

Estimated daily intake of biotin in hospital meals in Japan through dietary surveillance

Toshiaki WATANABE^{†, 1)}, Atsushi FUKUSHIMA¹⁾, Kimi MATSUMOTO¹⁾, Hiromi SAWAMURA²⁾ and Hidetaka MIZOHATA³⁾

¹⁾ *Department of Dietary Environment Analysis, School of Human Science and Environment, Himeji Institute of Technology, University of Hyogo**,

²⁾ *Department of Molecular Nutrition, School of Human Science and Environment, University of Hyogo***,

³⁾ *Department of Life Sciences, School of Human Sciences, Kobe Shoin Women's University****

Summary

We determined the biotin content in hospital meals using different dietary surveys such as “food group calculation survey” and “duplicate meals survey,” to estimate the dietary intake of biotin in patients. As for hospital meals, the daily average biotin contents for 10 days calculated by the food group calculation survey were 19.4 µg/day in meals for infants, 45.5 µg/day in meals for children, 52.8 µg/day in meals for pregnant/lactating women, and 56.7 µg/day in common meals for adults. The average contents of biotin for 10 days by the duplicate meals survey were 9.7 µg/day in meals for infants, 41.6 µg/day in meals for children, 34.2 µg/day in meals for pregnant/lactating women and 41.9 µg/day in common meals for adults. The common meals contained basic and standard cooking ingredients and balanced with Dietary Reference Intake (DRI) for adult patients. The average intake of biotin in common meals for adults in the hospital exceeded the Adequate Intake (AI) (50 µg/day) for adults via the food group calculation survey, but was not sufficient in meals for adults and pregnant/lactating women by the duplicate meals survey. It is necessary to be careful about the kinds of foods and the method of cooking the ingredients when the biotin intake for hospital patients is calculated using the hospital menu sheet.

Introduction

Biotin is one of the water-soluble vitamins which are contained relatively rich in the liver and the egg yolk¹⁾. This vitamin is associated with gluconeogenesis, amino acid metabolism, synthesis of fatty acid and energy production as a coenzyme of carboxylases *in vivo*²⁾. Biotin deficiency is rare in humans because biotin is widely distributed in various foods and is synthesized by the intestinal microflora. However, clinical signs such as dermatitis and hair loss appear when biotin becomes deficient. In the experimental animals, we found that the biotin deficiency during gestation induced congenital anomalies such as cleft palate and micrognathia in fetuses^{3,5)}. Also, urinary biotin decreases in the later stages of gestation in pregnant women and mice⁶⁾. It was recently reported that biotin deficiency is related to atopic dermatitis⁷⁾ and diabetes⁸⁾, and the organic acid, 3-hydroxyisovaleric acid, is a

sensitive biomarker of biotin deficiency⁹⁾.

In Japan, the recommended intake of biotin was set to 30 µg/day in “Recommended Dietary Allowances-Dietary Reference Intakes (DRIs)-” in 2000¹⁰⁾. Afterwards, the Adequate Intake (AI) of biotin for adults was set to 45 µg/day and 50 µg/day in “Dietary Reference Intakes for Japanese, 2005 version” and “2010 version,” respectively¹¹⁻¹³⁾. On the other hand, the use of biotin for Foods with Nutrient Function Claims (FNFC) was authorized in 2003, and as a health claim, biotin was regulated as a nutrient which helps maintain healthy skin and mucosa¹⁴⁾. The upper and lower limits are 500 and 10 µg/day, respectively. However, this vitamin is not yet permitted as an additive in any food. As biotin was not listed in the Standard Tables of Food Composition in Japan, 5th revised edition (Japanese Food Composition Table) until 2005¹⁵⁾, the biotin intake from meals or the requirement of biotin could not be estimated by diet surveys using the food composition table.

*. ** Address: 1-1-12 Shinzaike Honcho, Himeji, Hyogo 670-0092, Japan

*** Address: 1-2-1 Shinohara Obanoyamacho, Nada, Kobe, Hyogo 657-0015, Japan

[†] To whom correspondence should be addressed.

Tel.: +81-79-292-9325; Fax: +81-79-292-9325.

E-mail address: watanabe@shse.u-hyogo.ac.jp (T. Watanabe)

However, the biotin contents of limited foods (only 498 foods) were recently added to the Japanese Food Composition Table in 2011¹⁶⁾.

The AI of biotin for healthy adults was set to 30 µg/day in the DRIs in 1998 in the U.S.A., which is different from that in Japan¹⁷⁾. There are not many studies on the dietary intake of biotin in each country and a confirmed conclusion has not been obtained as yet. Also, evidence is limited on the nutritional state of micronutrients in hospital patients as well. The nutritional state frequently has serious problems in hospital and aged care settings. In hospitals, biotin deficiency is often induced by using total parenteral nutrition¹⁸⁾. Biotin deficiency is induced in infants fed with special therapeutic infant formulas that are only in Japan¹⁹⁾.

Parenteral nutrition-associated biotin deficiency has not been reported since the vitamin was routinely added to parenteral nutrition²⁰⁾. However, marginal biotin status associated with diffuse hair loss could still be prevalent^{21, 22)}. Thus, it is not well-known whether the intake of biotin is filled to maintain the requirement for patients. Also the biotin content of all foods is not listed in Japanese Food Composition Table as yet. Therefore, we determined the biotin content in hospital meals using different dietary surveys such as “food group calculation survey” and “duplicate meals survey,” to estimate the dietary intake of biotin in patients.

Materials and Methods

Meal sampling

Meals for patients in a hospital located in Himeji, Japan were nutritionally surveyed. Hospitals cater various kinds of meals for infants (aged 7 through 12 months), children, pregnant/lactating women, and adults. These meals were collected at breakfast, lunch and dinner, separately, for 10 days. A snack was also included in meals for children and pregnant/lactating women. The meal sampling periods were divided into two periods, from July 30 to August 10 and from August 21 to September 3, 2007. Total 120 meal samples were pretreated in accordance with the procedure manual set out in the Japanese Food Composition Table. After weighing meals and foods, all samples were homogenized into every meal and frozen at -40°C until shortly before use.

The biotin content of 500 foods and royal jelly was already determined in our previous studies and published on our website in English and Japanese²³⁾. The average content of biotin was calculated in 98 food groups according to the food categorization used in the Nation Health

and Nutrition Survey (NHNS) in Japan (group average of biotin content)²⁴⁾. The foods determined were selected by considering the characteristics of their food groups and our dietary habits among the 1,878 foods listed in the Japanese Food Composition Table (2011)¹⁶⁾. Also, the biotin content of 498 foods which are different from our 500 foods was newly listed in the Japanese Food Composition Table in 2011¹⁶⁾.

Food group calculation survey

Food group calculation is one of the diet surveys. This survey can be easily estimated, even the dietary intake of nutrients which are not listed in the food composition table, using a nutrient content of the limited foods determined. A dietary intake of biotin can be estimated using both the group average of biotin content and the food amount taken in food groups. So, based on the group averages of biotin content and the amount of the cooking ingredient classified in 98 food groups on the hospital menu sheet, daily intakes of biotin were estimated from the hospital meals for each day. As a comparative study, using the amount of foods taken in 98 food groups reported in the NHNS in Japan, 2011²⁴⁾, the daily intake of biotin for Japanese adults (males and females) was estimated from common meals for adults.

Duplicate meals survey

The duplicate meals survey can also calculate the intake of micronutrients which are not listed in the Japanese Food Composition Table. Four different kinds of hospital meals for infants, children, pregnant/lactating women and adults were obtained for 10 consecutive days. The dietary intake of biotin was estimated by determining the content of biotin in these hospital meals. It was compared to the food group calculation surveys shown above.

Biochemical analysis

The biotin content in the meals was determined by the microbiological assay using *Lactobacillus plantarum* ATCC 8014. Biotin contained in meals and foods is generally in parts in protein-bound form. Therefore, protein-bound biotin was hydrolyzed before biotin determination. 100 µL of each sample were transferred into microtubes and 100 µL of 2.25 M H₂SO₄ were added. After samples were hydrolyzed by autoclaving for 60 min at 121 °C (2 atoms), they were neutralized by adding approximately 100 µL of 4.5 M NaOH. The supernatant was used as a sample for biotin determination after centrifuging at 10,000 rpm for 5 min. The biotin content was expressed as µg/100 g.

Table 1 Estimation of biotin intake by the food group calculation survey using 98 food groups

Large classification ^a	Food group classification		Biotin contents ^d	Amount of food intake ^e	Biotin intake ^f		
	Middle classification ^b	Small classification ^c			males	females	
Cereals	Rice-processed products	Rice	1.9	381.8	7.4	5.1	
		Rice products	1.7	4.3	0.1	0.1	
	Wheat-processed products	Wheat	1.8	3.8	0.1	0.1	
		Bread	2.4	31.3	0.8	1.1	
		Sweet buns	2.9	6.1	0.2	0.1	
		Japanese noodles	1.2	46.5	0.5	0.5	
		Precooked noodles	1.4	5.8	0.1	0.1	
		Macaroni and spaghetti	2.7	9.3	0.3	0.3	
		Other wheat products	3.2	5.7	0.2	0.1	
	Other cereal-processed products	Buckwheat noodles and their products	2.5	7.9	0.2	0.2	
		Corn and its products	2.5	0.5	0.0	0.0	
Others		4.7	1.3	0.1	0.1		
Potatoes and Starches	Potato-processed products	Sweet potatoes and their products	3.5	5.4	0.2	0.3	
		Potatoes and their products	1.2	29.3	0.3	0.3	
		Others	1.5	17.6	0.3	0.3	
	Starch-processed products	Starches and their products	0.9	2.5	0.0	0.0	
Sugars and Sweeteners	Sugars and Sweeteners	Sugars and Sweeteners	3.1	6.4	0.2	0.2	
Pulses	Soybeans-processed products	Whole soybeans and their products	19.3	1.2	0.2	0.3	
		<i>Tofu</i>	7.5	33.6	2.5	2.1	
		<i>Abura-age</i> ^g	3.9	7.9	0.3	0.3	
		<i>Natto</i>	11.7	5.5	0.6	0.6	
		Other soybean products	8.3	3.7	0.3	0.5	
	Other pulses-processed	Others	9.0	1.3	0.1	0.2	
Nuts and Seeds	Nuts and Seeds	Nuts and Seeds	22.5	2.0	0.5	0.3	
Vegetables	Green and yellow vegetables	Tomatoes	1.7	13.9	0.2	0.2	
		Carrots	1.9	19.4	0.4	0.5	
		Spinach	3.3	16.7	0.6	0.5	
		Sweet peppers	1.3	4.0	0.1	0.0	
		Other green and yellow vegetables	5.3	31.4	1.7	1.9	
	Other vegetables	Cabbages	1.8	23.8	0.4	0.4	
		Cucumbers	1.9	9.8	0.2	0.2	
		<i>Daikon</i> ^h (Japanese radishes)	0.7	30.0	0.2	0.1	
		Onions	1.2	33.4	0.4	0.3	
		Chinese cabbage	2.1	21.2	0.4	0.4	
		Other vegetables	2.6	45.5	1.2	1.0	
		Vegetable juice	Vegetable juice	2.7	10.9	0.3	0.2
		Pickles	Pickles	0.5	3.9	0.0	0.0
	Fruits	Raw fruits	<i>Takuan-zuke</i> ⁱ and others	0.8	8.3	0.1	0.1
Strawberries			2.2	0.1	0.0	0.0	
Citrous fruits			0.7	18.6	0.1	0.2	
Bananas			2.5	14.6	0.4	0.4	
Apples			1.6	16.9	0.3	0.3	
Others			1.3	29.9	0.4	0.6	
Jam		Jam	0.5	1.0	0.0	0.0	
Fruit juices		Juice	0.7	13.6	0.1	0.1	
Mushrooms	Mushrooms	Mushrooms	9.3	15.0	1.4	1.4	
Algae	Algae	Algae	5.1	10.5	0.5	0.6	
Fishes and Shellfishes	Raw fishes and shellfishes	Horse mackerels and sardines	3.9	10.9	0.4	0.4	
		Salmons	8.1	4.2	0.3	0.3	
		Righteye flounders and sea breams	11.3	6.6	0.7	0.2	
		Tunas	2.6	5.4	0.1	0.1	
		Other fishes	3.1	9.2	0.3	0.7	
		Shellfishes	9.1	2.8	0.3	0.2	
		Cuttlefishes	3.9	5.2	0.2	0.1	
		Prawns, shrimps and crabs	2.4	6.0	0.1	0.1	
		Fishes and shellfishes-products	Dried products	10.6	16.2	1.7	1.4
	Canned products		8.0	2.3	0.2	0.1	
	<i>Tsukudani</i> ^j		12.8	0.2	0.0	0.0	
	Fish paste products		1.1	11.3	0.1	0.1	
	Fish hams and sausage		1.8	0.7	0.0	0.0	
	Meats	Animal meats	Beef	1.2	17.0	0.2	0.1
Pork			2.1	36.9	0.8	0.6	
Hams and sausages			3.9	14.8	0.6	0.5	
Other animal meat			—	0.2	—	—	
Poulties		Poulties	2.4	26.4	0.6	0.5	
Other poultries		—	0.1	—	—		
Offal meats	Offal meats	26.1	1.7	0.4	1.4		
Other meats	Whale meats	—	0.0	—	—		
Other meats and their products	—	0.0	—	—			

Eggs	Eggs	Eggs	17.9	37.6	6.7	5.8
Milks	Milk and dairy products	Milks	3.1	88.9	2.7	2.7
		Cheeses	2.6	2.4	0.1	0.1
		Fermented milks and lactobacillus beverages	1.4	20.9	0.3	0.5
		Others	1.7	9.2	0.2	0.1
		Other milks	—	0.3	—	—
Fats and Oils	Fats and Oils	Butters	0.6	1.0	0.0	0.0
		Margarines	0.1	1.1	0.0	0.0
		Vegetable fats and oils	0.0	9.0	0.0	0.0
		Animal fats	0.0	0.2	—	—
		Others	—	0.0	—	—
Confectioneries	Confectioneries	Traditional confectioneries	1.7	9.7	0.2	0.2
		Cakes, buns and pastries	4.9	5.8	0.3	0.4
		Biscuits	1.2	1.4	0.0	0.0
		Candies	1.3	0.3	0.0	0.0
		Others	3.8	5.5	0.2	0.3
Beverages	Alcoholic beverages	<i>Sake</i>	0.0	15.5	—	—
		Beers	1.0	98.0	1.0	0.2
		Others	0.4	52.4	0.2	0.0
	Other beverages	Teas	0.4	276.0	1.2	1.9
		Coffees and cocoas	2.1	123.4	2.6	1.1
Others		0.2	128.4	0.2	0.3	
Seasonings and Spices	Seasonings	Sauces	5.7	2.0	0.1	0.1
		<i>Soyu</i> ^k	12.7	15.8	2.0	1.6
		Edible salts	0.0	1.5	—	—
		Mayonnaise	0.5	3.2	0.0	0.0
		<i>Miso</i> ^l	9.4	10.9	1.0	1.4
		Others	1.5	64.8	1.0	1.2
	Spices	20.9	0.3	0.1	0.1	
Prepared Foods	—	—	13.6	—	—	
Total		all ages			52.0	45.6
		children ages 1-2 years old			27.3	24.0
		children ages 8-9 years old			42.8	41.5
		children ages 10-11 years old			46.6	43.4
		adults ages 18-29 years old			48.7	39.2

^a17 food group

^b33 food group

^c98 food group

^dµg/100 g

^eFor males in The National Health and Nutrition Survey (2011) (g/day)

^fµg/day

^gFried thin slices of pressed *tofu*

^hJapanese radishes

ⁱPickled with rice bran and salt

^jSimmered in soy sauce and sugar

^kSoy sauce

^lFermented soybean

Itaric: Japanese food name (If there are not English name)

Statistical analysis

Values are mean \pm SD. The Spearman's rank correlation test was used to compare the relationship in dietary intake of biotin between the food group calculation survey using the group average of biotin content and the duplicate meals survey. Statistical analysis of these data was performed using Statcel 2 (OMS Publishing, Saitama, Japan). Differences were considered significant as a probability level of $p < 0.05$ in all analysis.

Results

Daily intakes of biotin by the food group calculation survey

1) Biotin contents of foods

Other than royal jelly, 500 foods were classified into 98 food groups (the small classification) according to the categorization of the Japanese Food Composition Table¹⁶⁾ (Table 1). A group average of biotin content was calculated in 98 food groups. In this group average, the biotin content of meats (offal meats) was 26.1 µg/100 g, which was highest in the small classification, but the biotin content of vegetable fats and oils, Sake and edible salt was under the detectable limit.

2) Biotin intakes from meals surveyed in NHNS

The estimated daily intakes of biotin in males were 52.0 µg for all ages, 27.3 µg for children ages 1-2 years, 42.8 µg for children ages 8-9, and 48.7 µg for adults ages 18-29 from average biotin contents and food intakes of 98 food groups in the NHNS 2011 (Table 1). In females,

Table 2 Representative menu for children in the hospital

Meals	Dishes	Foods	Volume ^a	Biotin contents ^b	Biotin intake ^c
Breakfast	Bread	Mitsuami bread	70	2.9	2.0
	Milk	Milk	100	3.1	3.1
	Boiled egg	Egg	50	17.9	9.0
	Fruit	Orange	40	0.8	0.3
Lunch	Rice	Rice	130	1.9	2.5
	Hamburg steak	Hamburg steak	60	1.2	0.7
		Oil		2	0.0
	Aurora sauce	Soft flour	3	1.7	0.1
		Margarine	3	0.1	0.0
		Ketchup	15	2.5	0.4
		Mushrooms, canned	7	9.8	0.7
	Boiled vegetables	Young sweet corn, frozen	30	2.2	0.7
	Spaghetti	Spaghetti, dried	12	2.7	0.3
		Parsley	1	5.4	0.1
		Oil	2	0.0	0.0
	Salad	Cabbage	35	1.9	0.7
		Radish	7	2.2	0.2
		Cucumber	15	1.9	0.3
		French sauce, white	7	2.5	0.2
	Green potage	Green peas	15	2.2	0.3
		Potage sauce	7	2.2	0.2
		Milk	12	3.1	0.4
	Fruit	Delaware	40	1.5	0.6
	Milk	Milk	200	3.1	6.2
Snack	Almond jelly	Almond jelly sauce	10	1.5	0.2
		Milk	15	3.1	0.5
		Peach, canned	10	1.5	0.2
		Cherry, canned	5	1.5	0.1
Dinner	Rice ball	Rice	130	1.9	2.5
		Umechirimen	2	4.6	0.1
	Fish boiled in broth	Fish	60	6.6 ^d	4.0
		Daikon	50	0.2	0.1
	Ohitashi	Spinach	50	3.3	1.7
		Sesame	1	15.4	0.2
	Kentinjiru	Tofu	15	7.5	1.1
		Burdock	5	2.2	0.1
		Daikon	10	0.2	0.0
		Carrot	5	2.8	0.1
		Shiitake mushroom	5	9.8	0.5
		Welsh onions	2	5.4	0.1
		Sayaingen, frozen	2	2.2	0.0
	Fruit	Grapefruit	80	0.8	0.6
		Cherry, canned	5	1.5	0.1
		Suger	3	3.1	0.1
Milk	Milk	200	3.1	6.2	
			1528	—	47.0

^ag^bµg/100 g^cµg^dThe average of the group of Fish and Shellfishes in 17 food groups.

45.6 µg for all, 24.0 µg for children ages 1-2, 41.5 µg for children ages 8-9, 39.2 µg for adults, respectively.

The highest intake of biotin in men and women was 7.4 µg/day and 5.1 µg/day in rice, and was subsequently 6.7 µg/day and 5.8 µg/day in eggs, 2.7 µg/day and 2.7 µg/day in milk which were contributed 14.2% and 11.2%, 12.9% and 12.7% and 5.2% and 5.9% in daily total biotin intake, respectively. On the other hand, fats and oils such as vegetable oil did not contribute to the biotin intake at all.

3) Biotin intakes from meals shown in the hospital menu

As for hospital meals, the daily average biotin intakes for 10 days calculated from the group average biotin and the amounts of the cooking ingredients shown on the hospital menu sheets were 19.4 µg/day in meals for infants, 45.5 µg/day in meals for children, 52.8 µg/day in meals for pregnant/lactating women, and 56.7 µg/day in common meals for adults (Table 2) (Fig. 1). There was a large variation in daily biotin intakes in children by the kinds of

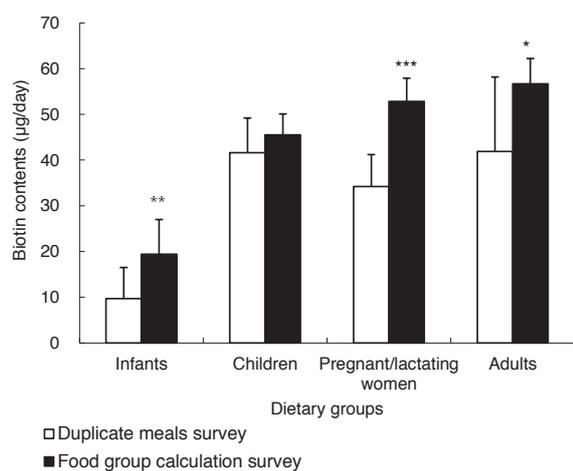


Fig. 1 Comparison of average biotin contents from hospital meals by the food group calculation survey and the duplicate meals survey. Values are means \pm SD, n = 10. Student's t test, * p < 0.05, ** p < 0.01, *** p < 0.001

cooking ingredients (Fig. 2a).

Daily intakes of biotin by the duplicate meals survey

1) Biotin intakes from meals in the hospital

The average biotin intakes for 10 days by the duplicate meals survey were 9.7 $\mu\text{g}/\text{day}$ in meals for infants, 41.6 $\mu\text{g}/\text{day}$ in meals for children, 34.2 $\mu\text{g}/\text{day}$ in meals for pregnant/lactating women and 41.9 $\mu\text{g}/\text{day}$ in common meals for adults. These meals contained basic and standard cooking ingredients but did not balance with the DRI for both adult patients and for pregnant/lactating women (Fig. 1). The daily biotin intakes in infants, children, pregnant/lactating women and adults varied greatly during 10 days (Fig. 2b).

2) Correlation between the food group calculation and duplicate surveys

There was a correlation in biotin contents of hospital meals between the food group calculation and duplicate meals surveys (Fig. 3). Both data were fitted to a smooth curve in regression analysis ($r = 0.89$, $P < 0.01$). The biotin contents of these meals determined by the duplicate meals survey was mainly lower than that calculated by the food group calculation survey. The difference between both surveys was especially large in hospital meals for children, pregnant/lactating women and adults. These hospital meals contained a large amount of eggs and milk.

Discussion

The average intake of biotin in men was 52.0 $\mu\text{g}/\text{day}$ (45.6 $\mu\text{g}/\text{day}$ in women) for all ages by the food group calculation survey (98 food groups) in Japan. These values were as close as 50 $\mu\text{g}/\text{day}$ of AI for adults in DRIs for

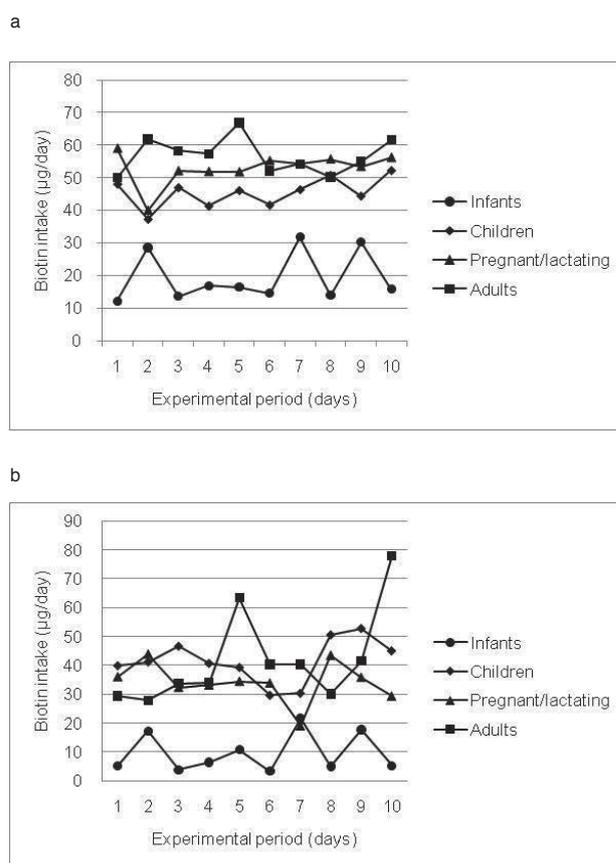


Fig. 2 Change of daily intake of biotin from hospital meals estimated by the food group calculation survey and the duplicate meals survey for 10 consecutive days. a, Food group calculation survey. b, Duplicate meals survey.

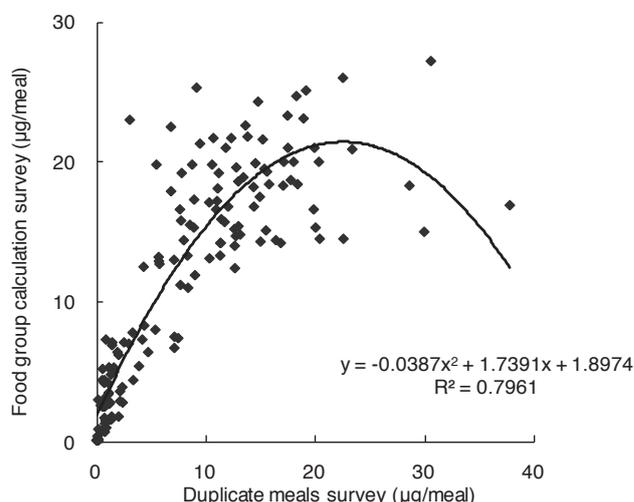


Fig. 3 Correlation in biotin contents of hospital meals between the food group calculation survey and the duplicate meals survey.

Japanese (2010 version)¹². On the other hand, the dietary intake of biotin in the hospital meals estimated by the food group calculation survey was 56.7 $\mu\text{g}/\text{day}$, on average for adults, which was higher than 41.9 $\mu\text{g}/\text{day}$ by the duplicate meals survey. Also, the estimation of dietary biotin intake for other hospital meals tended to be low in the du-

plicate meals survey. The biotin intake by the food group calculation survey exceeded the AI for all ages, but only exceeded the AI in children by the duplicate meals survey. This is because foods determined by the food group calculation survey are not cooked completely, which means there are raw food materials. It is suggested that biotin may be lost via cooking in hospital meals which was determined by the duplicate meals survey in this study.

Also, if there is a large variation in biotin content within the food group, it seems to cause a difference in biotin intake between both dietary surveys. For example, livers where the biotin content is high raises the mean of the food group "Pork"¹⁶⁾, and biotin intake rises in an appearance by the food group calculation survey when a lot of pork classified in the same food group is eaten. The number of foods used in this study is 500, which is only 27.0% of the 1,878 foods listed in the Japanese Food Composition Table. To increase the accuracy of estimation of biotin intake, the remaining foods, especially cooked and processed foods, should be determined.

The AI of biotin set in DRIs for Japanese (2010)¹²⁾ is 4 µg/day for infants ages 0 through 5 months and 10 µg/day for infants ages 6 through 11 months, which in US-DRIs is 5 µg/day for infants ages 0 through 6 months and 6 µg/day for infants ages 7 through 12 months¹⁷⁾. The biotin contents of infant meals in the hospitals in this study were 19.4 µg/day in the food group calculation survey, which exceeded the AI for infants ages 6 through 11 months, and the biotin content by the duplicate meals survey was 9.7 µg/day which was close to the AI for infants ages 6 through 11 months. As the number of foods in the meals catered in the hospitals is low for infants, the biotin intake largely depends on the kind of foods. The biotin content becomes high in the meals containing liver or egg yolk¹⁾. This may be also related to the daily variation of biotin content in the infant meals (CV = 69.8 in the duplicate meals survey; CV = 39.3 in the food group calculation survey).

The AI of biotin intake in DRIs for Japanese people is 20 µg/day for children ages 1 through 2 years and 25 µg/day for children ages 3 through 5 years¹²⁾, which in US-DRIs is 8 µg/day for children ages 1 through 3 years and 12 µg/day for children ages 4 through 8 years¹⁷⁾. The meals for children contain high contents of biotin, which are 45.5 µg/day in the food group calculation survey and 41.6 µg/day in the duplicate meals survey. The biotin contents in these meals by both diet surveys exceeded the AI. The reason why the biotin content of children meals is high may be because milk is contained in hospital meals. Fresh milk is especially served to children twice every

day. By the food group calculation survey, the biotin intake from two cups of milk (400 mL) is 12.4 µg, and the contribution is 27.3% in the total intake of biotin. Therefore, it is thought that the use of milk in the meals is a useful way to get biotin in infants and children in hospitals.

The AI of biotin is 50 µg/day for adults in Japan¹²⁾ and 30 µg/day in US¹⁷⁾. The food group calculation survey showed 56.7 µg/day in the biotin amount of the common meals. On the other hand, the duplicate meal survey was 41.9 µg/day, which is markedly lower than that by the food group calculation survey. The biotin amount per common meal for adults ranged from 27.4 to 78.0 µg/day with a large difference (Fig. 2b). Eggs (17.9 µg/100 g) are commonly included in hospital meals, which largely contribute to the biotin intake as well (12.9% in male adults).

The additional amount of 2 µg/day is set for pregnant women¹²⁾, and the AI in Japan¹²⁾ and US¹⁷⁾ is 52 and 30 µg/day for pregnant women, respectively. The biotin amount in meals for pregnant/lactating women was 52.8 µg/day by the food group calculation survey. On the other hand, the duplicate meals survey showed 34.2 µg/day. The kind of foods in meals for pregnant/lactating women is similar to infant meals in this hospital menu, but fresh milk is not included in these meals. A large variability of biotin intake between days was not seen in meals for pregnant/lactating women. It is suggested that a device to increase biotin intake from meals is necessary for these women.

The comparison of biotin intakes reported in other countries is shown in Table 3²⁵⁻³⁷⁾. In the early studies, Hoppner et al.²⁵⁾ first reported that the biotin intake in the common meals in Canada was 62 µg/day in the calculated value and 60 µg/day in the analyzed value. Lewis and Buss²⁸⁾ demonstrated that the average daily intake of biotin was 35 µg/day (37.5 µg/day including alcoholic drinks and confectionery) in 6,925 households surveyed in England. Afterwards, several studies demonstrated that biotin intake ranged from 25-35 µg/day in different of countries and dietary surveys. In studies over the past 10 years, authors³¹⁾ have reported that the biotin intake ranged from 29.8-33.3 µg/day in the middle and old ages in the Tohoku district of Japan in the duplicate meals survey. Also, the biotin intake ranges from 40-50 µg/day for adults by the duplicate meals survey and also by the total diet study (TDS)^{32, 35, 36)}. On the other hand, the biotin intake ranged from 50-60 µg/day for adults by the food group calculation survey³⁶⁾, which is higher than the duplicate meals and TDS surveys. Taking into consideration the result of two kinds of dietary surveys, it is thought that the variability

Table 3 Comparison of the dietary intake of biotin

References	Daily intake	Subjects	Notes
Hoppner et al. 1978 ²⁵⁾	62 60	Canadaca	calculation of composite diet analysis
Bull & Buss 1982 ²⁶⁾	35.5	7,277 housewives	calculation by the National Food Survey, 1979
Murphy & Calloway 1986 ²⁷⁾	39.9 ± 26.9 ^a	996 women (18-24 years old, USA)	24-hour recall (NHANES II)
Lewis & Buss 1988 ²⁸⁾	35	6,925 households	calculation by the National Food Survey, 1986
Bliss et al. 2000 ²⁹⁾	32 ± 12 ^a	32 subjects (25-85 years old, USA)	diet record for consecutive days
Iyengar et al. 2000 ³⁰⁾	35.5 ± 7.5 ^a	USA	total diet composites (FDA-TDS)
Watanabe et al. 2004 ³¹⁾	29.8-33.3	adults in Tohoku district, Japan	duplicate meals
Saitoh & Ushio 2004 ³²⁾	45.1	Tokyo, Japan	TDS (13 food groups)
Taniguchi et al. 2005 ³³⁾	109.8 92.3	men women	food group calculation survey (18 food groups, 100 foods)
Watanabe & Taniguchi 2006 ³⁴⁾	60.7	Tokyo, Japan	Re-analysis TDS (13 food groups)
Murakami et al. 2008 ³⁵⁾	70.1	Osaka City, Japan	TDS (13 food groups)
Watanabe & Taniguchi 2009 ³⁶⁾	51.1 54.5	all ages	TDS (18 food groups), 2005 food group calculation survey (98 food groups, 330 foods)
Watanabe et al. 2010 ³⁷⁾	51.0 52.4	all ages adults ages 24 years and older	TDS (18 food groups), 2006
Present study	9.7 ^b / 19.4 ^c 41.6 / 45.5 34.2 / 52.8 41.9 / 56.7 27.3 ^{c, d} / 24.0 ^{c, e} 42.8 / 41.5 46.6 / 43.4 48.7 / 39.2 52.0 / 45.6	infants (baby foods) children pregnant/lactating women adults children ages 1-2 years old children ages 8-9 years old children ages 10-11 years old adults ages 18-29 years old all ages	Hospital meals NHNS ^f

^amean ± SD^bduplicate meals survey^cfood group calculation survey (98 food groups, 500 foods)^dmales^efemales^f8,762 subjects (except infants)

of biotin intake in these studies is caused by difference in dietary survey, cooking loss, investigated country, and age distribution of population.

From these findings, it is demonstrated that the food group calculation survey is a useful method for the dietary intake of biotin. The merit of this survey makes it possible to easily calculate the dietary intake of biotin and to utilize the estimated dietary intake of biotin not listed in the Japanese Food Composition Table¹⁶⁾. The average intake of biotin in common meals for adults in the hospital is appropriate at 50 µg/day by the food group calculation survey, which is overestimated compared to that (the real intake) determined by the duplicate meals survey. It is necessary to be careful about the kinds of foods and the method of cooking the ingredients when the biotin intake for hospital patients is calculated using the hospital menu sheets.

Acknowledgements

This work was supported in part by Grants-in-Aid for Scientific Research (C) from the Department of Education

and Science, 2006-2008, (to T.W.) and a Grant-in-Aid from the University of Hyogo, 2005 and 2006, (to T.W.).

References

- 1) Pennington JAT (1989) Bowes and Church's Food Values of Portions Commonly Used 14th Edition, Lippincott Williams and Wilkins; Pennsylvania.
- 2) Zempleni J, Wijeratne SSK, Kuroishi T (2012) Biotin. In: Present Knowledge in Nutrition, 10th Edition, ed. Erdman JW, Jr, MacDonald IA, Zeisel SH, International Life Sciences Institute, Washington DC, pp.359-374.
- 3) Watanabe T (1983) Teratogenic effects of biotin deficiency in mice. J Nutr 113: 574-581.
- 4) Watanabe T, Endo A. (1989) Species and strain differences in teratogenic effects of biotin deficiency in rodents. J Nutr 119: 255-261.
- 5) Takechi R, Taniguchi A, Ebara S, Fukui T, Watanabe T (2008) Biotin deficiency affects the proliferation of human embryonic palatal mesenchymal cells in culture. J Nutr 138: 680-684.

- 6) Watanabe T, Oguchi K, Ebara S, Fukui T (2005) Measurement of 3-hydroxyisovaleric acid in urine of biotin-deficient infants and mice by HPLC. *J Nutr* 135: 615-618.
- 7) Makino Y, Osada K, Sone H, Sugiyama K, Komai M, Ito M, Tsunoda K, Furukawa Y (1999) Percutaneous absorption of biotin in healthy subjects and in atopic dermatitis patients. *J. Nutr. Sci. Vitaminol.* 45: 347-352.
- 8) Báez-Saldaña A, Zendejas-Ruiz I, Revilla-Monsalve C, Islas-Andrade S, Cárdenas A, Rojas-Ochoa A, Vilches A, Fernandez-Mejia C (2004) Effects of biotin on pyruvate carboxylase, acetyl-CoA carboxylase, propionyl-CoA carboxylase, and markers for glucose and lipid homeostasis in type 2 diabetic patients and nondiabetic subjects. *Am J Clin Nutr* 79: 238-243.
- 9) Mock DM, Henrich-Shell CL, Carnell N, Mock NI (2002) Indicators of marginal biotin deficiency and repletion in humans: validation of 3-Hydroxyisovaleric acid excretion and a leucine challenge. *Am J Clin Nutr* 76: 1061-1068.
- 10) Ministry of Health Labour, and Welfare (1999) Recommended Dietary Allowances for Japanese, 6th Revision. Dietary Reference Intakes. Tokyo, Dai-ichi-Shuppan Co.
- 11) Ministry of Health, Labour and Welfare: Dietary Reference Intakes for Japanese, 2005 version.
- 12) Ministry of Health, Labour and Welfare: Dietary Reference Intakes for Japanese, 2010 version. <http://www.mhlw.go.jp/bunya/kenkou/sessyu-kijun.html>.
- 13) Shibata K, Fukuwatari T, Imai E, Hayakawa T, Watanabe F, Takimoto H, Watanabe T, Umegaki K (2013) Dietary reference intakes for Japanese 2010: Water-soluble vitamins. *J Nutr Sci Vitaminol* 59: S67-S82.
- 14) Ministry of Health, Labour and Welfare: Food with Nutrient Function Claims (FNFC). <http://www.mhlw.go.jp/english/topics/foodsafety/fhc/01.html>
- 15) Resources Council of the Science and Technology Agency (2005) Standards tables of food composition in Japan, fifth revised and Enlarged Edition, Ministry of Finance Printing Bureau. Tokyo.
- 16) Ministry of Education, Culture, Sports, Science and Technology: Food Composition Database. <http://fooddb.mext.go.jp/>
- 17) Institute of Medicine (1998) Biotin. In. Dietary Reference Intakes for Thiamin, Riboflavin, Niacin, Vitamin B₆, Folate, Vitamin B₁₂, Pantothenic Acid, Biotin, and Choline. pp.374-389. National Academy Press, Washington, DC.
- 18) Velazquez A, Zamudio S, Baez A, Murguía-Corral R, Rangel-Peniche B, Carrasco A (1990) Indicators of biotin status: A study of patients on prolonged total parenteral nutrition. *Eur J Clin Nutr* 44: 11-16.
- 19) Watanabe T, Masaki T, Yuasa M, Morimoto M, Sawamura H (2013) Estimate of the dietary intake of biotin in infants prescribed special therapeutic infant formulas in Japan. *Int J Ana Bio-Sci.* (in press)
- 20) Daniells S, Hardy G (2010) Hair loss in long-term or home parenteral nutrition: are micronutrient deficiencies to blame? *Curr Opin Clin Nutr Metab Care* 13: 690-697.
- 21) Stratton SL, Henrich CL, Matthews NI, Bogusiewicz A, Dawson AM, Horvath TD, Owen SN, Boyesen G, Moran JH, Mock DM (2012) Marginal biotin deficiency can be induced experimentally in humans using a cost-effective outpatient design. *J Nutr* 142: 22-26.
- 22) Stratton SL, Horvath TD, Bogusiewicz A, Matthews NI, Henrich CL, Spencer HJ, Moran JH, Mock DM (2010) Plasma concentration of 3-hydroxyisovaleryl carnitine is an early and sensitive indicator of marginal biotin deficiency in humans. *Am J Clin Nutr* 92: 1399-1405.
- 23) University of Hyogo, School of Human Science and Nutrition (2008) Food Composition Tables (biotin content). <http://www.shse.u-hyogo.ac.jp/watanabe/biotin-english.htm>
- 24) Ministry of Health, Labour and Welfare (2011) The National Health and Nutrition Survey in Japan. <http://www.mhlw.go.jp/bunya/kenkou/eiyou07/> (in Japanese)
- 25) Hoppner K, Lampi B, Smith DC (1978) An appraisal of the daily intakes of vitamin B₁₂, pantothenic acid and biotin from a composite Canadian diet. *Can Inst Food Sci Technol J* 11: 71-74.
- 26) Bull NL, Buss DH (1982) Biotin, pantothenic acid and vitamin E in the British household food supply. *Hum Nutr Appl Nutr* 36: 190-196.
- 27) Murphy SP, Calloway DH (1986) Nutrient intakes of women in NHANES II, emphasizing trace minerals, fiber, and phytate. *Am Dietet Assoc* 86: 1366-1372.
- 28) Lewis J, Buss DH (1988) Trace nutrients. 5. Minerals and vitamins in the British household food supply. *Br J Nutr* 60: 413-424.
- 29) Bliss DZ, McLaughlin J, Jung HJ, Lowry A, Savik K,

- Jensen L (2000) Comparison of the nutritional composition of diets of persons with fecal incontinence and that of age- and gender-matched controls. *J Wound Ostomy Continence Nurs* 27: 90-97.
- 30) Iyengar GV, Wolf WR, Tanner JT, Morris (2000) Content of minor and trace elements, and organic nutrients in representative mixed total diet components from the USA. *Sci Total Environ* 256: 215-226.
- 31) Watanabe T, Ogushi M, Fukui T (2004) Biotin status in the middle-aged and elderly in Japan. *J Anal Bio-Sci* 27: 403-408. (in Japanese with English abstract)
- 32) Saitoh Y, Ushio F (2004) Estimate of the daily dietary intake of biotin, vitamin B6 and Niacin from the 1999 Tokyo total diet study. *Jpn J Nutr Diet* 62: 165-169. (in Japanese with English abstract)
- 33) Taniguchi A, Ooguchi M, Takechi R, Watanabe T (2005) Biotin content of foods in Japan. *J Jpn Soc Nutr Food Sci* 58: 185-198. (in Japanese with English abstract)
- 34) Watanabe T, Taniguchi A (2006) Study on the estimate of dietary intake of biotin by total diet study. *J Jpn Soc Clin Nutr* 27: 304-312. (in Japanese with English abstract)
- 35) Murakami T, Yamano T, Nakama A, Mori Y (2008) Estimation of dietary intake of biotin and its measurement uncertainty using total diet samples in Osaka, Japan. *J AOAC Int* 91: 1402-1408.
- 36) Watanabe T, Taniguchi A (2009) Estimation of dietary intake of biotin from the Japanese diet. *Vitamins (Japan)* 83: 461-468. (in Japanese with English abstract)
- 37) Watanabe T, Suemura K, Taniguchi A, Ebara S, Kimura S, Fukui T (2010) Dietary intake of seven B vitamins based on a total diet study in Japan. *J Nutr Sci Vitaminol* 56: 279-286.