MINI REVIEW



Time trend of cadmium intake in Korea

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Abstract

Objectives The aim of this study was to elucidate past and current levels of cadmium (Cd) intake among the general populations in Korea.

Methods For this purpose, publications reporting dietary intake of cadmium (Cd-D), cadmium concentration in blood (Cd-B) and that in urine (Cd-U) in Korea were retrieved through literature survey for a period from 1975 to 2015.

Results In practice, 9, 21 and 14 articles were available on Cd-D, Cd-B and Cd- U_{cr} (Cd-U as corrected for creatinine concentration), respectively. Linear regression analyses of the reported values as a function of years (i.e., the year when each survey was conducted) showed steady decreases in all of the three exposure markers of Cd-D, Cd-B and Cd- U_{cr} . Factors possibly contributing for the reduction were discussed including the government-set guideline of 0.2 mg/kg for rice and changes in food habits among general populations.

Conclusions There have been steady decreases in Cd-D, Cd-B and Cd-U_{cr}. The current estimates for Cd-D, Cd-B and Cd-U_{cr} were $6.0-7.4 \ \mu g/day$, $0.73-0.83 \ \mu g/L$ and $0.60-0.95 \ \mu g/g$ cr, respectively.

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Keywords Cadmium in diet \cdot Cadmium in blood \cdot Cadmium in urine \cdot General population \cdot Korea \cdot Time trend

Introduction

It was previously observed that cadmium (Cd) contents in rice to be consumed in Korea were among the high group in rice-dependent Asian areas [1, 2]. In fact, surveys by food duplicate collection [3] in two large cities in Korea in 2000 disclosed that the Cd intake was at the level of 21 μ g/day, which was the second highest among 13 areas surveyed in Asia [4]. Cd has insidious toxicities primarily on the renal tubules and then bone tissues after long-term exposure through daily foods [5–7]. Consumption of rice, a traditional staple food in Korea [8], indeed accounted for 23 % [9] to 25 % [10] of total dietary Cd intake. Thus, current dietary Cd intake levels in combination with chronological changes apparently deserve studies from environmental health viewpoints.

The present study was initiated to obtain an answer to this question of public health importance. The results will be presented in this article. A preliminary report has been published [11].

Materials and methods

All data were cited from existing articles, and approval of the study by an institutional review board was considered to be unnecessary.

Two databases of PubMed and KoreaMed (established by Korean Association of Medical Journal Editors) were searched for publications with two keywords of Korea and

Tabl	e 1	Dietary	cadmium	intal	ĸe
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References	Ref. no.	No. of cases studied	Cd-D (µg/day) (GM)	Cd-D (µg/kg BW/day) (GM)	Year of survey	Note
Yum et al.	[<mark>17</mark>]	NA ^a	70.53	1.12 ^b	1980	Estimate by market basket method
Moon et al.	[18]	18	14.4	0.257 ^b	1994	Women in a large city
Ibid.		38	24.8	0.442	1994	Women in a large city
Ibid.		17	18.49	0.330	1994	Women in a small city
Ibid.		34	22.48	0.401	1994	Women in a rural village
Ibid.		107	21.21	0.379	1994	Total
Moon et al.	[11]	NA ^a	17.02	$0.270^{\rm b}$	1995	Estimate
Moon et al.	[19]	38	11.2	0.457	2000	Children
Ibid.		38	16.7	0.3	2000	Mothers
Moon et al.	[11]	NA ^a	16.57	0.257 ^b	2001	Estimate
Oh et al.	[20]	30	12.16	0.203 ^c	2001	Estimate by market basket method
Watanabe et al.	[21]	33	7.6	0.4	2003-2004	Kindergarten children in Seoul
Ibid.		37	20.5	0.89	2003-2004	Kindergarten children in Jeju city
Ibid.		18	11.5	0.52	2003-2004	Kindergarten children in Jeju village A
Ibid.		20	11.6	0.54	2003-2004	Kindergarten children in Jeju village B
Ibid.		108	12.4	0.58	2003-2004	Kindergarten children, total
Moon et al.	[11]	NA ^a	16.08	0.257 ^b	2005	Estimate
Ibid.		NA ^a	14.82	0.235 ^b	2007	Estimate
kim et al.	[22]	457		0.34	2010	Food record by telephone call, 0-6 year-old children
Lim et al.	[23]	4867		0.18	2010-2011	Adult subjects
Huang et al.	[24]	239	6.53	$0.09^{\rm b}$	2011 ^d	Memory recall method ≥ 20 year-old men
Ibid.		404	5.36	0.09 ^b	2011 ^d	Memory recall method ≥ 20 year-old women
Ibid.		643	5.72	0.09 ^b	2011 ^d	Memory recall method ≥ 20 year-old men + women

^a NA not avaiable: Cd-D was estimated based on other food intake data such as national statistics and numbers of cases were not available

^b Adjusted for a body weight of 63 kg for a total of adult men and women [16]

^c Adjusted for a reported mean body weight of 56.4 kg for adult women [16]

^d Year of publication

cadmium. In practice, 22 and 25 articles, respectively, were identified as those with data on 1; cadmium in daily (24-h) foods (Cd-D in μ g/day or μ g/kg body weight/day), 2; cadmium in peripheral blood (Cd-B in μ g/L), or 3; cadmium in urine or preferably as cadmium in urine as corrected for creatinine concentration (Cd-U and Cd-U_{cr}). The articles retrieved through KoreaMed were limited to those either in English, or in Korean with an English abstract. However, publications from Korean government were written only in Korean. Other articles in English were also cited as necessary.

Smirnov's test for extreme value and Pearson's correlation coefficient were employed as necessary. In some articles, Cd-D, Cd-B or Cd-U_{cr} values were provided in terms of arithmetic means (AMs) and arithmetic standard deviations (ASDs) [or arithmetic standard errors (ASEs)], and the geometric mean values (GMs) were calculated by use of the moment method [12] for uniformity in the data compilation, considering that Cd-D, Cd-B and Cd-U_{cr} may distribute log-normally rather than normally. It should be noted that the reports cited were limited to those on areas where there was no known Cd pollution. In studies on Cd-exposed residents in vicinity of abandoned mines and non-exposed control subjects (e.g., [13–15]), the values for the controls were included in the analyses whereas the exposed cases were not.

Survey years were cited as described in the original publications. In case the survey year was not given, the year of publication was taken in the place.

In some reports, Cd-D was presented in terms of μ g/day. To convert to the μ g/kg body weight/day basis, an average body weight of 63 kg was assumed for a total of adult men (69.6 kg) and women (56.4 kg) [16].

In collecting Cd-D data on China, attention was paid to limit to rice-depending southern part of China. It is known that people in the northern half depend primarily on wheat rather than rice.

In calculating the estimates for 2000 and 2015 based on the regression equations established in this work, intercepts

Table 2 Cadmium in peripheral blood

References	Ref. no.	No. of cases studied	Cd-B (µg/L) (GM)	Year of survey	Note
Lee and Kim	[25]	34	2.82 ^a	1985	Urban residents
Ibid.		37	2.43 ^a	1985	Rural residents
Watanabe et al.	[26]	161	1.58	1986	Women in Masan
Hwang et al.	[27]	72	5.8	1987 ^b	Women
Watanabe et al.	[28]	42	1.54	1987 ^b	Male farmers
Ibid.		46	1.45	1987 ^b	Female farmers
Yeon et al.	[29]	160	1.6	1992 ^b	Men
Ibid.		243	1.4	1992 ^b	Women
Ibid.		403	1.5	1992 ^b	Men + women
Moon et al.	[18]	18	1.30	1994	Seoul
Ibid.		38	1.40	1994	Pusam
Ibid.		17	1.22	1994	Chunan
Ibid.		34	1.51	1994	Haman
Ibid.		107	1.39	1994	Total
Ohn et al.	[30]	35	1.82	1995	Rural residents; men
Ibid.		67	1.97	1995	Rural residents; women
Ibid.		102	1.92	1995	Rural residents; men $+$ women
Moon et al.	[19]	38	1.51	2000	Children
Ibid.		38	2.74	2000	Mothers
Kim et al.	[31]	66	1.21 ^a	2001	Seoul; men
Ibid.		78	1.66 ^a	2001	Seoul; women
Ibid.		144	1.43 ^a	2001	Seoul: men + women
Moon et al.	[33]	47	1.70	2007-2008	Coastal areas
Ibid.		53	1.72	2007-2008	Inland areas
Ibid.		100	1.71	2007-2008	Total
Son et al.; MHWFAK ^c	[32, 34]	800	0.92	2007-2008	Data from 2nd KNHEBE ^d ; men
Ibid.		1546	1.08	2007-2008	Data from 2nd KNHEBE ^d ; women
Ibid.		2346	1.02	2007-2008	Data from 2nd KNHEBE ^{d} ; men + women
Kim et al.	[35]	795	0.89 ^a	2008	Data from KNHANES ^e IV; men
Ibid.		914	1.02 ^a	2008	Data from KNHANES ^e IV women
Ibid.		1709	0.96 ^a	2008	Data from KNHANES ^e IV; men + women
Kim and Lee	[36]	997	1.49	2009	Data from KNHANES ^e III; men
Ibid.		1000	1.57	2009	Data from KNHANES ^e III;women
Ibid.		1997	1.53	2009	Data from KNHANES ^e III; men + women
Park and Lee	[37]	2127	0.753	2008-2010	Data from KNHANES ^e ; men
Ibid.		2395	1.085	2008-2010	Data from KNHANES ^e ;women
Ibid.		4523	0.919	2008-2010	Data from KNHANES ^e ; men + women
Hong et al.	[38]	776	0.92	2010	Data from KNHANES ^e ; men
Ibid.		1486	1.08	2010	Data from KNHANES ^e ;women
Ibid.		2262	1.02	2010	Data from KNHANES ^e ; men + women
Huang et al.	[24]	239	1.29	2011 ^b	\geq 20 year-old men
Ibid.	_ 4	404	0.82	2011 ^b	≥ 20 year-old women
Ibid.		643	1.22	2011 ^b	≥ 20 year-old men + women
Kim et al.	[39]	1669	1.01	2011	Adults
Kim et al.	[35]	795	0.89	2011 ^b	National average; men
Ibid.	_ 4	914	1.72	2011 ^b	National average; women
Ibid.		1709	1.02	2011 ^b	National average; men + women

Table 2 continued

References	Ref. no.	No. of cases studied	Cd-B (µg/L) (GM)	Year of survey	Note
Ha et al.	[40]	199	0.32	2011-2012	Data from KEHSCA ^f (6–19 years); men
Ibid.		152	0.29	2011-2012	Data from KEHSCA ^f (6–19 years); women
Ibid.		351	0.30	2011-2012	Data from KEHSCA ^f (6–19 years); men + women
Yang et al.	[15]	457	0.86	2015 ^b	0-4 year-old children

^a Originally the AM and ASD (or ASE) were given, from which GM was estimated by the moment method [12]

^b Year of publication in place of the survey year

^c MHWAK Ministry of Health, Welfare and Family Affaires, Korea

^d *KNHEBE* Korean National Human Exposure and Bio-monitoring Examination

e KNHANES Korean National Health and Nutrition Examination Survey

^f KEHSCA Korean Environmental Health Survey in Children and Adolescents

and slopes were taken down to several digits below the decimal points. This was necessary because the multipliers were in the order of thousand (i.e., 2000 and 2015).

Results

Data retrieved from 1980 to 2015 were summarized in three tables, one each for Cd-D (Table 1), Cd-B (Table 2) and Cd-U_{cr} (Table 3). The scale of studies were various when expressed by number of subjects studied, i.e., from less than 20 to nearly 5000. In some Cd-D studies, Cd-D was estimated taking advantage of e.g., national data on food consumption and the number of cases studied was not available (shown as NA in Table 1).

Time trend in Cd-D

Search for publication on Cd-D gave 9 articles including 3 Cd-D estimation reports [1, 11, 20]. The surveys were conducted in 10 individual regions together with one whole country survey. The articles are summarized in Table 1, in which GM Cd-D values were given both on per day and per kg body weight per day bases, together with years of survey and the number of cases studied. The first report was published in 1980 [17] and the latest one was in 2011 [24].

When the data collected were listed in the order of year of survey (Table 1), it became clear that no data were available for about 15 years from 1980 till 1994 (the first reported value was an estimate). Nevertheless, the two parameters of survey year and Cd-D closely related to each other (Fig. 1) when the correlation was examined for 1980 to 2011. The regression analysis showed a negative (i.e., <0) slope indicating a decreasing trend as a function of time (see Eq. 1 in Table 4 for the equation of the regression line). The correlation coefficient (r = -0.586) was significant (p < 0.01)

Time trend in Cd-B

In total, 21 articles were available on Cd-B (Table 2). Many of the cited articles reported Cd-B on more than one group of subjects, and accordingly Cd-B in 38 groups of people in non-polluted areas were available after exclusion of duplication in reporting such as men and women separately and also in combination.

The data were reported first in 1985 [25] and the reporting continued till present [15]. Perusal of data following the flow in time disclosed that there was an exceptionally high value of 5.8 µg/L reported in late 1980s [27] (Table 2), but the general trend of decrease was observable till recent years. The slope of the calculated regression line (Fig. 2) was negative and the correlation coefficient, r = -0.575, was statistically significant (p < 0.01) (Eq. 3 in Table 4) When the Cd-B of 5.8 µg/L reported in 1987 [27] (Table 2) was taken as an outlier to be excluded (p < 0.01 by Smirnov' test for extreme value), recalculation excluding the case gave an substantially greater absolute value for r of 10.6251 (with a negative slope of -0.0369 µg/L/year) to show closer clustering around the regression line although the slope became less steep (Eq. 4 in Table 4).

Time trend in Cd-U_{cr}

Search for Cd-U_{cr} was also rewarding, and 14 articles reporting Cd-U_{cr} values were identified. After exclusion of duplication cases (as described above), 28 groups with Cd-U_{cr} values were available (Table 3). Reading through the data following the time suggested that the values in excess of 2 μ g/g cr were found mostly in earlier years (e.g., before 2000) and typically in coastal areas such as Haman (e.g., [18, 33]). Reversely, Cd-U_{cr} was often around 1 μ g/g cr or below in the years after 2010. A regression analysis showed a time-dependent decrease also for Cd-U_{cr} (Eq. 5 in Table 4). Scattering around the regression line was however rather

References	Ref. No.	No. of cases studied	$Cd\text{-}U_{cr} \; (\mu\text{g/g cr}) \; (GM)$	Year of survey	Note
Hwang et al.	[27]	72	2.5 ^a	1987 ^b	Women
Lee and Kim	[41]	77	1.99	1992 ^b	No correction for CR
Yeon et al.	[29]	160	1.19	1992 ^b	Men
Ibid.		243	1.87	1992 ^b	Women
Ibid.		403	1.60	1992 ^b	Men + women
Moon et al.	[18]	18	2.25	1994	Seoul
Ibid.		38	2.16	1994	Pusam
Ibid.		17	1.40	1994	Chunan
Ibid.		34	3.05	1994	Haman
Ibid.		107	2.26	1994	Total
Lee et al.	[42]	426	0.74	1997	School children
Ibid.		250	1.29	1999	School children
Ibid.		249	1.48	2000	School children
Moon et al.	[19]	38	1.69	2000	Children
Ibid.		38	1.56	2000	Mothers
Watanabe et al.	[21]	33	2.54	2003-2004	Kindergarten children in Seoul
Ibid.		37	0.99	2003-2004	Kindergarten children in Jeju City
Ibid.		18	1.01	2003-2004	Kindergarten children in Jeju village A
Ibid.		20	0.88	2003-2004	Kindergarten children in Jeju village B
Ibid.		108	1.30	2003-2004	Kindergarten children, total
Chung et al.	[13]	88	0.88	2005 ^b	Control areas
MEK ^c	[43]	126	2.08 ^d	2007	Middle-sized city
Moon et al.	[33]	88	2.59	2007-2008	Coastal areas
Ibid.		180	1.81	2007-2008	Inland areas
Ibid.		268	2.04	2007-2008	Total
Lee et al.	[44]	5087	0.61	2008	KNSEPHB ^e ; men + women
Yun et al.	[14]	73	0.59	2010 ^b	Controls (Village B)
Ibid.		74	0.65	2010 ^b	Controls (Village D)
Ibid.		147	0.62	2010 ^b	Controls (Villages $B + D$)
Huang et al.	[24]	239	1.04	2011 ^b	≥ 20 year-old men
Ibid.		404	0.89	2011 ^b	≥ 20 year-old women
Ibid.		643	0.95	2011 ^b	≥ 20 year-old men + women
Yang et al.	[15]	457	1.40	2015 ^b	Controls

Table 3 Cadmium concentration in urine as corrected for creatinine concentration

^a Originally the AM and ASD (or ASE) were given, from which GM was estimated by the moment method [12]

^b Year of publication

^c MEK Ministry of Environment, Korea

^d μg/L

^e KNSEPHB Korea National Survey for Environmental Pollutants in Human Body in 2008

remarkable (Fig. 3) with r = -0.490, although the coefficient was statistically significant (p < 0.01).

Discussion

The present analyses of published reports on Cd-D in Korea clearly demonstrated that there has been steady reduction since 1980s (Table 1). The results of analyses for

time trends in Cd-B and Cd-U are also in agreement with the observation on Cd-D. It should be added that dietary Cd is an almost exclusive Cd exposure source for general populations [4]. A similar trend of time-depending decrease in Cd-D was reported also for Japan, a neighboring and another rice-depending country in East Asia [45, 46].

To confirm the above-stated observation, several statistical analyses were conducted in parallel. With regard to Table 4 Regression line

equations



Fig. 1 Time trend in cadmium in dietary intake. Each *dot* represents a GM value observed in one group of study subjects. The line in the middle is a calculated regression line (for equation, see Eq. 1 in Table 4), and the *two dotted curves* show the 95 % range

the databases, study years were not always available. Both the years of surveys and the years of publications were reported in 8, 8, and 13 articles on Cd-D, Cd-B and Cd-U_{cr}, respectively. The differences between the years of survey and the years of publication did not distribute normally or log-normally, and medians were 5.5, 5.5 and 3 years in the order. The years of surveys were tentatively estimated by subtracting the median values from the publication years for the reports in which no survey years were given. The survey years thus estimated were subjected (in combination with survey years reported in other publications) to regression analyses between survey years and Cd-exposure parameters (i.e., Cd-D, Cd-B or Cd-U_{cr}) (Case B in Table 4). The parameters (including slopes) of the regression lines thus obtained (Eqs. 6 to 10) did not differ

Y	Unit ^a	Equation no.	n ^b	Intercept (α)	Slope (β)	r	P for r
Case A ^c	(group-based an	alysis)					
Cd-D	µg/kg BW/day						
Total ca	ases	1	23	38.140	-0.0189	-0.586	p < 0.01
Selected	d cases ^f	2	18	21.025	-0.0103	-0.329	p > 0.05
Cd-B	μg/L	3	51	105.139	-0.0518	-0.575	p < 0.01
Cd-B ^g	μg/L	4	50	75.207	-0.0369	-0.625	p < 0.01
Cd-U _{cr}	μg/g cr	5	33	90.183	-0.0443	-0.490	p < 0.01
Case B ^d	(group-based an	alysis with adju	stment)				
Cd-D	µg/kg BW/day						
Total ca	ases	6	23	37.688	-0.0187	-0.522	p < 0.05
Selected	d cases ^f	7	18	11.702	-0.0057	-0.152	p > 0.05
Cd-B	μg/L	8	51	101.003	-0.0498	-0.583	p < 0.01
Cd-B ^g	μg/L	9	50	67.782	-0.0332	-0.585	p < 0.01
Cd-U _{cr}	μg/g cr	10	33	79.049	-0.0388	-0.426	p < 0.05
Case C ^e	(number-weight	ed analysis)					
Cd-D	µg/kg BW/day	11	7146	40.435	-0.0200	-0.630	p < 0.01
Cd-B	μg/L	12	35,385	81.492	-0.0400	-0.461	p < 0.01
Cd-B ^g	μg/L	13	35,313	60.503	-0.0296	-0.411	p < 0.01
Cd-U _{cr}	μg/g cr	14	10,260	84.707	-0.0417	-0.484	p < 0.01

In the equation $Y = \alpha + \beta X$, X is the year of survey and Y is the parameter shown in the table

^a Unit for Y

^b Number of groups in Case A and Case B, and number of individuals in Case C

^c Case A: the publication year is taken as the survey year when the survey year is not reported

^d Case B: in absence of report on the survey year, the survey year is estimated by subtracting 5.5, 5.5 and 3.0 years from the publication year in cases of Cd-D, Cd-B and Cd-Ucr, respectively. There is no significant difference (p > 0.05) in intercepts, slopes and correlation coefficients between Eqs. 1 and 6, Eqs. 2 and 7, Eqs. 3 and 8, Eqs. 4 and 9, and Eqs. 5 and 10

^e Case C: calculation was made as in Case A, except that the analysis is made in a number-weighted manner taking the number of individuals into consideration

^f Five studies are not included because the Cd-D is estimated taking advantage of e.g., national statistics of food consumptions and the number of participating subjects are not available

^g After exclusion of one group of extreme cases (p < 0.01 by Smirnov's test for extreme value)



Fig. 2 Time trend in cadmium concentration in peripheral blood. Notes are as under Fig. 1. For the regression line equation, see Eq. 3 in Table 4



Fig. 3 Time trend in cadmium concentration in urine as corrected for creatinine concentration. Notes are as under Fig. 1. For the regression line equation, see Eq. 5 in Table 4

significantly (p > 0.05) from the corresponding values for Eqs. 1 to 5 (Case A, in which the year of survey was taken as a surrogate of the pear of survey when necessary). Thus, the effects of the surrogate use (i.e., the use of publication years in place of survey years as necessary) should be small.

As the number of cases was various depending on the groups from less than 20 to nearly 5000 (Tables 1, 2, 3), number-weighted analyses were conducted in parallel (Case C in Table 4). In Cd-D, case numbers were not available in 2 studies [11, 17] for 5 estimated Cd-D, as stated above. Correspondingly, Case A analysis was conducted also with selected groups (i.e., excluding these studies) (Eq. 2 in Table 4) in addition to the analysis with all groups (Eq. 1) for better comparison with the results from Case C (Eq. 11). Accordingly, regression analysis for Cd-D was conducted also in Case C analysis excluding these studies [11, 17], the studies for which case numbers were missing (Eq. 11 in Table 4). In Cd-B, one study [27] gave extremely high GM Cd-B value and excluded as an outlier group. Thus, Case C analyses for Cd-B were conducted including and also excluding this outlier group (Eqs. 12 and 13).

Perusal of Table 4 revealed that both intercepts and slopes in Case C analyses differed to some extent from the corresponding results of group-based Case A and Case B analyses. Nevertheless, all slopes were negative with p < 0.01-0.05 for correlation coefficients to suggest time-dependent decreases.

There have been several factors which have been contributing to the reduction. Korea Food and Drug Administration set a guideline for Cd in rice at 0.2 mg/kg in 2012, and marketing of rice containing Cd at the concentration in excess of this guideline has been prohibited by law [47]. In addition, there has been a gradual reduction in rice consumption by general public due to changes in food habits. According to the national data, consumption of rice in 2005 and 2013 was 80.7 and 67.2 kg/person/year [48], indicating reduction by about 17 % in these 8 years (rice weight was assumedly for pre-cooked weight). From rice-cultivation technology side, efforts have been made to reduce Cd contents in rice by use of ameliorants, especially in Cdcontaminated rice fields in the vicinity of abandoned metal mines, e.g., [49, 50].

To compare with the past and current levels in Cd-D, Cd-B and Cd- U_{cr} in Korea, literature survey was made for the levels in other countries in 2000–2015 (Table 5). For this purpose, estimation for the levels in Korea in 2000 and 2015 was also presented, taking advantages of regression equations in Case A (Eqs. 1, 3 and 5), Case B (Eqs. 6, 8 and 10) and Case C analyses (Eqs. 11, 12 and 14) given in Table 4. Table 5 shows that regional differences were much more remarkable than time-dependent changes. Thus, it was apparent that Cd exposure has been higher in Asia than in Europe and in USA. Within Asia, all of Cd-D, Cd-B and Cd- U_{cr} were highest for Japan, and Korea is essentially the 2nd-highest, although there were substantial reductions in the estimates of three exposure parameters in 2015 as compared with the estimates in 2000.

In 2013, Joint FAO/WHO Expert Committee on Food Additives (JECFA) adopted 25 μ g/kg body weight/month as the provisional tolerable monthly intake (PTMI) for Cd [65]. Assuming that the average body weight for adult Korean population is 63 kg [16] and that a month consists of 30 days, 25 μ g/kg body weight/month is approximately equivalent to 52.5 μ g/day. Thus, the current estimate for Cd-D, 6.0–7.4 μ g/day (the bottom three rows in Table 5), is about 11–14 % of the PTMI.

There may be several limitations in the present study. The first point is on the instrumental methods for Cd determination. Early time studies used flame atomic absorption spectrometry, whereas later studies employed graphite furnace atomic absorption spectrometry and then inductively coupled plasma mass spectrometry. Compatibility between the last two methods was previously confirmed [60] but no information was available with regard to

Table 5	Selected	references	for	comparison	with	the	present	study
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References	Ref. no.	Year ^a	Study cites	Cd-D ^b (µg/day)	Cd-B ^b (µg/L)	$Cd-U^b_{cr}$ (µg/g cr)	Note
The present study		2000	Korea	25.2 ^{c,f}	1.55 ^c	1.58 ^c	Estimates ^c
Ibid.		Ibid.	Ibid.	24.5 ^{d,f}	1.48 ^d	1.53 ^d	Estimates ^d
Ibid.		Ibid.	Ibid.	24.9 ^{e,f}	1.43 ^e	1.23 ^e	Estimates ^e
Ikeda et al.	[4]	2000	South-east Asia	9.4	0.58	1.42	4 large cities
Ibid.		Ibid.	China	1.3	0.61	2.30	5 large cities in the southern half
Noonan et al.	[51]	2002	USA			0.17	
Jin et al.	[52]	2002	China		1.41	1.83	Zheijang Province (southern China)
Ezaki et al.	[53]	2003	Japan			1.26	
Jin et al.	[54]	2004	China		1.53	1.81	Zheijang Province (southern China)
Wang and Tian	[55]	2004	China			0.26 ^g	Fujian Province (southern China)
Åkesson et al.	[56]	2005	Sweeden			0.67 ^h	
Heitland and Köster	[57]	2006	Germany		0.38		
Coelho et al.	[58]	2007	Portugal		0.59 ^g		
McKelvey et al.	[59]	2007	USA		0.77		
Ikeda et al.	[60]	2011	Japan		1.23		
Morton et al.	[61]	2014	UK			0.17 ⁱ	
Ikeda et al.	[62]	2015	Japan	16.5			The latest estimate
Den Hond et al.	[63]	2015	Europe			$0.132 - 0.306^{j}$	Mothers
Mørck et al.	[64]	2015	Denmark			0.118	Mothers
The present study		2015	Korea	7.4 ^{c,f}	0.77 ^c	0.92 ^c	Estimates ^c
Ibid.		Ibid.	Ibid.	6.8 ^{d,f}	0.73 ^d	0.95 ^d	Estimates ^d
Ibid.		Ibid.	Ibid.	6.0 ^{e,f}	0.83 ^e	0.60 ^e	Estimates ^e

^a Year of publication

^b Geometric means unless otherwise specified

^c Estimates taking advantake of Eqs. 1, 3 and 5 in Table 4

^d Estimates taking advantake of Eqs. 6, 8 and 10 in Table 4

^e Estimates taking advantake of Eqs. 11, 12 and 14 in Table 4

^f An average body weight of 63 kg [16] is assumed for adult men and women in Korea

^g GM was clculated from AM and ASD by use of the moment method [12]

^h Median in μg/L

i Median in µg/g cr

^j The minimum and the maximum of GMs for 17 countries in Europe

the first method and other two methods. Secondly, only body weight was taken into consideration in evaluating the Cd-D values across men, women and children, despite the fact that children need energy (in terms of foods) not only for their daily life but for growth. It is also known through experiences that men used to take more rice (and therefore more Cd) than women. On this point, it should be added that preliminary analyses with adult populations only gave essentially the same decreasing trends in Cd-D, Cd-B and Cd-U_{cr} with the present analyses in which adult people and children were taken together (data not shown). Thirdly, a well-known confounding factor of smoking as a non-occupational source of Cd burden [6] was not taken into account. The original objectives of most studies cited were elucidation of Cd burdens on general Korean populations, and studies on selective evaluation of non-smokers were very limited. The fourth point is on the comparability of Cd-B and Cd-U_{cr} between adults and children. There is no confirming report if Cd-D intake at the same level gives comparable Cd-B both for adults and for children. The same point may need to be taken as a study limit when Cd-U_{cr} values for adults and children were taken together in trend analyses. It is known that aged people have higher Cd-U_{cr} possibly because Cd has accumulated in the kidney cortex in reflection of years in life [66]. Similarly but to the opposite direction, children may have lower Cd-U_{cr} because of shorter time for Cd accumulation. Therefore, it is one of the limitations of the present study that the data on Cd-D, Cd-B and Cd-U_{cr} for adults and children were intermingled.

In conclusion, literature survey for time trend in Cd-D made it clear that there has been a gradual decrease in the intake to the estimated current level of 6.0–7.4 µg/day or 0.09–0.12 µg/kg body weight/day (with assumption of 63 kg for body weight [16]), although Cd intake among general populations in Korea was once reported to be high in the past. Results of statistical analyses for trends in other two exposure parameters of Cd-B and Cd-U (as corrected for creatinine level) were in agreement with the above conclusion on Cd-D. The corresponding estimates for two exposure markers are 0.77–0.83 µg/L for Cd-B and 0.60–0.95 µg/g cr for Cd-U_{cr}.

Compliance with ethical standards

Conflicts of interest The authors declare that they have no conflicts of interest.

References

- Watanabe T, Nakatsuka H, Ikeda M. Cadmium and lead contents in rice available in various areas of Asia. Sci Total Environ. 1989;80:175–84.
- Watanabe T, Shimbo S, Moon C-S, Zhang Z-W, Ikeda M. Cadmium contents in rice samples from various areas in the world. Sci Total Environ. 1996;184:191–6.
- Acheson KJ, Campbell IT, Edholm OC, Miller DF, Stock MJ. The measurement of food and energy intake in man—an evaluation of some techniques. Am J Clin Nutr. 1990;33:147–54.
- Ikeda M, Zhang Z-W, Shimbo S, Watanabe T, Nakatsuka H, Moon C-S, et al. Urban population exposure to lead and cadmium in east and south-east Asia. Sci Total Environ. 2000;249:373–84.
- International Programme on Chemical Safety (IPCS). Environmental Health Criteria I34. Cadmium. Geneva: World Health Organization, 1992.
- International Programme on Chemical Safety (IPCS). Environmental Health Criteria 135. Cadmium—environmental aspects. Geneva: World Health Organization, 1992.
- Agency for Toxic Substance and Disease Registry (ATSDR), USA. Toxicological Profile for Cadmium 3.2.2.2 Health Effect-Renal effects. pp. 147–67, 2012.
- Kweon S, Kim Y, M-j Jang, Kim Y, Kim K, Choi S, et al. Data resource profile: The Korea National Health and Nutrition Examination Survey (KNHANES). Int J Epidemiol. 2014;43:69–77.
- Moon C-S, Zhang Z-W, Shimbo S, Watanabe T, Moon D-H, Lee C-U, et al. Dietary intake of cadmium and lead among the general population in Korea. Environ Res. 1995;71:46–54.
- Kim M, Wolt JD. Probabilistic risk assessment of dietary cadmium in the south Korean population. Food Addit Contam Part A Chem Anal Control Expo Risk Assess. 2011;28:62–70.
- Moon C-S, Lee CK, Lee JT, Kim JM, Paik J-M, Ikeda M. Time trends in dietary cadmium intake of Korean women. Toxicol Res. 2012;1:145–50.
- Sugita M, Tsuchiya K. Estimation of variation among individuals of biological half-times of cadmium calculated from accumulation data. Environ Res. 1995;68:31–7.

- Chung JH, Kang PS, Kim CY, Lee KS, Hwang TY, Kim GT, et al. Blood Pb, urinary Cd and health assessment of residents in the vicinity of abandoned mine in Gyeongsangbuk-do. Kor J Occup Environ Med. 2005;17:225–37 (in Korean with an English abstract).
- 14. Yun SH, Kim CY, Hwang TY, Won KC, Do JY, Lee SJ, et al. The concentration of cadmium in urine, and its role in health-risk. I Assessment of residents in the vicinity of abandoned mines in Gyeongsangbuk-do, Korea. Kor. J Occup Environ Med. 2010;22:351–61 (in Korean with an English abstract).
- Yang J, Kim E-G, Shin D-C, Jo S-J, Lim Y-W. Human exposure and risk assessment of cadmium for residents of abandoned metal mines in Korea. Environ Geochem Health. 2015;37:321–32.
- Korea Research Institute of Standard and Science. The 5th report on 'Size Korea' (2004) http://sizekorea.kats.go.kr/03_report/5th. asp (downloaded on 1 September, 2015).
- Yum Y-T, Bae E-S, Yun B-J. A study on the crops pollution with heavy metal. Kor J Prev Med. 1980;13:3–12 (in Korean with an English abstract).
- Moon C-S, Zhang Z-W, Shimbo S, Watanabe T, Lee C-U, Lee B-K, et al. Evaluation of urinary cadmium and lead as markers of background exposure of middle-aged women in Korea; dietary intake as an influential factor. Toxicol Lett. 1999;108:123–8.
- Moon C-S, Paik J-M, Choi C-S, Kim D-H, Ikeda M. Lead and cadmium levels in daily foods, blood and urine in children and their mothers in Korea. Int Arch Occup Environ Health. 2003;76:282–8.
- Oh E, Lee E-I, Lim H, Jang J-Y. Human multi-route exposure assessment for Korean volunteers. J Prev Med Public Health. 2006;39:53–8 (in Korean with an English abstract).
- Watanabe T, Kim ES, Yang HR, Moon SH, Nakatsuka H, Shimbo S, et al. Food intake survey of kindergarten children in Korea: Part 3 cadmium and lead burden. Environ Health Prev Med. 2015;20:307–13.
- Kim DW, Woo HD, Joo J, Park KS, Oh SY, Kwon H, et al. Estimated long-term dietary exposure to lead, cadmium, and mercury in young Korean children. Eur J Clin Nutr. 2014;68:1322–6.
- Lim J-A, Kwon H-J, Ha M, Kim H, Oh SY, Kim JS, et al. Korean research project on the integrated exposure assessment of hazardous substances for food safety. Environ Health Toxicol. 2015;30 (Article ID: e2015004).
- Huang M, Choi S-J, Kim D-W, Kim U-Y, Bae S-H, Yu S-D, et al. Evaluation of factors associated with cadmium exposure and kidney function in the general population. Environ Toxicol. 2011;8:563–70.
- 25. Lee S-S, Kim D-H. A comparison of gases and heavy metals in blood between urban and rural teenager. Kor J Prev Med. 1985;18:129–36 (in Korean with an English abstract).
- Watanabe T, Nakatsuka H, Seiji K, Inoue O, Cho K-S, Lee K-M, et al. Blood cadmium levels in populations of Masan, Korea, and Miyagi, Japan; an inter-regional comparison. Tox Lett. 1989;47:155–63.
- 27. Hwang ID, Ki NS, Lee JH, Park IS. A study on the heavy metal concentrations and their interrelationships in women's blood and urine in small towns. Kor J Prev Med. 1987;20:49–55 (in Korean with an English abstract).
- Watanabe T, Cha CW, Song DB, Ikeda M. Pb and Cd levels among Korean populations. Bull Environ Contam Toxicol. 1987;38:189–95.
- Yeon YY, Ahn KD, Lee BK. Blood and urine cadmium levels in non-exposed Korean to cadmium. Kor J Occup Environ Med. 1992;4:70–80 (in Korean with an English abstract).
- Ohn YH, Park JD, Choi BS, Hong YP, Chang JW. Blood cadmium and zinc and urinary N-acetyl-β-D-glucosaminidase

activity in rural residents not exposed to cadmium. Chung-Ang J Med. 1995;20:333–50 (in Korean with an English abstract).

- Kim H-H, Lim Y-W, Yang J-Y, Ho M-K, Shim D-C. Distribution of inorganic metals in blood of adults in urban area of Seoul, Korea. J Environ Toxicol. 2004;19:327–34 (in English with Korean abstract).
- Ministry of Health, Welfare and Family Affaires, Korea, 2007–2008. Korea National Health and Nutrition Examination Survey, 2007–2008 (in Korean).
- Moon C-S, Lee CK, Hong YS, Ikeda M. Higher cadmium burden in coastal areas than in inland areas in Korea; implications for seafood intake. Asia Pacific J Clin Nutr. 2014;23:219–24.
- 34. Son I-Y, Lee J, Park D, Lee J-T. Blood levels of lead, cadmium, and mercury in the Korean population: results from the Second Korean national human exposure and bio-monitoring examination. Environ Res. 2009;109:738–44.
- 35. Kim YA, Kim Y-N, Cho K-D, Kim MY, Kim EJ, Baek O-H, et al. Blood heavy metal concentrations of Korean adults by seafood consumption: using the fourth Korea National Health and Nutrition Examination Survey (KNHANESIV), 2008. Kor J Nutr. 2011;44:518–26 (in Korean with an English abstract).
- Kim N-S, Lee B-K. National estimates of blood lead, cadmium, and mercury levels in the Korean general adult population. Int Arch Occup Environ Health. 2011;84:53–63.
- 37. Park S, Lee B-K. Body fat percentage and hemoglobin levels are related to blood lead, cadmium, and mercury concentrations in Korean adult population (KNHANES 2008–2010). Biol Trace Elem Res. 2013;151:315–23.
- Hong S-Y, Lee B-K, Park J-D, Sakong J, Choi J-W, Moon J-D, et al. Blood cadmium concentration of residents living near abandoned metal mines in Korea. J Kor Med Sci. 2014;29:633–9.
- 39. Kim NH, Hyun YY, Lee K-B, Chang Y, Rhu S, Oh K-H, et al. Environmental heavy metal exposure and chronic kidney disease in the general population. J Kor Med Sci. 2015;30:272–7.
- 40. Ha M, Kwon H-J, Leem J-H, Kim H-C, Lee KJ, Park I, Lim Y-W, et al. Korean Environmental health survey in children and adolescents (KorEHS-C) survey design and pilot study results on selected exposure biomarkers. Int J Hyg Environ Health. 2014;17:260–70.
- Lee DH, Kim DH. Relationship of low blood cadmium and zinc to blood pressure. Kor J Prev Med. 1992;25:148–56 (in Korean with an English abstract).
- 42. Lee CR, Yoo CI, Lee JH, Lee H, Kim Y. Trend of the changes in levels of blood lead, urinary arsenic and urinary cadmium of children in Ulsan: 3-year follow-up study. Kor J Prev Med. 2001;34:166–74 (in Korean with an English abstract).
- 43. Ministry of Environment, Korea. Exposure level and biological monitoring of environmental hazards in residents (Gwangyang area), 2007 (**in Korean**).
- 44. Lee JW, Lee CK, Moon CS, Choi IJ, Lee KJ, Choi SM, et al. Korea national survey for environmental pollutants in human body 2008: heavy metals in the bold or urine of the Korean population. Int J Hyg Environ Health. 2012;215:449–57.
- 45. Ikeda M, Ezaki T, Tsukahara T, Moriguchi J. Dietary cadmium intake in polluted and non-polluted areas in Japan in the past and in the present. Int Arch Occup Environ Health. 2004;77:227–34.
- 46. Ikeda M, Watanabe T, Nakatsuka H, Moriguchi J, Sakuragi S, Ohashi F, et al. Cadmium exposure in general populations Japan: A review. Food Safety. 2015; 3:18–35.
- Korea Food and Drug Administration. Food code for Cd in rice. http://www.foodnara.go.kr/pollution/standard/pollution/pollution01. jsp (Cd standard in rice; in Korean). 2012 (downloaded on 1 September, 2015).
- Korean Agency of Statistics. Consumption of main agricultural products (kg/person/year); rice. 2014. http://www.index.go.kr/

potal/main/EachDtlPageDetail.do?idx_cd=2747. Downloaded on 1 Sept 2015 (in Korean).

- 49. Lee S-H, Lee J-S, Choi Y-J, Kim J-G. In situ stabilization of cadmium-, lead-, and zinc-contaminated soil using various amendments. Chemosphere. 2009;77:1069–75.
- Ok YS, Kim S-C, Kim D-K, Skousen JG, Lee J-S, Cheong Y-W, et al. Ameliorants to immobilize Cd in rice paddy soils contaminated by abandoned metal mines in Korea. Environ Geochem Health. 2011;33:23–30.
- Noonan CW, Sarasua SM, Compagna D, Kathman SJ, Lybarger JA, Mueller PW. Effects of exposure to low levels of environmental cadmium on renal biomarkers. Environ Health Perspect. 2002;110:151–5.
- 52. Jin T, Nordberg M, Frech W, Dumon X, Bernard A, T-t Ye, et al. Cadmium biomonitoring and renal dysfunction among a population environmentally exposed to cadmium from smeltering in China (ChinaCad). Biometals. 2002;15:397–410.
- 53. Ezaki T, Tsukahara T, Moriguchi J, Furuki K, Fukui Y, Ukai H, et al. No clear-cut evidence for cadmium-induced tubular dysfunction among over 10,000 women in the Japanese general population; a nationwide large-scale survey. Int Arch Occup Environ Health. 2003;76:186–96.
- Jin T, Kong Q, Ye T, Wu X, Nordberg GF. Renal dysfunction of cadmium-exposed workers residing In a cadmium-polluted environment. Biometals. 2004;17:513–8.
- 55. Wang X, Tian J. Health risk related to residential exposure to cadmium in Zhenhe County, China. Arch Environ Health. 2004;59:324–30.
- 56. Åkesson A, Lundh T, Vahter M, Biellerup P, Lidfedt J, Nerbrand C, et al. Tubular and glomerular kidney effects in Swedish women with low environmental cadmium exposure. Environ Health Perspect. 2005;113:1627–31.
- Heitland P, Köster HD. Biomonitoring of 37 trace elements in blood samples from inhabitants of northern Germany by ICP-MS. Trace Elem Med Biol. 2006;20:253–62.
- Coelho P, Silva S, Roma-Torres J, Costa C, Henriques A, Teixeira J, et al. Health impact of living near an abandoned mine case study: Jales mines. Int J Hyg Environ Health. 2007;210:399–402.
- McKelvey W, Gwynn C, Jeffery N, Kass D, Thorpe LE, Garg RK, et al. A biomonitoring study of lead, cadmium, and mercury in blood of New York city adults. Environ Health Perspect. 2007;115:1435–41.
- 60. Ikeda M, Ohashi F, Fukui Y, Sakuragi S, Moriguchi J. Cadmium, chromium, lead, manganese and nickel concentrations in blood of women in non-polluted areas in Japan, as determined by inductively coupled plasma-sector field-mass spectrometry. Int Arch Occup Environ Health. 2011;84:139–50.
- Morton J, Tan E, Leese E, Cocker J. Determination of 61 elements in samples collected from a non-occupationally exposed UK adult population. Toxicol Lett. 2014;231:179–93.
- Ikeda M, Nakatsuka H, Watanabe T, Shimbo S. Estimation of daily cadmium intake from cadmium in blood or cadmium in urine. Env Health Prev Med. 2015;20:455–59. doi:10.1007/ s12199-015-0479-x.
- 63. Den Hond E, Govarts E, Willems H, Smolders R, Gastelwyn L, Kolossa-Gehring M, et al. First steps toward harmonized human biomonitoring in Europe: demonstration project to perform human biomonitoring in a European scale. Environ Health Perspect. 2015;123:255–63.
- 64. Mørck TA, Nielsen F, Nielsen KSJ, Jensen JF, Hansen PW, Hansen AK, et al. The Danish contribution to the European DEMOCHPHES project: a description of cadmium, cotinine and mercury levels in Danish mother-child pairs and the perspectives of supplementary sampling and measurements. Environ Res. 2015;141:96–105.

- 65. Joint FAO/WHO Expert Committee on Food Additives (JECFA). Evaluation of the JECFA: Cadmium. 2013. http://apps.who.int/ food-additives-contaminants-jecfa-database/chemical.aspx?chem ID=1376 (Downloaded on 2 July 2015).
- 66. Moriguchi J, Ezaki T, Tsukahara T, Furuki K, Fukui Y, Okamoto S, et al. Comparative evaluation of four urinary tubular dysfunction markers, with special references to the effects of aging and correction for creatinine concentration. Toxicol Lett. 2003;143:279–90.