# Changes in the prevalence of anemia in Korean adolescents, 1998-2018 

Jun Young An, MD, Yoo Rha Hong, MD, Seom Gim Kong, MD<br>Department of Pediatrics, Kosin University Gospel Hospital, Kosin University College of Medicine, Busan, Korea

Background: Anemia is an important health problem affecting approximately $25 \%$ of the global population. Although its prevalence is decreasing worldwide, few studies have examined the prevalence of anemia in Korean adolescents.
Purpose: This study aimed to determine the prevalence of anemia in Korean adolescents over the past 21 years using data from the Korea National Health and Nutrition Examination Survey.
Methods: We investigated the prevalence of anemia in adolescents aged 10-18 years for the period 1998-2018 according to sex, age, residential area, and household income. The effects of menarche age and menstruation were examined in female adolescents for the period 2001-2018.
Results: Among the total 11,782 participants, the weighted prevalence of anemia was $4.0 \%$. The prevalence of anemia in male participants significantly decreased (from $3.0 \%$ to $0.5 \%$, $P<0.001$ ), whereas that in female participants did not change significantly over time (from $7.9 \%$ to $8.5 \%, P=0.054$ ). The average age at menarche was $12.4 \pm 0.0$ years; the prevalence of anemia in females increased from age 13 years onward. Multivariate analysis revealed that anemia was more common in female than male participants (odds ratio [OR], 9.88; 95\% confidence interval [CI], 7.19-13.57; $P<0.001$ ). In female adolescents, the prevalence of anemia increased with age (OR, 1.26 ; 95\% CI, 1.15-1.38, $P<0.001$ ) and was 3.6 times higher after than before menarche (OR, 3.65; 95\% CI, 2.21-6.05, $P<0.001$ ).
Conclusion: Over the 21-year study period, the prevalence of anemia decreased among male adolescents but did not change in female adolescents. In female adolescents, age and menarche were identified as significant risk factors that require continued attention.

Key words: Anemia, Prevalence, Adolescent, Menarche, Menstruation

## Key message

Question: Over the past 21 years, has the prevalence of anemia decreased among Korean adolescents?
Finding: The prevalence of anemia in boys aged 10-18 years decreased from $3.0 \%$ to $0.5 \%$ over the study period, whereas that in girls did not change significantly over time (increased from 7.9\% to 8.5\%).
Meaning: The prevalence of anemia in female adolescents remains high, requiring attention and efforts to improve it.

## Introduction

Anemia is a common health problem that has been identified in one quarter of the global population. ${ }^{1)}$ Prevalence of anemia is particularly high in preschool children and women of childbearing age. The World Health Organization (WHO) reported that the prevalence of anemia is $48 \%$ in preschool-aged children, $25 \%$ in school-aged children, and 30\% in nonpregnant women. ${ }^{1)}$ In growing children and adolescents, chronic anemia adversely affects growth and cognitive development. ${ }^{2)}$ In particular, iron deficiency anemia (IDA) is associated with low cognitive performance scores not only in infants, but also in school-aged children and adolescents. ${ }^{3,4)}$

Before 2000, several studies were conducted on the prevalence of anemia in adolescents in Korea. According to these studies, the prevalence of anemia in adolescents aged 10-24 years was 3.5\%$4.5 \%$ for males and $7.9 \%-17.1 \%$ for females ${ }^{5,6}$ In 2018, there was a large-scale study on the prevalence of anemia in Korean over the age of $15 . .^{7}$ The prevalence of anemia was $0.33 \%$ in males aged $15-29$ and $6.08 \%$ in females, showing a decrease in prevalence compared to that previously reported.' However, individuals younger than 15 years were not included in the study, and there was no further analysis of the clinical information.

The Korea National Health and Nutrition Examination Survey (KNHANES) has been conducted since 1998. This survey includes large-scale health data representing the Korean population. In this survey, blood tests are conducted on subjects

[^0]10 years of age or older. Using the KNHANES data from 1998 to 2018, we investigated the prevalence of anemia among adolescents at 10 years or older. In addition, we sought to identify factors associated with anemia.

## Methods

## 1. Study population

This study was conducted using data from the KNHANES conducted by the Korea Centers for Disease Control and Prevention (KCDC). The KNHANES is a project that seeks to improve health by investigating health behaviors, nutritional status, and prevalence of chronic diseases. The KNHANES was initiated in 1998, and we conducted this study based on data spanning 21 years between 1998 and 2018: KNHANES I (1998), KNHANES II (2001), KNHANES III (2005), KNHANES IV (2007-2009), KNHANES V (2010-2012), KNHANES VI (2013-2015), and KNHANES VII (2016-2018). The KNHANES was implemented once every 3 years from 1998 to 2005, and has been conducted annually since 2007. Since 2007, this survey was conducted with approval of the Institutional Review Board (IRB) of the KCDC (2007-02CON-04-P, 2008-04EXP-01-C, 2009-01CON-03-2C, 2010-02CON-21-C, 2011-02CON-06-C, 2012-01EXP-01-2C, 2013-07CON-03-4C, 2013-12EXP-035C, 2018-01-03-P-A). KNHANES was implemented without approval from 2015 to 2017 in accordance with the opinion of the IRB of the KCDC.

During the study period, 25,692 adolescents aged 10-18 years were included in the survey. Among them, we analyzed the data from 11,782 adolescents who underwent blood tests.

## 2. Study variables and definition

We investigated differences in age, sex, residential area, household income, height, weight, waist circumference, and body mass index (BMI) of the subjects in the KNHANES I to VII. Residential area was divided into urban and rural areas. When the administrative district of residential area was '-dong', the area was defined as an urban area, and '-eup/-myeon' was defined as a rural area. Household income was divided into 4th quartiles of low, middle-low, middle-high, and high. BMI status was defined by sex- and age-specific percentiles using the 2017 Korean National Growth Charts: underweight ( $<5$ th percentile), normal (5th $\leq \mathrm{BMI}<85$ th percentile), overweight ( 85 th $\leq \mathrm{BMI}<95$ th percentile), and obese ( $\geq 95$ th percentile). ${ }^{8)}$

The changes in hemoglobin $(\mathrm{Hb})$ level and the prevalence of anemia over time were investigated. According to the WHO guidelines, anemia was defined as a serum Hb level $<11.5 \mathrm{~g} / \mathrm{dL}$ in boys and girls aged $10-11$ years; level $<12 \mathrm{~g} / \mathrm{dL}$ in boys aged $12-14$ years and girls aged 12 years and over; and level $<13$ $\mathrm{g} / \mathrm{dL}$ in boys aged 15 years and over. ${ }^{9)}$ Anemia was classified according to severity, mean corpuscular volume (MCV), and mean corpuscular hemoglobin (MCH). The severity of anemia was characterized as mild for $\mathrm{Hb} 11 \mathrm{~g} / \mathrm{dL}$ or higher, moderate
for $8-11 \mathrm{~g} / \mathrm{dL}$, and severe for $\mathrm{Hb}<8 \mathrm{~g} / \mathrm{dL} .{ }^{9)}$ The normal range of MCV was defined as $78-90.5 \mathrm{fL}$ at age $10,80-93.5 \mathrm{fL}$ at age $11-14$, and $81-95.5 \mathrm{fL}$ at age $15-18 .{ }^{10)}$ The normal range of MCH was defined as $26.3-31.2 \mathrm{pg}$ at age $10,26.6-31.9 \mathrm{pg}$ at age $11-14$, and 27-32.8 pg at age 15-18. ${ }^{10}$
In 2001, questions were added to KNHANES to ask female participants about their age at menarche and current menstruation status. Among those who responded to the survey between 2001 and 2018, 4,730 female participants were interviewed. We investigated whether there was a difference in prevalence of anemia according to menarche, years after menarche and whether or not the participants were menstruating. Additionally, the risk factors for anemia were identified in male participants and in female participants with a menstrual history.

## 3. Statistical analysis

These data were extracted from a complex stratified sample, and were analyzed by applying weights, stratification variables, and cluster variables. The prevalence of anemia was calculated by weighted prevalence. For statistical methods, complex sample crosstabs for categorical variables and complex sample generalized linear regression for continuous variables were used. The risk factors for anemia were analyzed through logistic regression analysis. Statistical significance was defined by $P$ values $<0.05$. All statistical analyses were conducted using IBM SPSS Statistics ver. 25.0 (IBM Co., Armonk, NY, USA).

## Results

## 1. Characteristics of the subjects

A total of 11,782 participants with a mean age of $14.3 \pm 0.0$ years was included (Table 1). There were 6,250 male (53.3\%) and 5,532 female (46.7\%) participants. The proportion of urban residents was $83.8 \%$, which increased from $80.4 \%$ in 1998 to $88.5 \%$ in 2016-2018 ( $P=0.046$ ). The percentage of participants from low-income households decreased from 15.1\% in 1998 to $10.6 \%$ in 2016-2018. The percentage of those from highincome households increased from $26.9 \%$ in 1998 to $33.9 \%$ in 2016-2018 ( $P<0.001$ ). By survey year, there were differences in height, weight, waist circumference, and BMI of the subjects (Table 1).

## 2. Distribution of Hb levels

The mean Hb level of the subjects was $14.0 \pm 0.0 \mathrm{~g} / \mathrm{dL}$. The mean Hb level of males was $14.7 \pm 0.0 \mathrm{~g} / \mathrm{dL}$, and that of females was $13.2 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ (Fig. 1). The Hb level of males increased from $14.4 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ in 1998 to $14.9 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ in 2016-2018 ( $P<0.001$ ). According to age, the mean level of $13.5 \pm 0.0 \mathrm{~g}$ / dL at 10 years increased to $15.5 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ at 18 years for males ( $P<0.001$ ). In female adolescents, the Hb level increased slightly from $13.2 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ in 1998 to $13.3 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ in 2016-2018 ( $P<0.001$ ). The mean Hb level in females decreased with age; the level was highest, $13.6 \pm 0.0 \mathrm{~g} / \mathrm{dL}$, at 11 years and was $12.9 \pm 0.0$
$g / d L$ at 18 years $(P<0.001)$.

## 3. Prevalence of anemia in all subjects

Of all patients, 465 had anemia, a weighted prevalence of 4.0\% (Table 2). The prevalence of anemia was $0.9 \%$ in males and $7.5 \%$ in females. The change in anemia prevalence over time significantly decreased in males, from $3.0 \%$ in 1998 to $0.5 \%$ in 2016-2018 ( $P<0.001$ ). However, the prevalence of anemia in females did not change significantly over time, from 7.9\% in 1998 to $8.5 \%$ in 2016-2018 ( $P=0.054$ ). The prevalence of anemia in males decreased with age from $1.9 \%$ at 10 years to $0.4 \%$ at 18 years ( $P<0.001$ ). In females, the prevalence of anemia was $0.9 \%$ at 10 years of age and gradually increased from 13 years to $13.7 \%$ at 18 years $(P<0.001)$. There were differences in the prevalence of anemia between the age of 10-12 years and 13-18 years in both males and females. In female adolescents, the prevalence of anemia was $1.7 \%$, similar to that of males at the ages of $10-12$ years; but the prevalence was significantly higher, $9.6 \%$, at the ages of 13-18 years (Fig. 2).

There was no difference in anemia prevalence according to residential area. According to household income, the anemia prevalence was $5.4 \%$ for low household incomes and $2.8 \%$ for high household incomes. Therefore, the prevalence was sig.
nificantly higher when the household income was low ( $P=$ 0.001 ). There was no significant difference in males according to household income. However, there was a higher prevalence of anemia in females in the low-income group ( $P=0.002$ ). There was no difference in prevalence of anemia according to BMI status.

Regarding anemia severity, mild anemia was present in 73.4 $\%$, moderate anemia occurred in $25.7 \%$, and severely anemic individuals comprised $0.9 \%$ (Fig. 3). In males, $92.2 \%$ of cases were mild; in females $70.8 \%$ were mild and $28.2 \%$ were moderate ( $P<0.001$ ). According to the MCV of patients with anemia, $36.8 \%$ of cases were microcytic anemia, $61.8 \%$ were normocytic, and $1.4 \%$ were macrocytic. There was no significant difference between males and females ( $P=0.092$ ). According to the MCH of patients with anemia, $52.0 \%$ were hypochromic, $46.9 \%$ were normochromic, and $1.1 \%$ were hyperchromic anemic. Hypochromic anemia accounted for $34.6 \%$ of male cases and $54.4 \%$ of female cases. This difference was statistically significant ( $P=0.006$ ).

## 4. Prevalence of anemia according to menstruation in female adolescents

A total of 4,730 female participants was identified from a survey on menstrual history between 2001 and 2018. The

Table 1. General characteristics of the KNHANES I-VII participants

| Characteristic | Total | KNHANES |  |  |  |  |  |  | $P$ value |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  |  | I (1998) | II (2001) | III (2005) | IV (2007-2009) | V (2010-2012) | VI (2013-2015) | VII (2016-2018) |  |
| No. | 11,782 | 1,634 | 865 | 631 | 2,628 | 2,346 | 1,827 | 1,851 |  |
| Age (yr) | $14.3 \pm 0.0$ | $14.3 \pm 0.1$ | $14.7 \pm 0.1$ | $14.5 \pm 0.1$ | $14.1 \pm 0.1$ | $14.3 \pm 0.1$ | $14.4 \pm 0.1$ | $14.5 \pm 0.1$ | <0.001 |
| Sex (\%) |  |  |  |  |  |  |  |  | 0.936 |
| Male | 53.3 | 52.7 | 53.3 | 53.0 | 53.7 | 54.3 | 52.9 | 52.6 |  |
| Female | 46.7 | 47.3 | 46.7 | 47.0 | 46.3 | 45.7 | 47.1 | 47.4 |  |
| Residential region (\%) |  |  |  |  |  |  |  |  | 0.046 |
| Urban | 83.8 | 80.4 | 81.2 | 86.5 | 83.2 | 82.5 | 83.0 | 88.5 |  |
| Rural | 16.2 | 19.6 | 18.8 | 13.5 | 16.8 | 17.5 | 17.0 | 11.5 |  |
| Household income (\%) |  |  |  |  |  |  |  |  | <0.001 |
| Low | 13.0 | 15.1 | 17.5 | 15.7 | 11.8 | 14.6 | 15.9 | 10.6 |  |
| Middle-low | 25.8 | 24.3 | 22.9 | 29.1 | 23.6 | 29.0 | 27.3 | 23.7 |  |
| Middle-high | 31.5 | 33.8 | 29.3 | 29.1 | 32.0 | 29.3 | 33.7 | 31.8 |  |
| High | 29.7 | 26.9 | 30.2 | 26.1 | 32.6 | 27.2 | 28.0 | 33.9 |  |
| Body measurement |  |  |  |  |  |  |  |  |  |
| Height (cm) | 162.0土0.1 | $159.6 \pm 0.4$ | $163.5 \pm 0.4$ | $163.5 \pm 0.4$ | $161.2 \pm 0.3$ | $161.7 \pm 0.3$ | $161.9 \pm 0.3$ | $162.5 \pm 0.2$ | <0.001 |
| Weight (kg) | $55.3 \pm 0.2$ | $51.7 \pm 0.5$ | $57.0 \pm 0.6$ | $57.0 \pm 0.6$ | $54.2 \pm 0.4$ | $54.7 \pm 0.4$ | $56.2 \pm 0.4$ | $56.0 \pm 0.3$ | <0.001 |
| WC (cm) | $70.4 \pm 0.1$ | $69.0 \pm 0.3$ | $72.3 \pm 0.4$ | $70.8 \pm 0.5$ | $69.7 \pm 0.3$ | $69.5 \pm 0.3$ | 70.9 ${ }^{\text {0.3 }}$ | $70.5 \pm 0.2$ | <0.001 |
| BMI ( $\mathrm{kg} / \mathrm{m}^{2}$ ) | $20.8 \pm 0.0$ | $20.0 \pm 0.1$ | $21.2 \pm 0.2$ | $21.2 \pm 0.2$ | $20.6 \pm 0.1$ | $20.7 \pm 0.1$ | $21.2 \pm 0.1$ | $20.9 \pm 0.1$ | <0.001 |
| BMI status ${ }^{\text {a }}$ (\%) |  |  |  |  |  |  |  |  | <0.001 |
| Underweight | 6.7 | 7.9 | 5.0 | 5.0 | 7.3 | 6.7 | 5.6 | 7.7 |  |
| Normal | 72.8 | 77.8 | 73.5 | 73.5 | 72.5 | 73.3 | 72.2 | 70.3 |  |
| Overweight | 9.6 | 7.7 | 10.4 | 9.4 | 10.7 | 9.1 | 9.9 | 9.6 |  |
| Obese | 6.7 | 11.1 | 12.1 | 9.6 | 10.8 | 12.3 | 12.4 | 10.9 |  |

[^1]

Fig. 1. Mean hemoglobin $(\mathrm{Hb})$ level by age and sex. (A) The mean Hb level of male adolescents increased from $14.4 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ in 1998 to $14.9 \pm 0.0 \mathrm{~g} /$ dL in 2016-2018 ( $P<0.001$ ). With age, Hb increased from $13.5 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ to $15.5 \pm 0.0 \mathrm{~g} / \mathrm{dL}(P<0.001)$. (B) The mean Hb level of female adolescents decreased from $13.5 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ to $12.9 \pm 0.0 \mathrm{~g} / \mathrm{dL}$ with age ( $P<0.001$ ).
average age of menarche was $12.4 \pm 0.0$ years and decreased from $13.3 \pm 0.1$ in 2001 to $11.8 \pm 0.0$ in $2016-2018$ ( $P<0.001$ ). The proportion of girls with a menarcheal experience was $6.7 \%$ at the age of $10,28.8 \%$ at $11,55.2 \%$ at 12 , and above $80 \%$ after 13. In the 4,730 female participants, the proportion without menarcheal experience was $25.4 \%$; of the others, $12.6 \%$ were on their menstruation period, and $61.9 \%$ were not on their period at the time of survey. The prevalence of anemia was $1.8 \%$ in premenarcheal girls, $8.8 \%$ in those not on their period, and $11.7 \%$ of those on their period ( $P<0.001$ ). The mean number of years after menarche of all female participants was $2.5 \pm 0.1$ years while that in anemic female participants was $3.0 \pm 0.1$ years longer than the $2.0 \pm 0.0$ years of the nonanemic group ( $P<0.001$ ). The prevalence of anemia was $3.2 \%, 9.2 \%$, and $14.4 \%$ when the number of years after menarche was less than 1 year, between $1-5$ years, and more than 6 years, respectively ( $P<0.001$ ).

## 5. Factors associated with anemia

Table 3 shows the results of univariate and multivariate analyses of factors associated with anemia in adolescents. In the multivariate analysis of anemia in all adolescents, the odds

Table 2. Weighted prevalence of anemia among the participants

| Variable | Weighted prevalence of anemia (\%) |  |  |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Total | $P$ value | Male | $P$ value | Female | $P$ value |
| Total | 4.0 |  | 0.9 |  | 7.5 |  |
| Survey year |  |  |  |  |  |  |
| I (1998) | 5.3 | 0.003 | 3.0 | <0.001 | 7.9 | 0.054 |
| II (2001) | 4.5 |  | 0.9 |  | 8.6 |  |
| III (2005) | 5.6 |  | 1.0 |  | 10.6 |  |
| IV (2007-2009) | 3.3 |  | 0.6 |  | 6.5 |  |
| $V(2010-2012)$ | 4.0 |  | 1.2 |  | 7.4 |  |
| VI (2013-2015) | 2.7 |  | 0.1 |  | 5.6 |  |
| VII (2016-2018) | 4.3 |  | 0.5 |  | 8.5 |  |
| Age (yr) |  |  |  |  |  |  |
| 10 | 1.4 | <0.001 | 1.9 | <0.001 | 0.9 | <0.001 |
| 11 | 1.8 |  | 1.5 |  | 2.2 |  |
| 12 | 1.4 |  | 1.1 |  | 1.8 |  |
| 13 | 2.8 |  | 0.6 |  | 5.3 |  |
| 14 | 5.3 |  | 0.6 |  | 10.7 |  |
| 15 | 4.2 |  | 1.3 |  | 8.0 |  |
| 16 | 4.8 |  | 0.3 |  | 9.9 |  |
| 17 | 5.6 |  | 1.0 |  | 10.4 |  |
| 18 | 6.5 |  | 0.4 |  | 13.7 |  |
| Age group (yr) |  |  |  |  |  |  |
| 10-12 | 1.6 | <0.001 | 1.5 | 0.008 | 1.7 | <0.001 |
| 13-18 | 4.8 |  | 0.7 |  | 9.6 |  |
| Residential region |  |  |  |  |  |  |
| Urban | 3.9 | 0.238 | 0.8 | 0.144 | 7.4 | 0.603 |
| Rural | 4.5 |  | 1.3 |  | 7.9 |  |
| Household income |  |  |  |  |  |  |
| Low | 5.4 | 0.001 | 1.8 | 0.055 | 9.3 | 0.002 |
| Middle-low | 4.4 |  | 0.7 |  | 8.6 |  |
| Middle-high | 3.9 |  | 0.6 |  | 7.5 |  |
| High | 2.8 |  | 0.9 |  | 5.2 |  |
| BMI status ${ }^{\text {a }}$ |  |  |  |  |  |  |
| Underweight | 3.9 | 0.218 | 1.6 | 0.299 | 7.3 | 0.475 |
| Normal | 4.1 |  | 0.8 |  | 7.8 |  |
| Overweight | 3.4 |  | 1.0 |  | 6.5 |  |
| Obese | 2.9 |  | 0.7 |  | 5.9 |  |

Values are presented as weighted percentages. BMI, body mass index
$P$ values are calculated using complex sample crosstabs. Boldface indicates a statistically significant difference with $P<0.05$.
${ }^{\text {a) }}$ BMI status was defined by sex- and age-specific percentiles as underweight ( $<5$ th percentile), normal (5th $\leq$ BM $<85$ th percentile), overweight ( 85 th $\leq$ BMI $<95$ th percentile), and obese ( $\geq 95$ th percentile).
ratio (OR) for age was 1.22 (95\% confidence interval [CI], $1.17-1.27 ; P<0.001$ ), and the OR for female was 9.88 ( $95 \% \mathrm{CI}$, $7.19-13.57 ; P<0.001$ ). In addition, the prevalence of anemia in individuals with high household income was significantly lower than that in those with low household income (OR, 0.75 ; $95 \% \mathrm{CI}: 0.61-0.93 ; P=0.009$ ). Multivariate analysis in male adolescents showed that the prevalence of anemia decreased over time, with an OR for each 3 years of 0.76 ( $95 \% \mathrm{CI}, 0.66-0.87$; $P<0.001$ ), and the OR for age was 0.83 ( $95 \% \mathrm{CI}, 0.73-0.95$; $P=0.006$ ). In the univariate analysis of 4,730 female adolescents
with a menstrual history, there was a significant correlation between age, household income, menarche experience, years after menarche, and current menstrual status. In multivariate analysis, the risk of anemia increased $26 \%$ as the age increased by 1 year (OR, $1.26 ; 95 \% \mathrm{CI}, 1.15-1.38 ; P<0.001$ ). The prevalence of anemia was lower when the household income was high (OR, $0.75 ; 95 \% \mathrm{CI}, 0.59-0.95 ; P=0.020$ ) and BMI status was overweight (OR, $0.67 ; 95 \% \mathrm{CI}, 0.48-0.94 ; P=0.020$ ). In addition, the prevalence of anemia was 3.7 times higher in females with menarche experience (OR, 3.65; 95\% CI, 2.21$6.05 ; P<0.001$ ).

## Discussion

The prevalence of anemia is decreasing worldwide. DeMaeyer and Adiels-Tegman ${ }^{11)}$ reported a global prevalence of anemia of $30 \%$ in 1960-1985. In contrast, the prevalence of anemia


Fig. 2. Changes in the weighted prevalence of anemia by age and sex. Among male adolescents, the prevalence of anemia was $1.5 \%$ at age 10-12 years, with no difference among survey years ( $P=0.090$ ). The prevalence of anemia was $1.7 \%$ at age 13-18 years, and differences were seen among survey years ( $P<0.001$ ). Among female adolescents, the prevalence of anemia was $1.7 \%$ at age $10-12$ years ( $P=0.212$ ) versus $9.6 \%$ at age $13-18$ years ( $P=0.359$ ), with no difference among survey years.
between 1993 and 2005 according to the WHO decreased to $24.8 \% .^{1)}$ The Global Burden of Diseases, Injuries and Risk Factors (GBD) 2010 study reported a decrease in the prevalence of anemia from $40.2 \%$ to $32.9 \%$ between 1990-2010. ${ }^{12)}$ In this study, we showed a decrease in prevalence of anemia in Korean adolescents 10-18 years old from $5.3 \%$ in 1998 to $4.3 \%$ in 2016-2018. However, the prevalence of anemia was 9 times higher in female adolescents than males. In addition, the prevalence of anemia decreased in male adolescents over time, while there was no change in females.

In this study, the major risk factors for anemia in female participants were age and menarche. The prevalence of anemia in girls younger than 13 years of age was similar to that of same-age boys but increased from the age of 13 years. Similar to our results, the National Health and Nutrition Examination Surveys (NHANES) data from the United States in 2003-2012 showed that the prevalence of anemia increased from $2.1 \%$ to $5.1 \%$ and $7.1 \%$ in females aged 5-11, 12-14, and 15-29 years, respectively. ${ }^{13)}$

In our study, the average age of menarche was 12.4 years. The Hb levels were highest at 11 years of age and decreased from 12 years of age. The prevalence of anemia increased after the average menarcheal age. In other studies in Korea, the prevalence of anemia among 10 - to 18 -year-old female adolescents without menarche was $2.5 \%-3.3 \%$ compared to $10.6 \%-14.9 \%$ in those with menarcheal experience. ${ }^{14,15)}$ In a previous study, the menarcheal age was 13.8 years in Korean female adolescents. ${ }^{16}$ In our study, the age of menarche decreased from 13.3 to 11.8 years over the 21 years. In general, anemia is high in menstruating females older than 15 years. ${ }^{1)}$ However, anemia may develop in younger girls depending on age at menarche.

The main causes of anemia associated with menstruation include IDA and bleeding from obstetric or gynecologic problems. According to a GBD study, $45 \%-50 \%$ of the anemia in females aged 10-19 worldwide was caused by IDA, and 10\% was caused by obstetric and/or gynecologic problems. ${ }^{12)}$ In developed countries, there is a larger proportion of IDA as a cause of anemia than in developing countries. The Biomarkers Reflecting


Fig. 3. Anemia subtypes by severity (A), mean corpuscular volume (MCV) (B), and hemoglobin (MCH) (C).

Table 3. Logistic regression analysis of factors associated with anemia

| Variable | Univariate |  | $P$ value | Multivariate |  | $P$ value |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | OR | 95\% Cl |  | OR | 95\% Cl |  |
| Total |  |  |  |  |  |  |
| Survey year, 3 yr | 0.94 | 0.89-0.99 | 0.017 | 0.95 | 0.90-1.01 | 0.073 |
| Age (yr) | 1.20 | 1.16-1.25 | <0.001 | 1.22 | 1.17-1.27 | <0.001 |
| Female sex | 8.84 | 6.57-11.90 | <0.001 | 9.88 | 7.19-13.57 | <0.001 |
| Rural residence | 1.17 | 0.90-1.52 | 0.238 |  |  |  |
| High household income ${ }^{\text {a) }}$ | 0.70 | 0.58-0.86 | 0.001 | 0.75 | 0.61-0.93 | 0.009 |
| Height $z$ scores ${ }^{\text {b }}$ | 0.93 | 0.85-1.02 | 0.124 | 0.99 | 0.90-1.09 | 0.836 |
| Overweight ${ }^{\text {c }}$ | 0.76 | 0.57-1.00 | 0.046 | 0.77 | 0.58-1.03 | 0.076 |
| Male |  |  |  |  |  |  |
| Survey year, 3 yr | 0.73 | 0.64-0.84 | <0.001 | 0.76 | 0.66-0.87 | <0.001 |
| Age (yr) | 0.87 | 0.77-0.98 | 0.020 | 0.83 | 0.73-0.95 | 0.006 |
| Rural residence | 1.64 | 0.87-3.10 | 0.126 |  |  |  |
| High household income ${ }^{\text {a) }}$ | 0.70 | 0.41-1.20 | 0.198 |  |  |  |
| Height $z$ scores ${ }^{\text {b }}$ | 0.85 | 0.72-0.99 | 0.040 | 0.86 | 0.73-1.03 | 0.102 |
| Overweight ${ }^{\text {c }}$ | 0.98 | 0.47-2.05 | 0.959 | 1.12 | 0.52-2.41 | 0.780 |
| Female |  |  |  |  |  |  |
| Survey year, 3 yr | 0.97 | 0.90-1.05 | 0.444 |  |  |  |
| Age (yr) | 1.29 | 1.23-1.35 | <0.001 | 1.26 | 1.15-1.38 | <0.001 |
| Rural residence | 1.03 | 0.75-1.41 | 0.877 |  |  |  |
| High household income ${ }^{\text {a }}$ | 0.72 | 0.57-0.90 | 0.005 | 0.75 | 0.59-0.95 | 0.020 |
| Height zscores ${ }^{\text {b }}$ | 0.96 | 0.85-1.08 | 0.515 | 0.98 | 0.87-1.11 | 0.778 |
| Overweight) | 0.74 | 0.53-1.02 | 0.061 | 0.67 | 0.48-0.94 | 0.020 |
| Menarche experience | 5.49 | 3.56-8.47 | <0.001 | 3.65 | 2.21-6.05 | $<0.001$ |
| Years after menarche | 1.22 | 1.16-1.28 | <0.001 | 0.96 | 0.87-1.06 | 0.398 |
| Menstrual status ${ }^{\text {d) }}$ |  |  |  |  |  |  |
| Premenarche | Reference |  |  |  |  |  |
| Not menstruating | 5.17 | 3.34-8.00 | <0.001 |  |  |  |
| Menstruating | 7.11 | 4.28-11.80 | <0.001 |  |  |  |

OR, odds ratio; Cl , confidence interval.
$P$ values are calculated using logistic regression analysis.
Boldface indicates a statistically significant difference with $P<0.05$.
${ }^{\text {a) }}$ Household income above middle-high income. ${ }^{\text {b }}$ Height was converted to $z$ scores using the 2017 Korean National Growth Charts. ${ }^{\text {c }}$ When BMI is the 85th percentile or higher by sex- and age-specific percentiles. ${ }^{\text {d) }}$ Menstrual status at the time of the survey.

Inflammation and Nutritional Determinants of Anemia (BRINDA) project examined the prevalence of anemia among women aged $15-49$ years in 10 countries. ${ }^{17)}$ In the BRINDA project, IDA explained $71 \%$ of cases of anemia in low-infection burden countries compared to $35 \%$ in high-infection burden countries. Some reports suggest that IDA in women is associated with years after menarche. In the NHANES data, females aged 12-21 tended to have increased IDA risk when the time since menarche was longer than 3 years. ${ }^{18)}$ In women aged 22-49, the risk ratio of IDA was 1.9 when the years after menarche was longer than 25.

In our study, higher household income and overweight were associated with lower prevalence of anemia in female adolescents. According to the recent WHO report, ${ }^{1)}$ the prevalence of anemia was $42.7 \%$ in low-income countries, $25.6 \%$ in medium-income countries, and $9.1 \%$ in high-income countries. In the BRINDA project, ${ }^{17)}$ the prevalence of anemia was significantly different according to socioeconomic status in individual countries. In countries with low-infection burden, the prevalence of anemia
was $10.0 \%$ and $4.1 \%$ in low- and high-socioeconomic status individuals, respectively. Our research showed similar results to previous studies. In female adolescents only, there was a difference in the prevalence of anemia according to household income. The low prevalence of anemia in overweight females has been reported in other studies. ${ }^{19-21)}$ It is presumed to be related to nutrition, and one study reported higher iron intake in overweight. ${ }^{19)}$

This is a large-scale study that investigated the prevalence of anemia and its possible risk factors in Korean adolescents. Our results reflect representative data from Korea. However, this study did not include children younger than 10 years. In addition, the KNHANES has limitations in collecting detailed information; the KNHANES contains a wide range of data for investigating public health behaviors. Therefore, we were unable to analyze the causes of anemia in our study. In our study, normocytic anemia was the most common type in both males and females, and microcytic anemia and hypochromic anemia were higher in females than males. This is similar to other studies,
and it is speculated that the causes of anemia vary. ${ }^{7,22)}$ To reduce the prevalence of anemia, further studies are needed to determine the causes, and prevention and early diagnosis of disease are important.
As IDA is thought to be the major cause of anemia in female adolescents after menarche, there are recommendations for intake of iron-rich foods starting from the age of 12 and screening for IDA every 5-10 years. ${ }^{23)}$ Female adolescents are recommended to have a screening test for IDA every year in cases of heavy menstruation, bleeding, low iron intake, or past history of IDA. Patients with anemia who do not have iron deficiency or who do not respond to iron supplement therapy need to be actively evaluated for other causes. Although not common, diagnosis of hereditary spherocytosis and thalassemia is increasing in Korea, so attention should be paid to these conditions. ${ }^{24)}$

In conclusion, there was a significant decrease in the prevalence of anemia among male adolescents. In contrast, there was no change in the prevalence of anemia in female adolescents over the 21 years of our study. In female adolescents, age and menarcheal experience were identified as significant risk factors. Continuous attention to these factors is needed.

## Conflicts of interest

No potential conflicts of interest relevant to this article were reported.

ORCID
Yoo Rha Hong (Dittps://orcid.org/0000-0002-7673-070X
Seom Gim Kong (D) https://orcid.org/0000-0003-2361-2221

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[^0]:    Corresponding author: Seom Gim Kong, MD. Department of Pediatrics, Kosin University College of Medicine, 262 Gamcheon-ro, Seo-gu, Busan 49267, Korea EE-mail: ana313@hanmail.net, https://orcid.org/0000-0003-2361-2221
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[^1]:    Values are presented as mean $\pm$ standard error or weighted percentage.
    KNHANES, Korea National Health and Nutrition Examination Survey; WC, waist circumference; BMI, body mass index.
    $P$ values are calculated using complex sample crosstabs for percentage values and complex sample generalized linear regression for mean values. Boldface indicates a statistically significant difference with $P<0.05$.
    ${ }^{\text {a) }} \mathrm{BMI}$ status was defined as sex- and age-specific percentiles as underweight (<5th percentile), normal ( 5 th $\leq \mathrm{BMI}<85$ th percentile), overweight (85th $\leq \mathrm{BMI}$ $<95$ th percentile), and obese ( $\geq 95$ th percentile).

