RESEARCH Open Access

# Check for updates

# Comparative analysis of perinatal outcomes and birth defects amongst adolescent and older Ugandan mothers: evidence from a hospital-based surveillance database

Robert Serunjogi<sup>1\*</sup>, Linda Barlow-Mosha<sup>1</sup>, Daniel Mumpe-Mwanja<sup>1</sup>, Dhelia Williamson<sup>2</sup>, Diana Valencia<sup>2</sup>, Sarah C. Tinker<sup>2</sup>, Michelle R. Adler<sup>3</sup>, Joyce Namale-Matovu<sup>1</sup>, Dennis Kalibbala<sup>1</sup>, Jolly Nankunda<sup>4</sup>, Evelyn Nabunya<sup>5</sup>, Doreen Birabwa-Male<sup>5</sup>, Josaphat Byamuqisha<sup>5</sup> and Philippa Musoke<sup>1,4</sup>

#### **Abstract**

**Background:** Uganda has one of the highest adolescent pregnancy rates in sub-Saharan Africa. We compared the risk of adverse birth outcomes between adolescents (age 12–19 years) and mothers (age 20–34 years) in four urban hospitals.

**Methods:** Maternal demographics, HIV status, and birth outcomes of all live births, stillbirths, and spontaneous abortions delivered from August 2015 to December 2018 were extracted from a hospital-based birth defects surveillance database. Differences in the distributions of maternal and infant characteristics by maternal age groups were tested with Pearson's chi-square. Adjusted odds ratios (aORs) and 95% confidence intervals (CI) were calculated using logistic regression to compare the prevalence of adverse birth outcomes among adolescents to mothers 20–34 years.

**Results:** A total of 100,189 births were analyzed, with 11.1% among adolescent mothers and 89.0% among older mothers. Adolescent mothers had an increased risk of preterm delivery (aOR: 1.14; CI 1.06–1.23), low birth weight (aOR: 1.46; CI 1.34–1.59), and early neonatal deaths (aOR: 1.58; CI 1.23–2.02). Newborns of adolescent mothers had an increased risk of major external birth defects (aOR: 1.33; CI 1.02–1.76), specifically, gastroschisis (aOR: 3.20; CI 1.12–9.13) compared to mothers 20–34 years. The difference between the prevalence of gastroschisis among adolescent mothers (7.3 per 10,000 births; 95% CI 3.7–14.3) was statistically significant when compared to mothers 20–34 years (1.6 per 10,000 births; 95% CI 0.9–2.6).

**Conclusions:** This study found that adolescent mothers had an increased risk for several adverse birth outcomes compared to mothers 20–34 years, similar to findings in the region and globally. Interventions are needed to improve birth outcomes in this vulnerable population.

## **Plain English Summary**

Adolescent pregnancies are a global problem occurring in high-, middle-, and low-income countries with Uganda having one of the highest adolescent pregnancy rates in sub-Saharan Africa. We compared the risk of adverse

Full list of author information is available at the end of the article



© The Author(s) 2021. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third partial in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/. The Creative Commons Public Domain Dedication waiver (http://creativecommons.org/publicdomain/zero/1.0/) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

<sup>\*</sup>Correspondence: rserunjogi@mujhu.org

<sup>&</sup>lt;sup>1</sup> Makerere University-Johns Hopkins University Research Collaboration, P. O. Box 23491, Kampala, Uganda

Serunjogi et al. Reprod Health (2021) 18:56 Page 2 of 10

birth outcomes, including major external birth defects, between adolescents, (age 12–19 years) and mothers (age 20–34 years) in four urban hospitals.

All informative births, including live births, stillbirths, and spontaneous abortions; regardless of gestational age, delivered at four selected hospitals in Kampala from August 2015 to December 2018 were examined. Demographic data were obtained by midwives through maternal interviews and review of hospital patient notes.

Of the 100,189 births, 11.0% were among adolescent mothers and 89.0% among mothers (20–34 years). Adolescent mothers were more likely than mothers (20–34 years) to have an infant with preterm delivery, low birth weight, early neonatal death, and major external birth defects. Adolescent pregnancies were also associated with an increased risk of gastroschisis when compared to mothers (20–34 years).

In conclusion, this study found that adolescent mothers had an increased risk for several adverse birth outcomes compared to mothers 20–34 years. Research on the potential underlying causes or mechanisms for these adverse outcomes among adolescent births is necessary to identify possible interventions.

**Keywords:** Adolescent, Birth outcomes, Birth defects, Gastroschisis, Low birth weight, Early neonatal death, Preterm, Hospital-based surveillance, Sub-Saharan Africa, Uganda

### Introduction

Pregnancies among 15–19 year-old females account for 16 million (11%) births worldwide yet they contribute to 23% of the maternal disease burden attributed to pregnancy and childbirth [1, 2]. The highest prevalence of adolescent pregnancy is found in the sub-Saharan African region, with birth rates of 101 births per 1000 females aged 15–19 years in 2018, higher than the global adolescent birth rate of 44 per 1000 [3].

Uganda has one of the youngest populations in sub-Saharan Africa, with children and adolescents 12–19 years constituting more than half (55%) of the population in 2014 [4], and one of the highest adolescent pregnancy rates (25%) in sub-Saharan Africa [5]. Despite a decline in the fertility rate in Uganda from 6.9 in 2000 to 5.4 in 2016, and an increase in the use of modern contraception from 18% in 2000 to 35% in 2016, adolescent pregnancy remains a challenge with only 7.6% of adolescents having access to contraceptives [5]. Adolescents have also been reported to be less likely to prepare for birth and even be less knowledgeable about obstetric danger signs compared to older mothers who were not knowledgeable [6], potentially increasing the risk of adverse birth outcomes.

Although several studies have found a higher risk of adverse birth outcomes such as preterm birth, low birth weight (LBW), and early neonatal deaths (ENND), with adolescent births, [1, 7–13], some studies have not found an association for some adverse birth outcomes [14–19]. Some possible reasons for such differences in results could be the sample size or categorization of age groups among the adolescents and comparative age group. In addition, a systematic literature review and meta-analysis on adolescent childbearing in Sub-Saharan Africa by Gronvik et al. (2018) [19] showed that most studies were

primarily hospital or health clinic-based patient record reviews and therefore may not be representative of the general population.

Most studies [7, 10, 14, 19] that have reported birth outcomes among adolescent births in Sub-Saharan Africa have not reported the burden of major external birth defects among infants born to adolescents. There have also been a limited number of studies [21, 22] in the Sub-Saharan Africa region that have documented the prevalence and the risks of major external birth defects among adolescent births in comparison with births from mothers over 19 years of age. This may be as a result of limited data on these conditions probably as a result of inadequate birth defect registry systems [20].

Therefore, using a large dataset obtained from an ongoing hospital-based birth defect surveillance study, we compared the occurrence of adverse birth outcomes (preterm birth, LBW, and ENND), including the rates and prevalence of specific major external birth defects among adolescent mothers (12–19 years) and mothers (20–34 years) in Uganda, a low-middle income setting. The findings from this study would therefore be used as a benchmark for researchers and policymakers to understand the current estimate of the burden of adverse birth outcomes among adolescent births in a low-income Sub-Saharan African country.

#### **Methods**

We extracted and analyzed verified data collected between August 2015 and December 2018 from an ongoing birth defects surveillance system implemented at four major hospitals in Kampala, Uganda [23]. These hospitals have approximately 50,000 births annually, which make up more than 55% of all births in Kampala. The details of the birth defects surveillance system are described

Serunjogi et al. Reprod Health (2021) 18:56 Page 3 of 10

elsewhere [23]. Briefly, this birth defects surveillance system collected information from hospital records including: demographic (maternal age, delivery site), maternal health (maternal HIV status, obstetric history), and birth outcome (mode of delivery, pregnancy outcome, infant sex, gestational age, and infant examinations). Information on maternal HIV status and antiretroviral therapy was obtained from antenatal records and inpatient hospital records. Information on all live births, stillbirths, and spontaneous abortions was collected between the time of birth and discharge which usually occurs within the first 24 h after delivery [23]. Infants born outside the four hospitals and uninformative macerated stillbirths were not included in the surveillance system.

We defined adolescent births for this analysis as those occurring in women 12-19 years of age at delivery and the comparative group as births among women 20-34 years of age at delivery. There were no births to women younger than 12 years of age, and births of mothers  $\geq 35$  years of age were excluded because the risk for adverse obstetrical and perinatal outcomes has been shown to increase over age 34 [24, 25].

We defined gestational age as the interval between the date of delivery and the last menstrual period (LMP) in completed weeks; if the LMP was unknown or missing, a clinical estimate of gestational age was used, such as estimates from fundal height or abdominal ultrasound. We defined preterm delivery as live births occurring at gestations of less than 37 weeks. Low birth weight (LBW) was defined as an infant weighing less than 2,500 g measured within 24 h after birth using digital scales among term  $(\geq 37 \text{ weeks})$  live births. Early neonatal death (ENND) was defined as death among live neonates born at term during the first 48 h or before the mother was discharged from the hospital. Stillbirth was defined as a baby born with no signs of life at or after 28 weeks' gestation, while a spontaneous abortion was defined as fetal death at less than 28 weeks' gestation. Birth defects were confirmed through bedside examination by a physician and review of photographs, narrative descriptions, and or drawings by a birth defects expert who verified or reassigned the diagnosis code. Details of the birth defect ascertainment and classification have been described previously [23]. The definition of birth defects included in this analysis can be found in the Additional file 1.

Data were analyzed using STATA version 15 statistical software (StataCorp. 2017. College Station, TX: Stata-Corp LLC). Descriptive statistics of maternal and infant characteristics by maternal age group were calculated as frequencies and percentages, and the differences between proportions were tested with Pearson's chi-square test.

We used multivariable logistic regression analysis to estimate crude and adjusted odds ratios (cORs and aORs, respectively) along with their 95% confidence intervals (CIs) for the associations between adolescent births, and adverse birth outcomes with the 20-34 years age group as the reference. Separate multivariable logistic regression models were generated for preterm birth, LBW, ENND, each major birth defect category (neural tube defects, malformations of the eyes and ears, orofacial clefts, and malformations of the musculoskeletal system), and each of the 16 specific birth defects. The analysis of preterm birth was limited to live births; while that of LBW and ENND was limited to term live births. The following covariates were considered for adjustment: parity, mode of delivery, singleton/multiple delivery, number of antenatal visits, and initiation time of prenatal care. The specific covariates used in each model were selected based on previous studies [7, 14, 26-28], and excluded possible collider variables using directed acyclic graphs (DAGs) to evaluate confounding [29]. The variables considered in the models are shown in Table 1 and the final variables included in the models are listed as a footnote in Table 2.

Birth prevalence per 10,000 births for seven categories of major external birth defects and 16 specific birth defects [23] was calculated by each maternal age group along with 95% Wilson's CIs.

Those defects that were considered genetic, for example, anencephaly with Spina Bifida or OEIS complex (omphalocele-exstrophy-imperforate anus-spinal defects) which comprises a combination of defects including omphalocele, exstrophy of the cloaca, imperforate anus, and spinal defects, were included in prevalence estimates, but not be included in etiologic or risk factor analysis since the etiology of the defects are known or suspected [30, 31].

#### Raculto

A total of 96,938 pregnancies with 100,189 births among mothers 12 to 34 years of age were captured. Of these, 11,028 (11.0%) births were among adolescent mothers and 89,161 (89.0%) births were among mothers (20–34 years). The age distribution of the study population of all mothers aged 12 to 34 years is shown in Fig. 1.

Table 1 shows the maternal and infant characteristics by age group. The proportion of mothers with HIV infection was significantly lower in adolescent mothers (p<0.001) but a significantly higher proportion of HIV-infected adolescents had not been initiated on antiretroviral therapy (ART) by the time of delivery compared to mothers (20–34 years) (p<0.001). Adolescent mothers were less likely to have attended any antenatal care (ANC), attended the recommended four or more antenatal visits [32], or attended the first antenatal visit within the first trimester (p<0.001) compared to mothers (20–34 years). Also, adolescents were more likely to have been

Serunjogi et al. Reprod Health (2021) 18:56 Page 4 of 10

Table 1 Maternal and reproductive characteristics of adolescent mothers 12–19 and mothers 20–34 years of age

|   | Total, n (%)  | Maternal age, n (% | p-Value       |         |
|---|---------------|--------------------|---------------|---------|
|   |               | 12–19 years        | 20–34 years   |         |
| No. of births   | 100,189 (100) | 11,028 (11.0)      | 89,161 (89.0) | -       |
| No. of mothers  | 96,938 (100)  | 10,783 (11.1)      | 86,155 (88.9) | _       |
| Maternal age  |               |                    |               |         |
| Median; Inter-quartile range (IQR)                        | 25; 22–29     | 18; 18–19          | 26; 23–29     | -       |
| Hospital <sup>b</sup>                                     |               |                    |               |         |
| Lubaga  | 6410 (6.4)    | 134 (1.2)          | 6276 (7.0)    | < 0.001 |
| Mengo   | 7905 (7.9)    | 111 (1.0)          | 7794 (8.7)    |         |
| Nsambya   | 7531 (7.5)    | 99 (0.9)           | 7432 (8.3)    |         |
| Mulago national referral                                  | 78,343 (78.2) | 10,684 (96.9)      | 67,659 (75.9) |         |
| Maternal HIV status <sup>a</sup>                          |               |                    |               |         |
| Positive  | 8167 (8.4)    | 480 (4.5)          | 7687 (8.9)    | < 0.001 |
| Negative  | 88,631 (91.4) | 10,282 (95.3)      | 78,349 (91.0) |         |
| Unknown   | 140 (0.1)     | 21 (0.2)           | 119 (0.1)     |         |
| Maternal antiretroviral therapy (ART) at delivery $\beta$ |               |                    |               |         |
| Yes   | 7786 (95.3)   | 438 (91.3)         | 7348 (95.6)   | < 0.001 |
| No  | 381 (4.7)     | 42 (8.8)           | 339 (4.4)     |         |
| Maternal timing of initiation on $ART^\Pi$                |               |                    |               |         |
| Before conception   | 4161 (53.4)   | 133 (30.4)         | 4028 (54.8)   | < 0.001 |
| After conception  | 3625 (46.6)   | 305 (69.6)         | 3320 (45.2)   |         |
| Mother referred from other health center <sup>a</sup>     |               |                    |               |         |
| Yes   | 44,700 (46.1) | 7541 (69.9)        | 37,159 (43.1) | < 0.001 |
| No  | 52,238 (53.9) | 3242 (30.1)        | 48,996 (56.9) |         |
| Maternal parity <sup>a</sup>                              |               |                    |               |         |
| Primipara (1)   | 32,765 (33.8) | 9023 (83.7)        | 23,742 (27.6) | < 0.001 |
| Multipara (≥ 2)   | 64,173 (66.2) | 1760 (16.3)        | 62,413 (72.4) |         |
| Mode of delivery <sup>b</sup>                             |               |                    |               |         |
| Vaginal   | 68,756 (68.6) | 8575 (77.8)        | 60,181 (67.5) | < 0.001 |
| Caesarean section   | 31,433 (31.4) | 2453 (22.2)        | 28,980 (32.5) |         |
| Singleton/multiple deliveries <sup>b</sup>                |               |                    |               |         |
| Singleton   | 93,548 (93.4) | 10,516 (95.4)      | 83,032 (93.1) | < 0.001 |
| Multiple  | 6641 (6.6)    | 512 (4.6)          | 6129 (6.9)    |         |
| Received antenatal care (maternal) <sup>a</sup>           |               |                    |               |         |
| Yes   | 94,734 (97.7) | 10,403 (96.5)      | 84,331 (97.9) | < 0.001 |
| No  | 2204 (2.3)    | 380 (3.5)          | 1824 (2.1)    |         |
| Timing of first antenatal care (ANC) visit $^{a, \xi}$    |               |                    |               |         |
| ANC within 1st Trimester                                  | 6446 (7.9)    | 580 (6.6)          | 5866 (8.0)    | < 0.001 |
| ANC within 2nd Trimester                                  | 36,783 (44.9) | 3976 (45.3)        | 32,807 (44.8) |         |
| ANC within 3rd Trimester                                  | 38,696 (47.2) | 4217 (48.1)        | 34,479 (47.1) |         |
| Number of maternal antenatal visits <sup>a,π</sup>        |               |                    |               |         |
| No ANC Visit  | 2204 (2.3)    | 380 (3.5)          | 1824 (2.1)    | < 0.001 |
| 1–3 Visits  | 52,764 (54.6) | 6626 (61.5)        | 46,138 (53.7) |         |
| 4+Visits  | 41,731 (43.2) | 3761 (34.9)        | 37,970 (44.2) |         |

<sup>&</sup>lt;sup>a</sup> Denominator is the number of mothers

 $<sup>^{\</sup>rm b}~$  Denominator is the number of births

 $<sup>^{\</sup>beta}$  Denominator is the number of HIV positive mothers (n = 8167)

 $<sup>^{\</sup>rm II}\,$  Denominator is the number of HIV positive mothers on ART (n = 7786)

 $<sup>^{\</sup>xi}\,$  10,605 Mothers missing date of first ANC visit

 $<sup>^{\</sup>pi}$  239 Mothers missing the number of ANC visits

Serunjogi et al. Reprod Health (2021) 18:56 Page 5 of 10

Table 2 Comparison of perinatal outcomes between adolescent mothers 12-19 and mothers 20-34 years of age

| •                                | •             |                     |               |                  |                      | , -              |                      |
|----------------------------------|---------------|---------------------|---------------|------------------|----------------------|------------------|----------------------|
|                                  | Total, n (%)  | Maternal age, n (%) |               | cOR (95% CI)*    | p-Value <sup>c</sup> | aOR (95% CI) *   | p-value <sup>d</sup> |
|                                  |               | 12–19 Years         | 20-34 Years   |                  |                      |                  |                      |
| Gestational age <sup>a</sup>     |               |                     |               |                  |                      |                  |                      |
| < 37 weeks                       | 8564 (9.0)    | 1068 (10.2)         | 7496 (8.8)    | 1.18 (1.10-1.26) | < 0.001              | 1.14 (1.06–1.23) | 0.001                |
| ≥ 37 weeks                       | 86,839 (91.0) | 9358 (89.8)         | 77,481 (91.2) | 1                |                      | 1                |                      |
| Birth outcome                    |               |                     |               |                  |                      |                  |                      |
| Live birth                       | 95,403 (95.2) | 10,426 (94.5)       | 84,977 (95.3) | 1                |                      | 1                |                      |
| Stillbirth                       | 3102 (3.1)    | 359 (3.3)           | 2743 (3.1)    | 1.07 (0.95-1.19) | 0.258                | 1.08 (0.95-1.22) | 0.230                |
| Spontaneous Abortion             | 1684 (1.7)    | 243 (2.2)           | 1441 (1.6)    | 1.37 (1.19-1.58) | < 0.001              | 0.94 (0.83-1.11) | 0.488                |
| Infant birth weight (≥ 37 w      | reeks)a       |                     |               |                  |                      |                  |                      |
| < 2500 g                         | 6572 (7.6)    | 986 (10.5)          | 5586 (7.2)    | 1.51 (1.41-1.63) | < 0.001              | 1.46 (1.34-1.59) | < 0.001              |
| ≥ 2500 g                         | 80,267 (92.4) | 8372 (89.5)         | 71,895 (92.8) | 1                |                      | 1                |                      |
| ENND (≥ 37 weeks) <sup>a,b</sup> |               |                     |               |                  |                      |                  |                      |
| Yes                              | 441 (0.5)     | 82 (1.0)            | 359 (0.5)     | 1.96 (1.57-2.45) | < 0.001              | 1.58 (1.23-2.02) | < 0.001              |
| No                               | 82,159 (99.5) | 8511 (99.0)         | 73,648 (99.5) | 1                |                      | 1                |                      |
| Birth defect                     |               |                     |               |                  |                      |                  |                      |
| No                               | 99,674 (99.5) | 10,954 (99.3)       | 88,720 (99.5) | 1                |                      | 1                |                      |
| Yes¥                             | 515 (0.5)     | 74 (0.7)            | 441 (0.5)     | 1.36 (1.06–1.74) | 0.015                | 1.36 (1.02–1.80) | 0.032                |

<sup>&</sup>lt;sup>a</sup> Live births only (n= 95,403)

Gestational age model was restricted to live births only with adjustment for parity, mode of delivery, singleton/multiple deliveries, and number of antenatal visits

Birth outcome model was adjusted for parity, mode of delivery and number of antenatal visits

Early neonatal death model was restricted to full-term infants (gestation  $\ge$  37 weeks) and adjusted for parity, mode of delivery and number of antenatal visits Birth weight model was restricted to full-term infants (gestation  $\ge$  37 weeks) and adjusted for parity, mode of delivery, singleton/multiple deliveries and number of antenatal visits

Overall birth defect model was adjusted for parity, mode of delivery, singleton/multiple births and number of antenatal visits. \*Newborns with at least one of the sixteen major external birth defects of interest to the study

referred from another health center for delivery, contributing 70% of referred women. Adolescent mothers were also more likely than mothers (20-34 years) to be primipara, have vaginal deliveries, and have singleton deliveries (p < 0.001).

Adolescent mothers were significantly more likely than mothers (20–34 years) to have preterm (<37 weeks) live births (aOR: 1.14; 95% CI 1.06–1.23, p=0.001) (Table 2). Among live births delivered at term, adolescents were at higher risk of delivering a LBW infant (aOR: 1.46; 95% CI 1.34–1.59; p<0.001) and early neonatal death (aOR: 1.58; 95% CI 1.23–2.02; p<0.001) (Table 2). Adolescents were also more likely to have a spontaneous abortion (cOR:1.37 95% CI 1.19–1.58; p<0.001), but after adjusting for confounders the association was not statistically significant (Table 2).

Adolescent mothers had a higher prevalence of birth defects (67.1 per 10,000 births, 95% CI 53.5–84.2) compared to mothers (20–34 years) (49.7 per 10,000 births, 95% CI 45.3–54.5). The odds of major external birth

defects were higher among adolescents in comparison to mothers (20–34 years) (aOR: 1.36; 95% CI 1.02–1.80; p=0.032). Talipes equinovarus was the most prevalent major external birth defect among adolescent mothers (19.9 per 10,000 births; 95% CI 13.2–30.2) (Fig. 1) The prevalence estimates (per 10,000 births) of 10 birth defects (Encephalocele, microcephaly, anophthalmia; microphthalmia, all oral-facial clefts, talipes equinovarus, limb reduction defects, omphalocele, and gastroschisis) were higher among adolescent mothers, however, only the difference between the prevalence of gastroschisis among adolescent mothers (7.3 per 10,000 births; 95% CI 3.7–14.3) was statistically significant when compared to mothers (20–34 years) (1.6 per 10,000 births; 95% CI 0.9–2.6) (Fig. 2).

Adolescent mothers were significantly more likely to have an infant born with microcephaly and gastroschisis. However, after adjustment for parity and initiation time of prenatal care, only gastroschisis (aOR: 3.20; 95% CI 1.12–9.13) remained significantly associated with

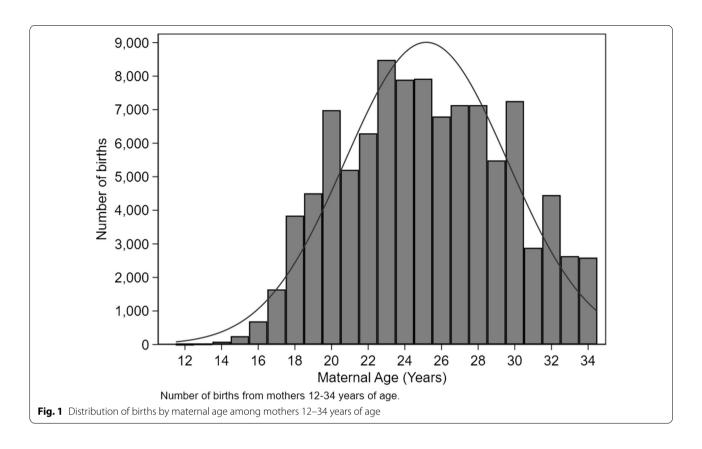
<sup>&</sup>lt;sup>b</sup> Early neonatal death (ENND); term births (n= 86,839)

c p-value for cOR

 $<sup>^{\</sup>rm d}\,$  p-value for aOR

<sup>\*</sup>The cOR (95% CI) and aOR (95% CI) were calculated with 20-34 years as the reference age group

Serunjogi et al. Reprod Health (2021) 18:56 Page 6 of 10



adolescent pregnancy (Table 3). Musculoskeletal defects (aOR: 1.69; 95% CI 1.15–2.50) and malformations of eyes and ears (aOR: 3.09; 95% CI 1.01–9.42) were also significantly higher among adolescent births compared to those from mothers (20–34 years) (Table 3).

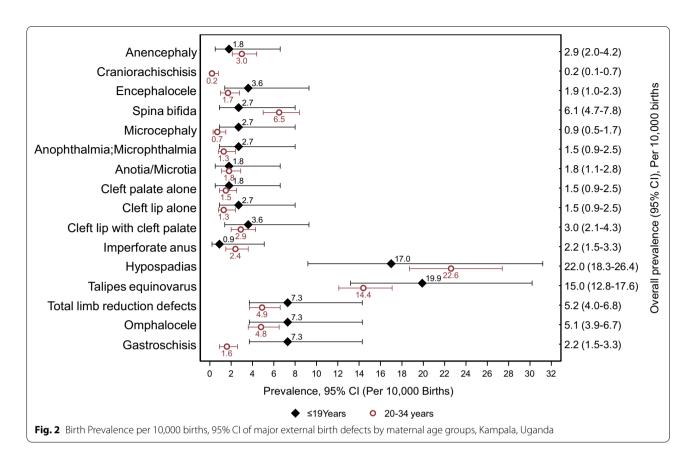
### Discussion

In this study, we observed that adolescent mothers were more likely to have an infant with the adverse birth outcome of preterm delivery, LBW, ENND, or a major external birth defect such as gastroschisis as compared to mothers 20-34 years. Previous studies have also found an increased risk for preterm delivery in adolescent births [7, 14, 19], which could be attributable to the maternalfetal competition for nutrients that arises when pregnancy coincides with continuing or incomplete growth in adolescents. Our study finds that adolescent mothers were more likely to deliver LBW babies, and is consistent with results from the Uganda Demographic Health Survey (UDHS) 2011 [33], and several other studies in sub-Saharan Africa [14, 34-36]. That UDHS also identified infants born with LBW to be at increased risk of neonatal death [37], highlighting the risks associated with LBW in this population. The LBW observed among infants born to adolescent mothers could have been due to factors such as inadequate maternal nutrition, or the related but distinct issue of inadequate weight gain during pregnancy [26], which were not assessed in our study.

Comparable to findings from a study exploring the impact of early motherhood on neonatal mortality in 45 low and middle-income countries [8], our study showed that ENNDs in full-term babies occurred more frequently among adolescent mothers. In contrast, a World Health Organization (WHO) multi-country survey across 29 countries in Africa, Asia, Latin America, and the Middle East found that ENND among infants born to adolescent mothers was not significantly different from mothers aged 20-24 years, after adjustment for gestational age and birth weight [7]. This difference may be related to restriction in the WHO study to mothers aged 24 years or younger who gave birth to an infant of at least 22 weeks' gestation as compared to mothers ≤ 34 years in our analysis, and the WHO study's classification of ENND as intra-hospital deaths that occurred within 7 days after birth as compared to deaths within 48 h in our analysis.

In this study, adolescent mothers were more likely to deliver a newborn with a birth defect when compared with mothers 20–34 years. These findings are consistent with findings from studies in North America and Europe [38, 39]. Our finding of a higher birth defects prevalence estimate (per 10,000 births) among adolescent mothers compared to older mothers is consistent with findings

Serunjogi et al. Reprod Health (2021) 18:56 Page 7 of 10



from a population-based prevalence study using data from EUROCAT congenital anomaly registers in 23 regions of Europe in 15 countries [39]. However, Zile and Villerusa et al. (2013), from a study based on data from the Medical Birth Register in Latvia differed showing that the prevalence of birth defects was instead higher for mothers aged 20–34 years as compared to adolescent mothers [40]. The difference could however be attributed to the fact that our study's prevalence estimates included births from all live births, stillbirths, and spontaneous abortions while Zile and Villerusa et al. (2013) included only live births and also included other defects/syndromes and chromosomal defects.

Although the number for some birth defects were small in our study, our findings suggest that gastroschisis was significantly higher among adolescent mothers when compared to mothers 20–34 years. The strong association between adolescents births with gastroschisis has also been reported by other studies [28, 38, 39, 41]. While comparing gastroschisis to other congenital anomalies, Given et al. (2017) reported sexually transmitted infections, and continuation of oral contraceptives in early pregnancy, as preventable risk factors [42]. However, we were not able to assess these factors in this study.

Our study also found that adolescent mothers were associated with increased odds of musculoskeletal defects as well as malformations of eyes and ears combined. In one retrospective cohort study in the United States of America, [38] Chen et al. (2007) found increased odds of musculoskeletal defects, however, the study included some other defects within the category, specifically, polydactyly/syndactyly/adactyly, diaphragmatic hernia, integumentary anomalies.

We also found that a significantly higher proportion of HIV-infected adolescents were not on ART at conception or delivery compared to women 20–34 years, which is consistent with findings from the Uganda Population-Based HIV Impact Household-based National Survey [43]. Maternal HIV infection has been shown to be associated with increased rates of adverse pregnancy outcomes such as LBW, prematurity, and ENND [44], and the lower prevalence of ART use among HIV-infected adolescents would further exacerbate the situation because it translates to a potential increased risk of MTCT of HIV among adolescents compared to mothers (20–34 years) justifying the need to strengthen services for this population [45].

Serunjogi et al. Reprod Health (2021) 18:56 Page 8 of 10

Table 3 Birth defects among adolescent mothers 12-19 and mothers 20-34 years of age

| ICD-10 RCPCH code <sup>a</sup> | Birth defects                | Number of defects       |     | cOR (95% CI)                   | aOR (95% CI) <sup>d</sup> | p-value |  |
|--------------------------------|------------------------------|-------------------------|-----|--------------------------------|---------------------------|---------|--|
|                                |                              | 12–19 years 20–34 years |     |                                |                           |         |  |
| Neural tube defects (NTD)*     |                              | 9                       | 95  | 0.77 (0.39–1.52)               | 0.63 (0.27–1.52)          | 0.311   |  |
| Q00.0                          | Anencephaly                  | 2                       | 27  | 0.60 (0.14-2.52)               | 0.64 (0.14-2.90)          | 0.559   |  |
| Q00.1                          | Craniorachischisis           | 0                       | 2   | na                             | na                        | na      |  |
| Q01.0-Q01.2, Q01.8-Q01.9       | Encephalocele                | 4                       | 11  | 2.94 (0.94-9.24)               | 1.43 (0.27-7.43)          | 0.673   |  |
| Q05.0-Q05.9                    | Spina bifida                 | 3                       | 56  | 0.43 (0.14-1.38)               | 0.40 (0.08-1.67)          | 0.202   |  |
| Q02                            | Microcephaly                 | 3                       | 6   | 4.04 (1.01–16.17) <sup>β</sup> | 4.54 (0.81-25.39)         | 0.085   |  |
| Malformations of eyes and ears |                              | 5                       | 28  | 1.44 (0.56-3.74)               | 3.09 (1.01-9.42)          | 0.047   |  |
| Q11-Q11.1; Q11.2               | Anophthalmia; Microphthalmia | 3                       | 12  | 2.02 (0.57-7.16)               | 3.21 (0.71-14.38)         | 0.128   |  |
| Q16.0; Q17.2                   | Anotia; Microtia             | 2                       | 16  | 1.01 (0.23-4.40)               | 2.94 (0.55-15.72)         | 0.206   |  |
| Orofacial clefts <sup>b</sup>  |                              | 9                       | 51  | 1.43 (0.70-2.90)               | 1.28 (0.57-2.91)          | 0.549   |  |
| Q35.1-Q35.9, Q38.5, Q87.0      | Cleft palate                 | 2                       | 13  | 1.24 (0.28-5.51)               | 0.71 (0.08-6.12)          | 0.752   |  |
| Q36.0, Q36.9                   | Cleft lip alone              | 3                       | 12  | 2.02 (0.57-7.16)               | 2.54 (0.59-11.50)         | 0.213   |  |
| Q37.0-Q37.9                    | Cleft lip + palate           | 4                       | 26  | 1.24 (0.43-3.56)               | 1.09 (0.35-3.41)          | 0.877   |  |
| Q42.3                          | Imperforate anus             | 1                       | 20  | 0.40 (0.05-3.01)               | 1.06 (0.12-9.08)          | 0.960   |  |
| Q54.0-Q54.3, Q54.8-Q54.9       | Hypospadias <sup>c</sup>     | 10                      | 104 | 0.75 (0.39-1.44)               | 0.63 (0.29-1.34)          | 0.230   |  |
| Musculoskeletal system*        |                              | 45                      | 214 | 1.70 (1.23–2.35) <sup>β</sup>  | 1.69 (1.15-2.50)          | 0.008   |  |
| Q66.0, Q66.8                   | Talipes equinovarus          | 22                      | 128 | 1.41 (0.89-2.22)               | 1.33 (0.77-2.30)          | 0.309   |  |
| Q71.0-Q73.8                    | Total limb reduction         | 8                       | 44  | 1.47 (0.69-3.12)               | 1.75 (0.67-4.56)          | 0.249   |  |
| Q79.2                          | Omphalocele                  | 8                       | 41  | 1.58 (0.74–3.37)               | 2.17 (0.92-5.18)          | 0.078   |  |
| Q79.3                          | Gastroschisis                | 8                       | 14  | $4.62 (1.93-11.02)^{\beta}$    | 3.20 (1.12–9.13)          | 0.030   |  |

<sup>\*</sup> Some infants had more than one type of defect in the neural tube defects and musculoskeletal system categories

## Study strengths and limitations

This study's strengths include a large sample size, which made it possible to assess the association between adolescent births and possible risk factors of adverse birth outcomes. In addition, our study used an active birth defects case ascertainment and collection of data to ensure accuracy and improved birth defect detection and reporting versus extraction of data from medical records. Also, the physical examination of newborns by trained staff and several levels of external birth defect review ensured consistent birth defect classification and coding.

Unlike other studies that only include live births [38, 40], this study included stillbirths, spontaneous abortions, and live births which minimized selection bias especially since some structural birth defects commonly occur among stillbirths thereby giving more accurate risks and birth prevalence estimates among the different age groups.

Study limitations include surveillance activities being conducted at four major urban hospitals located in the capital city and is not representative of adolescent births nationally [5]. However, since 55% of the births in Kampala were at these four hospitals, and one of them (Mulago National Referral Hospital) contributed 60.0% of the total births [23], they provide a fair representation of births nationally. Secondly, because infants were not followed post-discharge, we captured only ENND that occurred within 48 h of birth. The standard definition of ENND is death within seven days of delivery so infants that died between discharge and seven days of life was not accounted for, resulting in a possible underestimation of ENND.

In addition, this study did not control for several risk factors known to influence reproductive health outcomes such as social-economic status, level of education, tobacco smoking, alcohol drinking, maternal nutrition, and the use of folic acid since this information was not captured in the surveillance [28, 46].

Finally, it has been demonstrated that adolescents are not a homogeneous group, and therefore differ in their

a International Classification of Disease 10, Royal College of Paediatrics and Child Health (ICD-10 RCPCH), used to specifically clasify the types of defects included [49]

<sup>&</sup>lt;sup>b</sup> Excluded Q36.1 (medial Cleft lip) because it is suggestive of underlying brain malformation (holoprosencephaly) and chromosomal syndrome [50]

<sup>&</sup>lt;sup>β</sup> Statistically significant at p<0.05

na - Prevalence, cOR, aOR, and 95% confidence intervals that cannot be calculated

<sup>&</sup>lt;sup>c</sup> Denominator for males: N=51,922; 12-19 Years (n= 5,896); 20-34 Years (n= 46,026)

<sup>&</sup>lt;sup>d</sup> Covariates for the birth defect models: parity and initiation time of prenatal care

Serunjogi et al. Reprod Health (2021) 18:56 Page 9 of 10

emotional or cognitive development [47], and that categorizing adolescents into one age group could withhold full knowledge of the most vulnerable age groups associated with adverse birth outcomes. However, we lumped the adolescent age-group into one group of mothers less than 19 years of age because our study had small segregated sample sizes within the finer age group categories, especially in the 12–14-year-old group. Therefore, further research from this ongoing surveillance will seek to investigate the risk factors associated with the different adolescent age groups.

#### **Conclusion**

Our study is one of the few studies reporting adverse birth outcomes among adolescents in Uganda Our results corroborate previous findings in both developed and developing countries on birth outcomes and demonstrate that adolescent births are at risk for several neonatal adverse birth outcomes. With the growing population and high rates of adolescent births in Africa, the number of adverse birth outcomes is likely to increase and thereby remains a key public health concern [5].

Further research on individual, socio-cultural, environmental, economic, and health service-related factors are required to identify practicable and scalable measures to decrease adolescent pregnancy and to identify and reduce obstacles that discourage the use of qualified antenatal services, that would prevent or reduce adverse reproductive outcomes such as neonatal deaths, low birth weight, birth defects, and mother to child transmission of HIV. The establishment of dedicated adolescent-friendly antenatal care programs would help improve neonatal and adolescent health [48], and, better understand associated risk factors and the impact of younger maternal age on pregnancy outcomes. It is critical to monitor trends in birth outcomes and prevalence of major external birth defects across age groups to inform health-care policies and to plan for needed services among the affected population. Research on the potential underlying causes or mechanisms for these adverse outcomes among adolescent births is necessary to identify possible interventions.

#### Supplementary Information

The online version contains supplementary material available at https://doi.org/10.1186/s12978-021-01115-w.

Additional file 1. Definition of Birth defects

#### Abbreviations

cOR: Crudes odds ratio; aORs: Adjusted odds ratios; CI: Confidence intervals; LBW: Low birth weight; ENND: Early neonatal deaths; MTCT: Mother to child transmission; ANC: Antenatal care; ART: Antiretroviral therapy; WHO: World

Health Organization; CDC: Centers for Diseases Control and Prevention; HIV: Human immunodeficiency virus; LMP: Last menstrual period; US: United States; IQR: Inter-quartile range; NTD: Neural tube defects; ICD-10 RCPCH: International Classification of Disease 10, Royal College of Paediatrics and Child Health; OEIS: Omphalocele-exstrophy-imperforate anus-spinal defects.

#### Acknowledgements

The authors wish to thank all staff of the Birth Defects Surveillance study, and all management and staff of Mulago Referral Hospital, Mengo Hospital, St. Francis Hospital Nsambya, and Uganda Martyrs Hospital Lubaga who made the surveillance possible at their hospital. We also thank the management and administration of Makerere University-Johns Hopkins University Research Collaboration for all their support.

#### Authors' contributions

RS: took the lead in writing the manuscript and is accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. LBM, PM, DMM, SCT, DW, MRA, and DV: Substantial contributions to the conception or design of the work; or the acquisition, analysis, or interpretation of data for the work. They were involved in drafting the manuscript and revising it critically for important intellectual content. DK, JNM, JN, EN, DBM, and JB: were involved in drafting the manuscript and revising it critically for important intellectual content. All authors read and approved the final manuscript.

#### Funding

This research has been supported by the President's Emergency Plan for AIDS Relief through the United States Centers for Disease Control and Prevention under the terms of grants number: 1U10GH000487, 5U01GH000487, and 5U01GH0002171.

## Availability of data and materials

Not applicable

## Ethics approval and consent to participate

This surveillance study was approved by the Uganda National Council of Science and Technology (UNCST), (Ref: HS 1693); the Joint Clinical Research Centre institutional review board/Ethics committee (JCRC IRB), and the US Centers for Disease Control and Prevention Institutional Review Board (CDC IRB) (protocol # 6606.0). Consent to participate in the surveillance was waived by both IRBs (JCRC and CDC) because the surveillance involves no more than minimal risk to the participants. However, IRB-approved written informed consent was obtained for photographs of newborns with birth defects from their mothers or legal quardians.

## Consent for publication

Not applicable.

#### Competing interests

The authors declare that they have no competing interests.

#### **Author details**

<sup>1</sup> Makerere University-Johns Hopkins University Research Collaboration, P. O. Box 23491, Kampala, Uganda. <sup>2</sup> US Centers for Disease Control and Prevention (CDC), Atlanta, GA, USA. <sup>3</sup> US Centers for Disease Control and Prevention (CDC), Kampala, Uganda. <sup>4</sup> Department of Paediatrics and Child Health, College of Health Sciences, Makerere University, Kampala, Uganda. <sup>5</sup> Department of Obstetrics and Gynaecology, College of Health Sciences, Makerere University Kampala, Kampala, Uganda.

Received: 9 October 2020 Accepted: 22 February 2021 Published online: 04 March 2021

#### References

- WHO. World Health Organisation. Adolescent pregnancy. 2014. https:// apps.who.int/iris/rest/bitstreams/514257/retrieve. Accessed 17 Aug 2018.
- WHO. Why is giving special attention to adolescents important for achieving Millennium Development Goal 5? Adolescent pregnancy Fact Sheet: WHO/

Serunjogi et al. Reprod Health (2021) 18:56 Page 10 of 10

- MPS/08.14 2008; https://www.who.int/maternal\_child\_adolescent/event s/2008/mdg5/adolescent\_preg.pdf.
- Frolov P, Alali J, Klein MD. Clinical risk factors for gastroschisis and omphalocele in humans: a review of the literature. PediatrSurg Int. 2010;26(12):1135–48.
- Uganda Bureau of Statistics. Statistical Abstract. 2017. [341]. https://www. ubos.org/wp-content/uploads/publications/03\_20182017\_Statistica l\_Abstract.pdf. Accessed 11 Feb 2018.
- Uganda Bureau of Statistics (UBOS) and ICF International Inc, Uganda Demographic and Health Survey 2016. 2017, UBOS, and Rockville, Maryland, USA: UBOS and ICF International Inc: Kampala, Uganda.
- MuhumuzaKananura R, et al. Effect of a participatory multisectoral maternal and newborn intervention on birth preparedness and knowledge of maternal and newborn danger signs among women in Eastern Uganda: a quasi-experiment study. Glob Health Action. 2017;10(sup4):1362826.
- Ganchimeg T, et al. Pregnancy and childbirth outcomes among adolescent mothers: a World Health Organization multicountry study. BJOG. 2014;121(Suppl 1):40–8.
- Neal S, Channon AA, Chintsanya J. The impact of young maternal age at birth on neonatal mortality: evidence from 45 low and middle income countries. PLoS ONE. 2018;13(5):e0195731.
- Althabe F, et al. Adverse maternal and perinatal outcomes in adolescent pregnancies: The Global Network's Maternal Newborn Health Registry study. Reprod Health. 2015;12(Suppl 2):S8.
- Kongnyuy EJ, et al. Adverse perinatal outcomes of adolescent pregnancies in Cameroon. Matern Child Health J. 2008;12(2):149–54.
- Gortzak-Uzan L, et al. Teenage pregnancy: risk factors for adverse perinatal outcome. J MaternFetal Med. 2001;10(6):393–7.
- Igwegbe AO, Udigwe GO. Teenage pregnancy: still an obstetric risk. J Obstet Gynaecol. 2001;21(5):478–81.
- 13. Olausson PO, Cnattingius S, Haglund B. Teenage pregnancies and risk of late fetal death and infant mortality. Br J Obstet Gynaecol.
- Mombo-Ngoma G, et al. Young adolescent girls are at high risk for adverse pregnancy outcomes in sub-Saharan Africa: an observational multicountry study. BMJ Open. 2016;6(6):e011783.
- Usynina AA, et al. Adverse pregnancy outcomes among adolescents in northwest russia: a population registry-based study. Int J Environ Res Public Health. 2018;15(2):261.
- Zuckerman B, et al. Neonatal outcome: is adolescent pregnancy a risk factor? Pediatrics. 1983;71(4):489–93.
- Dillen J, Beijeren E, Roosmalen J. Perinatal outcome of primiparous teenagers in northern Namibia. Trop Doct. 2008;38:122.
- 18. Adam GK, et al. Maternal and perinatal outcome in teenage pregnancies in Sudan. Int J Gynaecol Obstet. 2009;105(2):170–1.
- Gronvik T, FossgardSandoy I. Complications associated with adolescent childbearing in Sub-Saharan Africa: a systematic literature review and meta-analysis. PLoS ONE. 2018;13(9):e0204327.
- Adane F, et al. Prevalence and associated factors of birth defects among newborns in sub-Saharan African countries: a systematic review and meta-analysis. Pan Afr Med J. 2020;36:19.
- Ajao AE, Adeoye IA. Prevalence, risk factors and outcome of congenital anomalies among neonatal admissions in OGBOMOSO, Nigeria. BMC Pediatr. 2019;19(1):88.
- Ndibazza J, et al. A description of congenital anomalies among infants in Entebbe, Uganda. Birth Defects Res A ClinMolTeratol. 2011;91(9):857–61.
- Mumpe-Mwanja D, et al. A hospital-based birth defects surveillance system in Kampala, Uganda. BMC Pregnancy Childbirth. 2019;19(1):372.
- Waldenström U, et al. Advanced maternal age increases the risk of very preterm birth, irrespective of parity: a population-based register study. BJOG Int J ObstetrGynaecol. 2017;124(8):1235–44.
- Goetzinger KR, et al. Advanced maternal age and the risk of major congenital anomalies. Am J Perinatol. 2017;34(3):217–22.
- Kang G, et al. Adverse effects of young maternal age on neonatal outcomes. Singapore Med J. 2015;56(3):157–63.
- Duong HT, et al. Is maternal parity an independent risk factor for birth defects? Birth Defects Res Part A, ClinMolTeratol. 2012;94(4):230–6.
- Gill SK, et al. Association between maternal age and birth defects of unknown etiology—United States, 1997–2007. Birth Defects Res Part A ClinMolTeratol. 2012;94(12):1010–8.

- Hernan MA, et al. Causal knowledge as a prerequisite for confounding evaluation: an application to birth defects epidemiology. Am J Epidemiol. 2002;155(2):176–84.
- Rasmussen SA, et al. Guidelines for case classification for the National Birth Defects Prevention Study. Birth Defects Res AClinMolTeratol. 2003;67(3):193–201.
- Feldkamp ML, et al. Etiology and clinical presentation of birth defects: population based study. BMJ. 2017;357:j2249.
- Ministry of Health, U. Uganda Clinical Guidelines 2016. 2016. 1118]. http://health.go.ug/sites/default/files/Uganda%20Clinical%20Guidelines%20 2016\_FINAL.pdf. Accessed 11 Feb 2018.
- Uganda Bureau of Statistics (UBOS) and ICF International Inc, Uganda Demographic and Health Survey 2011. 2012, UBOS and Calverton, Maryland: ICF International Inc: Kampala Uganda.
- 34. Kassa GM, et al. Adverse neonatal outcomes of adolescent pregnancy in Northwest Ethiopia. PLoS ONE. 2019;14(6):e0218259.
- Gebregzabiherher Y, et al. The prevalence and risk factors for low birth weight among term newborns in Adwa General Hospital, Northern Ethiopia. ObstetrGynecol Int. 2017;2017;7.
- Bater J, et al. Predictors of low birth weight and preterm birth in rural Uganda: findings from a birth cohort study. PLoS ONE. 2020;15(7):e0235626.
- Arunda MO, Agardh A, Asamoah BO. Survival of low birthweight neonates in Uganda: analysis of progress between 1995 and 2011. BMC Pregnancy Childbirth. 2018;18(1):189.
- 38. Chen XK, et al. Teenage pregnancy and congenital anomalies: which system is vulnerable? Hum Reprod. 2007;22(6):1730–5.
- Loane M, et al. Maternal age-specific risk of non-chromosomal anomalies. BJOG. 2009;116(8):1111–9.
- Zile I, Villerusa A. Maternal age-associated congenital anomalies among newborns: a retrospective study in Latvia. Medicina (Kaunas). 2013;49(1):29–35.
- 41. Reefhuis J, Honein MA. Maternal age and non-chromosomal birth defects, Atlanta–1968-2000: teenager or thirty-something, who is at risk? Birth Defects Res A ClinMolTeratol. 2004;70(9):572–9.
- Given JE, et al. Gastroschisis in Europe—a case-malformed-control study of medication and maternal illness during pregnancy as risk factors. PaediatrPerinatEpidemiol. 2017;31(6):549–59.
- Ministry of Health, Uganda. Uganda population-based HIV Impact assessment (UPHIA) 2016–2017: Final Report. UPHIA 2016–2017 July, 2019. https://phia.icap.columbia.edu/wp-content/uploads/2019/07/UPHIA \_Final\_Report\_Revise\_07.11.2019\_Final\_for-web.pdf. Accessed 24 Feb 2020.
- Xiao PL, et al. Association between maternal HIV infection and low birth weight and prematurity: a meta-analysis of cohort studies. BMC Pregnancy Childbirth. 2015;15:246.
- Ramraj T, et al. Adolescent access to care and risk of early mother-to-child HIV transmission. J Adolesc Health. 2018;62(4):434–43.
- 46. Tinker SC, et al. Challenges in studying modifiable risk factors for birth defects. CurrEpidemiol Rep. 2015;2(1):23–30.
- 47. World Health, O. Orientation programme on adolescent health for health care providers. Geneva: World Health Organization; 2006.
- WHO. Making health services adolescent friendly: developing national quality standards for adolescent friendly health services. 2012. https:// apps.who.int/iris/bitstream/handle/10665/75217/9789241503594\_eng. pdf;jsessionid=6391680A79868E5D0FAB89144F3466DF?sequence=1. Accessed 15 Feb 2020.
- Centers for Disease Control and Prevention. Use of Codes for Surveillance, Data Analysis and Presentation: Division of Birth Defects and Developmental Disabilities, NCBDDD; 2019. updated November 1, 2019. https:// www.cdc.gov/ncbddd/birthdefects/surveillancemanual/chapters/chapt er-5/chapter5-6.html. Accessed 24 Feb 2021.
- Kruszka P, Muenke M (2018) Syndromes associated with holoprosencephaly. Am J Med Genet C Semin Med Genet. 2018;178(2):229–37.

## **Publisher's Note**

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.