Epidemiology of anaemia in children, adolescent girls, and women in Bhutan

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Abstract
Anaemia inhibits health and development in Bhutan. We estimated anaemia prevalence and explored risk factors in children and women using data from Bhutan’s National Nutrition Survey 2015. Prevalence was calculated using life-stage-specific cut-offs adjusted for altitude and survey design. Risk factors were evaluated in modified Poisson regressions. Anaemia affected 42%, 29%, 36%, and 28% of children, adolescent girls, and non-pregnant and pregnant women, respectively. Risk of anaemia was greater in children who were younger (RR 2.0, 95% CI [1.7, 2.3] and RR 1.9, 95% CI [1.6, 2.3], respectively, for 12–23 and 6–11 vs. 24–59 months), male (1.2, 1.1–1.4, ref.: female), and stunted (1.2, 1.0–1.3, ref.: height-for-age ≥ −2z). Older (15–19 years) versus younger (10–14 years) adolescents were at higher risk (1.5, 1.2–1.8), as were adolescents living at home versus at school (1.2, 0.9–1.6) and those working versus studying (1.3, 1.0–1.7). Among adult women, anaemia risk increased with age (1.2, 1.0–1.4 and 1.3, 1.1–1.5, for 30–39 and 40–49, respectively, vs. 20–29 years) and was higher for women without schooling (1.1, 1.0–1.3, vs. primary schooling), who were unmarried or separated (1.4, 1.2–1.7 and 1.3, 1.1–1.6, respectively, vs. married), without a child <5 years (1.1, 1.0–1.3), and lacking improved sanitation (1.1, 1.0–1.3). High coverage of antenatal iron and folic acid supplementation may contribute to the lower prevalence of anaemia among pregnant women and women with young children. Expansion of iron supplementation programmes, fortification, and other strategies to improve dietary iron intake may reduce the prevalence of anaemia, but causes of anaemia other than iron deficiency (e.g., thalassemias) should also be investigated.

KEYWORDS
adolescent girls, anaemia, children, pregnant women, risk factors, South Asia

1 | INTRODUCTION

Anaemia continues to burden large segments of the global population. An estimated 29% of non-pregnant women and 43% of preschool-age children worldwide have anaemia (Stevens et al., 2013), and iron-deficiency anaemia is the single largest cause of years lived with disability in children and adolescents worldwide (Global Burden of Disease et al., 2017). Between 1990 and 2013, the global prevalence of anaemia declined by 21% (from 33.3% to 27.0%; Kassebaum, 2018), but progress has been uneven, with differences by region and greater
Anemia is a condition of insufficient erythrocyte haemoglobin content, in which persistent inadequate oxygen delivery impairs the function of body tissues and organs, with wide-ranging consequences. Anemia is associated with weakness, fatigue, reduced productivity, and inhibited immune function (Beard, 2001; Fernandez-Gaxiola & De-Regil, 2011). In pregnant women, anemia may cause low birthweight and preterm birth (Rasmussen, 2001) and increase the risk of maternal mortality due to haemorrhage and other delivery complications (Sanghvi, Harvey, & Wainwright, 2010). In children, anemia is associated with impaired cognitive development and possibly with motor development as well (Balarajan, Ramakrishnan, Özaltin, Shankar, & Subramanian, 2011; Sachdev, Gera, & Nestel, 2005). Iron deficiency is the most common cause of anemia (Kassebaum, 2018), but other causes include parasitic and other infections, chronic inflammatory conditions, deficiencies of micronutrients other than iron, and genetic causes (Bates, McKew, & Sarkinfada, 2007; Williams & Weatherall, 2012). The proportion of observed anemia attributable to iron deficiency is estimated to be just over 60% worldwide but differs by geographic region and country and by demographic group (Kassebaum, 2018), as well as with the degree of economic development and prevalence of inflammation (Petry et al., 2016). Recently reported prevalence of anemia in South and Southeast Asia is 58% in preschool-age children and 47% in non-pregnant women, despite substantial reductions in the past 15 years (Stevens et al., 2013). Prior to 2015, only two national surveys had described the burden of anemia in Bhutan (Bailey, Lubach, & Coe, 2004; Food and Nutrition Section Department of Public Health Ministry of Health, 2003), and no study had investigated the subnational distribution of anemia or its risk factors. Additionally, little data exist globally describing anemia prevalence and risk factors among adolescent girls. In 2015, a National Nutrition Survey (NNS) in Bhutan reported high rates of anemia among children, adolescent girls, and non-pregnant and pregnant women of reproductive age, despite indicators of relatively good household diet quality and child nutritional status (Campbell et al., 2018; Kang et al., 2018). In the present study, we aimed to determine the risk factors for anemia among each of the demographic groups included in Bhutan’s 2015 NNS.

2 METHODS

2.1 Study design

Data for the present analysis came from the 2015 NNS in Bhutan. The survey utilized a multistage sampling strategy to generate representative nutritional status estimates nationally, for three major regions (West, Central, and East) and for the urban and rural areas of the country. The survey report describes the design of the survey in detail (Nutrition Program, 2015). Briefly, two districts (Dzongkhags) were randomly selected from each of the three regions, within which smaller administrative areas (rural Chiwogs and urban enumeration areas) were randomly selected in proportion to the urban/rural makeup of the region. Within each small administrative area included in the sample, 12 households were randomly selected. In each selected household, a questionnaire was administered to determine household socio-economic characteristics and household composition. Additional questionnaires were administered to all pregnant women in the household and to the mother of the youngest child aged 0–23 months, and length/height and weight were measured for all children aged 0–59 months. Haemoglobin was measured with a Hemocue 301 (HemoCue, Brea, CA, USA) in eligible household members: children aged 6–59 months, adolescent girls aged 10–19 years, and pregnant and non-pregnant women aged 20–49 years. Adolescent girls from selected households who were attending school away from home were located at school and their haemoglobin measured. Altitude at each selected Chiwog or enumeration area was recorded. For adolescent girls residing in schools, the school’s altitude was recorded.

2.2 Data management

All participants with haemoglobin data were included in this analysis. Children, adolescent girls, and non-pregnant and pregnant women were analysed separately. Children included all boys and girls aged 6–59 months. Adolescent girls were non-pregnant girls aged 10–19 years. Non-pregnant women were those aged 20–49 years, and pregnant women were all currently pregnant women 15–49 years. For anemia prevalence estimates, non-pregnant women aged 15–19 years were included in both the adolescent girls and the non-pregnant adult women groups, for comparability with survey data from other countries for women of reproductive age. Women and adolescents with missing pregnancy status data were omitted from the analysis.

Participants with implausible haemoglobin values (<4 or ≥18 g/dl) were omitted from the analysis (n = 33 children and n = 20 adolescent girls and women; Sullivan, Mei, Grummer-Strawn, & Parvanta, 2008).
Haemoglobin values were adjusted for administrative area altitude with the Centers for Disease Control and Prevention (CDC) continuous adjustment equation (1989). Altitude-adjusted haemoglobin and anaemia variables were used for all analyses. Cut-offs for anaemia were based on World Health Organization (WHO) guidelines (WHO, 2011). For children, anaemia was defined as mild (Hb < 11 g/dl), moderate (Hb < 10 g/dl), and severe (Hb < 7 g/dl). For non-pregnant adolescents and adult women, mild was Hb < 12 g/dl or Hb < 11.5 g/dl for girls 10 or 11 years old; moderate was Hb < 11 g/dl; and severe was Hb < 8 g/dl. For pregnant women, cut-offs were mild (Hb < 11 g/dl), moderate (Hb < 10 g/dl), and severe (Hb < 7 g/dl).

Individual and household characteristics were measured by questionnaire. Improved sanitation was defined as the household having access to a private improved latrine, whereas improved water source was defined as having piped water into the house, the criterion set by the Government of Bhutan. A household wealth index was generated from household asset ownership and house structure variables according to a method previously developed by the Bhutan National Statistics Bureau. Food insecurity was assessed with a series of questions about the household’s experience over the previous 6 months and was dichotomized for this analysis as any—one or more questions answered in the affirmative—or none. A food consumption score was calculated based on reported household food group consumption in the week before the survey and rated as acceptable, borderline, or poor (World Food Programme, 2008); borderline and poor were combined for this analysis. Household-reported past-week consumption of flesh foods (meat, fish, and eggs) was also examined separately, with reported intake categorized as none, 1 to 6 of the past 7 days, or daily. Anthropometry was measured in all children 0–59 months of age in selected households. Weight was measured to the nearest 0.1 kg using a portable Seca 881 scale (Chino, CA, USA). Length or height was measured to the nearest 0.1 cm using a wooden measuring board (Nutrition Program, 2015).

Adolescent girls’ and women’s ages were self-reported in years, whereas children’s ages were calculated from maternal-reported date of birth. Adolescents’ and adults’ level of completed education was collected in categories: none, early learning, primary completed, high school completed, some university, diploma/certificate completed, monastic schooling, and non-formal education. For adult women, the categories were combined into a three-level scale: none or early learning only; primary, monastic, or informal schooling; high school, university, or diploma. For adolescents, three categories of education were considered: none, early learning, monastic, or informal schooling; primary school; and high school or greater. Women and adolescents reported their occupation in an open-ended question. For adolescents, the responses were grouped into student; farmer or other employment; and no employment. For adult women, occupation responses were grouped into student, farmer, other employment, and housewife/other. Women’s marital status was recorded as never married, living together, married, divorced, separated, or widow. For this analysis, living together and married were combined into one group labelled “married,” and divorced, separated, and widowed were combined into a “separated” group.

2.3 | Statistical analysis

National, regional, and urban/rural prevalence of anaemia was estimated for each demographic group using the “svy:” prefix to account for survey weights and clustering of observations from the design of the survey. For each group, prevalence of mild, moderate, and severe anaemia were estimated separately, as was prevalence of any anaemia with mild, moderate, and severe combined.

Models of individual and household factors predictive of anaemia were developed for the demographic groups separately using modified Poisson regression using the generalized linear model command with robust standard errors (Zou, 2004). For each demographic group, unadjusted regression models with dichotomous anaemia status as the outcome were generated for each potential risk factor. All variables significant at the p ≤ 0.10 level were then included in a multivariable model along with known anaemia risk factors: age, sex (child model only), region, urban/rural location, and household wealth index. Multivariable regression models were developed in the same way for each demographic group with independent variable continuous altitude-adjusted haemoglobin as a sensitivity analysis for the influence of the anaemia cut-offs on observed associations. A blockwise modelling approach was also applied for dichotomous and continuous outcomes, as a sensitivity analysis, to check that significant predictors in a fully adjusted model were not overlooked in a data-driven modelling approach (Engle-Stone et al., 2017). For this sensitivity analysis, blocks of risk factors were selected by referencing the conceptual framework proposed in Namaste et al. (2017). Survey-adjusted models were also developed using the “svy:” command with Poisson models in a further sensitivity analysis of the effect of the survey sampling strategy on the anaemia risk factors identified. All analyses were conducted in Stata 14.1 (StataCorp, College Station, TX, USA).

3 | RESULTS

Haemoglobin data were available for 1,083 children aged 6–59 months, 1,216 non-pregnant adolescent girls aged 10–19 years, 2,649 non-pregnant women aged 20–49 years, and 118 pregnant adolescents and women aged 15–49 years living in 2,478 households. Most children, adolescents, and women included in the sample lived in rural households (75%). Most households reported improved sanitation facilities (70%) and piped drinking water (86%). Household food insecurity and poor or borderline food consumption scores were reported infrequently (2% and 8%, respectively), and many households reported consuming flesh foods daily (68%). About 40% of households were located in administrative areas at altitudes between 1,000 and 2,000 m, and an additional 26% were located at altitudes ≥2,000 m.

3.1 | Anaemia prevalence

In Table 1, estimates of anaemia prevalence adjusted for survey design are presented. In all groups, anaemia prevalence was high. Among children aged 6–59 months, 42% were anaemic: 25% had mild anaemia
(10 g/dl ≤ Hb < 11 g/dl), 17% had moderate anaemia (7 g/dl ≤ Hb < 10 g/dl), and fewer than 1% had severe anaemia (Hb < 7 g/dl). The prevalence of anaemia among adolescent girls was 29%: 15% had mild anaemia (11 g/dl ≤ Hb < 12 g/dl); 11 g/dl ≤ Hb < 11.5 g/dl for girls 10–11 years), moderate anaemia (Hb < 11 g/dl), and severe anaemia (Hb < 8 g/dl). In non-pregnant women 15–49 yearsd for children, mild anaemia (Hb < 11 g/dl), moderate anaemia (Hb < 10 g/dl), and severe anaemia (Hb < 7 g/dl).

3.2 | Risk factors for anaemia

Table 2 presents multivariable models for anaemia risk factors in young children, in which child age, sex, and stunting status were associated with risk of anaemia. Children 6–11 and 12–23 months old (RR: 1.9, 95% CI: 1.6–2.3 and 2.0, 1.7–2.3, respectively, ref: 24–59 months) and boys (1.2, 1.1–1.4, ref: girls) were at greater risk of anaemia. Stunted children (1.2, 1.0–1.3, ref: height-for-age ≥ –2z) were also at a higher risk of anaemia. Odds ratios for anaemia in children with wasting and overweight were both positive but with 95% confidence intervals that included unity (1.1, 0.9–1.5 and 1.3, 0.9–1.8, respectively). Risk of anaemia differed somewhat by region of the country, with children in the West at somewhat higher risk of anaemia compared with those in the East (1.1, 1.0–1.3). Most household socioeconomic and diet quality markers were not associated with anaemia, although living in a household without improved sanitation or improved water was each associated with a trend towards increased risk of anaemia (1.1, 0.9–1.2 and 1.1, 0.9–1.3 for sanitation and water, respectively). Associations between infant and young child feeding indicators and anaemia status in children <2 years were evaluated, but no associations were observed (results not shown). In multivariable models with haemoglobin concentration as a continuous outcome, significant risk factors were similar to those in dichotomous anaemia models. In addition, having a mother with moderate anaemia was associated with 0.4 g/dl (–0.7 to –0.2) lower Hb concentration (not shown).

### Table 1: National and regional prevalence of anaemia in children, adolescents, and non-pregnant and pregnant women in Bhutan

| Demographic group | N   | Any | Mild | Moderate | Severe |
|-------------------|-----|-----|------|----------|--------|
| Children 6–59 monthsa | 1,083 | 42.3 (3.3) | 25.0 (1.4) | 16.9 (2.5) | 0.4 (0.3) |
| Adolescent girls 10–19 yearsb | 1,216 | 29.3 (3.6) | 15.1 (1.7) | 12.9 (1.9) | 1.3 (0.4) |
| Non-pregnant women 15–49 yearsd | 3,233 | 36.3 (2.1) | 18.0 (0.8) | 16.9 (1.3) | 1.4 (0.3) |
| Pregnant women 15–49 yearsa | 118 | 28.2 (3.4) | 17.9 (3.5) | 10.3 (3.2) | 0 (0) |

aEstimates adjusted for survey design using “svy:” commands.
bFor children, mild anaemia (Hb < 11 g/dl), moderate anaemia (Hb < 10 g/dl), and severe anaemia (Hb < 7 g/dl).
cFor non-pregnant adolescent girls, mild anaemia (Hb < 12 g/dl; Hb < 11.5 g/dl for girls 10–11 years), moderate anaemia (Hb < 11 g/dl), and severe anaemia (Hb < 8 g/dl).
dFor non-pregnant women, mild anaemia (Hb < 12 g/dl), moderate anaemia (Hb < 11 g/dl) and severe anaemia (Hb < 8 g/dl).
eFor pregnant women, mild anaemia (Hb < 12 g/dl), moderate anaemia (Hb < 11 g/dl), 17% had moderate anaemia (Hb < 10 g/dl), and 1% had severe anaemia (Hb < 7 g/dl).

### Table 2: Personal and household characteristics predictive of children’s risk of anaemia (n = 1,083)

| Characteristica | N   | Hb (g/dl), mean (SD) | RR (95% CI)b |
|-----------------|-----|----------------------|--------------|
| Region          |     |                      |              |
| East            | 481 | 11.5 (1.4)           | 1.0          |
| West            | 311 | 11.7 (1.2)           | 1.1 (1.0, 1.3) |
| Central         | 291 | 11.6 (1.2)           | 1.0 (0.8, 1.2) |
| Area            |     |                      |              |
| Urban           | 232 | 11.6 (1.2)           | 1.0          |
| Rural           | 851 | 11.6 (1.3)           | 0.9 (0.8, 1.2) |
| Wealth quintiles|     |                      |              |
| Highest         | 92  | 11.7 (1.2)           | 1.0          |
| High            | 145 | 11.5 (1.2)           | 1.0 (0.7, 1.4) |
| Medium          | 196 | 11.5 (1.4)           | 1.1 (0.8, 1.5) |
| Low             | 312 | 11.6 (1.3)           | 1.2 (0.9, 1.7) |
| Lowest          | 338 | 11.6 (1.4)           | 1.1 (0.8, 1.6) |
| Improved sanitation | |                      |              |
| Yes             | 721 | 11.6 (1.3)           | 1.0          |
| No              | 362 | 11.4 (1.3)           | 1.1 (0.9, 1.2) |
| Improved water  |     |                      |              |
| Yes             | 924 | 11.6 (1.3)           | 1.0          |
| No              | 159 | 11.4 (1.3)           | 1.1 (0.9, 1.3) |
| Child age, months |     |                      |              |
| 24–59           | 709 | 11.8 (1.2)           | 1.0          |
| 12–23           | 258 | 11.1 (1.3)           | 2.0 (1.7, 2.3) |
| 6–11            | 116 | 11.1 (1.3)           | 1.9 (1.6, 2.3) |
| Child sex       |     |                      |              |
| Female          | 575 | 11.7 (1.3)           | 1.0          |
| Male            | 508 | 11.4 (1.4)           | 1.2 (1.1, 1.4) |
| Stuntedc        |     |                      |              |
| No              | 799 | 11.6 (1.3)           | 1.0          |
| Yes             | 256 | 11.5 (1.3)           | 1.2 (1.0, 1.3) |
| Wastedc         |     |                      |              |
| No              | 1,017 | 11.6 (1.3)       | 1.0          |
| Yes             | 41  | 11.3 (1.3)           | 1.1 (0.9, 1.5) |
| Overweightc     |     |                      |              |
| No              | 1,059 | 11.6 (1.3)       | 1.0          |
| Yes             | 24  | 10.8 (1.3)           | 1.3 (0.9, 1.8) |

aOnly personal/household characteristics that were included in the multivariable model are listed in the table. Variables that were investigated but were not significant were as follows: household food insecurity, food consumption score, and meat consumption; maternal education, age, and anaemia; and child underweight.
bRisk ratios and confidence intervals generated with modified Poisson regression models with robust standard errors.
cDichotomous anthropometry indicator definitions: stunted: HAZ or LAZ < –2; wasted: WZL or WHZ < –2; overweight: WZL or WHZ > 2.
Table 3 presents risk factors for anaemia among adolescent girls, in whom older age (15–19 vs. 10–14 years) was associated with increased risk of anaemia (1.5, 1.2–1.8), as was working as a farmer or in another occupation (1.3, 1.0–1.7) versus being a student. Living in the West or Central region was associated with increased risk of anaemia (1.3, 1.1–1.7 and 1.3, 1.0–1.6 for West and Central, respectively, relative to East), and living at home was also associated with a greater anaemia risk relative to residing at school, although the confidence interval included 1 (1.2, 0.9–1.6). Household socio-economic status (SES) and diet quality indicators were not predictive of adolescent girls’ risk of anaemia. In models for continuous Hb, the only additional risk factor that emerged was the availability of improved water source in the household, which was surprisingly associated with 0.3 (−0.5 to −0.1) g/dl lower Hb (not shown).

Table 4 presents risk factors for anaemia among non-pregnant adult women. The risk of anaemia was associated with region, household sanitation, women’s age, level of education, marital status, and maternal status. Living in the West or Central region was associated with greater anaemia risk (1.5, 1.3–1.7 and 1.4, 1.2–1.6, respectively, relative to East), as was living in a house without an improved sanitation facility (1.1, 1.0–1.3). Risk of anaemia increased with age (1.2, 1.0–1.4 and 1.3, 1.1–1.5 in women ages 30–39 and 40–49 years, respectively, relative to women 20–29 years). Having no schooling was associated with increased risk of anaemia (1.1, 1.0–1.3) relative to those who attended primary school, but the risk of anaemia among women who completed high school was not statistically significantly different from those who attended primary school. Unmarried and separated/divorced/widowed women relative to married (1.4, 1.2–
TABLE 5 Characteristics predictive of pregnant women’s odds of anaemia (n = 118)

| Characteristica | n  | Hb (g/dl), mean (SD) | RR (95% CI)b |
|-----------------|----|---------------------|--------------|
| Region          |    |                     |              |
| East            | 47 | 12.2 (1.4)          | 1.0          |
| West            | 40 | 11.7 (1.2)          | 0.9 (0.5, 1.7)|
| Central         | 31 | 11.5 (1.4)          | 1.0 (0.5, 1.8)|
| Area            |    |                     |              |
| Urban           | 32 | 11.9 (1.7)          | 1.0          |
| Rural           | 86 | 11.7 (1.3)          | 0.6 (0.3, 1.5)|
| Wealth quintiles|    |                     |              |
| Highest         | 15 | 12.6 (1.6)          | 1.0          |
| High            | 23 | 12.0 (1.0)          | 0.7 (0.2, 2.9)|
| Medium          | 22 | 11.2 (1.4)          | 3.3 (1.1, 9.6)|
| Low             | 26 | 11.3 (1.1)          | 2.7 (0.9, 8.0)|
| Lowest          | 32 | 11.8 (1.4)          | 1.6 (0.5, 5.2)|
| Age, years      |    |                     |              |
| 15–19           | 9  | 12.0 (0.6)          | 0.4 (0.1, 2.5)|
| 20–29           | 67 | 11.8 (1.1)          | 1.0          |
| 30–39           | 34 | 12.6 (1.8)          | 1.0 (0.5, 2.0)|
| 40–49           | 8  | 11.9 (1.5)          | 3.6 (1.7, 7.6)|
| Trimester       |    |                     |              |
| 1               | 42 | 12.6 (1.6)          | 1.0          |
| 2               | 44 | 11.6 (1.2)          | 2.8 (1.2, 6.2)|
| 3               | 32 | 11.1 (1.1)          | 2.7 (1.3, 5.6)|

aOnly personal/household characteristics that were included in the multivariable model are listed in the table. Variables that were investigated but were not significant in unadjusted models were as follows: gravidity, receipt of any ANC, initiation of ANC in the first trimester of pregnancy and past 7-day iron and folic acid supplement consumption.

bRisk ratios and confidence intervals generated with modified Poisson regression models with robust standard errors.

1.7 and 1.3, 1.1–1.6, respectively) and those without a child under age 5 years (1.1, 1.0–1.3) were at greater risk of anaemia. Associations between risk factors and continuous Hb were similar (not shown).

Models for anaemia risk factors in pregnant women, shown in Table 5, were limited by the small sample size of pregnant women (n = 118). In multivariable models, only older age (3.6, 1.7–7.6 for those 0–49 vs. 20–29 years) and trimester of pregnancy (2.8, 1.2–6.2 and 2.7, 1.3–5.6 for second and third trimester, respectively, relative to first trimester) persisted as significant predictors of anaemia. Risk ratios for anaemia in lower wealth quintiles were also positive, although confidence intervals mostly included 1. No risk factors reached statistical significance in multivariable linear regressions for continuous Hb (not shown).

In models adjusted for survey design, magnitudes of risk associations generally persisted, but fewer confidence intervals excluded zero (Tables S3–S6).

4 | DISCUSSION

This national survey in 2015 revealed the prevalence of anaemia in Bhutan to be 42%, 29%, 36%, and 28% among preschool-age children, adolescent girls, non-pregnant women, and pregnant women, respectively. In children and pregnant women, most anaemia was mild, whereas among adolescent girls and non-pregnant women, about half of all anaemia cases were moderate in severity, although few cases of severe anaemia were observed. In 1985 and 2002, two national anaemia surveys among children under 5 years of age reported prevalence rates of 58% and 81%, respectively (Bailey et al., 2004). Although the former compares with anaemia rates in the region in 1990 (Kassebaum et al., 2014), reasons for the arguably implausible estimate from 2002 remain unclear. Our current finding of a decline to 42% in 2015, however, is consistent with a gradual trend of nutrition and health conditions that are improving in the country (Atwood, Napgal, Mbuya, & Laviolette, 2014; Tobgay, Dorji, Pelzom, & Gibbons, 2011). Trend data for adult women are less comparable, as the 1986 survey in Bhutan examined anaemia only in pregnant women whereas the 2002 survey reported anaemia only in non-pregnant women. Neither survey included adolescents. Nonetheless, the observed high prevalence of anaemia in children and women in 2015 reveals a continued need to better understand its causes and risk factors to improve prevention.

Anaemia prevalence rates reported here for preschoolers and adult women in Bhutan were intermediate within those reported from other countries in South and Southeast Asia over the past two decades (Table S1). For adolescent girls in the region, only Afghanistan has comparable national data, where 30% were reported to have anaemia (Ministry of Public Health of the Islamic Republic of Afghanistan and UNICEF, 2013).

Risk of anaemia varied with age—it was greater in younger than older preschoolers and increased with age in adolescent girls and in adult women—and was higher in the Central and Western than Eastern regions, associations that may help guide causal investigations and prevention. Dietary patterns were not associated with haemoglobin concentration in preschool children, a surprising finding given that dietary diversity remains inadequate among young Bhutanese children (Campbell et al., 2018). Recent analyses of Demographic and Health Survey data in Bangladesh also failed to find associations between anaemia and the WHO-defined indicator of a minimum acceptable diet (frequency and diversity of complementary foods; WHO, 2010) or reported intake of iron-rich foods among children aged 6–23 months (Rawat et al., 2014). A lack of association could reflect non-specificity of dietary quality indicators obtained in a cross-sectional design in revealing usual dietary patterns that increase risk of iron deficiency or could point to a higher burden of causes of anaemia other than iron deficiency.

Household socio-economic status was unrelated to risk of anaemia in preschool-age children. These findings contrast with those of studies in low-income settings of Bangladesh and Nepal, where SES was associated with risk of early childhood anaemia (Balarajan, Fawzi, & Subramanian, 2013; Khan, Awan, & Misu, 2016; Rohner et al., 2013), but agree with findings of population studies in Cambodia—& Subramanian, 2013; Khan, Awan, & Misu, 2016; Rohner et al., 2013) and UNICEF, 2013).
poorer, more rural areas who consume tube well water drawn from aquifers laden with iron relative to women in more urban areas who drink piped water (Merrill et al., 2011; Rahman et al., 2016). To our knowledge, no studies of groundwater iron in Bhutan have been conducted, but it could be an avenue for further investigation.

We did, however, observe the risk of anaemia to be higher among stunted preschool children; adolescent girls living at home rather than at school and those with employment rather than studying; and uneducated, unmarried or separated women, and mothers not having a preschool-aged child. These risk factors possibly reflect, respectively, chronic undernutrition underlying anaemia risk in children, adolescent girls being protected somewhat by school-based iron supplementation programmes that have been in place since 2004 (Ministry of Health and UNICEF, 2014), and perhaps durable effects of antenatal iron supplementation and other health system contact among women who had been pregnant recently. These observed relationships offer evidence of groups within the population who are at greater risk of anaemia, but modest risk ratios and the possibility of residual confounding should be considered.

The possibility that routine health care including iron-folic acid (IFA) supplementation during pregnancy reduced anaemia risk in pregnant women is supported by the observation of nearly 20% lower anaemia prevalence among pregnant than non-pregnant women (despite higher iron demands during pregnancy). This difference, combined with previously reported high national attendance at antenatal care (98%) and adherence to daily prenatal IFA supplementation (87%; Nutrition Program, 2015), suggests that regular IFA supplementation through the country’s antenatal care system is effectively protecting many women from anaemia during pregnancy and may be building iron stores that confer extended postpartum protection against iron deficiency anaemia. However, the fact that gestational anaemia is not fully prevented with extensive IFA coverage may suggest that either the supplemental iron being delivered is still inadequate, other hematric nutrient deficiencies are not being prevented (Fishman, Christian, & West, 2000) or causes other than iron deficiency contribute in many anaemia cases (e.g., thalassemia), as has been reported from Bangladesh (Merrill et al., 2012). Such causes are known to vary markedly by population (Balarajan et al., 2011), and the burdens of parasitic infections, other micronutrient deficiencies, and genetic haemoglobinopathies are not well described in Bhutan. Evidence from elsewhere in South and Southeast Asia, however, suggests that in certain subpopulations within the region, rates of hemoglobinopathies are very high (Fucharoen & Winichagoon, 2011; Hossain et al., 2017; Wah, Yi, Khin, Plabplueng, & Nuchnoi, 2017), appearing to warrant their assessment and public health management. Better understanding the contributions of thalassemia and other causes of anaemia to be examined across these multiple life stages.

The present study benefits from having analysed data from a nationally representative anaemia survey that allowed anaemia status to be linked to household and individual characteristics. Additionally, anaemia assessment in a large sample of young children, adolescent girls, and pregnant and non-pregnant women of reproductive age is an unusual feature of an NNS that enabled prevalence, trends, and risk factors for anaemia to be examined across these multiple life stages. The lack of status data on iron, other micronutrients, and inflammation limited our ability to differentiate causes of anaemia. Additionally, the small sample of pregnant women limited the risk factor analysis in that group.

Bhutan's 2015 NNS reveals a setting of surprisingly high anaemia prevalence given the relatively good nutritional status in children and women. New and enhanced programmes are needed to prevent anaemia in preschool-age children, adolescent girls, and women of reproductive age to protect the growth, development, and wellbeing of these vulnerable groups. Identification of few sociodemographic or nutritional/dietary risk factors for anaemia coupled with lack of data on iron deficiency, inflammation, and other aetologies of anaemia warrants further investigation into causes of anaemia in Bhutan. Such data combined with the present findings may be expected to better inform the design and implementation of anaemia control strategies in children, adolescent girls, and women of reproductive age.

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CONFLICTS OF INTEREST

The authors declare that they have no conflicts of interest.

CONTRIBUTIONS

RKC, VMA, YK, and KPW designed the study; VMA, LD, and VJ led the design and implementation of the NNS 2015; VMA, LD, VJ, JLW, SDG, and NH prepared and analysed the data for the survey report; RKC and YK conducted data analysis for this study; RKC wrote the manuscript. All authors read and approved the manuscript; KPW had primary responsibility for the final content.

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