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Adverse perinatal outcomes in 665,244 term and post-term deliveries—a Norwegian population-based study

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ABSTRACT

Objective: To assess the prevalence and risk of adverse perinatal outcomes in early-term $(37^{+0}-38^{+6} \text{ weeks})$, full-term $(39^{+0}-40^{+6} \text{ weeks})$, late-term $(41^{+0}-41^{+6} \text{ weeks})$, and post-term $(>42^{+0} \text{ weeks})$ deliveries with spontaneous labor onset.

Study design: A population-based cohort with data from the Medical Birth Registry Norway (MBRN) and Statistics Norway (SSB) was conducted. The study population consisted of 665,244 women with cephalic singleton live births at term or post-term with spontaneous labor onset during the period of 1999–2014 in Norway. Maternal, obstetric, and fetal characteristics were obtained from the MBRN. Maternal education data were obtained from the SSB. The prevalence rates of adverse perinatal outcomes for each gestational age (GA) group were estimated. Inter-group differences were detected with Chi square tests. Multivariable regression analysis adjusted for maternal age, educational level, smoking, parity, maternal diabetes, and preeclampsia was used to assess adverse outcome prevalence for early- late-, and post-term births compared to full-term births.

Results Deliveries at early-term were associated with an increased prevalence of neonatal jaundice, polyhydramnios, small for gestational age (SGA) status, respiratory support, and neonatal intensive care unit (NICU) admission compared with deliveries at GAs of 39–43 weeks (p < 0.001). Low 5-min Apgar scores and newborn antibiotic treatment occurred at an increased prevalence in both early-term and post-term infants, relative to the full-term group (p < 0.001). The prevalence of oligohydramnios, meconium-stained amniotic fluid, and newborn birth injuries increased with increasing GA.

Conclusions More perinatal morbidity was observed among early-term infants compared to infants with later term deliveries, underscoring the need for cautious management of low-risk early-term deliveries.

Keywords

- Adverse perinatal outcome
- Early-term delivery
- Population based register data
- Perinatal morbidity

Term delivery

INTRODUCTION

Traditionally, term pregnancy has been defined as a pregnancy length between 37⁺⁰ and 41⁺⁶ weeks (1). Pregnancies with durations outside of this range are associated with increased perinatal morbidity and mortality (2-7). In recent decades, there has been an increase in non-medically indicated planned deliveries, including labor inductions and caesarean sections, motivated by an intention to avoid post-term complications (8, 9). These increases have been associated with a leftward shift in gestational age (GA) at delivery (10).

Elective labor induction for all pregnant women at 39 weeks gestation has been discussed, though associated benefits for the infant are uncertain (9, 11). Few studies have investigated GA-related perinatal morbidity within the term period (2, 3, 12-15). Relevant published data are challenging to compare due to differing risk profile and GAs across studies (13-15). Neonatal outcomes after term caesarean delivery have been described (16-18), but may not generalize to vaginal delivery cases. The aim of this study was to assess the prevalence and risk of adverse perinatal outcomes in early-term ($37^{+0}-38^{+6}$ weeks), full-term ($39^{+0}-40^{+6}$ weeks), late-term ($41^{+0}-41^{+6}$ weeks), and post-term ($>42^{+0}$ weeks) deliveries with spontaneous labor onset.

MATERIALS AND METHODS

This population-based cohort study is part of The PURPLE Study approved by the Regional Committee for Medical Research Ethics in South-East Norway (2015/681) and the Institutional Personal Data Officer in Oslo University Hospital. All parts of the study followed Norwegian Health Research legislation. Data were obtained from the Medical Birth Registry Norway (MBRN) and Statistics Norway (SSB). The SSB is a central agency that produces official statistics, including of educational level. MBRN and SSB data were merged by applying a similar code scheme that does not reveal personal identification; researchers used anonymized data. The MBRN is a national health registry containing information on all pregnancies and deliveries in Norway since 1967, including homebirths. Information on maternal pre-pregnancy health, pregnancy complications, and obstetric interventions, as well as postnatal fetal/newborn outcomes, is recorded on a standardized form by attending midwives. Information on newborns admitted to a neonatal intensive care unit (NICU) is reported to the MBRN by pediatricians.

A study period of 1999–2014 was chosen because GA in the MBRN has been defined by a 2nd trimester ultrasound, conducted free of charge universally in Norway (>97% use), since 1999. Congenital malformations, multiple pregnancies, preterm deliveries (<37 weeks), breech deliveries, stillbirths, scheduled caesarean sections, and induced labor were excluded. Deliveries registered with a GA of \geq 44 weeks were excluded due to likely incorrect GA registration or estimation. The study population was comprised of 665,244 cephalic singleton live births with spontaneous labor onset, with gestational periods of 37^{+0} to 43^{+6} weeks during the years of 1999–2014 in Norway (Fig. 1).

Definition of variables and data collection methods

Respiratory support was defined as newborn treatment with continuous positive airway pressure or use of an endotracheal tube. NICU admissions included transfers to a NICU for observation or treatment for any reason. Antibiotic treatment included treatment with one or more doses of systemic antibiotics. Intracranial hemorrhage was confirmed by ultrasonography. Brachial plexus injures were diagnosed clinically almost exclusively. Clavicle fractures were identified by clinical and radiographic diagnosis. Neonatal jaundice due to hyperbilirubinemia was defined by a need for phototherapy according to Norwegian national guidelines (19). Facial nerve palsy was recognized based on facial asymmetry apparent during clinical examination of the newborn. These conditions were determined and diagnosed by an attending neonatologist and reported to the MBRN. Small for gestational age (SGA) was defined as a birth weight below the 5th percentile for GA, based on Norwegian fetal growth percentiles (20). A sensitivity analysis showed similarly increased risks of perinatal morbidity among SGA-infants defined by 10th, 5th and 3rd percentile cut-offs. Because the proportion of constitutionally small infants could be overestimated if we adopted the 10th percentile or underestimated with a 3rd percentile cut-off, we chose to use the 5th percentile cut-offf.

Low Apgar scores (<7 and <4 at 5 min) were assigned by the attending midwife in normal deliveries and by the neonatologist in complicated deliveries. Meconium-stained amniotic fluid was identified by foul-smelling or discolored amniotic fluid by medical staff after membrane rupture. Oligohydramnios (single deepest pocket of amniotic fluid <2 cm or amniotic fluid index \leq 5 cm) and polyhydramnios (single deepest pocket of amniotic fluid \geq 8 cm or amniotic fluid index \geq 24 cm) were diagnosed by obstetricians antenatally based on ultrasound criteria.

Spontaneous labor onset was defined as labor starting without intervention. The term-period was divided according to expert recommendations (21): early-term ($37^{+0}-38^{+6}$ weeks), full-term ($39^{+0}-40^{+6}$ weeks), late-term ($41^{+0}-41^{+6}$ weeks), and post-term ($>42^{+0}$ weeks). Preeclampsia was defined as a blood pressure $\ge 140/90$ with proteinuria.

Maternal education was used as a substitute measure of socioeconomic status and classified as: none (did not complete compulsory education), compulsory (1–10 years of schooling), secondary (11–13 years of schooling), higher education (Bachelor's degree), or highest education (graduate degree). Maternal smoking during the 1st trimester was categorized as no, sometimes, or daily. Maternal age was categorized into the following groups (in years): <20, 20–24, 25–29, 30–34, 35–39, and ≥40. Maternal diabetes was classified as none, Type 1, Type 2, or gestational.

Patient involvement

Patient subjects were not involved in study planning, design, recruitment, or implementation.

Descriptive analysis

The study population was stratified according to GA. Continuous data were categorized. Adverse perinatal outcome prevalence rates are presented as percentages. Chi square tests (significance level, 1%) were used to detect inter-group differences.

Regression analysis

Regression analysis was performed to determine odds ratios for adverse perinatal outcomes for each term period, with full-term ($39^{+0}-40^{+6}$ weeks) deliveries, defined according to expert recommendations (21), as the reference group. Adjusted odds ratios (aORs) were calculated in multivariable regression with maternal age, education, smoking, parity, maternal diabetes, and preeclampsia as covariates, with the full-term delivery (21) group as the reference group. The aORs are reported with 95% confidence intervals (CIs). Variables with an association p <0.01 were included in the multivariate analyses. Data analyses were performed in SPSS version 24 (IBM Corp., Armonk, NY).

Missing values

Maternal educational was not recorded for 4% of the deliveries. A missing information category was managed separately in the regression analysis. Missing smoking status cases (15.5%) were merged with the non-smoking group as in previous studies with MBRN data (22, 23). For all other variables, the missing information rate was <0.05%.

RESULTS

Study population characteristics

Demographic characteristics and obstetric outcomes of the study population are presented in Table 1. Polyhydramnios, neonatal jaundice, SGA, newborn respiratory support, and NICU admission were more prevalent for deliveries at 37 weeks than for deliveries beyond 37 weeks. The prevalence of meconium stained-amniotic fluid, oligohydramnios, and newborn birth injuries increased with GA (Table 2, Fig. 1, and Fig. 2). Low 5-min Apgar scores and newborn antibiotic treatment were most common among early-term and post-term born infants.

Regression analysis

The results of GA-specific regression analyses are reported in Table 3. After adjustment for maternal age, education, smoking, diabetes, preeclampsia, and parity, odds ratios for adverse perinatal outcomes remained similar to those in our crude analysis. Need for respiratory support (aOR 2.20, CI 1.93–2.50) and NICU admission (aOR 2.00, CI 1.90–2.10) were two-fold higher among infants born at 37 weeks than among infants born at 39–40 weeks. The odds ratios of SGA (aOR 1.24, CI 1.17–1.31) and of polyhydramnios (aOR 1.27, CI 1.12–1.45) were higher at week 37 than in weeks 39–40 (Table 3).

The odds ratios of oligohydramnios (aOR 2.9, CI 2.76–3.13) and of newborn birth injury were elevated for post-term infants, including a two- to three-fold higher prevalence of facial nerve palsy (aOR 2.57, CI 1.64–4.04) compared with that in infants born at 39–40 weeks (Table 3). Meconium-stained amniotic fluid was two-fold more common among infants born after 42 weeks (aOR 2.11, CI 2.06–2.17) compared with those born at 39–40 weeks (Table 3).

The odds ratios of low 5-min Apgar scores (<7 points aOR 1.81, CI 1.63–2.02; and <4 points aOR 1.61, CI 1.26–2.06) and of antibiotic treatment were higher for infants born at 42 weeks

(aOR 1.59, CI 1.47–1.72) than for infants born at 39–40 weeks. The odds ratios of neonatal jaundice decreased with increasing GA, with a three-fold higher prevalence being observed for infants born at 37 weeks (aOR 3.23, CI 3.10–3.37) compared to those born at 39–40 weeks (Table 3).

COMMENT

Respiratory support and NICU admission were more common among early-term infants than full-term infants, likely due to lung immaturity (2, 4, 13, 14). Errors in GA determination are unlikely in our study owing to ultrasound confirmation (24). NICU admissions of early-term infants could be associated with hypothermia and/or hypoglycemia (25), conditions for which we did not have available data, as well as with a need for antibiotic treatment. Increased prevalence of antibiotic treatment and NICU admission, relative to full-term infants, was also observed among post-term infants, as reported previously (26, 27). Some women giving birth in the early-term period are given antibiotics following prelabor membrane rupture, whereas post-term born infants are more likely to be given antibiotics due to meconium stained amniotic fluid, meconium aspiration, or chorioamnionitis (26). Linder et al. (12) found similar results in a low-risk population, whereas Leal et al. (28) reported increased antibiotic treatment of early-term newborns delivered following medically indicated labor inductions, but not for those born following spontaneous labor onset.

An association of polyhydramnios with preterm birth has been documented (29, 30). It is possible that uterine overdistention is a causative factor of polyhydramnios. Conversely, an unidentified cause of polyhydramnios could affect labor onset timing. High rates of oligohydramnios and meconium-stained amniotic fluid in post-term deliveries could be consequent to gradually increasing placental insufficiency with increasing GA (4, 6, 13).

Although the prevalence of intracranial hemorrhage was highest for the early-term delivery group, it was not significantly higher than that of the full-term reference group in our multivariable regression.

The prevalence of SGA < 5th percentile was also highest among infants in the early term group, suggesting that placental dysfunction could lead to labor onset. Although SGA is an indication for labor induction or planned caesarean, detection of SGA can be missed antenatally. Morken et al. (31), Gouyon et al. (2), and Eskes et al. (32) reported SGA prevalence rates for term and post-term deliveries that were similar to the present findings, whereas Pulver et al. (33) did not find an effect of GA at birth on SGA risk. The study populations in these prior studies, however, varied and the 10th percentile cut-off for SGA was applied in all of them. The reasons for increased prevalence of SGA with spontaneous labor onset at early term remain to be resolved.

Caughey et al. (4) and Cheng et al. (13) reported low Apgar score data similar to our data. Despite enrolling a study population similar to the present one, Linder et al. (12) found no GA-related differences in Apgar scores. Meanwhile, the prevalence of neonatal jaundice decreased with increasing GA in our study, consistent with previous reports (12, 34).

The present finding of birth injury prevalence being highest among post-term infants may be related to their size (4, 13). Comparison with previous studies is challenging, due to varying definitions of birth injuries, diagnosis groupings, and small sample sizes. Notwithstanding, Ojumah et al. (35) found that brachial plexus injuries and clavicle fractures were associated with large fetus size, shoulder dystocia, and vaginal breech deliveries. Meanwhile, newborn facial nerve palsy, a rare complication, has been associated with birth trauma, particularly with forceps delivery and birth weights over 3.5 kg (35, 36). Here, we observed higher

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forceps delivery rates with increasing GA, consistent with the supposition that later delivery is a major risk factor for this complication.

Early-term delivery can increase long-term health challenges for children, including respiratory complications (37), obstructive sleep apnea (38), and endocrine problems (39). Further research is needed to clarify the pathophysiological mechanisms mediating perinatal morbidity risk among infants with early-term deliveries.

Although labor induction has become common worldwide, there is not yet a consensus regarding the optimal timing (15, 40). It is well recognized that there is a need to balance risks associated with earlier GA delivery versus post-term complication risks, though doing so can be challenging.

This study had three notable strengths. Firstly, we analyzed a large dataset from a populationbased registry with robust, reliable data (41), which provides confidence regarding the generalizability of our findings. Secondly, GA was determined by second-trimester ultrasonography, a valid and precise means of determining pregnancy duration (24). Thirdly, because we focused our analysis on a low-risk study population, GA was an appropriate main exposure variable for our analysis of perinatal outcomes.

This study had two notable limitations. Firstly, the MBRN does not collect information on numbers of antenatal visits or antenatal hospital admissions. All pregnant women are offered free antenatal care in Norway, which has resulted in an equalization of antenatal care across socioeconomic backgrounds. In Norway, all high-risk pregnancies, including late- and postterm pregnancies, are referred to obstetricians in maternity units for follow-up. Secondly, health registry information is expected to include some recording errors, missing data, and underreporting for some variables. Underreporting of risk factors can contribute to underestimation and a dilution of potential effect sizes, but does not create false-positive

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associations. Regarding missing data, only 20% of our study population had a maternal body mass index record. Given that maternal obesity is associated with adverse perinatal outcomes (42), these missing data could be a potential confounder we could not investigate in this study.

Our study sample consisted entirely of births with spontaneous labor onset, which can be considered a low-risk study population because all high-risk pregnancy cases are referred for labor induction or planned caesarean section. Despite the low-risk profile of our subjects, more adverse perinatal outcomes were observed among early-term deliveries than later term deliveries, underscoring the importance of managing even low-risk early-term deliveries with caution. We advocate for vigilant monitoring of early-term labor and attentiveness to potential GA-related complications when undertaking a non-medically indicated delivery.

Contribution to authorship

GM was involved with study design development and planning, data analysis, interpretation of the results, and the writing and revising of the manuscript.

SR was involved with study design development and planning, data-analysis, interpretation of the results, and critical review of the manuscript.

AFJ contributed to the conceptualization and planning of the study, the interpretation of the results, and provided critical review of the manuscript.

KBS conducted data processing, participated in the interpretation of results, and provided critical comments on the manuscript.

LB conducted data processing, participated in the interpretation of results, and provided critical comments on the manuscript.

KL supervised the study and was involved with study design development and planning, as well as with data processing and analyses, interpretation of the results, and revising the manuscript.

All authors read and approved the final version of the manuscript

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Conflict of interest notification

Gulim Murzakanova: none. Sari Räisänen: none. Anne Flem Jacobsen: none. Kristina Baker Sole: none. Lisa Bjarkø: none. Katariina Laine: none.

Data sharing statement

No additional data available.

According to Norwegian research legislation and regulations governing the Medical Birth Registry and Statistics Norway, given the sensitive nature of the data, open sharing of data from these institutions is not allowed. Therefore, we are not able to share data outside our research group.

Ethical approval

This study protocol was approved by the Regional Committee for Medical Research Ethics in South East Norway in 2015 (2015/681).

REFERENCES

1. World Health Organization. ICD-10: International statistical classification of diseases and related health problems, 10th revision. Geneva: WHO;. 2004.

2. Gouyon JB, Vintejoux A, Sagot P, Burguet A, Quantin C, Ferdynus C. Neonatal outcome associated with singleton birth at 34-41 weeks of gestation. Int J Epidemiol. 2010;39(3):769-76.

3. Caughey AB, Musci TJ. Complications of term pregnancies beyond 37 weeks of gestation. Obstet Gynecol. 2004;103(1):57-62.

4. Caughey AB, Washington AE, Laros RK, Jr. Neonatal complications of term pregnancy: rates by gestational age increase in a continuous, not threshold, fashion. Am J Obstet Gynecol. 2005;192(1):185-90.

5. Muglu J, Rather H, Arroyo-Manzano D, Bhattacharya S, Balchin I, Khalil A, et al. Risks of stillbirth and neonatal death with advancing gestation at term: A systematic review and meta-analysis of cohort studies of 15 million pregnancies. PLoS Med. 2019;16(7):e1002838.

6. Olesen AW, Westergaard JG, Olsen J. Perinatal and maternal complications related to postterm delivery: a national register-based study, 1978-1993. Am J Obstet Gynecol. 2003;189(1):222-7.

7. Maoz O, Wainstock T, Sheiner E, Walfisch A. Immediate perinatal outcomes of postterm deliveries. J Matern Fetal Neonatal Med. 2019;32(11):1847-52.

8. Kozhimannil KB, Macheras M, Lorch SA. Trends in childbirth before 39 weeks' gestation without medical indication. Med Care. 2014;52(7):649-57.

9. Murthy K, Grobman WA, Lee TA, Holl JL. Trends in induction of labor at early-term gestation. Am J Obstet Gynecol. 2011;204(5):435.e1-6.

10. Donahue SM, Kleinman KP, Gillman MW, Oken E. Trends in birth weight and gestational length among singleton term births in the United States: 1990-2005. Obstet Gynecol. 2010;115(2 Pt 1):357-64.

11. Glantz JC. Why not induce everyone at 39 weeks? Birth. 2017;44(2):97-100.

12. Linder N, Hiersch L, Fridman E, Lubin D, Kouadio F, Berkowicz N, et al. The effect of gestational age on neonatal outcome in low-risk singleton term deliveries. J Matern Fetal Neonatal Med. 2015;28(3):297-302.

13. Cheng YW, Nicholson JM, Nakagawa S, Bruckner TA, Washington AE, Caughey AB. Perinatal outcomes in low-risk term pregnancies: do they differ by week of gestation? Am J Obstet Gynecol. 2008;199(4):370.e1-7.

14. Clark SL, Miller DD, Belfort MA, Dildy GA, Frye DK, Meyers JA. Neonatal and maternal outcomes associated with elective term delivery. Am J Obstet Gynecol. 2009;200(2):156.e1-4.

15. Souter V, Painter I, Sitcov K, Caughey AB. Maternal and newborn outcomes with elective induction of labor at term. Am J Obstet Gynecol. 2019;220(3):273.e1-.e11.

16. Tita AT, Landon MB, Spong CY, Lai Y, Leveno KJ, Varner MW, et al. Timing of elective repeat cesarean delivery at term and neonatal outcomes. N Engl J Med. 2009;360(2):111-20.

17. Zanardo V, Simbi AK, Franzoi M, Solda G, Salvadori A, Trevisanuto D. Neonatal respiratory morbidity risk and mode of delivery at term: influence of timing of elective caesarean delivery. Acta Paediatr. 2004;93(5):643-7.

18. Wilmink FA, Hukkelhoven CW, Mol BW, van der Post JA, Steegers EA, Papatsonis DN. Neonatal outcome following elective cesarean section of twin pregnancies beyond 35 weeks of gestation. Am J Obstet Gynecol. 2012;207(6):480.e1-7.

19. Bratlid D, Nakstad B, Hansen TW. National guidelines for treatment of jaundice in the newborn. Acta Paediatr. 2011;100(4):499-505.

20. Gjessing HK, Grottum P, Okland I, Eik-Nes SH. Fetal size monitoring and birthweight prediction: a new population-based approach. Ultrasound Obstet Gynecol. 2017;49(4):500-7.

21. Spong CY. Defining "term" pregnancy: recommendations from the Defining "Term" Pregnancy Workgroup. JAMA. 2013;309(23):2445-6.

22. Sole KB, Staff AC, Laine K. The association of maternal country of birth and education with hypertensive disorders of pregnancy: a population-based study of 960 516 deliveries in Norway. Acta Obstet Gynecol Scand. 2018.

23. Ebbing C, Rasmussen S, Skjaerven R, Irgens LM. Risk factors for recurrence of hypertensive disorders of pregnancy, a population-based cohort study. Acta Obstet Gynecol Scand. 2017;96(2):243-50.

24. Dietz PM, England LJ, Callaghan WM, Pearl M, Wier ML, Kharrazi M. A comparison of LMP-based and ultrasound-based estimates of gestational age using linked California livebirth and prenatal screening records. Paediatr Perinat Epidemiol. 2007;21 Suppl 2:62-71.

25. Engle WA. Morbidity and mortality in late preterm and early term newborns: a continuum. Clin Perinatol. 2011;38(3):493-516.

26. Caughey AB, Stotland NE, Washington AE, Escobar GJ. Maternal and obstetric complications of pregnancy are associated with increasing gestational age at term. Am J Obstet Gynecol. 2007;196(2):155.e1-6.

27. De Los Santos-Garate AM, Villa-Guillen M, Villanueva-Garcia D, Vallejos-Ruiz ML, Murguia-Peniche MT. Perinatal morbidity and mortality in late-term and post-term pregnancy. NEOSANO perinatal network's experience in Mexico. J Perinatol. 2011;31(12):789-93.

28. Leal MDC, Esteves-Pereira AP, Nakamura-Pereira M, Domingues R, Dias MAB, Moreira ME, et al. Burden of early-term birth on adverse infant outcomes: a population-based cohort study in Brazil. BMJ open. 2017;7(12):e017789.

29. Odibo IN, Newville TM, Ounpraseuth ST, Dixon M, Lutgendorf MA, Foglia LM, et al. Idiopathic polyhydramnios: persistence across gestation and impact on pregnancy outcomes. Eur J Obstet Gynecol Reprod Biol. 2016;199:175-8.

30. Taskin S, Pabuccu EG, Kanmaz AG, Kahraman K, Kurtay G. Perinatal outcomes of idiopathic polyhydramnios. Interventional medicine & applied science. 2013;5(1):21-5.

31. Morken NH, Klungsoyr K, Skjaerven R. Perinatal mortality by gestational week and size at birth in singleton pregnancies at and beyond term: a nationwide population-based cohort study. BMC Pregnancy Childbirth. 2014;14:172.

32. Eskes M, Waelput AJM, Scherjon SA, Bergman KA, Abu-Hanna A, Ravelli ACJ. Small for gestational age and perinatal mortality at term: An audit in a Dutch national cohort study. Eur J Obstet Gynecol Reprod Biol. 2017;215:62-7.

33. Pulver LS, Guest-Warnick G, Stoddard GJ, Byington CL, Young PC. Weight for gestational age affects the mortality of late preterm infants. Pediatrics. 2009;123(6):e1072-7.
34. Norman M, Aberg K, Holmsten K, Weibel V, Ekeus C. Predicting Nonhemolytic

Neonatal Hyperbilirubinemia. Pediatrics. 2015;136(6):1087-94.

35. Ojumah N, Ramdhan RC, Wilson C, Loukas M, Oskouian RJ, Tubbs RS. Neurological Neonatal Birth Injuries: A Literature Review. Cureus. 2017;9(12):e1938.

36. Ciorba A, Corazzi V, Conz V, Bianchini C, Aimoni C. Facial nerve paralysis in children. World journal of clinical cases. 2015;3(12):973-9.

37. Walfisch A, Beharier O, Wainstock T, Sergienko R, Landau D, Sheiner E. Early-term deliveries as an independent risk factor for long-term respiratory morbidity of the offspring. Pediatr Pulmonol. 2017;52(2):198-204.

38. Walfisch A, Wainstock T, Beharier O, Landau D, Sheiner E. Early Term Deliveries and the Risk of Pediatric Obstructive Sleep Apnoea in the Offspring. Paediatr Perinat Epidemiol. 2017;31(2):149-56.

39. Paz Levy D, Sheiner E, Wainstock T, Sergienko R, Landau D, Walfisch A. Evidence that children born at early term (37-38 6/7 weeks) are at increased risk for diabetes and obesity-related disorders. Am J Obstet Gynecol. 2017;217(5):588.e1-.e11.

40. Stock SJ, Ferguson E, Duffy A, Ford I, Chalmers J, Norman JE. Outcomes of elective induction of labour compared with expectant management: population based study. BMJ (Clinical research ed). 2012;344:e2838.

41. Langhoff-Roos J, Krebs L, Klungsoyr K, Bjarnadottir RI, Kallen K, Tapper AM, et al. The Nordic medical birth registers--a potential goldmine for clinical research. Acta Obstet Gynecol Scand. 2014;93(2):132-7.

42. Lynch TA, Malshe A, Colihan S, Meyers J, Li D, Holloman C, et al. Impact of Maternal Obesity on Perinatal Outcomes in Preterm Prelabor Rupture of Membranes >/=34 Weeks. Am J Perinatol. 2019.

FIGURE LEGENDS

Figure 1. Flow-chart of study population enrollment and methodology.

Figure 2. Gestational age-specific prevalence (%) of adverse perinatal outcomes. Cephalic singleton live births at term and post-term with spontaneous labor onset were included (N = 665,244); cases involving major anomalies were excluded.

Figure 3. Gestational age-specific prevalence (%) of amniotic fluid abnormalities. Cephalic singleton live births at term and post-term with spontaneous labor onset were included (N = 665,244); cases involving major anomalies were excluded

TABLES

Table 1 Summary of patient demographics and obstetric outcomes by GA group, including variables used in the regression analysis, in a study population of term and post-term cephalic singleton live births with spontaneous labor onset (N = 665,244).

Table 2 Adverse perinatal outcomes by GA group in study sample (N = 665,244).

Table 3 Regression analyses of adverse perinatal outcomes in relation to GA grouping (N = 665,244).

Figure 1 Flow-chart of the study population enrollment and methodology

All deliveries in Norway between 1999-2014 N= 930 881 (\geq 22 gestation or \geq 500 grams)

Exclusion criteria

1 Multiple pregnancies	-16 447
2 Preterm deliveries <37+0	- 49 597
3 Deliveries >43+6	- 646
4 Major anomalies	-5027
3 Breech deliveries	-33 849
5 Scheduled caesarean section	- 40 615
6 Induced labour	- 118 914
7 Stillbirths	- 542

Study population: 665 244 cephalic singleton live births with spontaneous labour onset, 37+0 - 43+6 weeks of gestation

Descriptive statistics: univariate analysis, Chi square test

Multivariate regression analysis: adjusted for maternal age, parity, education, smoking, diabetes type 1, diabetes type 2, gestational diabetes, preeclampsia

Figure 2. Gestational age-specific prevalence (%) of adverse perinatal outcomes. Cephalic singleton live births at term and post-term with spontaneous labor onset were included (N = 665,244); cases involving major anomalies were excluded.



NICU=Neonatal intensive care unit SGA=Small for gestational age

Figure 3. Gestational age-specific prevalence (%) of amniotic fluid abnormalities. Cephalic singleton live births at term and post-term with spontaneous labor onset were included (N = 665,244); cases involving major anomalies were excluded



Table 1 Summary of patient demographics and obstetric outcomes by GA group, including variables used in the regression analysis, in a study population of term and post-term cephalic singleton live births with spontaneous labor onset (N = 665,244).

	Week 37 (N=26634)	Week 38 (N=71765)	Week 39 (N=165166)	Week 40 (N=225478)	Week 41 (N=146196)	Week 42 (N=29161)	Week 43 (N=844)
	% (n)	% (n)	% (n)	% (n)	% (n)	% (n)	% (n)
Maternal age ¹							
- < 20	3.0 (803)	2.7 (1922)	2.4 (3997)	2.2 (4932)	2.0 (2926)	2.1 (620)	4.1 (35)
- 20-24	17.3 (4604)	17.1 (12253)	16.0 (26490)	15.1 (34043)	14.6 (21321)	14.6 (4270)	17.9 (151)
- 25-29	33.2 (8844)	33.6 (24116)	33.9 (56045)	33.8 (76236)	33.3 (48672)	32.8 (9554)	33.3 (281)
- 30-34	30.6 (8163)	31.5 (22575)	32.5 (53688)	33.2 (74924)	33.8 (49432)	34.0 (9918)	30.8 (260)
- 35-39	13.5 (3593)	13.0 (9315)	13.0 (21482)	13.6 (30719)	14.3 (20815)	14.4 (4210)	11.4 (96)
- ≥ 40	2.4 (627)	2.2 (1580)	2.1 (3448)	2.0 (4615)	2.0 (2959)	2.0 (586)	2.5 (21)
Parity							
- Para 0	44.9 (11962)	40.7 (29225)	39.4 (65138)	40.4 (91110)	43.7 (63948)	49.1 (14329)	47.6 (402)
- Para 1	34.5 (9176)	37.5 (26937)	39.0 (64496)	38.5 (86771)	36.1 (52749)	31.9 (9309)	28.7 (242)
- Para ≥ 2	20.6 (5496)	21.7 (15603)	21.5 (35532)	21.1 (47597)	20.2 (29499)	18.9 (5523)	23.7 (200)
Education ²							
- compulsory (1-10years	20.7 (5263)	19.4 (13244)	17.5 (27711)	16.0 (34780)	15.1 (21356)	15.7 (4416)	22.5 (178)
- secondary (11-13 years	32.5 (8251)	31.9 (21832)	31.3 (49450)	30.7 (66672)	30.4 (42938)	31.6 (8898)	35.3 (279)
-higher (Bachelor)	36.5 (9254)	37.6 (25709)	39.1 (61755)	40.3 (87441)	41.0 (57866)	40.1 (11285)	32.4 (256)
-highest (Graduate degree)	10.3 (2620)	11.1 (7607)	12.1 (19083)	13.0 (28120)	13.4 (18852)	12.5 (3527)	9.7 (77)
Smoking							
- no	85.1 (22660)	85.9 (61656)	86.9 (143585)	87.4 (197057)	87.5 (127899)	85.5 (24919)	81.4 (687)
- sometimes	1.5 (402)	1.5 (1045)	1.5 (2521)	1.6 (3590)	1.6 (2373)	1.8 (538)	2.5 (21)
- daily	13.4 (3572)	12.6 (9064)	11.5 (19060)	11.0 (24831)	10.9 (15924)	12.7 (3704)	16.1 (136)
Diabetes							
- type 1	0.8 (217)	0.4 (318)	0.2 (342)	0.1 (216)	0.1 (66)	0.1 (14)	0.1 (1)
- type 2	0.4 (105)	0.3 (201)	0.2 (320)	0.1 (337)	0.1 (170)	0.1 (24)	0.2 (2)
- gestational	1.9 (504)	1.6 (1118)	1.2 (1982)	0.9 (1990)	0.5 (768)	0.3 (95)	0.2 (2)
Preeclampsia	3,1 (813)	2,0 (1402)	1,4 (2252)	1,1 (2439)	1,0 (1436)	0,9 (251)	0,9 (8)
Instrumental delivery							
- forceps	1.0 (265)	0.9 (664)	1.0 (1687)	1.2 (2803)	1.7 (2424)	2.2 (655)	2.2 (19)
- vacuum extraction	5.8 (1542)	5.9 (4269)	6.5 (10769)	8.1 (18191)	10.3 (15049)	12.5 (3659)	9.4 (79)
Emergency caesarean section	5.2 (1377)	3.7 (2658)	3.6 (5905)	4.5 (10130)	7.0 (10220)	10.9 (3166)	9.6 (81)

¹Maternal age -missing values 43/665201 ²Educational level-missing values 26524/638720

Note: No cases with major anomalies were included in the study sample.

Table 2 Adverse perina	atal outcomes by GA gr	roup in study samp	ble (N = $665,244$).
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	Week 37	Week 38	Week 39	Week 40	Week 41	Week 42	Week 43	
	(N=26634)	(N=71765)	(N=165166)	(N=225478)	(N=146196)	(N=29161)	(N=844)	
Stillbirth/Livebirth	69/26634	93/71765	118/165166	143/225478	83/146196	34/29161	2/844	
Adverse perinatal outcomes	% (n)	% (n)	% (n)	% (n)	% (n)	% (n)	% (n)	P-value
Meconium-stained amniotic fluid	5.47 (1456)	8.41 (6035)	13.41 (22143)	18.73 (42231)	24.52 (35840)	29.72 (8666)	28.08 (237)	0.001
Respiratory support ¹	1.08 (287)	0.49 (350)	0.39 (639)	0.50 (1127)	0.63 (920)	0.66 (193)	0	0.001
5-min Apgar < 7	0.93 (247)	0.63 (449)	0.61 (1012)	0.74 (1667)	1.08 (1572)	1.36 (395)	0.48 (4)	0.001
5-min Apgar < 4	0.23 (61)	0.14 (103)	0.13 (220)	0.16 (358)	0.22 (317)	0.26 (76)	0	0.001
Admission to NICU ²	8.0 (2006)	4.5 (3043)	3.5 (5541)	3.8 (8140)	4.8 (6604)	6.5 (1753)	5.4 (40)	0.001
Treatment of the newborn with antibiotics, ≥ 1 dose	2.09 (551)	1.38 (987)	1.36 (2251)	1.72 (3889)	2.31 (3376)	2.66 (777)	1.30 (11)	0.001
SGA ³ , below 5 th percentile	6.0 (1594)	5.2 (3745)	4.8 (7907)	4.4 (9990)	4.3 (6347)	4.6 (1332)	13.3 (112)	0.001
Intracranial haemorrhage	0.09 (23)	0.05 (33)	0.05 (78)	0.07 (151)	0.06 (93)	0.07 (19)	0	0.0047
Birth injures								
Brachial plexus injury	0.17 (46)	0.16 (115)	0.17 (279)	0.18 (413)	0.22 (328)	0.32 (92)	0.12(1)	0.001
Clavicle fracture	0.23 (61)	0.22 (160)	0.23 (385)	0.26 (581)	0.33 (482)	0.45 (132)	0.47 (4)	0.001
Facial nerve palsy	0.05 (12)	0.03 (23)	0.03 (44)	0.03 (67)	0.04 (53)	0.08 (22)	0.12 (1)	0.001
Neonatal jaundice	11.61 (3092)	6.21 (4459)	4.23 (6989)	3.51 (7919)	3.13 (4579)	3.62 (1057)	3.91 (33)	0.001
Oligohydramnios	1.18 (314)	1.14 (817)	1.34 (2211)	1.69 (3816)	2.71 (3967)	4.40 (1284)	4.38 (37)	0.001
Polyhydramnios	1.07 (286)	0.83 (597)	0.80 (1329)	0.78 (1766)	0.89 (1307)	0.79 (229)	0.36 (3)	0.001

¹ The need for continuous positive airway pressure or for an endotracheal tube ² NICU=Neonatal intensive care unit ³ SGA=Small for gestational age

	Wee	ek 37	Wee	ek 38	Week 39-40	Week 41		$GA \ge 42$	
Adverse perinatal	Crude OR	aOR	Crude OR	aOR	Ref	Crude OR	aOR	Crude OR	aOR
outcomes	(95% CI)	(95% CI)	(95% CI)	(95% CI)		(95% CI)	(95% CI)	(95% CI)	(95% CI)
Meconium-stained	0.29	0.28	0.47	0.45	Ref	1.65	1.65	2.14	2.11
amniotic fluid	(0.28-0.31)	(0.26-0.29)	(0.45-0.48)	(0.44-0.46)		(1.62-1.67)	(1.63-1.67)	(2.08-2.20)	(2.06-2.17)
Respiratory support 1	2.40	2.20	1.08	1.06	Ref	1.39	1.39	1.43	1.34
	(2.12-2.72)	(1.93-2.50)	(0.96-1.21)	(0.95-1.20)		(1.29-1.51)	(1.28-1.51)	(1.23-1.66)	(1.15-1.56)
5-min Apgar < 7	1.36	1.25	0.91	0.89	Ref	1.57	1.55	1.95	1.81
	(1.19-1.55)	(1.09-1.43)	(0.83-1.01)	(0.81-0.99)		(1.48-1.68)	(1.46-1.66)	(1.76-2.17)	(1.63-2.02)
5-min Apgar < 4	1.55	1.39	0.97	0.95	Ref	1.47	1.46	1.71	1.61
	(1.19-2.02)	(1.05-1.83)	(0.79-1.20)	(0.77-1.18)		(1.28-1.68)	(1.27-1.68)	(1.35-2.18)	(1.26-2.06)
Admission to NICU ²	2.26	2.00	1.22	1.15	Ref	1.31	1.31	1.81	1.73
	(2.15-2.37)	(1.90-2.10)	(1.18-1.27)	(1.11-1.20)		(1.27-1.35)	(1.27-1.35)	(1.72-1.90)	(1.64-1.82)
Antibiotic treatment,	1.32	1.22	0.87	0.86	Ref	1.48	1.44	1.69	1.59
≥1 dose	(1.21-1.45)	(1.11-1.33)	(0.82-0.93)	(0.80-0.92)		(1.42-1.54)	(1.38-1.51)	(1.57-1.82)	(1.47-1.72)
SGA ³ , below 5 th	1.33	1.24	1.15	1.12	Ref	0.95	0.91	1.05	0.95
percentile	(1.26-1.40)	(1.17-1.31)	(1.11-1.19)	(1.08-1.16)		(0.92-0.97)	(0.89-0.94)	(1.0-1.11)	(0.90-1.01)
Intracranial	1.47	1.32	0.78	0.77	Ref	1.09	1.08	1.08	0.85
haemorrhage	(0.96-2.26)	(0.85-2.05)	(0.54-1.13)	(0.53-1.12)		(0.85-1.38)	(0.85-1.37)	(0.68-1.73)	(0.51-1.41)
Birth injures									
Brachial plexus injury	0.98	0.88	0.90	0.91	Ref	1.27	1.28	1.75	1.78
	(0.72-1.31)	(0.65-1.21)	(0.74-1.10)	(0.74-1.11)		(1.11-1.45)	(1.11-1.46)	(1.41-2.18)	(1.42-2.22)
Clavicle fracture	0.93	0.88	0.90	0.87		1.33	1.33	1.84	1.84
	(0.72-1.20)	(0.68-1.14)	(0.76-1.07)	(0.74-1.04)		(1.20-1.49)	(1.19-1.49)	(1.53-2.20)	(1.53-2.21)
Facial nerve palsy	1.59	1.46	1.13	1.12		1.28	1.270	2.70	2.57
	(0.87-2.88)	(0.80-2.66)	(0.72-1.77)	(0.71-1.75)		(0.92-1.77)	(0.91-1.76)	(1.72-4.23)	(1.64-4.04)
Neonatal jaundice	3.31	3.23	1.67	1.67	Ref	0.82	0.79	0.95	0.89
	(3.18-3.45)	(3.10-3.37)	(1.61-1.73)	(1.61-1.73)		(0.79-0.84)	(0.765-0.82)	(0.89-1.01)	(0.83-0.94)
Oligohydramnios	0.76	0.71	0.74	0.72	Ref	1.78	1.78	2.94	2.9
	(0.68-0.85)	(0.63-0.80)	(0.68-0.79)	(0.66-0.77)		(1.71-1.85)	(1.71-1.86)	(2.77-3.12)	(2.76-3.13)
Polyhydramnios	1.36	1.27	1.05	1.01	Ref	1.13	1.17	0.98	1.06
	(1.20-1.54)	(1.12-1.45)	(0.96-1.15)	(0.93-1.11)		(1.06-1.21)	(1.10-1.25)	(0.85-1.12)	(0.92-1.21)

Table 3 Regression analyses of adverse perinatal outcomes in relation to GA grouping (N = 665,244).

Adjusted for maternal age, parity, education, smoking, diabetes type 1, diabetes type 2, gestational diabetes, preeclampsia

¹ The need for continuous positive airway pressure or for an endotracheal tube

² NICU=neonatal intensive care unit

³ SGA=Small for gestational age