ACCEPTED VERSION

Sheila Skeaff, Ying Zhao, Robert Gibson, Maria Makrides, Shao Jia Zhou Iodine status in pre-school children prior to mandatory iodine fortification in Australia

Maternal and Child Nutrition, 2014; 10(2):304-312

© 2012 Blackwell Publishing Ltd

This is the peer reviewed version of the following article: Sheila Skeaff, Ying Zhao, Robert Gibson, Maria Makrides, Shao Jia Zhou Iodine status in pre-school children prior to mandatory iodine fortification in Australia Maternal and Child Nutrition, 2014; 10(2):304-312, which has been published in final form at http://dx.doi.org/10.1111/j.1740-8709.2012.00419.x. This article may be used for non-commercial purposes in accordance with Wiley Terms and Conditions for Self-Archiving.

PERMISSIONS

http://olabout.wiley.com/WileyCDA/Section/id-820227.html

Publishing in a subscription based journal

Accepted Version (postprint)

Self-archiving of the accepted version is subject to an embargo period of 12-24 months. The embargo period is 12 months for scientific, technical, and medical (STM) journals and 24 months for social science and humanities (SSH) journals following publication of the final article.

The accepted version may be placed on:

the author's personal website

- the author's company/institutional repository or archive
- certain not for profit subject-based repositories such as PubMed Central as listed below

Articles may be deposited into repositories on acceptance, but access to the article is subject to the embargo period.

The version posted must include the following notice on the first page:

"This is the peer reviewed version of the following article: [FULL CITE], which has been published in final form at [Link to final article using the DOI]. This article may be used for non-commercial purposes in accordance with Wiley Terms and Conditions for Self-Archiving."

21 August 2015

1 2 Running title: Iodine status of preschool children in Australia 3 Sheila Skeaff^{*}, Ying Zhao^{*}, Robert Gibson[†], Maria Makrides[¥], Shao Jia Zhou[¥] 4 5 ^{*} Department of Human Nutrition, University of Otago, 6 7 PO Box 56, Dunedin, New Zealand 8 9 ⁺ School of Agriculture, Food & Wine, 10 University of Adelaide. 11 Waite Campus 12 Waite Road, Glen Osmond, SA 5064 13 14 Australia 15 [¥]Women's and Children's Health Research Institute & 16 School of Paediatrics & Reproductive Health, 17 University of Adelaide. 18 72 King William Road, North Adelaide, SA 5006 19 Australia 20 21 22 Corresponding Author: Shao Jia Zhou 23 Women's and Children's Health Research Institute 24 25 72 King William Road, North Adelaide Australia, SA 5006 26 Phone: +61 8 8161 6849 27 28 Fax: +61 8 8239 0267

Iodine status in preschool children prior to mandatory iodine fortification in Australia

- 29 Email: <u>jo.zhou@adelaide.edu.au</u>
- 30
- 31

32 Acknowledgements

We thank Heather Garreffa and Helen Loudis for their clinical, administrative and technicalsupport.

35

36 Authors' contributions

37 S.S., S.J.Z., R.A.G. and M.M. designed the study; S.J.Z. and M.M. conducted the study; Y. Z.

and S. S. analyzed the urine iodine concentration and assessed iodine intake; S.S. drafted the

39 manuscript and all authors read and approved the final manuscript.

40

41 Conflict of interest statement

This project was funded by Wyeth Nutritional International Inc. Data collection, analysis andinterpretation were conducted independent of the funding body.

44 R.A.G. (ID: 519324) and M.M. (ID: 565000) were supported by a National Health & Medical

45 Research Council Senior Research Fellowship.

46 Author disclosures: S.J.Z., S.S. & Y.Z. have no conflict of interest. R.A.G. has been serving on

- 47 the scientific advisory boards for Nestle and Fonterra. M.M. has been serving on the scientific
- 48 advisory boards for Nestle, Fonterra and Nutricia. Associated honoraria for M.M. and R.A.G. are
- 49 paid to their institutions to support conference travel and continuing education for postgraduate
- 50 students and early career researchers.

51

52 Source of Funding

- 53 This study was funded by Wyeth Nutritional International Inc. Data collection, analysis and
- 54 interpretation were conducted independent of the funding body.

55 Abstract

The iodine status of children between the ages of 5 and 15 years has been routinely assessed in 56 many countries, but few studies have examined iodine status in preschool children. We 57 conducted a cross sectional study of preschool children living in Adelaide, South Australia, 58 between 2005 and 2007. Children 1-5 years old were identified using a unique sampling 59 strategy to ensure that the study population was representative. A 3-day weighed diet record, 60 blood sample and urine sample were obtained from each child. The median urinary iodine 61 62 concentration (UIC) of the children (n=279) was 129 µg/L indicating iodine sufficiency (normal range 100-199 μ g/L) but 35% of the children had a UIC <100 μ g/L. The median 63 64 thyroglobulin concentration of children (n=217) was 24 µg/L and thyroglobulin concentration declined with increasing age (p=0.024). The mean daily iodine intake was 76 µg. The intake of 65 iodine was lower than expected and highlights difficulties in accurately assessing iodine 66 67 intakes. Further studies are needed to monitor dietary changes and iodine status in this age group since the implementation of mandatory fortification of bread with iodised salt in 68 Australia in 2009. 69

70

71 Key Words (up to 6):

72 iodine deficiency, urinary iodine concentration, thyroglobulin, children, Australia

73

75 Introduction

An adequate amount of iodine is required in the diet for the synthesis of the thyroid 76 hormones tri-iodothyronine (T3) and thyroxine (T4) which in turn are needed for normal 77 growth and mental development. Pregnancy is the most critical time for the development of the 78 central nervous system because neurodevelopment is rapid during this period. The brain 79 continues to develop after birth (Cameron 2008), hence a good supply of thyroid hormones is 80 81 needed throughout childhood for optimal neurodevelopment as well as normal growth and 82 metabolism. Although the iodine status of children between the ages of 5 and 15 years has been assessed in many countries, there are few studies worldwide that have measured the iodine 83 84 status of preschool children. The diets of young children typically contain only small quantities of iodine-rich foods such as fish and seafood and dietary guidelines for children in this age 85 group often discourage the addition of salt (including iodised salt) to home prepared and 86 87 manufactured foods (National Health & Medical Research Council 2003). Given this, preschool children could be at higher risk of iodine deficiency than school-age children. 88

Australia has a history of iodine deficiency particularly in the eastern and southern states. 89 The introduction of iodised salt and the use of iodophors in the dairy industry were attributed 90 to eliminating iodine deficiency in the first half of the 20th century. However, in the late 91 1990's, Gunton et al. (1999) reported iodine deficiency in hospital patients in Sydney with 92 similar findings observed in subsequent studies in pregnant women (Guttikonda et al. 2002; 93 Burgess et al. 2007) and children (McDonnell et al. 2003; Hamrosi et al. 2005;) in Melbourne 94 and Tasmania. In 2003, a large study conducted by Li et al. of 8-10 year old schoolchildren 95 living in five mainland states confirmed mild iodine deficiency had re-emerged in the eastern 96 states (Li et al. 2006). To our knowledge, there have been no studies that have investigated the 97

98 iodine status of preschool children in Australia, a sub-group of the population that is often99 overlooked.

The most commonly accepted method of assessing iodine status of a population is to 100 determine the urinary iodine concentration (UIC) from a casual or spot urine sample, with a 101 UIC >100 μ g/L indicative of adequate iodine status. Although this cut-off has only been 102 validated in school aged children, it is still recommended for use in younger children (WHO et 103 al. 2007). In addition to UIC, the volume of the thyroid gland can be measured by 104 ultrasonography, with a value $> 97^{\text{th}}$ percentile indicating goiter but there are no published 105 reference values for thyroid volume in children under the age of 5 years (Zimmermann et al. 106 107 2001). In addition, there is growing interest in the measurement of thyroglobulin (Tg), a sensitive biochemical index of iodine status (Vejbjerg et al. 2009) as serum Tg concentrations 108 increase when the thyroid gland enlarges but Tg has not been routinely reported in children. 109 110 Using a combination of biomarkers including UIC and Tg, this paper reports on the iodine status of preschool children, who participated in a comprehensive survey assessing nutritional 111 status in this age group from a representative sample of Australian children (Zhou et al. 2012). 112

113

114 Subjects and Methods

This was a cross-sectional study of a representative sample of children living in Adelaide, South Australia, conducted between September 2005 and July 2007. Demographic data, a 3day weighed diet record, blood sample and urine sample were collected from each child. The study received ethical approval from the Human Research Ethics Committee at the Women's & Children's Health Network, Adelaide, South Australia. The nature of the study was fully explained to the caregivers of the children and informed written consent was obtained from allcaregivers.

There were 2132 Census Collection Districts (CCD) in Adelaide, each containing 220 122 dwellings. Each CCD was assigned an Advantage-Disadvantage Index by the Australian 123 Bureau of Statistics as an index of socioeconomic status (SES); these were stratified to low, 124 medium, and high SES. A stratified random sampling technique was used to select CCDs to 125 126 obtain a representative population sample, using a door-knocking protocol (Karr et al. 1996; 127 Soh et al. 2002). In brief, an address start point and direction was randomly determined for each identified CCD. From each start point, two trained research assistants knocked on the 128 129 doors of households to identify if there was a child aged between 1-5 years in the household. If no one was home, households were visited up to three times on a different day (including a 130 weekend day) and at different times from the previous visits. CCDs from each of the SES strata 131 132 were visited according to the random selection until the required number of children (n=100)from each SES strata was obtained. Children were excluded if they were diagnosed with 133 congenital or metabolic disorders (e.g. diabetes mellitus, cystic fibrosis) that required 134 specialised dietary intervention, or were hospitalised in the last 6 months before the study, or 135 were immunosuppressed. If more than one child was eligible per household, the one with the 136 earliest birth date was selected to participate in the study. Consenting families were asked to 137 138 attend a clinic appointment with their child for assessment.

The weight and length/height of each child, wearing light clothing without shoes using standardized procedures (WHO 2008), was measured during the clinic appointment. Weightfor-age (WAZ) and weight-for-height Z-scores (WHZ) were calculated using WHO ANTHRO 2005 software version 3 (WHO Geneva, Switzerland).

Parents were asked to complete 3-day weighed record of their child's food intake on 3 143 consecutive days, including one weekend day. Parents were given detailed instructions on how 144 to complete the food record and were provided with sample food diaries as examples. Parents 145 were supplied with a weighing scale and metric cups and spoons, and were asked to measure 146 and record everything their child ate and drank during the study period, both in the home and 147 away from home. For dishes made at home, detailed recipes were obtained, and for commercial 148 149 packaged foods, brand name was recorded and packaging retained to check for nutritional composition. Parents were instructed to weigh and record the amount of food and drink served 150 to the child as well record the weight of any uneaten food, which was subtracted from the 151 152 served amount to obtain the actual amount eaten. Completed diet records were collected from homes and checked by a dietitian. Phone calls were made to clarify any ambiguities or missing 153 information. Food records were entered and analysed using a dietary analysis package 154 155 (FoodWorks Professional, Xyris Pty Ltd, QLD) with the NUTTAB 2006 database (Food Standards Australia New Zealand 2006) included in the software. Iodine contents of foods that 156 are the main sources of dietary iodine in the diets of preschool children including eggs, dairy 157 products and seafood were obtained from the NUTTAB 2006 database. Some foods such as 158 biscuits and other non-core foods in the NUTTAB 2006 database did not contain iodine values 159 and the iodine contents of these food items were substituted using values from the New 160 Zealand Food Composition Database (Crop & Food Research 2006), an approach used by 161 Food Standards Australia New Zealand to assess iodine intake of Australian for Mandatory 162 iodine fortification (Australian Institute of Health and Welfare 2011). For foods that were of a 163 similar nature, the same iodine content was used for each food. 164

A non-fasting blood sample was obtained from each child via venepuncture by a trained 165 paediatric phlebotomist. Blood samples were processed within 3 hours of collection and serum 166 samples were stored at -80°C before being couriered with dry ice to the analytical laboratory 167 for testing. The concentration of serum Tg was determined using a radioimmunoassay by 168 EndoLab, Christchurch Hospital, Christchurch. The Tg assay has an analytical detection limit 169 of 0.1 µg/L and accuracy checked using CRM 457 (Institute for Reference Materials and 170 171 Measurement, Geel, Belgium). The inter-assay CV was 25% at 0.2 µg/L, 8% at 40.4 µg/L, and 5% at 333 µg/L. Intra-assay CVs were 5% at 0.2 µg/L, 2% at 40.4 µg/L, and 2% at 333 µg/L. 172 Samples were screened for serum antibodies to Tg (TgAb) as these antibodies can interfere 173 174 with serum Tg determination.

During the clinic appointment, the parents of toilet trained children were asked to take children to the toilet and collect a sample of urine. For children who were not toilet trained, a urine sample was collected via a paediatric collection bag (Liberty, Implex Pty Ltd, Heathcare SA) *in situ*. If a urine sample was unable to be obtained during the clinic appointment, parents were instructed to collect a urine sample at home and place the sample in a fridge, which was collected within 3 weeks. For all children, 5-10 mL of urine was then decanted into a 50 mL sterilized urine collection container and stored at -20 °C until analysis.

Urine samples were sent by courier to the Department of Human Nutrition, University of Otago, and analysed (Y.Z.) using a modification of the method of Pino et al (Pino *et al.* 1998). A certified urine standard sample (Seronorm, Sero AS, Asker, Norway) and an internal pooled urine sample was included with each batch of urine samples in the analysis. The mean concentration of Seronorm was 139µg/L (95% CI: 133, 145µg/L) compared with the certified value of $141\mu g/L$ (95% CI: 132, 150 $\mu g/L$). The coefficient of variation for the Seronorm and pooled urine sample was 4.2% (n=74) and 4.7% (n=70), respectively.

Stata 11.1 (STATA Corporation, College Station, Texas, USA) was used for statistical 189 analyses. Children were divided into four age groups: 12-24 months, >24-36 months, >36-48 190 months, >48-60 months for all analyses. Descriptive statistics, including median and 191 interquartile range (IQR), were used to summarize UIC and Tg. UIC was log transformed to 192 193 improve normality for subsequent statistical analyses. Univariate regression was conducted to assess the association between UIC and age, sex, WHZ, SES, total energy intake and iodine 194 intake. Any variable with a p < 0.20 in the univariate model was included in the multivariate 195 196 model to examine predictors of UIC. Univariate regression was also undertaken to identify food groups that contributed to iodine intake, and those food groups with a p < 0.20 were 197 198 included in the multivariate model. Tests were two-sided and statistical significance set at p < 199 0.05.

200

201 **Results**

A total of 13,272 households were visited from 54 CCDs and 9,464 households answered 202 the door. There were 573 eligible children from the 54 CCDs visited and 300 children, 100 203 children from each SES strata, consented to take part in the survey. Diet records were obtained 204 from 297 children, however, 8 were excluded because the child was still breastfed and it was 205 not possible to estimate the quantity of breast milk consumed. Urine was collected from 279 206 207 children and serum samples were available for Tg determination from 217 children. There was complete data on dietary intake, UIC and Tg for 202 children. Of these children, 96% were 208 born in Australia, 48% were first born, 52% were boys, and their mean (SD) was 0.54 (0.98) 209

for WAZ and 0.71 (0.99) for WHZ, respectively. There were no significant differences between children with or without complete data with regard to gender (p=0.547), SES (p=0.680) and UIC (p=0.453).

The median UIC (IQR) for the children was 129 (78 to 202) µg/L, 35% of children had a 213 UIC below 100 µg/L and 11% had a UIC above 300 µg/L, which indicates that these children 214 were iodine sufficient and iodine intake was adequate but not excessive (WHO et al. 2007). 215 216 Univariate and multivariate linear regression found that only dietary iodine was significantly 217 associated with UIC (Table 2), such that every 10 µg increase in the intake of dietary iodine, increased UIC by 4%. The median (IQR) Tg concentration was 24 (16, 35) µg/L and 96% 218 219 (208/217) of children had a Tg concentration >10 µg/L. Tg concentration declined with increasing age (p=0.025) (Table 1). Only 1 child tested positive for Tg-Ab. 220

The mean daily energy intake of the children was 5142 (95% CI: 4992 to 5293) kJ and 221 222 energy intakes increased with age (p<0.001) as follows: 4421 (95% CI: 4187, 4656) kJ/d for 12-24 months, 5051 (95% CI: 4787, 5315) kJ/d for >24-36 monhs, 5264 (95% CI: 4969, 5559) 223 kJ/d for >36-48 months, and 6029 (95% CI: 5734, 6324) kJ/d for >48-60 months. The energy 224 contribution as a percentage from carbohydrate, fat, and protein was 49 % (95% CI: 48, 50), 34 225 % (95% CI: 33, 35), and 17 % (95% CI: 17, 18), respectively. The mean daily iodine intake 226 was 76 (95% CI: 73, 80) µg and iodine intakes declined with age (p=0.003) as follows: 84 227 (95% CI: 76, 90) µg for 1-2 years, 78 (95% CI: 71, 85) µg for 2-3 years, 71 (95% CI: 64, 78) 228 µg for 3-4 years, and 71 (95% CI: 63,78) µg for 4-5 years. The main sources of dietary iodine 229 in the diet of these children were dairy products (28.4%) and bakery products (22.0%). 230

231

232 **Discussion**

233 This is the first study to assess the iodine status of preschool children in Australia and one of a handful of studies worldwide involving this age group (Delange et al. 2001; Pouessel et al. 234 2003; Heydon et al. 2009). The median UIC of 129 µg/L and less than 50% (i.e 35%) of 235 children with UIC <100 μ g/L indicates adequate iodine status in these children according to the 236 WHO/UNICEF/ICCIDD criteria. The iodine status of a representative sample of Australian 237 school children aged 8-10 year old living in the same state (i.e. South Australia) was reported 238 239 to be 101 µg/L (Li et al. 2006). The higher UIC observed in our children may reflect higher 240 dairy product consumption in younger children, a variable found to be significantly associated with UIC. The consumption of dairy products has been shown to decline with age in Australian 241 242 children (Department of Health and Aging et al. 2008).

The overall concentration of Tg in these children (24 μ g/L) and the 26 μ g/L observed in 12-243 24 month olds was lower than a median Tg of 35 µg/L reported in 12-24 month old Canadian 244 245 children, a population classified as iodine sufficient (Djemli et al. 2004). There are no recommended cut-offs for Tg to classify iodine status of children. However, the normal 246 reference range for Tg concentration in children 5-14 years old is 4-40 µg/L (Zimmermann et 247 al. 2003). A low UIC (i.e. <100 µg/L) can increase thyroid volume and subsequently the 248 concentration of Tg. A median UIC <100 μ g/L and a median Tg concentration >10 μ g/L have 249 both been set as indicators of iodine deficiency (WHO et al. 2001). As these indices have only 250 been validated in school-age children, this may explain the discrepancy between a median UIC 251 that classifies the children in our study as iodine sufficient, but a Tg concentration that 252 categorises the children as iodine deficient. We did find that Tg concentration declined with 253 age and a similar finding has been observed in Canadian children aged 0-17 years (Djemli et 254

al. 2004). If Tg concentration is to be used as an index of iodine status, the development of
age-specific cut-offs to categorise iodine status are needed.

It is difficult to accurately assess iodine intake and as a result few studies have measured 257 iodine intakes for a number of reasons. Firstly, the contribution of iodine from iodised salt, 258 used at the table or in cooking, is difficult to estimate. Secondly, many national food 259 composition databases (e.g. USA) do not include information on the iodine content of foods. 260 261 Thirdly, even within the same country, varying soil iodine contents, practices in food processing (i.e. use of iodates in bread or iodophors in the dairy industry) and animal rearing 262 (i.e. use of iodised salt licks or iodine supplemented feeds) can result in fluctuations in the 263 264 iodine content of foods. The children in this study had energy intakes that met recommendations for this age group (Department of Health and Aging et al. 2006) and the 265 percent of total energy from macronutrients were within the Acceptable Macronutrient 266 267 Distribution Range (Institute of Medicine 2005). The iodine intake, however, was lower than the 2007 Australian Children's Nutrition Survey, the only other Australian study that has 268 assessed iodine intake of preschool Australian children; the survey, reported that 24-36 month 269 old children had an iodine intake of 126 µg/day (Department of Health and Aging et al. 2008), 270 however, no biomarkers of iodine status were included in the 2007 Survey. The discrepancy in 271 iodine intake between our study and that reported in the 2007 Children's Nutrition Survey is 272 likely due to a number of factors. The different dietary assessment methodologies between the 273 2007 Children's Nutrition Survey (24 hour recall) and our study (3-day weighed food record) 274 may partly contribute to the discrepancy as 24 hour recalls have been shown to overestimate 275 energy intake of infants and toddlers compared with 3-day weighed food records (Fisher et al. 276 2008). We did not collect information on iodised salt use in cooking or at the table, which may 277

278 have also contributed to the lower iodine intake, however, widespread use of iodised salt was uncommon in Australia at the time of study. In addition, the NUTTAB 2006 Australian food 279 composition database, the most up-to-date database available at the time of the study, was used 280 to determine iodine intakes in our study, while the 2007 Children's Nutrition Survey used a 281 modified food composition database specifically developed for use in the 2007 Survey. 282 Furthermore, in some instances foods missing from the 2006 Australian food composition 283 284 database were replaced with foods from the 2006 New Zealand food composition database, which typically would have lower iodine content because New Zealand has lower soil iodine 285 content than mainland Australia. A comparison of the same foods in the 2006, 2007, and 2010 286 287 Australian databases shown in Table 3, illustrates how national databases can change with regard to iodine contents in food within the same country (FSANZ 2006; FSANZ 2007; 288 FSANZ 2010), but also highlights the differences in iodine contents of foods between countries 289 290 including New Zealand (Crop and Food Research 2006) and the UK (Food Standards Agency 2002). 291

In response to concerns about the re-emergence of iodine deficiency in Australia, the 292 addition of iodised salt to most commercial bread products became mandatory in 2009. 293 Australian preschool children consume ~60-80 g/day of bread and bread products (Australian 294 Bureau of Statistics 1999). We estimate that the consumption of fortified bread would increase 295 the median UIC in these preschool children from ~130 μ g/L to ~160 μ g/L, still within 100-199 296 μ g/L range considered a safe, adequate intake of iodine. However, we also estimate that bread 297 fortification will increase the percentage of children with UIC > 300 μ g/L, a level associated 298 with excessive intake of iodine, from ~10% to ~14 % in this study population. Thus, the 299 addition of iodine to foods in Australia needs to be monitored, as there is growing evidence 300

that iodine excess (i.e. UIC >300 μ g/L) can be also associated with adverse health effects such as hypothyroidism (Laurberg *et al.* 2006). The Tolerable Upper Limit (UL) intake for iodine in Australia is 200 μ g/day for 1-3 year old children and 300 μ g/day for 4-8 year olds (Department of Health and Aging *et al.* 2006). Concern that some children would exceed the UL was the primary reason that iodine fortification was limited to breads rather than being added to a number of other staple foods (FSANZ 2008).

307 The strengths of this study were the use of a door-knocking strategy to identify a representative sample of children, and the collection of both biochemical and dietary measures 308 of iodine status. Based on UIC, these preschool children had adequate iodine status, and the 309 310 accompanying Tg data provides information on Tg concentration in iodine sufficient preschool children for future reference. The iodine intake of the children was lower than expected and 311 312 highlights the inherent difficulties in assessing dietary iodine, particularly with regard to the 313 iodine content of foods in food composition databases. Despite this limitation, it is important to measure iodine intakes in order to identify foods and food groups that are good sources of 314 iodine in the diets of preschool children, as changes in dietary patterns do occur which may 315 impact on iodine status. The mandatory fortification of bread with iodised salt means that bread 316 and bread products are now likely to make the largest contribution to total iodine intakes in 317 preschool children. Further studies are needed to monitor dietary changes and iodine status in 318 319 this age group since the implementation of mandatory fortification.

320

321

322

Key Messages

- Preschool children are often overlooked in surveys assessing iodine status, despite their relative high dietary requirements for iodine.
- Our study indicates adequate iodine status in preschool children prior to mandatory iodine fortification in Australia.

323 **References**

- Australian Bureau of Statistics. (1999) 1995 National Nutrition Survey: Foods Eaten. Australian
 Bureau of Statistics: Canberra.
- Australian Institute of Health and Welfare. (2011) Mandatory folic acid and iodine fortification
- in Australia and New Zealand: Baseline report for monitoring. Australian Institute of Healthand Welfare: Canberra.
- 329 Burgess J.R., Seal J.A., Stilwell G.M., Reynolds P.J., Taylor E.R. & Parameswaran V. (2007) A
- case for universal salt iodisation to correct iodine deficiency in pregnancy: another salutary

lesson from Tasmania. *Medical Journal of Australia* **186**, 574-576.

- Cameron N. (2008) The biology of growth. In DJP Barker, RL Bergmann & PL Ogra (Ed.), The
- window of opportunity: Pre-pregnancy to 24 months of age. Nestle Nutrition Institute,Switzerland.
- Crop and Food Research. (2006) New Zealand Food Composition Tables. Crop and Food
 Research: Palmerston North.
- 337 Delange F., Wolff P., Gnat D., Dramaix M., Pilchen M. & Vertongen F. (2001) Iodine deficiency
- during infancy and early childhood in Belgium: does it pose a risk to brain development?

European Journal of Pediatrics **160**, 251-254.

- 340 Department of Health and Aging, Australian Food and Grocery Council & Department of
- 341 Agriculture and Fisheries. (2008) 2007 Australian National Children's Nutrition and Physical
- 342 Activity Survey. Department of Health and Aging: Canberra.
- 343 Department of Health and Aging, National Health and Medical Research Council & Ministry of
- Health (2006) Nutrient Reference Values for Australia and New Zealand.

- 345 Djemli A., Van Vliet G., Belgoudi J., Lambert M. & Delvin E.E. (2004) Reference intervals for
- 346 free thyroxine, total triiodothyronine, thyrotropin and thyroglobulin for Quebec newborns,
- 347 children and teenagers.*Clinical Biochemistry* **37**, 328-330.
- 348 Fisher J.O., Butte N.F., Mendoza P.M., Wilson T.A., Hodges E.A., Reidy K.C. & Deming D.
- 349 (2008) Overestimation of infant and toddler energy intake by 24-h recall compared with
- weighed food records. *American Journal of Clinical Nutrition* **88**, 407-415.
- 351 Food Standards Agency. (2002). McCance and Widdowson's The Composition of Foods
- 352 Integrated Dataset. The Controller of Her Majesty's Stationery Office: London.
- 353 Food Standards Australia New Zealand. (2006) NUTTAB 2006 (Australian NUTritionTABles).
- Food Standards Australia New Zealand: Canberra.
- 355 Food Standards Australia New Zealand. (2007) NUTTAB 2007 (Australian NUTritionTABles).
- 356 Food Standards Australia New Zealand: Canberra.
- 357 Food Standards Australia New Zealand. (2008) P1003 Mandatory Fortification of Iodine in
- 358 Australia. Food Standards Australia New Zealand: Canberra.
- 359 Food Standards Australia New Zealand. (2010) NUTTAB 2010 (Australian NUTritionTABles).
- 360 Food Standards Australia New Zealand: Canberra.
- 361 Guttikonda K., Burgess J.R., Hynes K., Boyages S., Byth K. & Parameswaran V. (2002)
- 362 Recurrent iodine deficiency in Tasmania, Australia: A salutary lesson in sustainable iodine
- 363 prophylaxis and its monitoring. Journal of Clinical Endocrinology & Metabolism 87, 2809-
- 364 2815.
- Hamrosi M., Wallace E. & Riley M. (2005) Iodine status in pregnant women living in
- 366 Melbourne differs by ethnic group. *Asia Pacific Journal of Clinical Nutrition* **14**, 27-31.

| 367 | Heydon E.E., Thomson C.D., Mann J., Williams S.M., Skeaff S.A., Sherpa K.T. & Heydon J.L. |
|-----|---|
| 368 | (2009) Iodine status in a Sherpa community in a village of the Khumbu region of Nepal. |
| 369 | Public Health Nutrition 12, 1431-1436. |
| 370 | Institute of Medicine. (2005) Dietary reference intakes for energy, carbohydrate, fiber, fat, fatty |
| 371 | acids, cholesterol, protein and amino acids. National Academies Press: Washington, DC. |
| 372 | Karr M., Alperstein G. & Causer J. (1996) Iron status and anaemia in children in Sydney. |
| 373 | Australian and New Zealand Journal of Public Health 20, 618-622. |
| 374 | Laurberg P., Jorgensen A., Perrild H., Ovesen L., Knudsen N., Bulow Pedersen I., Rasmussen |
| 375 | L.B., Carle A. &Vejbjerg P. (2006) The Danish investigation on iodine intake and thyroid |
| 376 | disease, DanThyr status and perspectives. European Journal of Endocrinology 155, 219-228. |
| 377 | Li M., Eastman C.J., Waite K.V., Ma G., Zacharin M.R., Topliss D.J., Harding P.E., Walsh J.P., |
| 378 | Ward L.C., Mortimer R.H., Mackenzie E.J., Byth K. & Doyle Z. (2006) Are Australian |
| 379 | children iodine deficient? Results of the Australian National Iodine Nutrition Study. Medical |
| 380 | Journal of Australia 184, 165-169. |
| 381 | McDonnell C.M., Harris M. & Zacharin M.R. (2003) Iodine deficiency and goitre in |
| 382 | schoolchildren in Melbourne, 2001. Medical Journal of Australia 178, 159-162. |
| 383 | National Health & Medical Research Council. (2003) Dietary Guidelines for Children and |
| 384 | Adolescents in Australia incorporating the Infant Feeding Guidelines for Health Workers. |
| 385 | Commonwealth of Australia: Canberra. |
| 386 | Pino S., Fang S.L. & Braverman L.E. (1998) Ammonium persulfate: a new and safe method for |
| 387 | measuring urinary iodine by ammonium persulfate oxidation. Experimental and Clinical |
| 388 | Endocrinology and Diabetes 106, S22-27. |

| 390 | status and risk factors for iodine deficiency in infants and children of the french North |
|-----|---|
| 391 | department. Achives de Pediatrie 10, 96-101. |
| 392 | Soh P., Ferguson E.L., Mckenzie J.E., Skeaff S., Parnell W.R. & Gibson R.S. (2002) Dietary |
| 393 | intakes of 6-24 month-old urban South Island New Zealand children in relation to |
| 394 | biochemical iron status. Public Health Nutrition 5, 339-346. |
| 395 | Vejbjerg P., Knudsen N., Perrild H., Laurberg P., Carlé A., Pedersen I., Rasmussen L.B., Ovesen |
| 396 | L. &Jørgensen T. (2009) Thyroglobulin as a marker of iodine nutrition status in the general |
| 397 | population. European Journal of Endocrinology 161, 475-481. |
| 398 | World Health Organization. (2008) Training course on child growth assessment. World Health |
| 399 | Organization: Geneva. |
| 400 | World Health Organization, United Nations Children's Fund (UNICEF) & International Council |
| 401 | for the Control of Iodine DeficiencyDisorders (ICCIDD). (2001) Assessment of iodine |
| 402 | deficiency disorders and monitoring their elimination: a guide for programme managers. |
| 403 | World Health Organization: Geneva. |

Pouessel G., Bouarfa K., Soudan B., Sauvage J., Gottrand F. & Turck D. (2003) Iodine nutritional

- 404 World Health Organization, United Nations Children's Fund (UNICEF) & International Council
- 405 for the Control of Iodine DeficiencyDisorders (ICCIDD). (2007). Assessment of iodine
- 406 deficiency disorders and monitoring their elimination: a guide for programme managers (third
- 407 edition). World Health Organization: Geneva.

- 408 Zhou S.J., Gibson R.A., Gibson R.S. & Makrides M. (2012) Nutrient intakes and status of
- 409 preschool children in Adelaide. *Medical Journal of Australia (in Press)*.
- 410 Zimmermann M.B., Molinari L., Spehl M., Weidinger-Toth J., Podoba J., Hess S. & Delange F.
- 411 (2001) Toward a consensus on reference values for thyroid volume in iodine-replete

| 412 | schoolchildren: results of a workshop on inter-observer and inter-equipment variation in |
|-----|--|
| 413 | sonographic measurement of thyroid volume. European Journal of Endocrinology 144, 213- |
| 414 | 220. |
| 415 | Zimmermann M.B., Moretti D., Chaouki N. & Torresani T. (2003) Development of a dried |
| 416 | whole-blood spot thyroglobulin assay and its evaluation as an indicator of thyroid status in |
| 417 | goitrous children receiving iodised salt. American Journal of Clinical Nutrition 77, 1453- |
| 418 | 1458. |
| 419 | |