# **RESEARCH NOTE**





# Excess urinary iodine concentration and thyroid dysfunction among school age children of eastern Nepal: a matter of concern

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# Abstract

**Objectives:** Deficiency as well as excess dietary iodine is associated with several thyroid disorders including Grave's disease and goitre. Previously, cross sectional studies conducted among school children in Nepal showed high prevalence of iodine deficiency. In contrast, recently, few studies have revealed emerging trends of excess urinary iodine concentration in children. This paper, reports excess urinary iodine excretion and thyroid dysfunction among school age children from eastern Nepal.

**Results:** It was a community based cross sectional study in which we measured urinary iodine excretion levels among school age children at baseline and after educational program. The educational program consisted of audio-visual and pamphlets on thyroid health. We also screened them for thyroid function status by physical examination and measuring serum thyroid hormones. Our results show that 34.4% of the children had excess urinary iodine concentration above the WHO recommended levels. Overall, 3.2% of the children were identified to have thyroid dysfunction. Urinary iodine concentration was significantly different between types of salt used and between salt iodine content categories.

Keywords: Urinary iodine, Salt iodine, Deficiency, Children, Thyroid, Excess, Nepal

# Introduction

Iodine deficiency disorders (IDD) are still major public health problems in developing countries although significant progress has been made towards their control strategies [1]. Urinary iodine is a sensitive marker of recent dietary iodine intake but not of thyroid dysfunction. Spot urine sample is generally recommended for population based studies [2–4]. Total goitre rate in Nepal has dramatically reduced to 0.4% in 2007 from 55% in 1965 [5]. Although iodine deficiency (ID) and prevalence of goitre have been well evidenced cases, association between excess iodine intake and thyroid disease have been found [6]. Few recently conducted studies in Nepal have demonstrated increasing trends of excess urinary iodine concentration (UIC) in children [6, 7]. In addition, latest Nepal

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National Micronutrient Status Survey 2016 has revealed that median urinary iodine concentration (mUIC) in 6-9 years children was above 300 µg/L [8]. This report along with few other studies point out possibility of chronic excess iodine intake in Nepalese children. Acute iodine excess in new-borns can cause hypothyroidism but limited data are available on the effects of chronic high intake of iodine in thyroid function [9]. The current study was aimed to assess iodine status among school age children (SAC) and correlate it with salt iodine content (SIC), type of salt used and gender of the children. Our other aim was to check improvement of urinary iodine excretion (UIE) levels after educational program.

# Main text Methods

This community based study approached 1042 SAC from twelve government schools. These schools were selected randomly from six Village Development Committees (VDC) of the Udaypur District, Province no. 1, Nepal

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such that two schools were included from each VDC. The study commenced in October 2015 and completed in July 2017. Spot urine samples and household salt samples were collected from 946 children. Blood samples were obtained from 245 children. The urine samples were used to determine UIC whereas salt samples were used to assess household SIC. The salt samples were classified as open type and packet (iodized) type. UIC was determined by the ammonium persulfate digestion microplate (APDM) method using the Sandell-Kolthoff reaction in a specially designed apparatus, sealing cassette (Hitachi, Japan) [10] in microplate format. SIC was determined by the iodometric titration method. Blood samples were analysed for thyroid hormones (free thyroxine and thyroid stimulating hormone). Thyroid stimulating hormone (TSH) and free thyroxine (fT4) were measured using the HUMAELISA kit (Human Diagnostics, Germany) following the manufacturer's manual. Thyroid function status was determined according to the reference range provided in the kit, which reported TSH 0.39-6.16 mIU/L and fT4 0.8-2.2 ng/dL. Sub clinical hypothyroidism is considered if TSH>6.16 mIU/L with fT4 in the normal range whereas subclinical hyperthyroidism is considered if TSH < 0.39 mIU/L with fT4 in the normal range. Ethical approval for this research was taken from Institutional Review Committee, B. P. Koirala Institute of Health Sciences (BPKIHS) (IRC/655/015). Informed consent was taken from school teacher, head teacher and concerned guardians before samples were collected. To compare mean difference between two groups, Mann Whitney U test was used while for comparing three groups, Kruskal Wallis test was used.

# Educational program

After the baseline survey, a series of educational programs were organized targeting the schools. The educational program consisted of an awareness of iodine and thyroid through pamphlets, posters and audio-visual aids. Information on iodized salts use and its importance

Table 2	Comparison	of	UIC	within	SIC,	gender	and	type
of salt u	sed							

Variables	Categories	Mean rank	p-value
UIC	Salt iodine conten	0.000 <sup>a</sup>	
	0	191.02	
	<15	241.54	
	>15	485.43	
UIC	Gender		0.640 <sup>b</sup>
	Воу	469.16	
	Girl	477.47	
UIC	Type of salt used		0.000 <sup>b</sup>
	Open salt	169.79	
	Packet salt	483.52	

<sup>a</sup> Kruskal Wallis test

<sup>b</sup> Mann-whitney U test

in health was provided to the SAC. The school teachers were trained on iodine, thyroid diseases, its preliminary identification in community setting and the importance of iodized salt. These programmes were organized twice in each school. After 1 year of the educational program, we again assessed UIC in the same cohort of school children, achieving a total of 311 urine samples.

# Results

Results showed that mUIC among the children at baseline was 231 µg/L (IQR: 156–333) which lies in the more than adequate level. We found that 9.8% of the SAC had iodine deficiency; followed by 23.5% of the SAC who had more than adequate UIC. Most importantly, 34.4% of children had excess UIC (i.e.  $\geq$  300 µg/L). Table 1 provides the full details. Open type salt was used by 9.3% of the children while majority of them used iodized salt distributed by Salt Trading Corporation Nepal. Salt iodine content (SIC) determination showed that 11.8% of the salt samples had iodine content of 0–15 ppm whereas 88.2% contained more than 15 PPM. As presented in Table 2, UIC was not significantly different between boys

	Table 1	UIC among the school children ( $n = 946$ ,	before and $n = 311$ after educ	ational program
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Urinary iodine concentration (UIC) status	Before educational	program	After educational program		
	Frequency (n)	Percentage	Frequency (n)	Percentage	
Severe ID (<20 μg/L)	4	0.4	2	0.6	
Mild ID (20–49 µg/L)	22	2.3	12	3.9	
Moderate ID (50–99 µg/L)	67	7.1	47	15.1	
Adequate (100–199 μg/L)	306	32.3	94	30.2	
More than adequate (200–299 µg/L)	222	23.5	54	17.4	
Excess ( $\geq$ 300 µg/L)	325	34.4	102	32.8	
Total	946	100	311	100	

and girls (p=0.64), but significantly different between the SIC categories (p < 0.05) and between type of salt used (p < 0.05). As shown in Fig. 1, 3.2% of the SAC had thyroid dysfunction with either hypothyroidism or hyperthyroidism. The assessment of thyroid hormones revealed that 1.6% of the SAC had subclinical hyperthyroidism and one child had overt hyperthyroidism. Overt and subclinical hypothyroidisms were identified in two children and one child respectively. Further, urinary iodine measurement after the educational program revealed that the proportion of excess UIC slightly decreased to 32.8% from baseline of 34.4%. After one year of educational program, mUIC in school children (n=311) was 201 µg/L, having found to decrease from baseline of 231 µg/L.

# Discussion

The present study highlighted excess urinary iodine excretion levels in SAC from eastern Nepal. Sufficient sample size is one of the big strengths of our study. In addition, physical examination of the children and biochemical test further strengthen our results. Similar to our findings, two studies from Nepal have reported high prevalence of excess UIC among school children with one study reporting median UIC of 291.8 µg/L (IQR 181.3–411.5 µg/L [7, 9]. A survey conducted in Nepal in 2007 demonstrated mUIC of 203 µg/L [11]. A cross sectional study from India in 2013 found a similar mUIC of  $200 \ \mu g/L$  [12]. This implies that the population is in transition phase from iodine deficiency to iodine sufficiency. The latest Iodine Global Network report has highlighted several countries around the world to be categorized as having excess iodine status in population [11].

The time has now come to monitor the iodine status at national level and adjust policies accordingly. Together with our current results, consistent findings of excess



UIC among school children imply chronic exposure to high dietary iodine intake. A reason for this may be due to poor awareness among SAC on the optimal consumption of iodized salt. Secondly, awareness regarding consumption of iodized salt among the households need to be improved with regard to optimal intake. In addition, poor monitoring of salt distribution and household salt consumption after initiation of salt iodization program by the government is probably a significant factor underlying this outcome. This is a matter of concern as high intake of dietary iodine could result in several thyroid disorders. This warrants a further large scale cross sectional as well as cohort studies comprising several districts to assess the iodine status of the population of the country.

# Conclusion

Based on our results, we conclude that there is a high prevalence of excess urinary iodine concentration among school age children of eastern Nepal. Since almost 88% of the household salt samples tested were found to have SIC of more than 15 PPM, we suggest the young children and their family to consume optimal salt in their diet. This may prevent them from iodine induced thyroid disorders. Also, the study recommends to explore further regarding excess iodine status in nationally representative population. Our findings suggest that programs and policies should be developed to explore and address this issue further.

# Limitations

The study has a number of limitations. First, we could not collect blood samples from all the participating school children due to refusal from their parents. Second, reassessment of urinary iodine concentration after educational program could only enrol about 30% baseline population, which might limit its generalization. Third, free tri-iodothyronine (fT3) concentration was not measured in the blood samples.

#### Abbreviations

UIC: urinary iodine concentration; SIC: salt iodine content; UIE: urinary iodine excretion; SAC: school age children; IDD: iodine deficiency disorder; VDC: village development committee; APDM: ammonium persulfate digestion microplate; BPKIHS: B. P. Koirala Institute of Health Sciences.

## Acknowledgements

Not applicable.

# Authors' contributions

BG and NB designed the study. NB was awarded the grant. MKT, BG and BT carried out the field study, data collection and laboratory analysis. MKT, BG, ML and DB prepared the manuscript in combination. All authors read and approved the final manuscript.

## Funding

We would like to sincerely thank Rotary Foundation for the project grant (Grant Number GG1412765). The funding body had no role in design of study, data collection, analysis and interpretation of data.

## Availability of data and materials

The data set used and analysed in this study are available from the corresponding author upon reasonable request.

#### Ethics approval and consent to participate

Ethical approval for this study was taken from Institutional Review Committee, BPKIHS (IRC/655/015). Prior written consent was taken from the guardian/ teacher before enrolling them into the study.

#### Consent to publish

Not applicable.

# Competing interests

The authors declare that they have no competing interests.

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Received: 23 January 2019 Accepted: 22 May 2019 Published online: 27 May 2019

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