing DS were excluded when calculating the iodine intakes of participants who reported their use $(n = 39)$ to avoid biasing the summary statistics toward high values inconsistent with the overall distributions. Participants with UICs $>1000 \mu g/L$ (n = 106) were excluded from the correlation analyses to avoid biasing the results toward nonrepresentative data pairs at the extreme high end of the UIC distribution. The iodine concentration of DW was assumed to be 9.2 $\mu g/L$, the median of values reported in several studies. $P < 0.0001$ for all comparisons. DS, dietary supplements; DW, drinking water; S_C , salt used in cooking; S_T , salt used at the table; UIC, urinary iodine concentration.

² Pearson correlation coefficient, $P < 0.05$.

³Excluding pregnant females.

dietary screening was performed before enrollment (33, 34). Another study found that because of large day-to-day variability in dietary iodine intake, ≥ 10 d of spot urine sampling were needed to reach 20% precision and to develop a reliable marker of individual iodine status in women (35). Substantial within-person variability in iodine excretion between two 24-h urine collections 4–11 d apart (25% for blacks and 17% for other races) and among timed spot urines (23–34%) has been reported (34).

Another important reason why UIC predicted a higher percentage of the population with "inadequate" iodine status than the percentage predicted by total usual dietary iodine intake is that the intake data were adjusted for within-person variability whereas the UIC data were not. Basing the prevalence of inadequacy on unadjusted UIC leads to overestimation because the distribution of single UIC values measured in single spot samples is broader than the distribution of usual UIC $(7, 23)$. One study found that when the UIC was adjusted for populationspecific intra- and interindividual variability, the percentage of women with values $\leq 63 \mu g/L$ decreased from 41% to 7% (24). Other studies have also suggested that the wide variability in the iodine content of some foods and their frequency of consumption may contribute to the magnitude of the intraindividual variation in UICs observed in single spot urine samples. Under some circumstances, the day-to-day variation in UICs could be 30–40% (24, 35, 36). In one study, when \geq 2 spot urine samples were collected, thereby allowing a variability adjustment to be made to the UIC, the distribution tightened; this, in turn, reduced the estimated percentage of the population with UICs \leq 50 μ g/L from 33% to 19% and those with UICs \geq 100 μ g/L from 21% to 17% (37). In a population with suboptimal iodine status and diets less varied than the typical Western diet, variability adjustment of the UIC likewise decreased the prevalences of both iodine inadequacy and iodine excess (38).

The calculated prevalence of iodine excess was very high for children \leq 9 y of age, whether based on UIC or iodine intake, leading to questions about the validity of both the ULs and the UL-like UIC cutoffs for young children. In all subgroups composed of individuals aged \geq 9 y, including pregnant females, the estimated prevalence of iodine excess based on total usual dietary iodine intake was considerably lower than that based on UIC. Following on the discussion above, overestimation of iodine excess may also be a consequence of the use of unadjusted UIC values measured in single spot samples. Furthermore, EAR-like and UL-like UIC cutoffs should be validated against a more reliable biomarker of individual iodine status than UIC (24), assuming that such a biomarker becomes available.

It has been noted by others that collecting multiple measures from a subject at the individual level decreases intraindividual variability and may allow more meaningful analysis of associations between diet-related risk factors and various biomarkers of risk (39). By estimating the prevalences of iodine inadequacy and excess among subjects who provided both UIC and dietary recall data on a single day, we sought to provide a more accurate assessment than what could be provided by either UIC data or iodine intake data alone.

The fact that the correlation between UIC and iodine intake is moderately weak for pregnant females and weaker still for all other subgroups indicates that the 2 assessment methods do not always identify the same individuals with either low or high intakes. It is likely that neither a single spot urine sample nor a single 24-h dietary recall is sufficient to screen individuals for iodine inadequacy and excess; repeated measurements would be needed for this purpose. However, if properly adjusted for within-person variability, either method should be appropriate for estimating the population prevalences of inadequacy and excess.

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