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a register-based cohort study

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BMJ Open Educational disparities in perinatal health in Denmark in the first decade of the 21st century: a register-based cohort study

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ABSTRACT

Objective To investigate socioeconomic differences in six perinatal health outcomes in Denmark in the first decade of the 21st century.

Design A population-based cohort study. Setting Danish national registries.

Participants A total of 646 829 live born children and 3076 stillborn children (≥22+0 weeks of gestation) born in Denmark from 2000 to 2009. We excluded children with implausible relations between birth weight and gestational age (n=644), children without information on maternal country of origin (n=138) and implausible values of maternal year of birth (n=36).

Main outcome measures We investigated the following perinatal health outcomes: stillbirth, neonatal and postneonatal mortality, small-for-destational age, preterm birth grated into moderate preterm, very preterm and extremely preterm, and congenital anomalies registered in the first year of life.

Results Maternal educational level was inversely associated with all adverse perinatal outcomes. For all examined outcomes, the risk association displayed a clear gradient across the educational levels. The associations remained after adjustment for maternal age, maternal country of origin and maternal year of birth. Compared with mothers with vocational education, mothers with more than 15 years of education had an adjusted risk ratio for stillbirth of 0.64(95% CI 0.56 to 0.72). The corresponding adjusted risk ratios for neonatal mortality, postneonatal mortality, congenital anomalies, moderate preterm birth and small-for-gestational age were, respectively, 0.79(95% CI 0.67 to 0.93), 0.57(95% CI 0.42 to 0.78), 0.87(95% CI 0.83 to 0.91), 0.80(95% CI 0.77 to 0.83) and 0.83(95% CI 0.81 to 0.85).

Conclusion Substantial educational inequalities in perinatal health were still present in Denmark in the first decade of the 21st century.

INTRODUCTION

The Nordic countries are generally regarded as egalitarian and have a low prevalence of adverse perinatal outcomes. However in the 1980s and 1990s, strong socioeconomic gradients in adverse perinatal outcomes

Strengths and limitations of this study

- ► Danish national registries with high degree of completeness and quality were used in this study.
- The large study population enabled investigation of rare perinatal outcomes.
- The grading of maternal education into five categories enabled detailed investigation of the education-
- Congenital anomalies are not registered for stillbirths and spontaneous abortions during the study period and as a consequence we could only estimate risk among live born children.

were observed in Denmark, Finland, Norway and Sweden. Several studies documented socioeconomic inequalities in respectively stillbirth, preterm birth, birth weight, infant mortality in all of the Nordic countries¹⁻⁴ but studies have been inconclusive with regard to congenital anomalies.^{5–8}

During the first decade of the 21st century, economic inequality rose in Denmark and elsewhere, 9-11 but it is unclear what impact this has had on inequalities in perinatal health. Perinatal health measures, such as infant mortality, are used internationally as indicators of population health status. In addition, perinatal outcomes such as preterm birth and fetal growth are predictive at the individual level for health later in life. Several studies have found that low birth weight is related to the risk of chronic diseases in adulthood, such as type 2 diabetes and cardiovascular disease. $^{12\,13}$ The health condition of the mother affects perinatal outcomes, thereby transferring socioeconomic inequality in health from one generation to the next. 13

Socioeconomic position refers to the economic and social indicators that influence what positions individuals or groups hold within the structure of a society.¹⁴ In this



study, educational level has been chosen as a measure of socioeconomic position because education is a favourable indicator of socioeconomic position in early adult life. Furthermore, education is associated to future occupational status and income. ¹⁵

The aim of this study was to describe the associations between maternal educational level and risk of adverse perinatal outcomes in Denmark during the first decade of the 21st century. The adverse perinatal outcomes of interest in this study were stillbirth, preterm birth, small-for-gestational age (SGA), congenital anomalies, neonatal mortality and postneonatal mortality.

METHODS

We did a longitudinal register linkage study of all live births and stillbirths in Denmark from 2000 to 2009, as recorded in the Danish Medical Birth Register. Information on maternal educational level from the Population's Education register¹⁶ was linked to information on perinatal health outcomes from the Danish Medical Birth Register,¹⁷ the Danish National Patient Register¹⁸ and the Danish Register of Causes of Death.¹⁹ The Danish system of unique person identifiers was used to link individuals across the different registers in an anonymised dataset in Statistics Denmark which could be assessed via a vitual private network connection from Department of Public Health, University of Copenhagen.

Study populations

The study population consisted of all live born and stillborn children born in Denmark during the study period. Children were excluded from the study population as illustrated in figure 1. Children whose mothers were born after 1999 were excluded. Furthermore, we excluded children with implausible relations between gestational age and birth weight defined according to the method described by Alexander *et al.*²⁰ After the exclusions, study population 1 consisted of a total of 649 905 live and stillborn children. Study population 2 consisted of 646 829 live born when stillborn children were excluded from study population 1. In the analysis of SGA, children with missing information on birth weight (n=4388) were not included.

Patient and public involvement

As this is a whole population register-based study, we did not recruit individuals. Patients and public were not involved in the development of the research question, the design or conduct of this study. We planned to disseminate the results of this study through open-access publication.

Study variables

The main exposure variable of interest was maternal educational level, defined as the highest educational level attained or expected (based on ongoing education) of the mother at the year of birth categorised into five categories: primary education (≤9 years), secondary

education (high school 10-12 years), vocational education (10-12 years), short-cycle and medium-cycle higher education (12-15 years), long-cycle higher education (>15 years) and missing. Since the education register is based on reports from Danish educational institutions, the category of missing is composed of individuals who are not registered at any education in Denmark.

Information on birth weight and the outcomes preterm birth and stillbirth were obtained from the Medical Birth Register. SGA was defined as birth weight falling below the 10th percentile of birth weight according to sex-specific intrauterine growth curves presented by Marsàl et al.²¹ Preterm birth was divided into extremely preterm birth defined as birth ≥22+0-27+6 weeks, very preterm birth defined as birth ≥28+0-31+6 and moderately preterm birth defined as birth ≥32+0-36+6. The variable stillbirth was defined as fetal death at or after 22 completed gestational weeks. In Denmark, the threshold between spontaneous abortions and stillbirths was changed in 2004 from 28 completed weeks of gestation to 22 completed weeks of gestation. All pregnancies from 2000 to 2003 for which a spontaneous abortion was registered after 22 completed weeks of gestation were recoded into stillbirths. Information on infant mortality was obtained from The Central Person Register and The Danish Register of Causes of Death. Neonatal mortality was defined as death of a live born child before or at day 27 after birth, and postneonatal mortality was defined as death of a live born child at day 28 through day 365 after birth. Information on congenital anomalies was obtained from the Danish National Patient Register. The variable congenital anomalies registered in the first year of life were defined according to EuroCAT's definitions excluding minor anomalies according to EuroCAT criteria.²²

Other potential confounders in the analysis were the maternal year of birth (<1965, 1965-69, 1970-74, 1975-79, ≥1980), maternal age at the time of delivery (<25 years, 25-29 years, 30-34 years, 35-39 years, ≥40 years), offspring's year of birth (2000-01, 2002-03, 2004-05, 2006-07, 2008-09) and country of origin of the mother categorised as Denmark, 'Other western country' (Andorra, Australia, Canada, Iceland, Liechtenstein, Monaco, New Zealand, Norway, San Marino, Switzerland, the USA, Vatican City State and all European Union member countries except Denmark) or 'Non-western country' (all remaining countries).

Statistics

Initially, we calculated the prevalence for the six perinatal outcomes by maternal educational level. To investigate the associations between maternal educational level and the perinatal outcomes, we used multiple logistic regression analyses, and the analysis populations were restricted to children with information on maternal educational level. The a priori chosen model for adjustment included the covariates maternal age, maternal country of origin and maternal year of birth as these were considered potential confounders. The variable year of birth of the

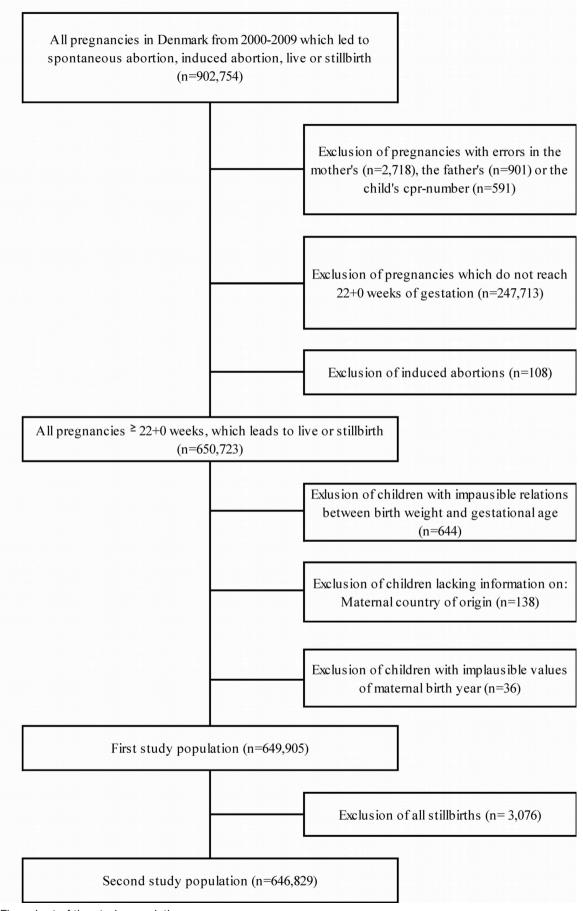


Figure 1 Flow chart of the study populations.

Table 1 Maternal age, year of birth and country of origin, and offspring's year of birth by educational level, all children born in Denmark, 2000–2009

	All children n=649905	Primary education n=106129	Vocational education n=222744	Secondary education n=46026	Short-cycle and medium- cycle higher education n=185417	Long-cycle higher education n=94168	Missing n=15421
	%	%	%	%	%	%	%
Maternal age							
<25	12.3	32.6	12.2	17.8	3.8	2.3	20.0
25–29	33.5	30.9	37.4	31.2	25.4	26.2	34.6
30–34	36.3	23.0	34.5	32.6	42.1	46.9	28.0
35–39	15.3	11.0	13.8	15.8	16.4	21.1	14.0
≥40	2.6	2.4	2.1	2.6	2.7	3.5	3.4
Maternal year of	f birth						
<1965	4.2	4.1	3.8	4.6	4.3	4.8	4.5
1965–69	16.5	12.3	16.4	17.6	17.0	20.7	12.6
1970–74	33.5	24.0	33.0	32.4	37.6	39.6	24.2
1975–79	30.6	28.8	32.0	27.9	31.8	28.3	31.1
≥1980	15.2	30.8	14.8	17.6	9.8	6.7	27.7
Maternal countr	y of origin						
Denmark	86.2	74.7	91.0	77.6	93.2	92.0	10.7
Western countries	3.1	1.3	1.9	3.8	2.6	4.3	26.3
Non-western countries	10.7	24.0	7.2	18.6	4.3	3.7	63.0
Year of birth (offspring)							
2000–2001	20.5	24.2	22.3	23.7	17.9	16.0	20.2
2002–2003	19.9	21.1	20.9	22.1	18.8	18.0	14.9
2004–2005	20.0	19.6	20.1	20.1	20.6	20.0	12.6
2006–2007	20.0	18.0	19.1	18.2	21.5	22.2	21.0
2008–2009	19.7	17.1	17.7	15.9	21.2	23.8	31.3

offspring was not included in the model since the variables maternal age at the time of delivery and maternal year of birth in combination practically describe the variable. Unadjusted and adjusted risk ratios (RR) were presented with a 95% CI. Additionally, we fitted two models to test robustness of the findings: one adjusted for maternal age only and the second including maternal age, parity and country of origin, and maternal year of birth as covariates. However, since the findings from the two models fitted to test robustness were similar to findings from the other adjusted model, their findings were not shown. All statistical analyses were conducted in SAS V.9.4.

RESULTS

The distributions of maternal educational level by maternal age, maternal year of birth, maternal country of origin and calendar year of birth are presented in table 1. In Denmark, maternal completed and ongoing educational level increased during the study period as shown

in table 1. The proportion of mothers born outside Denmark was higher among mothers with missing educational level.

The prevalence of stillbirth, neonatal mortality, postneonatal mortality, congenital anomalies, SGA and three degrees of preterm birth, respectively, by maternal educational level are shown in table 2. Women with the shortest education had a higher prevalence of all of the adverse perinatal outcomes compared with women with higher educational level. The group of women with unknown educational level had the highest prevalence of stillbirth and SGA compared with the women with known educational level.

Table 3 shows the relative risks, expressed as adjusted and unadjusted RR, of respectively stillbirth, neonatal mortality, postneonatal mortality, congenital anomalies, SGA and moderately preterm birth, very preterm birth and extremely preterm birth, with mothers with vocational education as the comparison group. Compared

Table 2 Prevalence of adverse perinatal outcome by educational level, Denmark, 2000–2009

	Stillbirth	Neonatal mortality	Postneonatal mortality	Congenital anomalies		Moderate preterm birth 32+0-36+6	Very preterm birth 28+0-31+6	Extreme preterm birth <28+0
	n=3076	n=1688	n=648	n=19449	n=59979	n=36610	n=4670	n=1843
	Per 1000	Per 1000	Per 1000	Per 1000	Per 1000	Per 1000	Per 1000	Per 1000
Primary education	6.7	3.6	1.9	34.9	125.9	64.5	8.3	4.1
Vocational education	4.8	2.7	1.0	30.7	95.1	60.0	7.8	3.0
Secondary education	4.0	2.5	1.0	28.2	94.2	52.7	7.0	2.4
Short-cycle and medium-cycle higher education	4.3	2.2	0.7	28.7	77.9	53.7	6.6	2.3
Long-cycle higher education	3.1	2.1	0.5	27.1	77.9	48.8	6.3	2.4
Missing	7.1	3.0	1.2	28.9	126.5	51.2	5.9	3.1

SGA, small-for-gestational age.

with mothers with vocational education, mothers with primary education had a statistically significant increased RR for stillbirth, neonatal mortality, postneonatal mortality, congenital anomalies, moderate preterm birth, very preterm birth, extreme preterm birth and SGA. Women with short-cycle and medium-cycle higher education and women with long-cycle higher education had a statistically significant lower risk of all the six perinatal outcomes compared with women with vocational education. These results indicated consistently inverse educational gradients in the risk of adverse perinatal outcomes. The gradients were consistent after adjusting for maternal age, maternal country of origin and maternal year of birth. The estimates were almost identical in analyses with adjustment for maternal age only and with adjustment for maternal age, maternal country of origin, maternal year of birth and parity (results not shown).

DISCUSSION

In a population register study of perinatal outcomes in Denmark, we found consistent inverse educational gradients for all perinatal outcomes in the study period, with higher rates of adverse outcomes in children of women with lower educational levels. Adjustments for maternal age, country of origin, parity and maternal year of birth did not change the gradients.

An educational gradient in SGA was found in this study. This result is consistent with previous Nordic studies which have reported socioeconomic inequalities in SGA. Socioeconomic differences have been found in preterm birth as an overall outcome and in preterm birth categorised according to gestational age. This study found an educational gradient in moderate, very and extreme preterm birth. Consistent with previous studies, 2-34 an educational gradient was found in stillbirth in this study. This study observed educational gradients in neonatal and postneonatal mortality. Socioeconomic

inequalities have been found in earlier Nordic studies in neonatal mortality 28 32 $^{35-38}$ and also in postneonatal mortality. 32 35 36 38

Previous Nordic studies that investigated the association between socioeconomic position and congenital anomalies reported inconsistent results.⁵⁻⁸ In accordance with a previous Danish study,⁶ this study found a statistically significant educational gradient in congenital anomalies. Both studies used maternal educational level as a measure of socioeconomic position whereas the other studies used occupation.^{5 8} or a combination of occupation and education.⁷ The differences in findings regarding socioeconomic differences in congenital anomalies are likely to be related to the use of different socioeconomic measures.

We used population-covering register individual-based data with minimal sources of selection and information bias. The data from the Danish Medical Birth Register used in this study are of high quality.¹⁷ The size of the study population enabled investigation of relatively rare perinatal outcomes like postneonatal mortality. Information on education was collected for administrative purposes, and according to the documentation from Statistics Denmark, the education variable was a so-called high-quality variable. 39 A limitation of this study is that the Danish registers only obtained information on congenital anomalies for live births. The prevalence of congenital anomalies is presumably higher among stillbirths, spontaneous and induced abortions than among live births. 40 Consequently, the socioeconomic gradient in congenital anomalies observed in this study could be underestimated. In order to understand the true association between socioeconomic position and congenital anomalies, it is essential to include information on congenital anomalies for live births, stillbirths, induced abortions and spontaneous abortions.

In contrast to several previous studies, we did not adjust for lifestyle factors such as smoking, alcohol intake, BMI

Table 3	Adverse	perinatal	outcome l	by maternal
education	nal level,	all births,	Denmark,	2000-2009

educational level, a	RR	95 % CI	aRR	95 % CI
Stillbirth		30 /0 01	uiii	30 /0 01
Stillbirth Primary education	1.39	1,26 to 1,53	1.34	1.21 to 1.48
Vocational education	1.00		1.00	13 1.40
Secondary education	0.84	0.71 to 0.98	0.81	0.69 to 0.94
Short-cycle and medium-cycle higher education	0.89	0.81 to 0.97	0.89	0.81 to 0.98
Long-cycle higher education	0.65	0.57 to 0.74	0.64	0.56 to 0.72
Neonatal mortality				
Primary education	1.33	1.16 to 1.51	1.29	1.12 to 1.47
Vocational education	1.00		1.00	
Secondary education	0.92	0.76 to 1.13	0.90	0.73 to 1.10
Short-cycle and medium-cycle higher education	0.79	0.70 to 0.90	0.80	0.71 to 0.92
Long-cycle higher education	0.79	0.67 to 0.93	0.79	0.67 to 0.93
Postneonatal mortality	*			
Primary education	1.93	1.59 to 2.35	1.81	1.47 to 2.22
Vocational education	1.00		1.00	
Secondary education	1.02	0.74 to 1.41	0.98	0.71 to 1.36
Short-cycle and medium-cycle higher education	0.74	0.59 to 0.92	0.76	0.61 to 0.95
Long-cycle higher education	0.55	0.41 to 0.75	0.57	0.42 to 0.78
Congenital anomalies				
Primary education	1.14	1.09 to 1.18	1.13	1.08 to 1.18
Vocational education	1.00		1.00	
Secondary education	0.92	0.86 to 0.98	0.91	0.86 to 0.97
Short-cycle and medium-cycle higher education	0.94	0.90 to 0.97	0.93	0.90 to 0.97
Long-cycle higher education	0.88	0.84 to 0.92	0.87	0.83 to 0.91
Moderate preterm birth, 32+0 to 36+6†				
Primary education	1.07	1.04 to 1.10	1.11	1.08 to 1.14
Vocational education	1.00		1.00	
Secondary education	0.88	0.84 to 0.92	0.89	0.86 to 0.93
Short-cycle and medium-cycle higher education	0.89	0.87 to 0.92	0.89	0.86 to 0.91
				Continued

Con	tinu	lec

Table 3 Continued					
	RR	95 % CI	aRR	95% CI	
Long-cycle higher education	0.81	0.79 to 0.84	0.80	0.77 to 0.83	
Very preterm birth, 28+0 to 31+6†					
Primary education	1.06	0.98 to 1.15	1.10	1.01 to 1.20	
Vocational education	1.00		1.00		
Secondary education	0.89	0.79 to 1.01	0.91	0.81 to 1.03	
Short-cycle and medium-cycle higher education	0.83	0.77 to 0.90	0.82	0.76 to 0.89	
Long-cycle higher education	0.80	0.73 to 0.88	0.78	0.71 to 0.86	
Extreme preterm birth, <28+0†					
Primary education	1.36	1.20 to 1.54	1.35	1.19 to 1.54	
Vocational education	1.00		1.00		
Secondary education	0.79	0.64 to 0.96	0.78	0.63 to 0.96	
Short-cycle and medium-cycle higher education	0.77	0.68 to 0.87	0.76	0.67 to 0.86	
Long-cycle higher education	0.78	0.67 to 0.91	0.76	0.65 to 0.89	
SGA					
Primary education	1.32	1.30 to 1.35	1.25	1.22 to 1.28	
Vocational education	1.00		1.00		
Secondary education	0.99	0.96 to 1.02	0.96	0.93 to 0.99	
Short-cycle and medium-cycle higher education	0.82	0.80 to 0.84	0.83	0.82 to 0.85	
Long-cycle higher education	0.82	0.80 to 0.84	0.83	0.81 to 0.85	

*Compared with children who were alive 365 days after birth. †Compared with children born from 37+0-41+6. aRR, risk ratio adjusted for maternal age, maternal country of origin and maternal year of birth; RR, crude risk ratio; SGA, small-forgestational age.

and physical activity. In this study, we were interested in estimating the overall effect of maternal educational level on adverse perinatal outcome and as a consequence we did not adjust for lifestyle factors as these were believed to mediate the association. It could be debated whether maternal age is a mediator or a confounder in the association. In this study, we assumed that maternal age affects maternal education since young women have not had the opportunity to undergo as much education as older women. As a consequence, we considered maternal age as a confounder. Both singletons and multiples were included in the analyses as we did not think the associations between maternal education and the adverse perinatal outcomes differed between singletons and multiples

when the estimates were adjusted for maternal age and other potential confounders.

The general level of education in Denmark increased during the study period. The proportion of women who attained primary school and no further education declined slightly during the study period. However, educational inequality in adverse perinatal outcomes was still observed. A contributing explanation of the inequality is likely to be found in a possible selection of women with the lowest educational level.³⁷ The group of women with primary education becomes more highly selected and thereby more socially vulnerable, ⁴¹ and this may contribute to the higher risk of adverse perinatal outcomes. The educational distribution forwomen who give birth changes over time this makes generalisations to other cohorts difficult. Nevertheless, socioeconomic differences in perinatal outcomes have been observed in different countries and in different time periods.⁴

Educational inequality in perinatal outcomes persists in the 21st century. A possible part of the explanation of this is an overall healthier lifestyle among women with higher socioeconomic position. Certain lifestyle risk factors are more common among lower socioeconomic groups than among higher socioeconomic groups. For instance, the prevalence of smokers is highest among women with lower educational level in Denmark, ⁴² and smoking is associated with a higher risk of several adverse perinatal outcomes. 40 To investigate the mediating factors between socioeconomic position and adverse perinatal outcomes was beyond the scope of this article. However, in order to reduce socioeconomic inequality in perinatal health, future studies should focus on the mediating mechanisms in the association of socioeconomic position and adverse perinatal outcome.

CONCLUSION

Socioeconomic inequality in perinatal health was observed in Denmark in the first decade of the 21st century. Maternal educational gradients were evident in the rare but serious outcomes such as congenital anomalies and fetal and infant mortality. The socioeconomically patterned risk of preterm birth and growth restriction is worrying because it may be the first tracks on the way to socioeconomic inequalities in health later in life. Interventions to diminish socioeconomic inequality in the earliest phases of life are still needed.

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Contributors A-MNA, JBA and JFB designed the study. JAB and JFB analysed the data with assistance from A-MNA, AVH and LHM. JBA and JFB interpreted the data and prepared the first draft of the manuscript which was revised by A-MNA, AVH and LHM. All authors read and approved the final manuscript.

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Competing interests None declared.

Patient consent Not required.

Ethics approval According to Danish legislation, no ethical permission is required for register-based research; however, this study was approved by the local data protection authorities.

Provenance and peer review Not commissioned; externally peer reviewed.

Data sharing statement The data that support the findings of this study are available from Statistics Denmark but restrictions apply to the availability of these data which were used under licence for the current study, and so are not publicly available. Data are however available from the authors upon reasonable request and with permission of Statistics Denmark.

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