

**Original Paper** 

# **Evaluating the Urinary Iodine Concentration in Operating Room Personnel After Using Povidone-iodine Hand Scrub**





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# **ABSTRACT**

**Introduction:** Studies have shown conflicting results regarding the skin absorption of iodine from povidone-iodine solution (Betadine 7.5%) and its effects on the health of operating room personnel.

**Objective:** This study aimed to measure the urinary iodine concentration (UIC) among the operating room personnel of a hospital in Isfahan, Iran, after using povidone-iodine scrub (Betadine 7.5%).

**Materials and Methods:** This descriptive-analytical study was conducted among operating room personnel of a selected hospital in Isfahan, Iran. Using a convenience sampling method, 100 participants were selected and divided into two groups: Povidone-iodine scrub (n=50) and alcohol-based scrub (n=50). The scrubs were used for one week, three times per day, each for three minutes. One week after the study, morning urine samples were collected to measure the UIC using the Sandell-Kolthoff reaction method. The Shapiro-Wilk test assessed the normality of data distribution, and statistical comparisons between the two groups were conducted using the Mann-Whitney U test and chi-square test.

**Results:** The mean age of the povidone-iodine scrub group and the alcohol-based scrub group were  $31.9\pm6.29$  and  $31.7\pm5.59$  years, respectively. Also, 94% in the povidone-iodine scrub group and 86% in the alcohol-based scrub group were female. No significant differences were observed between the two groups in age, work experience, gender, educational level, scrubbing frequency, or number of shifts per week. The results showed a significantly higher UIC in the povidone-iodine scrub group compared to the alcohol-based scrub group (290.0±0.73 vs 106.8±27.88 μg/L; P=0.001). Additionally, the prevalence of iodine poisoning (>300 μg/L) was higher in the povidone-iodine group (8% vs 0%; P=0.001). Furthermore, in the povidone-iodine scrub group, the UIC was significantly higher in individuals with hand scrubbing frequency ≥10 per week compared to those with fewer frequency (316.3±54.20 vs 196.5±54.00 μg/L, P=0.001) and in those who worked ≥6 shifts per week (299.3±6.9 vs 206.2±81.6 μg/L; P=0.011).

**Conclusion:** This study suggests that hand scrubbing with povidone-iodine (Betadine 7.5%) can increase the UIC and the risk of iodine poisoning in operating room personnel. Therefore, it is recommended to consider alternative solutions, such as alcohol-based solutions, for hand scrubbing in operating rooms.

## Keywords:

Operating rooms, Povidone-lodine, Hand scrub, lodine, Urine

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# **Highlights**

- Hand scrubbing with povidone-iodine can increase urinary iodine concentration in operating room personnel.
- Hand scrubbing with povidone-iodine can increase the risk of iodine poisoning in operating room personnel.
- Urinary iodine concentration in the povidone-iodine scrub group was significantly higher than in the alcohol-based scrub group.

# **Plain Language Summary**

This study demonstrated that urinary iodine concentration was significantly higher in the group using povidone-iodine scrub compared to the group using alcohol-based scrub. In this study, a large proportion of individuals in the povidone-iodine group had very high iodine levels, indicative of iodine toxicity, while none in the control group exhibited such levels. A statistically significant correlation was observed between the frequency of scrubbing and increased urinary iodine concentration. The findings also indicated that more work shifts were associated with increased iodine concentration. These results suggest that operating room personnel should utilize safer alternatives, such as alcohol-based solutions.

# Introduction

ffective hand scrubbing and disinfection are crucial, particularly before wearing sterile gloves for surgery, as a study found that surgical gloves have perforations in 5.9% of cases [1, 2]. There are two main types of hand sanitizers, including alcoholic and non-alcoholic (povidone-iodine, hexachlorophene, Chlorhexidine gluconate and Hibitane) [3-5]. The materials used for hand sanitizing, particularly in surgical operations, are diverse. Povidone-iodine solution (7.5% betadine) is the most well-known and widely used solution [6]. As a topical antiseptic agent in the form of an iodophor, povidone-iodine reduces the irritating effects of iodine [7]. One of its key characteristics is its high bactericidal activity, which is capable of killing a broad spectrum of microorganisms within 3-5 minutes due to the presence of iodine [8]. However, excessive iodine consumption can lead to poisoning in individuals, with a tolerable intake level of 1100 µg/day for people aged >19 years [9]. The complications and symptoms of iodine poisoning are diverse, ranging from gastrointestinal symptoms (headache, dizziness and nausea) to metabolic and hormonal disorders, including thyroid gland disorders [10, 11].

Despite the limited and negligible side effects of povidone-iodine (7.5% betadine) in case of prolonged use (approximately ≥3 hours), this substance has demonstrated acceptable efficacy [11, 12]. Consequently, it remains a widely used agent in surgical rooms for hand sanitizing [13]. Povidone-iodine exhibits a broad-

spectrum antimicrobial effect, displaying high sensitivity against gram-negative and gram-positive bacteria, acidophiles, fungi, amoebic cysts and spores [14-16]. Research has shown that prolonged skin exposure to iodine (≥3 hours) can result in limited absorption [17, 18]. Additionally, the widespread use of amiodarone and iodinated radiological contrast agents has been found to expose the body to high doses of exogenous iodine [7].

The routine use of povidone-iodine solution may be a potential cause of iodine poisoning in operating room personnel, as evidenced by elevated urinary iodine levels [19]. A study revealed that the average urinary iodine level one week after using povidone-iodine for sanitizing was significantly higher than the pre-use period [20]. Furthermore, some studies have suggested that povidone-iodine solution can be absorbed through the skin or mucous membrane due to its high iodine content [17, 18]. However, contradictory results have been reported in other studies [17, 21]. For instance, a study found that the urinary iodine level of operating room personnel in Turkey was at an average level, despite the regular use of povidone-iodine for hand sanitizing, and it did not show elevated levels [21]. This finding highlights the conflicting evidence concerning the skin absorption of iodine from povidone-iodine solutions and its potential impact on the health of operating room personnel [17, 20, 21]. This study aimed to measure the urinary iodine concentration (UIC) in operating room personnel of a hospital in Isfahan, Iran, who used povidone-iodine solution (betadine 7.5%) for hand scrubbing.



### **Materials and Methods**

This is a descriptive-analytical study. A total of 100 operating room personnel (technologists and scrub nurses) were selected from a hospital affiliated to Isfahan University of Medical Sciences in 2022, using a convenience sampling method. The inclusion criteria were consent to participate, no wounds on hand skin up to 5 cm above the elbow, self-report of no skin dermatitis, not receiving iodinated contrast agents during the study, no history of taking certain medications or supplements related to iodine, no history of iodine deficiency or thyroid disorders, no history of thyroidectomy and not following a special diet that includes excessive seafood (≥3 times a week). The exclusion criteria were unwillingness to continue participation, development of skin wounds or scratches during the study, and inconsistent use of povidone-iodine and alcohol solutions for scrubbing. To determine the sample size, the formula for estimating proportions was used [18]. By considering a maximum estimation error of 0.05 at a confidence level of 95%  $(Z1-\alpha/2=1.96)$  and a precision (d) of 7.5%, the minimum sample size was calculated to be 100 (50 per group). The study hospital has two operating rooms. We selected 50 personnel from one operating room and assigned them to the povidone-iodine scrub group, while 50 personnel from the second operating room were assigned to the alcohol-based scrub group. The work-related and demographic data were collected using a form surveying age, gender, work experience, educational level, frequency of scrubbing per week, and number of shifts per week. The UIC was measured for both groups using the Sandell-Kolthoff reaction method. This method is popular for its accuracy and sensitivity in measuring urinary iodine, showing reliable results [17, 18].

Before the study, participants were trained on the standard surgical scrub method, which involved scrubbing the four surfaces of each finger, the outside of the fifth finger, the back of the hand, the palm, from the little finger to the thumb, from the wrist to the arm (three times), and up to 5 cm above the elbow [22]. The betadine used during the study was made by a pharmaceutical company that had a contract with the hospital (The participants, the person responsible for measuring the UIC in the laboratory and the researchers did not have information about the name of the company). The betadine was poured into the betadine storage containers in the scrub rooms by the person in charge of operating room equipment. In both groups, scrubbing was performed for one week, three times a day, each for three minutes, using 3-5 mL of povidone-iodine or

alcoholic solution. All procedures were carried out under the supervision of the researcher. Participants were instructed to maintain their usual diet and avoid using chemical or herbal medications during this period.

One week after the study, morning urine samples were collected from the participants on a single day. Prior to sampling, comprehensive instructions on urine sampling procedures (initially washing the external urinary tract with water and then collecting mid-stream urine) were provided by the researcher. These instructions were given individually and face-to-face in a private area within the operating rooms. If any confusion arose, further explanations were provided to the participants. Mmidstream urine samples were placed in deionized test tubes with lids shut tightly, placed in cool boxes, maintained at a temperature of 2-8 °C, and transported to the Comprehensive Research Laboratory of Isfahan University of Medical Sciences immediately. The samples were acidified and stored under -20 °C until analysis. The UIC was subsequently measured by the modified Sandell-Kolthoff method using a microplate technique, adhering to the recommendations of the centers for disease control and prevention. The measurement was done by a laboratory expert, and the researcher was responsible for ensuring the accuracy of this measurement. The urine samples were initially dissolved at room temperature and then incubated for 30 minutes with 0.9 M ammonium persulfate at 95 °C using a thermal cycler. After cooling, 50 µL of the processed urine samples were placed into the wells of a microplate. Subsequently, 100 µL of a 0.05 M arsenious acid solution was added to each well and mixed properly. In the final step, 50 µL of a 0.019 M ceric ammonium sulfate solution was added. After a 30-min incubation at 25 °C, absorbance was measured at 405 nm. The UIC was calculated by plotting a standard curve and reported in micrograms per liter (μg/L) [17, 23]. A UIC≤300 μg/L shows normal range, and >300 μg/L indicates a potential risk for abnormal thyroid function [17].

Data analysis was performed in SPSS software, version 25 (IBM Corporation, Chicago, IL). The normality of data distribution was assessed using the Shapiro-Wilk test. Quantitative data were presented as Mean±SD or median ±95% interquartile range. The differences between groups were analyzed using the Chi-square test or Mann-Whitney U-test (for non-parametric variables). The significance level was set at 0.05.



Table 1. Work-related and demographic information of the participants

		No.		
Variab	les	Gro	P*	
		Povidone iodine	idone iodine Alcohol-based	
Age (y)	18-29	20(40)	24(48)	0.420
	30-59	30(60)	26(52)	0.420
Gender	Male	3(6)	7(14)	0.182
	Female	47(94)	43(86)	0.102
Work experience (y)	≥5	39(78)	39(78)	0.999
	>5	11(22)	11(22)	0.555
Educational level	Bachelor's degree	46(92)	47(94)	0.695
	Masters' degree	4(8)	3(6)	0.093
Frequency of scrubbing (per week)	<10	11(22)	13(26)	0.640
	≥10	39(78)	37(74)	0.640
Number of shifts (per week)	<6	5(10)	6(12)	0.749
	≥6	45(90)	44(88)	0.743

<sup>\*</sup>Chi-square test

# Results

A comparison of the povidone-iodine scrub group with the alcohol-based scrub group revealed no significant differences in terms of age (31.9 $\pm$ 6.29 vs 31.7 $\pm$ 5.59 years), work experience (3.90 $\pm$ 1 vs 4.08 $\pm$ 1.69 years), gender (94% vs 86% female), educational level (92% vs 94% with a bachelor's degree), frequency of scrubbing (78% vs 74% with >6 shifts per week) and shifts per week (Table 1). However, the UIC was significantly higher in the povidone-iodine scrub group (289.9 $\pm$ 73.19 vs 106.6 $\pm$ 27.88  $\mu$ g/L; P=0.001).These findings were shown in Table2. Furthermore, the UIC at a level indicating excessive iodine in-

take (>300  $\mu$ g/L) was significantly higher in the povidone-iodine scrub group compared to the alcohol-based scrub group (8% vs 0%; P=0.001) that was shown in Table 3.

In the povidone-iodine scrub group, the UIC was significantly higher in females (296.3±70.82 vs 190.1±0.95 µg/L; P=0.032), in those who scrubbed  $\geq$ 10 times per week (316.3±54.20 vs 196.5±52.54 µg/L; P=0.001), and in those who worked  $\geq$ 6 shifts per week (299.3±66.9 vs 206.2±81.6 µg/L; P=0.011). However, no significant differences were found in the UIC based on age, work experience, or educational level. In the alcohol-based scrub group, no significant differences were observed in the UIC based on any factors (Table 4).

 $\textbf{Table 2.} \ \textbf{The mean UICs in the study groups}$ 

Comme	UIC (μg/L)		
Groups	Min	Max	Mean±SD
Povidone iodine	159.4	400.1	289.97±73.19
Alcohol-based	50.4	114.85	106.82±27.88
P*		0.001	

<sup>\*</sup>Mann-Whitney U test

UIC: Urinary Iodine concentratio.



Table 3. The UIC in the study groups categorized based on frequency

	No. (	P*	
UIC (μg/L)	Group		
	Povidone iodine	Alcohol-based	
Normal level (≤300)	46(92)	50(100)	0.041
Excessive intake level (>300)	4(8)	0	0.001

<sup>\*</sup>Mann-Whitney U test

UIC: Urinary Iodine concentration.

#### Discussion

This study demonstrated that, among operating room personnel, the UIC in the povidone-iodine scrub group was significantly higher than in the alcohol-based scrub group. A similar study reported that rinsing the mouth with povidone-iodine increased the serum iodine level by 2-3 times and the UIC by 5 times [20]. Our results are also in line with previous studies, which showed that using hand sanitizers containing iodine increased UIC and suggested that operating room personnel should take

necessary measures to control iodine exposure [17, 21]. Our findings differed from these studies; we compared an alcohol-based solution with povidone-iodine, while these two studies only examined povidone-iodine. A laboratory study by Nesvadbova et al. demonstrated that iodine in povidone-iodine solution can be absorbed through skin cells to a significant extent [18]. The authors suggested that frequent professional contact with the povidone-iodine solution as a skin scrub can lead to iodine penetration through the skin. Our study's findings are in agreement with the results of this study.

Table 4. The UICs in the study groups based on work-related and demographic factors

		UIC (µg/L)			
Variables	Povidone Iodine Scrub Group		Alcohol-based Scrub Group		
		Mean±SD	P*	Mean±SD	P*
Ago (v)	18-29	283.67±77.65	0.692	107.05±29.32	0.930
Age (y)	30-59	294.17±71.10	0.692	106.60±27.08	
	Male	190.17±0.95	0.032	65.40±5.73	0.061
Gender	Female	296.34±70.82		92.56±23.78	
Made our oil or or (a)	≥5	298.13±71.21	0.770	105.72±26.86	0.264
Work experience (y)	>5	292.94±83.46	0.779	110.72±32.33	0.361
51 11	Bachelor's degree	293.80±73.56		106.54±28.67	0.713
Educational level	Masters' degree	245.97±59.29	0.224	111.17±10.05	
[	<10	196.48±52.54	0.001	64.71±7.60	0.094
Frequency of scrubbing (per week)	≥10	316.34±54.20	0.001	121.61±13.16	
Number of shifts (source-1)	<6	206.20±81.61	0.011	59.28±5.35	0.063
Number of shifts (per week)	≥6	299.28±66.89	0.011	113.30±22.89	

<sup>\*</sup>Mann-Whitney U test

UIC: Urinary Iodine concentration.



Our study revealed that 80.0% of operating room personnel in the povidone-iodine scrub group had extremely high intake levels, indicating iodine poisoning, whereas none of the participants in the control group had excessive intake levels. Idiz et al. in Turkey reported that 7.5% of surgical staff who scrubbed with povidoneiodine had iodine poisoning [17]. Our findings are consistent with their findings. However, Erdoğan et al. in Turkey reported a higher prevalence of excessive intake level (approximately 39%) among operating room staff who scrubbed with povidone-iodine [21], which is nearly five times the prevalence observed in our study. This discrepancy may be due to difference in the follow-up periods; in our study, the UIC was assessed after one week, whereas Erdoğan et al. [21] evaluated it after three months, which may have led to a cumulative increase in UIC.

Our study revealed a significant difference in the UIC based on the frequency of scrubbing in the povidoneiodine scrub group. Those with ≥10 scrubs per week exhibited significantly higher UICs compared to others in this group. Nesvadbova et al. in Italy demonstrated a significant statistical relationship between UIC and the number of povidone-iodine scrubs, where surgical staff performing ≥20 scrubs per week displaying higher UIC than others [18]. Our findings are consistent with their results. In our study, in the povidone-iodine scrub group, those with ≥ 10 shifts per week exhibited significantly higher UIC compared to those with <10 shifts per week. Similar to our findings, Idiz et al. in Turkey reported that operating room staff who worked at least four consecutive shifts per week and used povidone-iodine for hand scrubbing had higher UIC [17]. In contrast, Michalaki et al. in Greece found no significant correlation between UIC and the number of working days among operating room staff using povidone-iodine scrub [24]. This discrepancy may be due to differences in UIC measurement methods and sample sizes among the studies.

This study had some limitations. The UIC was not measured before the study, which can potentially affect the results. The sampling was done by a convenience method, which may lead to selection bias. Other factors, such as dietary habits, were not fully examined which may affect the study results. Also, the generalizability of the results should be done with caution. Further interventional studies are recommended.

Based on the findings, we recommend the operating room personnel to use alternative hand scrubbing solutions, such as alcohol-based solutions. The use of povidone-iodine scrub can increase UIC in operating room

personnel, particularly in those at risk of iodine toxicity or with a history of thyroid disorders, such as pregnant women. These individuals may need to consider alternative hand scrubbing solutions or reduce their frequency of povidone-iodine scrub use.

## **Ethical Considerations**

#### **Compliance with ethical guidelines**

This study obtained ethical approval from the Ethics Committee of Isfahan University of Medical Sciences, Isfahan, Iran (Code: IR.ARI.MUI.REC.1401.047). Informed written consent was obtained from all participants after describing the study objectives. They were free to leave the study at any time.

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## **Authors' contributions**

Conceptualization, study design, review & editing: Fateme Karbalai and Soheila Bakhtiari; Methodology: Soheila Bakhtiari and Farzaneh Jafari; Sampling, writing the original daft: Fateme Karbalai; Data analysis: Fateme Karbalai and Farzaneh Jafari; Final approval: All authors.

#### **Conflict of interest**

The authors declared no conflict of interest.

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