Iodine Deficiency Disorders Continue to be a Problem in the Asia Pacific Region

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Abstract

Iodine deficiency disorders (IDD) due to inadequate dietary iodine is one of the major non-communicable diseases yet to be eliminated from our world, particularly in the Asia Pacific region. There has been remarkable progress in reducing iodine deficiency over the past two decades, with 50 more countries declaring successful elimination mainly due to effective implementation of universal salt iodization (USI) programs. The most devastating—but often the least visible—effect of IDD is damage to the central nervous system, causing a spectrum of impairment from mild loss of intelligence quotient (IQ), to severe, irreversible brain damage called neurological cretinism.

The majority of nations in the vast expanse of the Asia Pacific region, encompassing small Pacific island states to the most populous countries in the world, continue to suffer from the health and socio-economic consequences of IDD. Contrary to popular belief, IDD is not confined to remote mountainous regions and economically disadvantaged populations, but may also affect wealthy, industrialized societies such as Australia and New Zealand. Japan and China are examples of countries relatively free of IDD; the former because of a diet naturally rich in iodine, and the latter due to the implementation of a sustainable USI program. While USI is the preferred strategy for sustainable elimination of IDD, it is not always practicable. As such, countries need to consider other methods of iodine supplementation or food fortification.

Keywords: iodine deficiency, goitre, hypothyroidism

INTRODUCTION

The World Health Organization (WHO) has recently released updated global and country population statistics on the prevalence of iodine deficiency. In 2005, the World Health Assembly (WHA) resolved that member states should monitor and report the iodine nutrition status of their respective populations every 3 years. The new WHO global estimates of iodine status include statistics from 115 out of 193 WHO member states and cover over 90% of the world’s population of children. Overall, it is estimated that 30% of these children, numbering approximately 241 million, have inadequate iodine intake. More than half of these children come from Southeast Asia and Africa. In 2011, there were 30 known iodine deficient countries. This is a conservative figure as many smaller countries have not provided any data on which an assessment can be made.

The vast geographical expanse of Southeast Asia and the Pacific region—including more than 40 sovereign states ranging from tiny Pacific islands to some of the most populous nations on earth, such as India, Indonesia and China—poses a challenge in identifying and addressing the problems of IDD in this part of the world. Poverty and social disadvantage characterize many of these underdeveloped countries. While IDD most commonly coexists with poor social and economic development, geographic isolation and mountainous terrain, it is a myth that it is restricted to countries with such conditions. The United Nations Children’s Fund (UNICEF) has estimated that there are more than 20 million babies born annually within the Asia Pacific region who are not protected from iodine deficiency. The majority of these babies are born in India, Bangladesh and western China. Despite the persistence of iodine deficiency in many parts of our world, we need to emphasize that major improvements in iodine nutrition have been made, and that our current efforts are making a difference, albeit more slowly than expected.

The Health Impact of Endemic Iodine Deficiency

Iodine deficiency in humans and animals causes a wide spectrum of disorders, collectively known as Iodine Deficiency Disorders (IDD), affecting brain and skeletal growth and metabolism from conception until the end of life (Table 1). For quite some time, iodine deficiency was considered synonymous with endemic goiter because this manifestation is easily recognized, and in the child (but not in mature adults), it is readily reversed by iodine supplementation given to correct the deficiency (Figure 1).
One of the most puzzling questions has been how and when brain damage occurs from iodine deficiency. While congenital hypothyroidism, due mainly to under-development or agenesis of the thyroid gland, is known to cause congenital myxedematous cretinism in the affected child, this clinical picture is not seen very often in populations where iodine deficiency is severe. In contrast, children born to mothers who were severely iodine deficient during pregnancy manifested a totally different clinical picture of a severe neurological disorder that has been termed neurological cretinism (Figure 2). This manifestation is very different from myxedematous cretinism. We now know that moderate to severe maternal iodine deficiency results in maternal and fetal hypothyroxinemia in the first and second trimesters, causing impaired development of the fetal central nervous system which, depending on severity, may vary from a mild decrease in IQ to severe neurological cretinism. Persistence of severe iodine deficiency and hypothyroidism during infancy causes myxedematous features in the child (Figure 3).

It is commonly stated that iodine deficiency is the most common cause of preventable or impaired mental development worldwide. It has been well-documented that children born into families living in areas of moderate to severe iodine deficiency may lose up to 13.5 IQ points as a consequence of maternal iodine deprivation during pregnancy. It is this fact that makes iodine deficiency one of the more important and devastating non-communicable disorders in our world. The effects of IDD are a consequence of inadequate thyroid hormone production and action on hormone-sensitive target tissues. There is no evidence for an independent physiological function for iodine. In mammals, the sole function of iodine is to be incorporated into the thyroid hormone molecule through thyroid gland biosynthesis.

Table 1. Health consequences of iodine deficiency disorders at different life stages

<table>
<thead>
<tr>
<th>Life stages</th>
<th>Health consequences of iodine deficiency</th>
</tr>
</thead>
</table>
| Fetus and neonate | Spontaneous abortion  
Premature delivery and stillbirth  
Congenital anomalies  
Increase perinatal mortality  
Neurological cretinism |
| Child and adolescent | Myxedematous cretinism  
Neurological cretinism  
Impaired mental function; decreased IQ  
Hypothyroidism  
Delayed physical development  
Goiter |
| Adult | Hypothyroidism  
Impaired mental function  
Goiter  
Iodine-induced hyperthyroidism (in nodular goiter) |
| All ages | Goiter  
Hypothyroidism  
Increased susceptibility to nuclear radiation |

*Adapted from: Li M, Eastman CJ. Nat Rev Endocrinol 2012;8:434-40; and Hetzel BS, Lancet 1983;2:1126-8.*
Social and Economic Impact

It is well-established that moderate to severe iodine deficiency in a population has a negative impact on the health and well-being of that population, decreases productivity within the population, and is a barrier to socio-economic development. The International Child Development Steering Group has identified iodine deficiency as one of four key global health factors for impaired child development and where the need for intervention is most urgent. Whenever the negative impact of micronutrient deficiencies on child development is examined, iodine deficiency usually figures at the top or near the top of the list of causes. Even economists attending the Copenhagen Consensus in 2008 ranked USI as third on their list of priorities to confront and overcome in their listing of the 10 biggest challenges in advancing global welfare. Unfortunately, hard evidence with good cost-benefit analyses from randomized, controlled trials is lacking. These studies need to be done to provide quantitative data on the correction of this nutritional deficiency.

Indicators for Assessing Iodine Deficiency and its Elimination

The WHO has provided a number of criteria as guidance for the assessment of iodine deficiency and to determine whether IDD elimination goals have been met. Total goiter prevalence rate (TGR) in schoolchildren was the traditional method for assessing iodine deficiency in a population. While TGR provides information on the effect of long-term iodine deficiency in a community, it lags behind changes in iodine nutritional status and does not provide information on the current situation. It reflects what has happened in the past, but not what is happening in the present. As such, TGR is no longer used as a primary tool. For this reason, the definition of iodine deficiency is based primarily on population median urinary iodine concentration (MUID) estimations. Approximately 90% of absorbed iodine is excreted in the urine within a day or two of absorption, and the MUID is an excellent proxy indicator of iodine intake. The MUID is usually determined from a statistical, representative sample of schoolchildren. While school-based surveys are the easiest to conduct, the major drawback to using iodine excretion in children as a proxy for assessing iodine deficiency in a population is that it is unable to assess iodine nutrition in the most vulnerable population groups, particularly pregnant and lactating women, infants and pre-school children. In the general population, the MUID should be between 100 and 199 μg/L, while in pregnant women it should be in the range of 150 to 249 μg/L. Other biological indicators include monitoring of neonatal TSH levels, usually as part of national congenital hypothyroid screening programs; and more recently, measurement of serum thyroglobulin in blood spots taken from samples of the population. Household iodized salt usage is a useful indicator in rural populations, where most ingested salt is added at the household level and is not derived from processed food or salty condiments added to food.

Changing Epidemiology of Iodine Deficiency

Within the Asia Pacific region, IDD has been recognized as a serious public health problem at various times in India, Pakistan, Bangladesh, Indonesia, Myanmar (Burma), Cambodia, Thailand, China (western provinces including Tibet), Mongolia, North Korea (DPRK), Laos, Malaysia, Papua New Guinea, East Timor and the Philippines (Table 2). This is not an exhaustive list, as national statistics are not available for most countries despite the WHA resolution that binds all member states to collect data and report back to the WHA. For this discussion we have excluded an analysis and discussion of sub-continental Asian countries such as India, Bangladesh and Pakistan. In some of these countries in the Asia Pacific region, serious iodine deficiency is limited to remotely located pockets of the population that have limited access to iodized salt on a sustainable basis (Table 2).

Despite the difficulties, there has been considerable improvement in household iodized salt coverage in most of these countries over the past decade, giving hope to the realization of the goal of sustainable IDD control within the next decade. It should be noted that populations from Korea and Japan have liberal dietary iodine mainly from a high intake of iodine-containing seaweed products.

### Table 2. Current status of iodine nutrition in the Asia Pacific region

<table>
<thead>
<tr>
<th>Country</th>
<th>Status</th>
<th>iodine sufficient</th>
<th>iodine deficient</th>
<th>Unknown</th>
</tr>
</thead>
<tbody>
<tr>
<td>People’s Republic of China</td>
<td>Vietnam</td>
<td></td>
<td></td>
<td>Myanmar (Burma)</td>
</tr>
<tr>
<td>Japan</td>
<td>Indonesia</td>
<td>(variable)</td>
<td></td>
<td>Laos</td>
</tr>
<tr>
<td>South Korea</td>
<td>Democratic People’s Republic of Korea (severe)</td>
<td></td>
<td></td>
<td>Most Pacific islands</td>
</tr>
<tr>
<td>Sabah and Sarawak</td>
<td>Malaysia</td>
<td>(variable)</td>
<td></td>
<td>Papua New Guinea</td>
</tr>
<tr>
<td>Cambodia</td>
<td>Philippines</td>
<td>(variable)</td>
<td></td>
<td>Solomon Islands</td>
</tr>
<tr>
<td>Australia (recent)</td>
<td>Thailand</td>
<td>(variable)</td>
<td></td>
<td>Samoa</td>
</tr>
<tr>
<td>New Zealand (recent)</td>
<td>East Timor</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Taiwan</td>
<td>India</td>
<td>(variable)</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mongolia</td>
<td>Bangladesh</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Fiji</td>
<td>New Caledonia</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Bhutan</td>
<td>Vanuatu</td>
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<td></td>
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</tr>
</tbody>
</table>

Myths about Iodine Deficiency

There have been sporadic reports of goiter being endemic in the South Pacific, particularly in parts of Vanuatu, New Caledonia, Fiji, Samoa and some of the other smaller islands such as Tuvalu. Because many of the smaller countries in the Asia Pacific region are island states, with large expanses of coastline and presumed easy access to seafood, it has erroneously been assumed that iodine deficiency is not a significant endemic problem in the Pacific. As a consequence, little or no data on IDD has been collected in many of these poorer, smaller nations, but some of the best clinical research on IDD has been done in this part of the world. For example, data from...
surveys in the highlands of Papua New Guinea more than 50 years ago documented a cause-and-effect relationship between severe iodine deficiency in mothers and neurological cretinism in the offspring. A controlled therapeutic trial of iodized oil given to mothers early in pregnancy showed, for the first time, that endemic cretinism was a preventable disorder.13

A survey in Fiji in 1994 reported a goiter rate of just under 50% in schoolchildren and 15% in pregnant women. The MUIC of 26 μg/L in schoolchildren confirmed moderately severe iodine deficiency.14 This has now been addressed through a national iodized salt program. In a recent study on the island of Tanna, we were able to demonstrate moderate iodine deficiency in school-age children with a median urinary iodine excretion of 49 μg/L and increased thyroid volumes in 30% of the children, when compared with international reference ranges.15 It is still unknown how widespread and severe iodine deficiency is within the Pacific island nations, as studies have been few and far between, and we are lacking data other than sporadic observational records. Surveys are now underway in Samoa in the South Pacific and Guam in the North Pacific to determine the iodine nutritional status of these areas.

Another myth is that iodine deficiency only occurs in remote, mountainous areas populated by disadvantaged people who do not have access to a diverse and nutritious diet. In contrast, there has been widespread disregard for the crucial role of iodine nutrition in people living in advanced industrial nations.16 Illustrative examples of this neglect include Australia, New Zealand and the United Kingdom.3,4 Until the early 1990s, the populations of Australia and New Zealand were known to be iodine replete because of the widespread use of iodized salt, and bountiful supplies of iodine in milk and dairy products from iodine leaching into the milk from iodophore sanitizers used in the dairy industry from the 1960s onward.17 A national study of school-age children performed in Australia in 2003/2004 demonstrated mild iodine deficiency with a MUIC of 96 μg/L, 50% lower than data obtained from Australian children a decade before.18 Surveys in schoolchildren in New Zealand and the UK have yielded similar results and have caught public health authorities unaware of the development of iodine deficiency in their populations.19,20 The cause for the re-emergence of iodine deficiency in these developed countries has been a consequence of changes in work practices in the dairy industry, where iodophores have been replaced by other non-iodine containing, less expensive sanitizers.21 This illustrates the point that optimal sustainable iodine nutrition cannot be based on adventitious sources of iodine in the diet.

USI and Other Strategies to Meet Global Iodine Requirements

More than half a century has passed since the WHO first recommended iodization of all food salt for the population in areas where iodine deficiency was known to exist. Mandatory universal salt iodization (USI) represents the most effective public health strategy, where safety and efficacy can be assured, with a demonstrated need for the nutrient in the population.22 This strategy requires the iodization of all salt for human and animal consumption at a sufficient level—usually 20 to 40 mg iodine per kg salt—to ensure an adequate intake for all members of the population. However, USI must be underpinned by national legislation, as we know that voluntary systems are not sustainable strategies for eradicating iodine deficiency in a population.22 Salt is the ideal vehicle for iodine as it is an essential nutrient consumed consistently on a day-to-day basis. Iodization of salt is simple, inexpensive and cost-effective, amounting to less than one cent per day for an individual.

There are some fundamental differences implementing USI in the developing versus the developed world. In the developing world, most edible salt is consumed by direct addition to food being preserved or cooked and/or added to the food at the table. This is not true for many countries in the Indochina region (Thailand, Laos, Cambodia and Vietnam), where a good proportion of daily salt intake comes from salty condiments rather than the direct use of salt in the home. In contrast, in the developed world, most ingested salt—estimated in the vicinity of 80% of total daily salt—comes from processed food and is not added at the household level. Therefore, unless iodized salt is used in food processing and preparation, the quantity of iodine derived from the discretionary salt intake will be inadequate to sustain optimal iodine nutrition. Other sources will be necessary. Voluntary fortification of salt and other foods has many limitations and fewer benefits.22 Furthermore, the current worldwide push to decrease salt intake to prevent cardiovascular disease presents an entirely new challenge in addressing iodine deficiency in both developing and developed countries.

Iodine Supplementation

The provision of iodine supplements to individuals or whole populations has long been a controversial issue, and is not generally recommended. Iodine supplementation is a useful but an expensive, inefficient and unsustainable strategy for preventing iodine deficiency. The major reasons for opposition to this strategy include poor population coverage, poor compliance within the population, expense, and lack of sustainability.22 It is now generally agreed that iodine supplementation has a place during pregnancy and while breastfeeding in populations where USI has not been implemented or has failed.23 Daily supplements of 150 to 250 μg for all pregnant and lactating women have been recommended by the Endocrine Society, the American Thyroid Association, and the National Health and Medical Research Council in Australia. For pregnant and lactating women living in remote areas where daily iodine supplements are not practicable, depot preparations of iodized oil (containing
400 mg of potassium iodate) are given orally on an annual basis. This is effective in preventing neurological cretinism, but the lack of randomized, controlled trials leave us with inadequate data to provide definitive statistics on efficacy and safety. Therefore, iodized oil should only be used in situations where daily iodine supplementation cannot be implemented.

**Iodine Excess**

Iodine excess, like iodine deficiency, may also have adverse effects in a population. Complications include goiter, hyperthyroidism, hypothyroidism, and an increased prevalence of thyroid autoimmunity. Iodine excess disorders have been described in China and Japan, but are otherwise rare in the Asia-Pacific region. While the adverse effects of iodine deficiency far outweigh the adverse effects of iodine excess, programs designed to eradicate iodine deficiency should seek to establish optimal iodine nutrition throughout the whole population, and not operate on the basis that because iodine is “good for you, more is better.”

**What are the lessons that we have learned in these developing countries?**

**Lessons Learned**

We have learned the following from our studies and observations on developing countries:

1. There is a lack of national data on the extent and severity of iodine deficiency in most countries in the Asia Pacific region, because representative surveys in the population have not been undertaken.
2. Framing the message of iodine deficiency around brain damage, loss of IQ and impaired mental and physical development is far more powerful and persuasive than advocating iodine prophylaxis programs for endemic goiter.
3. Advocacy at the government level is essential to influence those that have the power to make decisions. IDD is not simply a public health problem, but a “whole of government” issue affecting social and economic development. Obtaining and maintaining high-level political commitment to eliminating IDD is essential.
4. USI (mandatory fortification of all edible salt with iodine) must be accepted at a national level as the most appropriate vehicle for normalizing iodine nutrition within a community. Acceptance of USI must be followed by implementation of legislation and regulations underpinning this initiative.
5. Multi-sectoral involvement is a prerequisite for success. The salt industry is as an important player as the health sector in achieving USI, and must accept responsibility to ensure the supply of iodized salt.
6. Education of the public is necessary to create demand for iodized salt and sustainability of optimal iodine intake.
7. A vital component of successful outcome is the implementation of a program for regular monitoring of the iodine intake of the population, and the quality assurance of iodized salt at the production and retail levels, with mandatory public reporting of these data.
8. Where iodized salt cannot be provided to all citizens, particularly children and women of childbearing age, an iodized oil supplementation program should be employed as an interim strategy. This is an effective way of protecting the newborn, and the growth and development of infants and children.

In summary, iodine deficiency remains one of the most prevalent and devastating non-communicable disorders in our world, with the potential to profoundly threaten the health and socio-economic development of almost one-third of the population on our planet. Within the Asia Pacific region, iodine deficiency is highly prevalent. More attention is needed in gathering national data to determine its prevalence and to formulate effective public health policy for the development and implementation of sustainable IDD elimination programs.

**References**


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Iodine deficiency remains one of the most critical public health issues, affecting social and economic development. Obtaining and maintaining high-level political commitment to eliminating IDD is essential. Advocacy at the government level is necessary to influence those who have the power to make decisions. IDD is not simply a public health problem, but an economic one, and the development of public health policies must accept responsibility to ensure the supply of food and water that are safe for consumption and nutrition within a community. Acceptance of USI and the role of the private sector in the achievement of USI is a key factor for the development and implementation of sustainable IDD elimination programs.

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References


