## Review

() Check for updates

# The association between dietary sodium intake and obesity in adults by sodium intake assessment methods: a review of systematic reviews and re-meta-analysis

### Jounghee Lee <sup>(D)</sup> <sup>1</sup>, Cheongmin Sohn <sup>(D)</sup> <sup>2</sup>, Oh-Yoen Kim <sup>(D)</sup> <sup>3</sup>, Young-Min Lee <sup>(D)</sup> <sup>4</sup>, Mi Ock Yoon <sup>(D)</sup> <sup>5</sup>, and Myoungsook Lee <sup>(D)</sup> <sup>6§</sup>

<sup>1</sup>Department of Food and Nutrition, Kunsan National University, Gunsan 54150, Korea <sup>2</sup>Department of Food and Nutrition, Wonkwang University, Iksan 54538, Korea <sup>3</sup>Department of Food Science and Nutrition, Dong-A University, Busan 49315, Korea <sup>4</sup>Department of Practical Science Education, Gyeongin National University of Education, Incheon 21044, Korea <sup>5</sup>Nutrition Information Center, Korean Nutrition Society, Seoul 04376, Korea

<sup>6</sup>Department of Food and Nutrition, School of Bio-Health Convergence, Health & Wellness College, Sungshin Women's University, Seoul 01133, Korea

## ABSTRACT

**BACKGROUND/OBJECTIVES:** The scientific evidence of a sodium-obesity association is limited by sodium intake assessments. Our specific aim is to synthesize the association between dietary sodium intake and obesity across the sodium intake assessments as evidenced by systematic reviews in adults.

**SUBJECTS/METHODS:** A systematic search identified systematic reviews comparing the association of dietary sodium intakes with obesity-related outcomes such as body mass index (BMI), body weight, waist circumference, and risk of (abdominal) obesity. We searched PubMed on October 24, 2022. To assess the Risk of Bias in Systematic Reviews (ROBIS), we employed the ROBIS tool.

**RESULTS:** This review included 3 systematic reviews, consisting of 39 unique observational studies (35 cross-sectional studies and 4 longitudinal studies) and 15 randomized controlled trials (RCTs). We found consistently positive associations between dietary sodium intake and obesity-related outcomes in cross-sectional studies. Studies that used 24-h urine collection indicated a greater BMI for those with higher sodium intake (mean difference =  $2.27 \text{ kg/m}^2$ ; 95% confidence interval [CI], 1.59-2.51; P < 0.001;  $I^2 = 77\%$ ) compared to studies that used spot urine (mean difference =  $1.34 \text{ kg/m}^2$ ; 95% CI, 1.13-1.55; P < 0.001;  $I^2 = 95\%$ ) and dietary methods (mean difference =  $0.85 \text{ kg/m}^2$ ; 95% CI, 0.1-1.51; P < 0.05;  $I^2 = 95\%$ ).

**CONCLUSIONS:** Quantitative synthesis of the systematic reviews has shown that crosssectional associations between dietary sodium intake and obesity outcomes were substantially different across the sodium intake assessments. We need more high-quality prospective cohort studies and RCTs using 24-h urine collection to examine the causal effects of sodium intake on obesity.

Keywords: Sodium; obesity; systematic review; meta-analysis; association

## OPEN ACCESS

 Received:
 Dec 7, 2022

 Revised:
 Dec 22, 2022

 Accepted:
 Jan 17, 2023

 Published online:
 Jan 20, 2023

#### <sup>§</sup>Corresponding Author: Myoungsook Lee

Department of Food and Nutrition, School of Bio-Health Convergence, Health & Wellness College, Sungshin Women's University, 55 Dobong-ro 76ga-gil, Gangbuk-gu, Seoul 01133, Korea.

Tel. +82-2-920-7211 Fax. +82-2-920-2076 Email. mlee@sungshin.ac.kr

©2023 The Korean Nutrition Society and the Korean Society of Community Nutrition This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (https:// creativecommons.org/licenses/by-nc/4.0/) which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

#### ORCID iDs

Jounghee Lee https://orcid.org/0000-0001-8240-7602 Cheongmin Sohn https://orcid.org/0000-0003-0529-7037 Oh-Yoen Kim https://orcid.org/0000-0001-9262-3309 Young-Min Lee https://orcid.org/0000-0002-3335-2067 Mi Ock Yoon https://orcid.org/0000-0003-1404-9158 Myoungsook Lee https://orcid.org/0000-0003-1344-6979 Nutrition Research and



#### Funding

This work was supported by the Korea Institute of Planning and Evaluation for Technology in Food, Agriculture and Forestry (IPET) through the High Value-added Food Technology Development Program, funded by the Ministry of Agriculture, Food and Rural Affairs (MAFRA) (321029-05).

#### **Conflict of Interest**

The authors declare no potential conflicts of interests.

#### **Author Contributions**

Conceptualization: Lee M, Sohn C, Kim OY, Lee J; Data curation: Lee YM, Lee J; Formal analysis: Lee J; Funding acquisition: Lee M; Investigation: Lee YM, Lee J; Methodology: Lee M, Lee J; Project administration: Yoon MO; Supervision: Lee M; Writing - original draft: Lee YM, Lee J; Writing - review & editing: Lee M, Sohn C, Kim OY, Lee YM, Yoon MO, Lee J.

## INTRODUCTION

Obesity is one of the world's major epidemic challenges. The global prevalence of obesity approximately tripled between 1975 and 2016 [1]. Adult obesity has been associated with various complications such as cardiovascular disease, type 2 diabetes mellitus, coronary heart disease, certain types of cancer, and obstructive sleep apnea [2]. Obesity is a multifactorial disease affected by: (1) dietary factors (e.g., energy intake that exceeds energy needs, high consumption of calorie-dense, nutrient-poor food); (2) low physical activity; and (3) unhealthy psychological conditions (e.g., depression, stress) [3]. Although high energy intake is the major factor leading to obesity, evidence is emerging that high sodium intake leads to additional weight gain in adults [4-6]. Therefore, research is needed to systematically review individual studies and examine associations between dietary sodium intake and obesity.

Multiple methods were utilized to estimate dietary sodium intake, such as dietary methods, 24-h urine, spot urine, and overnight urine. While 24-h urine collection is considered the gold standard for assessing dietary sodium intake, the studies often employed other methods that were less time-consuming or burdensome. However, these other methods are known to be less accurate with regard to reflecting the sodium intake of individuals [7]. Some methods of measuring dietary sodium intake could underestimate or overestimate obesity outcomes. Consequently, it is critical to determine how sodium intake methods could differ when it comes to associations between dietary sodium intake and obesity.

We conducted a review of systematic reviews to identify and synthesize the scientific evidence regarding the association between dietary sodium intake and obesity in adults. Additionally, we performed subgroup analyses of primary studies in the included systematic reviews to examine whether the association of sodium and obesity differs by dietary sodium intake assessment method.

## SUBJECTS AND METHODS

#### **Data sources and searches**

The search strategy was developed based on a protocol of the earlier systematic review [8]. We conducted the data searches for systematic reviews with or without meta-analysis in the PubMed database (from inception to October 24, 2022) (**Supplementary Table 1**). Obesity-related outcomes were included in the search terms, such as body mass index (BMI), body weight, waist circumference, and abdominal obesity.

#### Study eligibility criteria

We included systematic reviews with randomized controlled trials (RCTs) and observational studies. To be included, systematic reviews had to contain information about adult subjects (over 18 yrs). Additionally, systematic reviews had to include dietary sodium intake with a quantitative assessment and different levels of sodium intake or compare the effect of dietary sodium intake based on different levels of intake. Systematic reviews describing a dietary pattern/index indicating a higher intake of sodium, but not indicating the specific levels of sodium intake, were ineligible. Systematic reviews had to have reported outcomes related to obesity (i.e., BMI, body weight, risk of overweight/obesity or abdominal obesity, and waist circumference). Finally, we included systematic reviews with or without meta-analysis, but excluded literature reviews, critical reviews, and review of systematic reviews.



#### **Study selection process**

Two researchers independently screened the abstract of all citations according to the established inclusion and exclusion criteria. To conduct the abstract screening, 2 researchers used the online software Abstrackr separately. Then, 2 researchers independently implemented the full-text screening according to the study eligibility criteria.

#### Data extraction and study quality assessment

We employed the Risk of Bias in Systematic Reviews (ROBIS) tool to assess the quality of the included systematic reviews. The ROBIS tool involves the assessment of 4 domains: study eligibility criteria; identification and selection of studies; data collection and study appraisal; and synthesis and findings [9]. Finally, an overall evaluation was rated as low, unclear, or high by considering the following issues: the interpretation of findings addressed all the concerns identified in the above domains; the relevance of the included studies to the review's research question; and the emphasizing of the results based on statistical significance. The risk of bias of each systematic review was independently evaluated by 2 researchers. Any discrepancies were resolved by discussion.

#### **Quantitative synthesis**

We performed re-meta-analyses of the individual studies included in the earlier systematic reviews to examine the association between dietary sodium and obesity. Due to significant clinical and methodological heterogeneity (e.g., various levels of sodium intake, different characteristics of participants), we used a random effects model for the meta-analyses. To examine the degree of statistical heterogeneity, we utilized the I<sup>2</sup> index. The cut-off values were 25% for low, 50% for moderate, and 75% for high heterogeneity for the descriptive purpose [10]. Meta-analyses were conducted in STATA/SE 17 (Stata Corp., College Station, TX, USA). Less than 0.05 of 2-tailed *P*-values were considered statistically significant.

#### **Narrative synthesis**

We narratively described the findings of the systematic review without meta-analysis in this research.

## RESULTS

#### **Characteristics of individual reviews**

The initial search generated 1,069 citations. Then, we conducted the abstract screening and selected 21 possibly relevant articles for full-text screening. Finally, we selected 3 systematic reviews (2 systematic reviews with meta-analyses and 1 systematic review without meta-analysis) that met the inclusion criteria. The study selection process is shown in **Fig. 1**. Included 3 systematic reviews were published between 2016 and 2021 [11-13]. **Table 1** shows a summary of review characteristics. One systematic review included RCTs and observational studies [11], while 2 systematic reviews included only observational studies [12,13]. Moosavian *et al.* [12] performed a meta-analysis of studies with the inclusion of both children and adults, while Grimes *et al.* [11] implemented a meta-analysis among children and adults separately [11]. Because the specific aim of this study was to investigate the association between sodium intake and adiposity outcomes across sodium intake assessments, we analyzed subgroup analyses with the use of the individual studies only in adults included in the past 2 systematic reviews [11,12]. The risk of bias for observational studies was evaluated by a modified version of the Newcastle-Ottawa Scale in the included 2 systematic reviews



Table 1. Characteristics of individual reviews

Author/Year	Review aim	Search strategy	Included studies in meta-analysis or narrative synthesis for this review	PICOS	Risk of bias
Grimes et al. (2021) [11]	To investigate the association between dietary sodium intake and adiposity	MEDLINE Complete (EBSCO Host), CINAHL (EBSCO Host), Scopus, Embase and CENTRAL	<ul> <li>Cross-sectional studies (n = 25)</li> <li>Longitudinal studies (n = 4)</li> <li>RCTs (n = 15)</li> </ul>	<ul> <li>Population: humans</li> <li>Intervention/exposure: high intake of sodium (observational study)/sodium-reduced diet (RCT)</li> <li>Comparator: low intake of sodium (observational study)/usual or control diet (RCT)</li> <li>Outcome: BMI, body weight, abdominal adiposity, body composition</li> <li>Study type: observational study, RCT</li> </ul>	<ul> <li>A modified version of the NOS for observational studies</li> <li>A modified version of the Cochrane Collaboration risk of bias for RCTs</li> </ul>
Moosavian et al. (2017) [12]	To examine the association between sodium intake and obesity	MEDLINE, Google Scholar, and Scopus	Cross-sectional studies (n = 14)	<ul> <li>Population: humans</li> <li>Intervention/exposure: high intake of sodium</li> <li>Comparator: low intake of sodium</li> <li>Outcome: weight, waist circumference, BMI, overweight, obesity, fat mass and adiposity</li> <li>Study type: observational study</li> </ul>	A modified version of the NOS
Kang et al. (2016) [13]	To identify the association between sodium intake and obesity	KMbase, KoreaMed, NDSL, DBpia, RISS, KISS	Cross-sectional studies (n = 5)	<ul> <li>Population: Koreans</li> <li>Intervention/exposure: high intake of sodium</li> <li>Comparator: low intake of sodium</li> <li>Outcome: overweight/obesity, waist circumference, body fat</li> <li>Study type: observational study</li> </ul>	NI

PICOS, Population, Intervention/exposure, Comparator, Outcome, and Study type; NI, no information; RCT, randomized controlled trial; NOS, Newcastle-Ottawa Scale; BMI, body mass index; CINAHL, Cumulative Index to Nursing and Allied Health Literature; CENTRAL, Cochrane Central Register of Controlled Trials; NDSL, National Discovery for Science Library; RISS, Research Information Sharing Service; KISS, Korean Information Service System.



Fig. 1. Literature search and study selection process.

DASH, Dietary Approaches to Stop Hypertension; SR, systematic review.

[11,12]. The risk of bias for RCTs was evaluated using a modified version of the Cochrane Collaboration in one systematic review [11]. However, the remaining systematic review did not include information related to the risk of bias for the included studies [13].

The results of the ROBIS evaluation are shown in **Table 2**. Regarding study eligibility criteria (domain 1), a clear definition of the search criteria revealed that all included systematic reviews were at low risk [11,12]. However, concern regarding methods used to identify and select studies (domain 2) was considered by Kang *et al.* [13], who did not include methods beyond database searching to identify relevant studies [13]. In the domain of data



#### Dietary sodium intake and obesity in adults

Table 2. Risk of bias using the R	Risk of Bias in Systematic Reviews tool
-----------------------------------	---

Author/Year	Study eligibility criteria	Identification and selection of studies	Data collection and study appraisal	Synthesis and findings	Risk of bias in the review
Grimes et al. (2021) [11]	Low	Low	Low	Low	Low
Moosavian <i>et αl</i> . (2017) [12]	Low	Low	Low	Low	Low
Kang et al. (2016) [13]	Low	High	High	NA <sup>1)</sup>	High

NA, not applicable.

<sup>1)</sup>Because the population of interest was limited to Koreans, there were not enough studies to implement quantitative synthesis.

collection (domain 3), 2 systematic reviews described sufficient study characteristics extracted by independent reviewers [11,12], while the remaining systematic review was not reported with regard to this [13]. Because an adequate tool to assess the risk of bias was used in only 2 systematic reviews, Kang *et al.* [13] raised an additional concern regarding quality assessment for the included studies. Overall, one systematic review was rated as being at high risk of bias [13].

#### **Meta-analysis findings**

#### BMI: cross-sectional studies

We performed a subgroup analysis for 28 cross-sectional studies that reported an association between dietary sodium intake and BMI (**Fig. 2**). The included unique studies utilized: (1) 24-h urine in 11 studies [14-24]; (2) spot urine in 9 studies [25-33]; (3) dietary methods in 7 studies [34-40]; and (4) overnight urine in 1 study [41]. The results from this pooled analysis (38 effect sizes) indicated that BMI was significantly greater among adults in the highest intake group of sodium compared to those in the lowest intake group of sodium (BMI mean difference =  $1.52 \text{ kg/m}^2$ ; 95% confidence interval [CI], 1.24-1.80; P < 0.001;  $I^2 = 97\%$ ), presenting considerable heterogeneity. When we implemented the subgroup analysis by sodium intake assessment method, the greatest BMI was seen in studies that used 24-h urine to examine sodium intake (BMI mean difference =  $2.27 \text{ kg/m}^2$ ; 95% CI, 1.59-2.51; P < 0.001;  $I^2$ 



**Fig. 2.** Pooled mean difference in body mass index by sodium intake assessment. CI, confidence interval; REML, Restricted Maximum Likelihood.

(continued to the next page)



Dietary methods						
Eufinger 2012		-	-		1.00 [ -0.66, 2.66]	1.58
Jablonski 2019	1		-		1.80 [ -1.03, 4.63]	0.77
Murata 2010 - Female					0.20 [ -0.02, 0.42]	3.41
Murata 2010 - Male					0.10 [ -0.11, 0.31]	3.42
Radhika 2007			-	F .	2.70 [ 2.11, 3.29]	3.02
Sharma 2014		-	H		0.80 [ 0.24, 1.36]	3.07
Shikata 2006					0.30 [ -0.04, 0.64]	3.32
Zhao 2014		-			0.80 [ 0.10, 1.50]	2.87
Heterogeneity: $\tau^2 = 0.73$ , $I^2 = 94.64\%$ , $H^2 = 18.67$					0.85 [ 0.18, 1.51]	
Test of $\theta_i = \theta_j$ : Q(7) = 73.33, p = 0.00						
Test of $\theta$ = 0: z = 2.49, p = 0.01						
Overnight urine						
Yokokawa 2016					1.00 [ 0.48, 1.52]	3.11
Heterogeneity: $\tau^2 = 0.00$ , $I^2 = .\%$ , $H^2 = .$					1.00 [ 0.48, 1.52]	
Test of $\theta_i = \theta_j$ : Q(0) = 0.00, p = .			4			
Test of θ = 0: z = 3.75, p = 0.00						
Spot urine						
Han 2014					1.10 [ 0.82, 1.38]	3.37
Han 2017 - Female					0.90 [ -0.06, 1.86]	2.48
Han 2017 - Male		_			1.20 [ 0.34, 2.06]	2.63
Huh 2015 - Female					1.40 [ 1.14, 1.66]	3.39
Huh 2015 - Male			T		0.60 [ 0.35, 0.85]	3.40
Lee 2015			1		0.90 [ 0.57, 1.23]	3.33
Oh 2015					1.50 [ 1.36, 1.64]	3.46
Petermann-Rocha 2019					2.00 [ 1.58, 2.42]	3.24
Pfister 2014 - Female					1.60 [ 1.35, 1.85]	3.39
Pfister 2014 - Male					1.20 [ 0.98, 1.42]	3.42
Watanabe 2019			-		1.40 [ 1.05, 1.75]	3.31
Welsh 2019 - Male					1.70 [ 1.64, 1.76]	3.48
Welsh 2019- Female					1.60 [ 1.53, 1.67]	3.48
Heterogeneity: $\tau^2 = 0.12$ , $I^2 = 94.68\%$ , $H^2 = 18.79$			4		1.34 [ 1.13, 1.55]	
Test of $\theta_i = \theta_i$ : Q(12) = 127.54, p = 0.00						
Test of $\theta$ = 0: z = 12.50, p = 0.00						
Overall			4		1 52 [ 1 24 1 80]	
Heterogeneity: $r^2 = 0.60$ $l^2 = 97.38\%$ $H^2 = 38.18$			Υ.		1.02 [ 1.24, 1.00]	
Test of $\theta_{1} = \theta_{1}^{*} O(37) = 775.96 \text{ n} = 0.00$						
Test of $\theta = 0$ ; $z = 10.55$ , $p = 0.00$						
1000 010 - 0.2 - 10.00, p - 0.00						
Test of group differences: $Q_b(3) = 20.20$ , p = 0.00	0.00		0.00			
	-2.00	0.00	2.00	4.00	6.00	
and an offering DEM medal						

Random-effects REML model



= 77%), followed by: (1) spot urine (BMI mean difference =  $1.34 \text{ kg/m}^2$ ; 95% CI, 1.13-1.55; *P* < 0;001; I<sup>2</sup> = 95%); (2) overnight urine (BMI mean difference =  $1.00 \text{ kg/m}^2$ ; 95% CI, 0.48-1.52; *P* < 0.001); and (3) dietary methods (BMI mean difference =  $0.85 \text{ kg/m}^2$ ; 95% CI, 0.18-1.51; *P* < 0.05; I<sup>2</sup> = 95%).



#### Risk of overweight/obesity: cross-sectional studies

We implemented a meta-analysis of 5 cross-sectional studies (8 effect sizes) (**Fig. 3**) [19,26,31,42,43]. The findings from this pooled analysis show that higher sodium intake was significantly associated with an increased risk of overweight or obesity (odds ratio [OR], 1.75; 95% CI, 1.40–2.18, P < 0.001; I<sup>2</sup> = 72%), presenting substantial heterogeneity. In a subgroup analysis by the sodium intake assessment, the risk of overweight/obesity was not significantly different across the group. A higher level of dietary sodium intake was significantly associated with a risk of overweight/obesity in studies that used the 24-h urine (OR, 2.44; 95% CI, 1.67–3.56; P < 0.001; I<sup>2</sup> = 59%) [19,42] and dietary methods (OR, 1.48; 95% CI, 1.17–1.88; P < 0.001; I<sup>2</sup> = 16%) [43]. However, in studies that used spot urine, there was no association between dietary sodium intake and the risk of overweight/obesity [26,31].

#### Body weight: cross-sectional studies

The findings from a meta-analysis of 7 cross-sectional studies (12 effect sizes) indicated that body weight was significantly greater among adults in the highest group of dietary sodium intake (body weight mean difference = 8.28 kg; 95% CI, 6.69–9.86; P < 0.001;  $I^2 = 66\%$ ), presenting substantial heterogeneity (**Fig. 4**) [15,16,18,19,21,34,41]. The results of a



Random-effects REML model

Fig. 3. Pooled risk of overweight or obesity by sodium intake assessment. OR, odds ratio; CI, confidence interval; REML, Restricted Maximum Likelihood.



subgroup analysis showed that body weight was greatest in studies that used 24-h urine (body weight mean difference = 9.00 kg; 95% CI, 7.53–10.48; P < 0.001;  $I^2 = 46\%$ ) [15,16,18,19,21], followed by those that used overnight urine (body weight mean difference = 5.00 kg; 95% CI, 3.46–6.54; P < 0.001) [41]. Studies that used dietary methods showed that there was an insignificant relationship between dietary salt intake and body weight [34].

Waist circumference: cross-sectional studies

We performed a meta-analysis of 10 cross-sectional studies (13 effect sizes) (**Fig. 5**) [15,17,19,23,25,27-30,37]. The results of this meta-analysis showed that higher sodium intake was significantly associated with an increase in waist circumference (waist circumference mean difference = 4.99 cm; 95% CI, 4.00–5.99; P < 0.001;  $I^2 = 89\%$ ), presenting considerable heterogeneity. In a subgroup analysis by sodium intake assessment, waist circumference was greater in studies that used 24-h urine (waist circumference mean difference = 6.34 cm; 95% CI, 5.27–7.42; P < 0.001;  $I^2 = 0\%$ ) [15,17,19,23] compared to those that utilized spot urine



Fig. 4. Pooled mean difference in body weight by sodium intake assessment. CI, confidence interval; REML, Restricted Maximum Likelihood.



Study								Effec with 9	t si 5%	ze Cl	Weight (%)
24-hr urine	Τ		1								
Hoffman 2009 - Female			_	-		-		6.80 [ 2.4	4,	11.16]	3.54
Hoffman 2009 - Male		-	_	-		_		6.00 [ 1.6	6,	10.34]	3.56
Ma 2015			_	-	-			7.80 [ 3.6	4,	11.96]	3.77
Nam 2017 - Female		-	_	-	2	_		6.20 [ 2.6	5,	9.75]	4.59
Nam 2017 - Male				-	14			5.50 [ 2.6	7,	8.33]	5.82
Yan 2016				-				6.40 [ 4.9	8,	7.82]	8.94
Heterogeneity: $\tau^2 = 0.00$ , $I^2 = 0.00\%$ , $H^2 = 1.00$				•				6.34 [ 5.2	7,	7.42]	
Test of $\theta_i = \theta_j$ : Q(5) = 0.89, p = 0.97											
Test of $\theta$ = 0: z = 11.56, p = 0.00											
Dietary methods											
Radhika 2007				-	-			7.70 [ 6.1	3,	9.27]	8.60
Heterogeneity: $\tau^2 = 0.00$ , $I^2 = .\%$ , $H^2 = .$								7.70 [ 6.1	3,	9.27]	
Test of $\theta_i = \theta_j$ : Q(0) = 0.00, p = .											
Test of $\theta$ = 0: z = 9.63, p = 0.00											
Spot urine											
Han 2014								3.50 [ 2.6	7,	4.33]	10.14
Huh 2015 - Female								4.20 [ 3.4	7,	4.93]	10.31
Huh 2015 - Male		-						2.20 [ 1.4	5,	2.95]	10.28
Lee 2015		-	F					3.30 [ 2.3	2,	4.28]	9.88
Oh 2015								4.70 [ 4.2	9,	5.11]	10.72
Petermann-Rocha 2019			-					5.70 [ 4.7	1,	6.69]	9.86
Heterogeneity: $\tau^2$ = 1.27, $I^2$ = 90.37%, $H^2$ = 10.39								3.93 [ 2.9	7,	4.89]	
Test of $\theta_i = \theta_j$ : Q(5) = 47.78, p = 0.00											
Test of $\theta$ = 0: z = 8.04, p = 0.00											
Overall								4.99 [ 4.0	0,	5.99]	
Heterogeneity: $\tau^2 = 2.37$ , $I^2 = 89.47\%$ , $H^2 = 9.49$								•			
Test of $\theta_i = \theta_i$ : Q(12) = 81.99, p = 0.00											
Test of θ = 0: z = 9.81, p = 0.00											
Test of group differences: $Q_b(2) = 20.23$ , p = 0.00											
	ó		5	5	ŝ	10	1	5			
Random-effects REML model											

Fig. 5. Pooled mean difference in waist circumference by sodium intake assessment. Cl, confidence interval; REML, Restricted Maximum Likelihood.

(waist circumference mean difference = 3.93 cm; 95% CI, 2.97–4.89; P < 0.001; I<sup>2</sup> = 90%) [25,27-29]. Additionally, one study utilizing the dietary method showed that higher sodium intake is significantly associated with greater waist circumference (waist circumference mean difference = 7.70 cm; 95% CI, 6.13–9.27; P < 0.001) [37].

#### Abdominal obesity: cross-sectional studies

We employed a meta-analysis of 5 cross-sectional studies (7 effect sizes) to investigate the association between dietary sodium intake and abdominal obesity (**Fig. 6**) [19,27,42,44,45]. The findings from this pooled analysis showed that higher sodium intake was significantly associated with an increased risk of abdominal obesity (OR, 2.05; 95% CI, 1.72–2.44; P < 0.001;  $I^2 = 67\%$ ), presenting substantial heterogeneity. Subgroup analysis by sodium intake assessment showed that the association between dietary sodium intake and the risk of



Study		Odds ratio with 95% CI	Weight (%)
24-hr urine			
Ge 2015		2.32 [ 1.82, 2.96]	16.62
Nam 2017 - Female		3.00 [ 1.49, 6.06]	4.99
Nam 2017 - Male		3.49 [ 2.08, 5.86]	7.83
Zhao 2019		2.07 [ 1.74, 2.46]	19.70
Heterogeneity: $\tau^2 = 0.01$ , $I^2 = 29.42\%$ , $H^2 = 1.42$	-	2.34 [ 1.94, 2.81]	
Test of $\theta_i = \theta_j$ : Q(3) = 4.36, p = 0.23			
Test of $\theta$ = 0: z = 8.99, p = 0.00			
Dietary methods			
Zhang 2018		1.78 [ 1.29, 2.45]	13.46
Heterogeneity: $\tau^2 = 0.00$ , $I^2 = .\%$ , $H^2 = .$		1.78 [ 1.29, 2.45]	
Test of $\theta_i = \theta_j$ : Q(0) = 0.00, p = .			
Test of $\theta$ = 0: z = 3.52, p = 0.00			
Spot urine			
Huh 2015 Female		2.01 [ 1.67, 2.42]	19.07
Huh 2015 Male		1.46 [ 1.19, 1.79]	18.33
Heterogeneity: $\tau^2 = 0.04$ , $I^2 = 80.43\%$ , $H^2 = 5.11$		1.72 [ 1.26, 2.35]	
Test of $\theta_i = \theta_j$ : Q(1) = 5.11, p = 0.02			
Test of $\theta$ = 0: z = 3.38, p = 0.00			
Overall	+	2.05 [ 1.72, 2.44]	
Heterogeneity: $\tau^2 = 0.03$ , $I^2 = 66.52\%$ , $H^2 = 2.99$			
Test of $\theta_i = \theta_j$ : Q(6) = 16.81, p = 0.01			
Test of θ = 0: z = 7.97, p = 0.00			
Test of group differences: $Q_b(2) = 3.85$ , p = 0.15	1 2 3 4		

Random-effects REML model

Fig. 6. Pooled risk of abdominal obesity by sodium intake assessment.

OR, odds ratio; CI, confidence interval; REML, Restricted Maximum Likelihood.

abdominal obesity was not significantly different among groups. There was a significant association between dietary sodium intake and the risk of abdominal obesity in all 3 groups: (1) 24-h urine (OR, 2.34; 95% CI, 1.94–2.81; P < 0.001;  $I^2 = 29\%$ ) [19,42,44]; (2) dietary methods (OR, 1.78; 95% CI, 1.29–2.45; P < 0.001) [45]; and (3) spot urine (OR, 1.72; 95% CI, 1.26–2.35; P < 0.001) [27].

#### Body weight: RCTs

The prior systematic review included 15 RCTs with durations ranging from 3 mon to 4 yrs. Included RCTs were conducted in various countries: (1) 4 in the USA; (2) 3 in the UK; (3) 3 in Australia; and (4) 1 in the Netherlands, 1 in Italy, 1 in Belgium, 1 in Japan, and 1 in China. Ten RCTs employed nutrition education or nutrition counseling and behavioral strategies to reduce sodium intake. Grimes *et al.* [11] performed a pooled analysis of RCTs to examine the effects of a reduced sodium diet on body weight. The findings of 15 RCTs reported that reduced-sodium diets compared to a usual diet or control diet showed non-significant effects on body weight [46-60].



Author/Year	Study types (sodium intake assessment)	Outcomes	Narrative synthesis			
Grimes et al. (2021) [11]	Longitudinal studies	Body weight	No change in body weight in 4 studies			
	• Larsen et al. (2013) (24-h urine) [4]					
	• Ard et al. (2004) (24-h urine) [61]					
	• Sakaki <i>et al</i> . (2014) (diet history) [62]					
	• Takahashi et al. (2006) (24-h urine) [63]					
	Longitudinal study	Waist circumstance	No change in waist circumference in 1 study			
	• Larsen <i>et al</i> . (2013) (24-h urine) [4]					
	Longitudinal study	Body fat	Per 100 mmol/d increase in sodium → Significant increase in body fat in 1 study			
	• Larsen et al. (2013) (24-h urine) [4]					
	Longitudinal study	Fat free mass	Per 100 mmol/d increase in sodium → Significan			
	• Larsen <i>et αl</i> . (2013) (24-h urine) [4]		decrease in fat free mass in 1 study			
Kang et al. (2016) [13]	Cross-sectional study • Song et al. (2013) (dietary method) [43]	Risk of being overweight	<ul> <li>Males: higher sodium intake → Significantly increased risk of being overweight</li> </ul>			
			<ul> <li>Females: no association</li> </ul>			
	Cross-sectional study • Kim et al. (2015) (dietary method) [64]	Risk of obesity	<ul> <li>Males: ≥ 4 g vs. &lt; 2 g of sodium intake → Significantly increased risk of obesity</li> </ul>			
			• Females: no association			
	Cross-sectional study	Waist circumference	Higher sodium intake was significantly associated			
	• Lim and Yang (2014) (dietary method) [65]		with greater waist circumference			
	Cross-sectional study	Risk of abdominal obesity	Higher sodium intake was significantly associated			
	• Oh et al. (2015) (24-h urine) [66]	, ,	with a higher risk of abdominal obesity			
	Cross-sectional study	Body fat	Higher sodium intake was significantly associated			
	• Oh et al. (2015) (spot urine) [29]		with higher body fat			

Table 3. Results of individual reviews that state a narrative synthesis for obesity-related outcomes

#### Narrative synthesis

#### Obesity-related outcomes: longitudinal studies

Grimes *et al.* [11] included 4 longitudinal studies to examine the association between dietary sodium intake and obesity outcomes. Four studies showed that higher sodium intake did not change body weight [4,61-63] (**Table 3**). One study indicated that higher sodium intake did not increase waist circumference [4]. However, higher sodium intake significantly increased body fat [4] but decreased fat free mass per a 100 mmol/d sodium increase [4].

*Obesity-related outcomes: a systematic review including only Korean studies* Kang *et al.* [13] included only Korean studies that examined the association between dietary sodium intake and obesity-related outcomes. Two studies suggested that higher sodium intake increased the risk of overweight/obesity in males but not in females [43,64]. Higher sodium intake was significantly associated with: (1) greater waist circumference in one study [65]; (2) increased risk of abdominal obesity in one study [66]; and (3) increased body fat in one study [29].

## DISCUSSION

The main findings from this review of systematic reviews and re-meta-analysis indicated consistently positive associations between dietary sodium intake and obesity-related outcomes in cross-sectional studies. Additionally, subgroup analyses showed that the associations between dietary sodium intake and obesity outcomes (i.e., BMI, body weight, and waist circumference) were significantly different across sodium intake assessments. Compared to studies utilizing 24-h urine collection, the other sodium intake assessments (spot or overnight urine and dietary methods) tend to underestimate the sodium-obesity association in cross-sectional studies. This review of the systematic reviews provides useful



information to detect the difference in the association between sodium intake and obesityrelated outcomes by sodium intake assessment.

The possible biological mechanisms could explain this positive association between dietary sodium intake and obesity. First, salt treatment increased adipogenesis/lipogenesis and proinflammatory adipocytokine secretion. However, it decreased lipolysis in a dose-dependent manner in adipocytes, suggesting inflammatory adipogenesis as a possible mechanism [67]. Second, endogenous fructose production was identified as the primary mechanism regarding salt-induced obesity in high-salt-fed mice [68]. High-salt-diet-fed rats were also associated with increased adipose tissue mass by enhancing insulin-stimulated glucose uptake and lipogenic capacity [69,70]. Third, a high-salt diet (18 g/day) increased significantly fasting ghrelin levels compared to a normal salt diet (3 g/day) in normotensive subjects, indicating that increased ghrelin induced obesity by stimulating appetite, fat accumulation, and glucose intolerance [71]. In addition, increased salt intake (6 g/day) decreased significantly diet-induced thermogenesis compared to the placebo group, which contributes to increased weight gain [72].

To identify the association between sodium intake and obesity, it is critical to use a valid, reliable tool for assessing sodium intake. Twenty-four urine collection is known to be the gold standard method for the assessment of dietary sodium intake [7], as about 90% of ingested sodium is excreted in urine within 24 h [73]. Because participants would carry a considerable burden with 24-h urine collection, some researchers tend to utilize more convenient or affordable alternatives such as spot or overnight urine and dietary assessments. However, this dietary assessment intake is limited to reflecting dietary sodium intake due to the variable of sodium content from food sources (e.g., home-cooked foods, restaurant foods, or processed foods), the omission of discretionary salt use, and the under-reporting of problems, especially for obese participants [7]. Spot or overnight urine samples are influenced by many non-dietary sources. Short-term urine samples would varied sodium content due to hydration status, body standing position, time of day, natriuretic or diuretic substance (e.g., caffeine), and disease status (e.g., renal disease and cardiac diseases) [74,75]. Consequently, short-term urine collection (i.e., spot urine or overnight urine) and dietary assessment methods could not replace 24-h urine.

Major scientific organizations have raised concerns about research methods for sodium intake assessments. Spot and short-duration timed urine collection methods show systematic errors in estimating an individual's sodium intake [76,77]. The International Consortium for Quality Research on Dietary Sodium/Salt (TRUE) recommends that single spot or short-duration timed urine collections not be used for assessments of dietary sodium intake of individuals especially related to health outcomes [74]. Consequently, it is critical to identify the association between sodium intake and obesity outcomes by utilizing a valid sodium assessment tool such as 24-h urine collection.

We concluded that there is a cross-sectional association between sodium intake and adiposity outcomes. These cross-sectional associations between dietary sodium and obesity were significantly different across sodium intake assessments. Studies that used spot or overnight urine tended to underestimate the sodium-obesity association compared to those utilizing 24-h urine collection. The limitation of this review is that we used only the PubMed database to search for relevant studies. It is recommended that more databases be used to conduct comprehensive searches. Another limitation is that some observational studies did not control



for confounding factors for obesity-related outcomes. To identify the causal effects of sodium intake on obesity, we need more high-quality RCTs with the provision of a sodium-reduced diet or foods and a large number of prospective cohort studies that use 24-h urine collection.

## SUPPLEMENTARY MATERIAL

#### Supplementary Table 1

Search strategy: PubMed (search date: 10/24/2022)

**Click here to view** 

## REFERENCES

- NCD Risk Factor Collaboration (NCD-RisC). Trends in adult body-mass index in 200 countries from 1975 to 2014: a pooled analysis of 1698 population-based measurement studies with 19-2 million participants. Lancet 2016;387:1377-96.
   PUBMED | CROSSREF
- Ansari S, Haboubi H, Haboubi N. Adult obesity complications: challenges and clinical impact. Ther Adv Endocrinol Metab 2020;11:2042018820934955.
   PUBMED | CROSSREF
- 3. Hruby A, Hu FB. The epidemiology of obesity: a big picture. Pharmacoeconomics 2015;33:673-89. PUBMED | CROSSREF
- Larsen SC, Ängquist L, Sørensen TI, Heitmann BL. 24h urinary sodium excretion and subsequent change in weight, waist circumference and body composition. PLoS One 2013;8:e69689.
   PUBMED L CROSSREF
- Yi SS, Firestone MJ, Beasley JM. Independent associations of sodium intake with measures of body size and predictive body fatness. Obesity (Silver Spring) 2015;23:20-3.
   PUBMED | CROSSREF
- 6. Yoon YS, Oh SW. Sodium density and obesity; the Korea National Health and Nutrition Examination Survey 2007-2010. Eur J Clin Nutr 2013;67:141-6.

## PUBMED | CROSSREF

- 7. McLean RM. Measuring population sodium intake: a review of methods. Nutrients 2014;6:4651-62. PUBMED | CROSSREF
- Grimes CA, Bolhuis DP, He FJ, Nowson CA. Dietary sodium intake and overweight and obesity in children and adults: a protocol for a systematic review and meta-analysis. Syst Rev 2016;5:7.
   PUBMED | CROSSREF
- Whiting P, Savović J, Higgins JP, Caldwell DM, Reeves BC, Shea B, Davies P, Kleijnen J, Churchill R; ROBIS group. ROBIS: a new tool to assess risk of bias in systematic reviews was developed. J Clin Epidemiol 2016;69:225-34.
   PUBMED | CROSSREF
- Higgins JP, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in meta-analyses. BMJ 2003:327:557-60.

- Grimes CA, Bolton KA, Booth AB, Khokhar D, Service C, He FH, Nowson CA. The association between dietary sodium intake, adiposity and sugar-sweetened beverages in children and adults: a systematic review and meta-analysis. Br J Nutr 2021;126:409-27.
   PUBMED | CROSSREF
- Moosavian SP, Haghighatdoost F, Surkan PJ, Azadbakht L. Salt and obesity: a systematic review and metaanalysis of observational studies. Int J Food Sci Nutr 2017;68:265-77.
   PUBMED | CROSSREF
- Kang YJ, Wang HW, Cheon SY, Lee HJ, Hwang KM, Yoon HS. Associations of obesity and dyslipidemia with intake of sodium, fat, and sugar among Koreans: a qualitative systematic review. Clin Nutr Res 2016;5:290-304.
   PUBMED | CROSSREF



- Baudrand R, Campino C, Carvajal CA, Olivieri O, Guidi G, Faccini G, Vöhringer PA, Cerda J, Owen G, Kalergis AM, et al. High sodium intake is associated with increased glucocorticoid production, insulin resistance and metabolic syndrome. Clin Endocrinol (Oxf) 2014;80:677-84.
   PUBMED | CROSSREF
- 15. Hoffmann IS, Cubeddu LX. Salt and the metabolic syndrome. Nutr Metab Cardiovasc Dis 2009;19:123-8. PUBMED | CROSSREF
- Hulthén L, Aurell M, Klingberg S, Hallenberg E, Lorentzon M, Ohlsson C. Salt intake in young Swedish men. Public Health Nutr 2010;13:601-5.

  PUBMED | CROSSREF
- 17. Ma Y, He FJ, MacGregor GA. High salt intake: independent risk factor for obesity? Hypertension 2015;66:843-9.

- Madhavan S, Alderman MH. Ethnicity and the relationship of sodium intake to blood pressure. J Hypertens 1994;12:97-103.
   PUBMED | CROSSREF
- Nam GE, Kim SM, Choi MK, Heo YR, Hyun TS, Lyu ES, Oh SY, Park HR, Ro HK, Han K, et al. Association between 24-h urinary sodium excretion and obesity in Korean adults: a multicenter study. Nutrition 2017;41:113-9.
   PUBMED | CROSSREF
- Polonia JJ, Magalhaes MT, Senra D, Barbosa L, Silva JA, Ribeiro SM. Association of 24-h urinary salt excretion with central haemodynamics and assessment of food categories contributing to salt consumption in Portuguese patients with hypertension. Blood Press Monit 2013;18:303-10.
   PUBMED | CROSSREF
- Vega-Vega O, Fonseca-Correa JI, Mendoza-De la Garza A, Rincón-Pedrero R, Espinosa-Cuevas A, Baeza-Arias Y, Dary O, Herrero-Bervera B, Nieves-Anaya I, Correa-Rotter R. Contemporary dietary intake: too much sodium, not enough potassium, yet sufficient iodine: the SALMEX Cohort results. Nutrients 2018;10:816.
  - PUBMED | CROSSREF
- Verhave JC, Hillege HL, Burgerhof JG, Janssen WM, Gansevoort RT, Navis GJ, de Zeeuw D, de Jong PE; PREVEND Study Group. Sodium intake affects urinary albumin excretion especially in overweight subjects. J Intern Med 2004;256:324-30.
   PUBMED | CROSSREF
- Yan L, Guo X, Wang H, Zhang J, Tang J, Lu Z, Cai X, Liu L, Gracely EJ, Ma J. Population-based association between urinary excretion of sodium, potassium and its ratio with albuminuria in Chinese. Asia Pac J Clin Nutr 2016;25:785-97.
- Rhee MY, Kim JH, Kim YS, Chung JW, Bae JH, Nah DY, Kim YK, Lee MM, Lim CY, Byun JE, et al. High sodium intake in women with metabolic syndrome. Korean Circ J 2014;44:30-6.
   PUBMED | CROSSREF
- Han SY, Hong JW, Noh JH, Kim DJ. Association of the estimated 24-h urinary sodium excretion with albuminuria in adult Koreans: the 2011 Korea National Health and Nutrition Examination Survey. PLoS One 2014;9:e109073.
   PUBMED | CROSSREF
- Han W, Hu Y, Tang Y, Xue F, Hou L, Liang S, Zhang B, Wang W, Asaiti K, Pang H, et al. Relationship between urinary sodium with blood pressure and hypertension among a Kazakh community population in Xinjiang, China. J Hum Hypertens 2017;31:333-40.
   PUBMED | CROSSREF
- Huh JH, Lim JS, Lee MY, Chung CH, Shin JY. Gender-specific association between urinary sodium excretion and body composition: analysis of the 2008-2010 Korean National Health and Nutrition Examination Surveys. Metabolism 2015;64:837-44.
   PUBMED | CROSSREF
- Lee SK, Kim JS, Kim SH, Kim YH, Lim HE, Kim EJ, Park CG, Cho GY, Kim J, Baik I, et al. Sodium excretion and cardiovascular structure and function in the nonhypertensive population: the Korean Genome and Epidemiology Study. Am J Hypertens 2015;28:1010-6.
   PUBMED | CROSSREF
- Oh SW, Han KH, Han SY, Koo HS, Kim S, Chin HJ. Association of sodium excretion with metabolic syndrome, insulin resistance, and body fat. Medicine (Baltimore) 2015;94:e1650.
   PUBMED | CROSSREF
- 30. Petermann-Rocha F, Sillars A, Brown R, Sweeney L, Troncoso C, García-Hermoso A, Leiva AM, Martínez MA, Diaz-Martínez X, Poblete-Valderrama F, et al. Sociodemographic patterns of urine sodium



excretion and its association with hypertension in Chile: a cross-sectional analysis. Public Health Nutr 2019;22:2012-21.

PUBMED | CROSSREF

- 31. Watanabe S, Konta T, Ichikawa K, Watanabe M, Ishizawa K, Ueno Y, Yamashita H, Kavama T, Kubota I. The association between urinary sodium excretion and blood pressure in a community-based population: the Yamagata (Takahata) study. Clin Exp Nephrol 2019;23:380-6. PUBMED | CROSSREF
- 32. Pfister R, Michels G, Sharp SJ, Luben R, Wareham NJ, Khaw KT. Estimated urinary sodium excretion and risk of heart failure in men and women in the EPIC-Norfolk study. Eur J Heart Fail 2014;16:394-402. PUBMED | CROSSREF
- 33. Welsh CE, Welsh P, Jhund P, Delles C, Celis-Morales C, Lewsey JD, Gray S, Lyall D, Iliodromiti S, Gill JM, et al. Urinary sodium excretion, blood pressure, and risk of future cardiovascular disease and mortality in subjects without prior cardiovascular disease. Hypertension 2019;73:1202-9. PUBMED | CROSSREF
- 34. Eufinger SC, Votaw J, Faber T, Ziegler TR, Goldberg J, Bremner JD, Vaccarino V. Habitual dietary sodium intake is inversely associated with coronary flow reserve in middle-aged male twins. Am J Clin Nutr 2012:95:572-9. PUBMED | CROSSREF
- 35. Jablonski KL, Gates PE, Pierce GL, Seals DR. Low dietary sodium intake is associated with enhanced vascular endothelial function in middle-aged and older adults with elevated systolic blood pressure. Ther Adv Cardiovasc Dis 2009;3:347-56.

#### PUBMED | CROSSREF

- 36. Murata A, Fujino Y, Pham TM, Kubo T, Mizoue T, Tokui N, Matsuda S, Yoshimura T. Prospective cohort study evaluating the relationship between salted food intake and gastrointestinal tract cancer mortality in Japan. Asia Pac J Clin Nutr 2010;19:564-71. PUBMED
- 37. Radhika G, Sathya RM, Sudha V, Ganesan A, Mohan V. Dietary salt intake and hypertension in an urban south Indian population -- [CURES - 53]. J Assoc Physicians India 2007;55:405-11. PURMED
- 38. Sharma S, McFann K, Chonchol M, Kendrick J. Dietary sodium and potassium intake is not associated with elevated blood pressure in US adults with no prior history of hypertension. J Clin Hypertens (Greenwich) 2014;16:418-23. PUBMED | CROSSREF
- 39. Shikata K, Kiyohara Y, Kubo M, Yonemoto K, Ninomiya T, Shirota T, Tanizaki Y, Doi Y, Tanaka K, Oishi Y, et al. A prospective study of dietary salt intake and gastric cancer incidence in a defined Japanese population: the Hisayama study. Int J Cancer 2006;119:196-201. PUBMED | CROSSREF
- 40. Zhao X, Yin X, Li X, Yan LL, Lam CT, Li S, He F, Xie W, Sang B, Luobu G, et al. Using a low-sodium, highpotassium salt substitute to reduce blood pressure among Tibetans with high blood pressure: a patientblinded randomized controlled trial. PLoS One 2014;9:e110131. PUBMED | CROSSREF
- 41. Yokokawa H, Yuasa M, Nedsuwan S, Moolphate S, Fukuda H, Kitajima T, Minematsu K, Tanimura S, Marui E. Daily salt intake estimated by overnight urine collections indicates a high cardiovascular disease risk in Thailand. Asia Pac J Clin Nutr 2016;25:39-45. PUBMED | CROSSREF
- 42. Zhao L, Cogswell ME, Yang Q, Zhang Z, Onufrak S, Jackson SL, Chen TC, Loria CM, Wang CY, Wright JD, et al. Association of usual 24-h sodium excretion with measures of adiposity among adults in the United States: NHANES, 2014. Am J Clin Nutr 2019;109:139-47. PUBMED | CROSSREF
- 43. Song HJ, Cho YG, Lee HJ. Dietary sodium intake and prevalence of overweight in adults. Metabolism 2013;62:703-8.

PUBMED | CROSSREF

- 44. Ge Z, Guo X, Chen X, Tang J, Yan L, Ren J, Zhang J, Lu Z, Dong J, Xu J, et al. Association between 24 h urinary sodium and potassium excretion and the metabolic syndrome in Chinese adults: the Shandong and Ministry of Health Action on Salt and Hypertension (SMASH) study. Br J Nutr 2015;113:996-1002. PUBMED | CROSSREF
- 45. Zhang X, Wang J, Li J, Yu Y, Song Y. A positive association between dietary sodium intake and obesity and central obesity: results from the National Health and Nutrition Examination Survey 1999-2006. Nutr Res 2018;55:33-44.



- 46. Appel LJ, Espeland MA, Easter L, Wilson AC, Folmar S, Lacy CR. Effects of reduced sodium intake on hypertension control in older individuals: results from the Trial of Nonpharmacologic Interventions in the Elderly (TONE). Arch Intern Med 2001;161:685-93. PUBMED | CROSSREF
- Bulpitt CJ, Daymond M, Bulpitt PF, Ferrier G, Harrison R, Lewis PJ, Dollery CT. Is low salt dietary advice a useful therapy in hypertensive patients with poorly controlled blood pressure? Ann Clin Res 1984;16 Suppl 43:143-9.
- Geleijnse JM, Witteman JC, Bak AA, den Breeijen JH, Grobbee DE. Reduction in blood pressure with a low sodium, high potassium, high magnesium salt in older subjects with mild to moderate hypertension. BMJ 1994;309:436-40.
   PUBMED | CROSSREF
- Kumanyika SK, Hebert PR, Cutler JA, Lasser VI, Sugars CP, Steffen-Batey L, Brewer AA, Cameron M, Shepek LD, Cook NR, et al. Feasibility and efficacy of sodium reduction in the Trials of Hypertension Prevention, phase I. Hypertension 1993;22:502-12.
- Nowson CA, Wattanapenpaiboon N, Pachett A. Low-sodium dietary approaches to stop hypertension-type diet including lean red meat lowers blood pressure in postmenopausal women. Nutr Res 2009;29:8-18.
   PUBMED | CROSSREF
- Petersen KS, Torpy DJ, Chapman IM, Guha S, Clifton PM, Turner K, Keogh JB. Food label education does not reduce sodium intake in people with type 2 diabetes mellitus. A randomised controlled trial. Appetite 2013;68:147-51.
  - PUBMED | CROSSREF
- Beard TC, Cooke HM, Gray WR, Barge R. Randomised controlled trial of a no-added-sodium diet for mild hypertension. Lancet 1982;2:455-8.
   PUBMED | CROSSREF
- Dodson PM, Beevers M, Hallworth R, Webberley MJ, Fletcher RF, Taylor KG. Sodium restriction and blood pressure in hypertensive type II diabetics: randomised blind controlled and crossover studies of moderate sodium restriction and sodium supplementation. BMJ 1989;298:227-30.
   PUBMED | CROSSREF
- 54. Gilleran G, O'Leary M, Bartlett WA, Vinall H, Jones AF, Dodson PM. Effects of dietary sodium substitution with potassium and magnesium in hypertensive type II diabetics: a randomised blind controlled parallel study. J Hum Hypertens 1996;10:517-21. PUBMED
- 55. He FJ, Wu Y, Feng XX, Ma J, Ma Y, Wang H, Zhang J, Yuan J, Lin CP, Nowson C, et al. School based education programme to reduce salt intake in children and their families (School-EduSalt): cluster randomised controlled trial. BMJ 2015;350:h770. PUBMED | CROSSREF
- 56. Nouvenne A, Meschi T, Prati B, Guerra A, Allegri F, Vezzoli G, Soldati L, Gambaro G, Maggiore U, Borghi L. Effects of a low-salt diet on idiopathic hypercalciuria in calcium-oxalate stone formers: a 3-mo randomized controlled trial. Am J Clin Nutr 2010;91:565-70. PUBMED | CROSSREF
- Hypertension Prevention Trial Research Group. The Hypertension Prevention Trial: three-year effects of dietary changes on blood pressure. Arch Intern Med 1990;150:153-62.
   PUBMED | CROSSREF
- Staessen J, Bulpitt CJ, Fagard R, Joossens JV, Lijnen P, Amery A. Salt intake and blood pressure in the general population: a controlled intervention trial in two towns. J Hypertens 1988;6:965-73.
   PUBMED | CROSSREF
- Takahashi Y, Sasaki S, Okubo S, Hayashi M, Tsugane S. Blood pressure change in a free-living populationbased dietary modification study in Japan. J Hypertens 2006;24:451-8.
   PUBMED | CROSSREF
- 60. The Trials of Hypertension Prevention Collaborative Research Group. Effects of weight loss and sodium reduction intervention on blood pressure and hypertension incidence in overweight people with high-normal blood pressure. The Trials of Hypertension Prevention, phase II. Arch Intern Med 1997;157:657-67. PUBMED | CROSSREF
- Ard JD, Coffman CJ, Lin PH, Svetkey LP. One-year follow-up study of blood pressure and dietary patterns in dietary approaches to stop hypertension (DASH)-sodium participants. Am J Hypertens 2004;17:1156-62.
   PUBMED | CROSSREF
- Sakaki M, Tsuchihashi T, Arakawa K. Characteristics of the hypertensive patients with good and poor compliance to long-term salt restriction. Clin Exp Hypertens 2014;36:92-6.
   PUBMED | CROSSREF



- 63. Takahashi Y, Sasaki S, Okubo S, Hayashi M, Tsugane S. Maintenance of a low-sodium, high-carotene and -vitamin C diet after a 1-year dietary intervention: the Hiraka dietary intervention follow-up study. Prev Med 2006;43:14-9.
  PUBMED | CROSSREF
- 64. Kim J, Lim G, Kang S, Lee K, Park T, Kim J. The relationship between daily sodium intake and obesity in Korean adults. Korean J Health Promot 2015;15:175-84.
- Lim S, Yang S. Association between dietary sodium intake and abdominal obesity in pre-diabetes Korean adults. J Korean Soc Food Sci Nutr 2014;43:763-71.
- 66. Oh H, Kim H, Jun D, Lee S. Associations between 24-hour urine sodium excretion level and obesityrelated metabolic risk factors. Korean J Community Nutr 2015;20:460-7.
- Lee M, Sorn SR, Lee Y, Kang I. Salt induces adipogenesis/lipogenesis and inflammatory adipocytokines secretion in adipocytes. Int J Mol Sci 2019;20:160.
   PUBMED | CROSSREF
- 68. Lanaspa MA, Kuwabara M, Andres-Hernando A, Li N, Cicerchi C, Jensen T, Orlicky DJ, Roncal-Jimenez CA, Ishimoto T, Nakagawa T, et al. High salt intake causes leptin resistance and obesity in mice by stimulating endogenous fructose production and metabolism. Proc Natl Acad Sci U S A 2018;115:3138-43. PUBMED | CROSSREF
- Fonseca-Alaniz MH, Takada J, Andreotti S, de Campos TB, Campaña AB, Borges-Silva CN, Lima FB. High sodium intake enhances insulin-stimulated glucose uptake in rat epididymal adipose tissue. Obesity (Silver Spring) 2008;16:1186-92.
- Fonseca-Alaniz MH, Brito LC, Borges-Silva CN, Takada J, Andreotti S, Lima FB. High dietary sodium intake increases white adipose tissue mass and plasma leptin in rats. Obesity (Silver Spring) 2007;15:2200-8.

- Zhang Y, Li F, Liu FQ, Chu C, Wang Y, Wang D, Guo TS, Wang JK, Guan GC, Ren KY, et al. Elevation of fasting ghrelin in healthy human subjects consuming a high-salt diet: a novel mechanism of obesity? Nutrients 2016;8:323.
   PUBMED | CROSSREF
- 72. Mähler A, Klamer S, Maifeld A, Bartolomaeus H, Markó L, Chen CY, Forslund SK, Boschmann M, Müller DN, Wilck N. Increased salt intake decreases diet-induced thermogenesis in healthy volunteers: a randomized placebo-controlled study. Nutrients 2022;14:253. PUBMED | CROSSREF
- 73. Cogswell ME, Maalouf J, Elliott P, Loria CM, Patel S, Bowman BA. Use of urine biomarkers to assess sodium intake: challenges and opportunities. Annu Rev Nutr 2015;35:349-87. PUBMED | CROSSREF
- 74. Campbell NR, He FJ, Tan M, Cappuccio FP, Neal B, Woodward M, Cogswell ME, McLean R, Arcand J, MacGregor G, et al. The International Consortium for Quality Research on Dietary Sodium/Salt (TRUE) position statement on the use of 24-hour, spot, and short duration (<24 hours) timed urine collections to assess dietary sodium intake. J Clin Hypertens (Greenwich) 2019;21:700-9.</p>
  PUBMED | CROSSREF
- 75. Bie P. Mechanisms of sodium balance: total body sodium, surrogate variables, and renal sodium excretion. Am J Physiol Regul Integr Comp Physiol 2018;315:R945-62.
  PUBMED | CROSSREF
- Huang L, Crino M, Wu JH, Woodward M, Barzi F, Land MA, McLean R, Webster J, Enkhtungalag B, Neal B. Mean population salt intake estimated from 24-h urine samples and spot urine samples: a systematic review and meta-analysis. Int J Epidemiol 2016;45:239-50.
   PUBMED | CROSSREF
- 77. Ji C, Sykes L, Paul C, Dary O, Legetic B, Campbell NR, Cappuccio FP; Sub-Group for Research and Surveillance of the PAHO–WHO Regional Expert Group for Cardiovascular Disease Prevention Through Population-wide Dietary Salt Reduction. Systematic review of studies comparing 24-hour and spot urine collections for estimating population salt intake. Rev Panam Salud Publica 2012;32:307-15. PUBMED | CROSSREF