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Gestational diabetes mellitus by maternal country of birth and length of residence in immigrant women in Norway

Short running title: Gestational diabetes in immigrant women

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What's new

- Immigrant background is associated with gestational diabetes mellitus (GDM), and our study contributes with knowledge on GDM disparities.
- Immigrant women from a number of countries had substantially higher odds of being diagnosed with GDM compared with non-immigrant women.
- The number of women diagnosed with GDM increased with longer length of residence in Norway among immigrant women.
- This study highlights the need of tailored interventions where the differences in GDM prevalence are taken into consideration, in maternity care for the heterogeneous group of immigrant women.

ABSTRACT

Aims Immigrant women are at higher risk for gestational diabetes mellitus (GDM) than non-immigrant women. This study described the prevalence of GDM in immigrant women by maternal country of birth and examined the associations between immigrants' length of residence in Norway and GDM.

Methods This Norwegian national population-based study included 192 892 pregnancies to immigrant- and 1 116 954 pregnancies to non-immigrant women giving birth during the period 1990-2013. Associations were reported as odds ratios (ORs) with 95% confidence intervals (CIs) using logistic regression models, adjusted for year of delivery, maternal age, marital status, health region, parity, education and income.

Results The prevalence and adjusted OR [CI] for GDM were substantially higher in immigrant women from Bangladesh (7.4%, OR 8.38 [5.41, 12.97]), Sri Lanka (6.3%, OR 7.60 [6.71, 8.60]), Pakistan (4.3%, OR 5.47 [4.90, 6.11]), India (4.4%, OR 5.18 [4.30, 6.24]) and Morocco (4.3%, OR 4.35 [3.63, 5.20]) compared to non-immigrants (prevalence 0.8%). Overall, GDM prevalence increased from 1.3% (OR 1.25 [1.14, 1.36]) to 3.3% (OR 2.55 [2.39, 2.71]) after nine years of residence in immigrants compared to non-immigrant women. This association was particularly strong for women from South Asia.

Conclusions GDM prevalence varied substantially between countries of maternal birth and was particularly high in immigrants from Asian countries. GDM appeared to increase with longer length of residence in certain immigrant groups.

Keywords

Gestational diabetes mellitus, immigrant women, length of residence, population-based study

Introduction

Gestational Diabetes Mellitus (GDM) is defined as any degree of glucose intolerance with onset, or first recognition during pregnancy (1,2). This definition therefore includes hyperglycaemia that is induced by the pregnancy, and undiagnosed diabetes prior to the pregnancy. The condition is associated with caesarean section, shoulder dystocia, macrosomia, and large for gestational age offspring (3), in addition to later development of type 2 diabetes mellitus in mothers (4).

Moreover, women who have experienced GDM appear to have a modestly increased risk of miscarriage in the subsequent pregnancy (5).

The prevalence of GDM has increased over the years (6,7), and established risk factors for GDM include higher maternal age, parity and BMI (8). In addition, several studies, including a meta-analysis, showed that immigrant background is a strong independent risk factor for GDM (7,9,10). However, most previous studies on GDM prevalence are either small or have typically reported GDM data according to large heterogeneous groups, potentially masking a great deal of variation in immigrant subgroups. Therefore, our knowledge of precisely which subgroups are at increased GDM risk is limited, suggesting updated GDM data by maternal country of birth in a large study sample.

Several risk factors for GDM vary with length of residence in the receiving countries. In particular, longer length of residence in receiving countries is associated with increased BMI (11), higher smoking rates (12), and increased risk of preeclampsia (13). Despite this, there is limited knowledge of whether long-term immigrant residents have different risks of GDM than short-term residents. More knowledge on how the association of length of residence with GDM appears in immigrant women from a range of different countries or regions of maternal birth when taking age, parity, and other risk factors into account, may contribute with important knowledge of which immigrant groups might be of specific concern and in need of additional support from maternity caregivers.

The aim of this study was to describe the prevalence of GDM in immigrant women by maternal country- and region of birth compared to Norwegian-born non-immigrants and to examine the associations between the immigrants' length of residence in Norway and GDM.

Participants and methods

This study used data from the Medical Birth Registry of Norway (MBRN) (14) and Statistics Norway (SSB) (15) for the period 1990-2013. The MBRN contains maternal and offspring information on personal, administrative and health-related aspects for all live- and stillbirths from 16 weeks of gestation (14). The SSB provided the sociodemographic and migration-related data

for the study sample. The data from MBRN and SSB were merged by using the unique national identity number, which is assigned to each individual who lives or settles in Norway (16).

The total number of pregnancies recorded in the MBRN between the years 1990 and 2013 was 1 415 666. Of these, we analysed 1 309 846 pregnancies to 711 677 Norwegian-born women with Norwegian-born parents (non-immigrants) and foreign-born women with foreign-born parents (1st generation immigrant women). We a priori excluded 23 855 pregnancies to women with multiple pregnancies, 389 pregnancies to women whose information on maternal country of birth was missing, and another 8625 pregnancies to women with any pre-gestational diabetes. This study focused on GDM in 1st generation immigrant women who had two foreign-born parents.

Therefore, to reduce heterogeneity between groups being compared, we excluded 6362 pregnancies to 2nd generation immigrant women (Norwegian-born with two foreign-born parents), 44 019 pregnancies to Norwegian-born with a mixed ethnic background (Norwegian-born with one foreign-born parent), 9 194 pregnancies to foreign-born with one Norwegian-born parent, and 13 376 pregnancies to foreign-born women with two Norwegian-born parents (Fig. 1). Also, as we lacked information on newly arrived asylum seekers, paperless immigrants and tourists giving birth in Norway, these groups were not included in the study.

Antenatal care for healthy pregnant women in Norway is carried out either by the midwives alone, by the General Practitioners (GP) alone, or in combination with the GP and the midwives, in primary care (17). During the study period, GDM was defined according to WHO (1999) as fasting plasma glucose of ≥ 7.0 mmol/L or plasma glucose ≥ 7.8 mmol/L two hours after intake of 75 g glucose (oral glucose tolerance test (OGTT)) (18). From 2009, the National Clinical Guideline for diabetes recommended an OGTT to high-risk women (family history of diabetes, previous GDM, BMI > 27 kg/m², age > 38 years, ethnic minorities from countries outside Europe with high prevalence of diabetes) (19). National Guidelines for Antenatal Care recommend that immigrant pregnant women from Asia and Africa should be routinely screened for GDM (17,20-22). We used variables for GDM as defined by MBRN.

Maternal country of birth as registered by SSB was analysed directly as a categorical variable including the 42 countries with the largest number of women, i.e., those countries contributing to most pregnancies during the study period. The remaining countries (n=158) were grouped and analysed as "Other countries". Region of maternal birth was classified according to the seven Global Burden of Disease (GBD) super regions (23): (i) Central Europe, Eastern Europe, and

Central Asia; (ii) High-income countries; (iii) Latin America and Caribbean; (iv) North Africa and Middle East; (v) South Asia; (vi) Southeast Asia, East Asia, and Oceania; and (vii) Sub-Saharan Africa.

The registration date according to the National Registry in Norway is the official date of immigration (24). The immigrant women's length of residence in Norway was calculated as the year of childbirth registered in the MBRN minus the year of immigration. The year of immigration was abstracted from the SSB and was the year when the individuals obtained their national identity number registered in the National Registry. Length of residence was analysed as a categorical variable: <2; 2-3; 4-5; 6-7; 8-9; and >9 years (categories were chosen a priori). Non-immigrant women were coded with a separate code and used as reference category for this variable.

Other variables collected from the MBRN included the year the women gave birth (continuous variable), maternal age at delivery (continuous variable), parity (0, 1, 2, 3 \geq 4 previous births), health region in Norway (South/East, West, Middle, North) and marital status at birth (married/cohabiting, not married/cohabiting). Variables collected from the SSB included maternal educational level (no education, primary school, secondary school, university/ college) and maternal income (calculated quartiles for 1990-2013). Maternal age, parity and socioeconomic status have previously been shown to be associated with GDM (7,8) and were included as adjustment variables in the regression analyses. We also adjusted for health region (South/East, West, Middle and North) in Norway to adjust for a potential geographical difference in GDM prevalence.

Statistical Analyses

We used logistic regression analysis to estimate the association of maternal country and region of birth with GDM, as well as the associations between length of residence in immigrant women and GDM. In all analyses, the reference group was non-immigrant women and results were reported as odds ratios (ORs) with 95% confidence intervals (CIs). To account for the dependency among pregnancies by the same mother, we used robust standard errors that allowed for within-mother clustering (25).

Analyses were performed crude and with adjustments for year of birth, maternal age at birth, parity, health region in Norway, marital status at birth, education and maternal income. To account for non-linear associations, year of birth and maternal age were incorporated in the regression models as polynomial quadratic terms in the regression models. Because GDM increased during the study period and immigrants immigrated on different time points, year of birth was an important adjustment variable for the current analysis.

Due to missing data (see Table 1) on health region, education and maternal income, a multiple imputation strategy by predictive mean matching was applied to produce ten multiply imputed datasets. The imputation model included GDM, length of residence, maternal country of birth, year of birth, maternal age at birth, marital status at birth, health region, parity, education and income. The combination of ORs with 95% CIs across imputed datasets was calculated using Rubin's combination rules. All analyses were performed using R 3.4.2 for Windows (The R Foundation for Statistical Computing, Vienna, Austria) and Stata IC version 14 (Stata Statistical Software, College Station, TX, USA).

Ethical approval

This study was approved by the Regional Committee for Medical Research Ethics the 7th of June 2018, in an overarching project that was approved 10th of September 2014 (2014/1278/REK South-East).

Results

Of the 1 309 846 included pregnancies, 192 892 were to immigrant women and 1 116 954 to non-immigrant women. In both groups, the women were on average 29 years old and approximately 92% were married or cohabiting at the time of delivery (Table 1). Immigrant women had a higher parity than the non-immigrant women and they had on average a length of residence in Norway of 6 years (SD±6) at the time of delivery. Overall, the prevalence of GDM was 2.2% in pregnancies to the immigrant women and 0.8% in pregnancies to the non-immigrant women (Table 1).

The prevalence of GDM increased in both immigrant- and non-immigrant women from 1990 to 2013 (Table S1). The strongest increase in GDM prevalence was seen during the period 2009-

2013 in both groups, but there was a consistently higher prevalence of GDM in immigrant women compared to non-immigrant women throughout the study period.

The prevalence of GDM varied substantially across maternal region- (Table 2) and country of birth (Fig.2). When examining the odds of GDM by the seven GBD super regions, women from the South Asia region (OR 5.51, CI 5.07, 5.98), the Southeast Asia, East Asia and Oceania region (OR 3.00, CI 2.80, 3.21) and the North Africa and Middle East region (OR 2.28, CI 2.11, 2.47) had more than two-fold higher adjusted OR for GDM compared to non-immigrant women.

When comparing the odds of GDM between the 42 maternal countries of birth and non-immigrant women (Fig. 2), the highest ORs for GDM were 8.38 (95% CI 5.41, 12.97) in immigrant women from Bangladesh, 7.60 (CI 6.71, 8.60) in those from Sri Lanka, 5.47 (CI 4.90, 6.11) in those from Pakistan, 5.18 (CI 4.30, 6.24) in those from India, and 4.35 (CI 3.63, 5.20) in those from Morocco. In addition, immigrant women from Ghana, Algeria, Cuba, Gambia, Nigeria, the Philippines, and China had substantially higher OR for GDM, relative to non-immigrant women. The lowest adjusted OR for GDM was in immigrant women from Lithuania (0.31, 95% CI 0.19, 0.50) and Sweden (0.79, 95% CI 0.64, 0.98) (Fig. 2).

The prevalence of GDM increased substantially with the immigrants' length of residence (Table 2). Compared with non-immigrant women, the adjusted OR of GDM in the immigrant group overall increased from 1.25 (CI 1.14, 1.36) for those who gave birth <2 years after arrival to Norway to 2.55 (CI 2.39, 2.71) for those who gave birth >9 years after arrival to Norway (Table 2).

When investigating the relationship between the immigrants' length of residence and GDM by the seven GBD super regions, there was a strong increase in GDM prevalence in women from the South Asia region, the Southeast Asia, East Asia and Oceania region and women from the North Africa and Middle East region (Fig. 3). GDM prevalence, unadjusted and adjusted ORs for the association between the immigrant's length of residence and GDM by GBD super region are available in Table S2.

Discussion

The prevalence of GDM varied substantially across maternal countries of birth, and immigrant women from Bangladesh, Sri Lanka, Pakistan, India and Morocco had the highest OR of GDM compared to the Norwegian-born non-immigrant women. At regional levels, women from the South Asia region, the Southeast Asia, East Asia and Oceania region, and the North Africa and Middle East region had the highest OR of GDM. Moreover, the OR of GDM appeared to increase with longer length of residence in immigrant women from these regions.

A strength of this national population-based study is that the large sample size (including 1 309 846 pregnancies) allowed examining GDM prevalence in pregnancies of immigrant women from a number of maternal countries and regions of birth. Our study also contributed with unique information on the relationship between maternal length of residence and GDM. Using length of residence as an exposure, we were able to distinguish GDM prevalence between long-term immigrant residents from short-term residents, an association that has been lacking in studies of GDM and immigrants (9). However, due to the exclusions of multiple pregnancies and certain immigrant groups (other than 1st generation immigrant women and non-immigrant women), results cannot be generalized to second generation immigrant women, those women who had a mixed parental background, or women with multiple pregnancies (Fig. 1). Another limitation of this study was the amount of missing values in education and income for immigrant women. We used well-established multiple imputation strategies to impute plausible data for these adjustment variables, however, due to the high proportion of missing data, we cannot rule out that skewness in the imputed values for these variables might have occurred.

The overall prevalence of GDM in our study (0.8% in Norwegian-born non-immigrants and 2.2% in immigrant women) is lower than that reported in other GDM studies from Nordic countries (6,10,26). A lower GDM prevalence found in the MBRN compared with that observed in the other studies could reflect a variation in the diagnostic criteria used, characteristics of the study population, or a variation in the coding of GDM. However, GDM could also be subject to some under-reporting especially during the early periods when GDM screening and awareness were low.

The National Clinical Guidelines for Diabetes from 2009 acknowledged the increased risk of type 2 diabetes and GDM in our largest groups of immigrant women, and recommended to perform OGTT in pregnant women from these regions. Awareness about the high risk of diabetes, particularly in South Asians (27), also in our population in Norway, increased gradually from

about the year 2000. We have reason to believe that the uptake of screening and subsequent OGTT in general differed somewhat between health regions and health personnel groups. We cannot rule out that relatively more South Asians than other groups were offered an OGTT, due to their higher risk. However, other immigrant groups (from Africa and Middle East) have generally higher BMI and obesity rates and would probably have been offered an OGTT based on this well-known risk factor.

Consistent with Urquia et al. (28), we found a higher adjusted odds ratio for GDM in immigrant women from South Asia compared with non-immigrant women. Moreover, our findings of a higher GDM prevalence in immigrant women from Bangladesh, Sri Lanka, Pakistan, India and Morocco agree with findings of a recent population-based study from Denmark (10) and a study among South Asian immigrants in New Jersey, United States (29). In our study, we further showed that immigrant women from a number of additional countries are at substantial increased OR for GDM, including women from Ghana, Algeria, Cuba, Gambia, Nigeria, the Philippines and China (Fig. 2), essentially in line with the screening recommendations by the Norwegian National Guidelines for Antenatal Care (17,20-22).

Our finding that the prevalence of GDM increased with longer length of residence in Norway in immigrant women is consistent with Sørbye et al. (30) in descriptive analysis in an overlapping period (1990-2009) using the same data material as we do. We further showed that GDM increased with length of residence in women from South Asia, Southeast Asia, East Asia and Oceania, and North Africa and Middle East, suggesting that several immigrant groups may be more vulnerable to GDM than others with length of residence in Norway.

A positive relationship between length of residence and GDM was also reported in the recent study by Nielsen et al. from Denmark (10), an association partly explained by maternal age and BMI. Unfortunately, information on pre-pregnancy BMI in MBRN was only available for a limited period from 2008 onwards in our study. With additional missing data from 2008 onwards, adjustment for BMI would yield substantially reduced study sample and thereby also reduced generalizability of findings. Therefore, we did not include BMI as covariate in our study. Results from our study and Nielsen et al.'s study (10) give emphasis on the future need of research examining mechanisms involved in the association between length of residence and GDM in women from different maternal countries of birth.

Chen et al. (31) examined the influence of acculturation (country of birth, language spoken at home and length of time in the United States) and reported that acculturation was inversely associated with GDM in Asian American women in Los Angeles, and discussed that acculturation could have a protective role (31). Sørbye et al. (30) reported an association between non-spontaneous preterm deliveries and longer length of residence in immigrant women, and the authors discussed the possibility of a higher detection rate of risk conditions among immigrants with longer length of residence due to for instance improved language skills (30). There is a lack of knowledge about positive and negative consequences of acculturation in women in reproductive age with specific focus on GDM. Nevertheless, Commodore-Mensah et al. (32) reported that immigrants residing ≥ 10 years in the United States were more likely to be obese, have diabetes and being hypertensive compare with people with < 10 years of residence, and emphasized the need of research examining primary care and health screening in newer immigrants versus long-term immigrants (32). Also, Hawkins et al. (12) reported that immigrant women with longer length of residence were more likely to smoke during pregnancy than those with short length of residence. Moreover, a change into an unhealthier dietary pattern in South Asians after migration to Europe is previously discussed in a review study (33). However, the authors emphasized that dietary change with regard to migration is complex (33), and dietary patterns should be further examined according to length of residence in immigrant women.

Immigrant women from a number of countries, including Bangladesh, Sri Lanka, Pakistan, India, and Morocco, have a substantially higher GDM prevalence compared to Norwegian-born non-immigrant women. That GDM increased with longer length of residence, particularly among immigrant women from South Asia, underscores the need for further investigation into the risk factors of GDM in immigrant women from these maternal birth regions. Our study highlights the importance of tailored interventions to the heterogeneous group of immigrant women in reproductive age, where the differences in GDM prevalence are taken into consideration.

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Conflicts of Interest

None declared

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Table 1. Sociodemographic, immigrant- and birth-related information on pregnancies in Norway between 1990 and 2013 (n = 1 309 846)

	Pregnancies to non-immigrant women*		Pregnancies to immigrant women†	
N	1 116 954		192 892	
Year of birth, n (%)				
1990-1993	208 105	(18.6)	17 157	(8.9)
1994-1998	250 311	(22.4)	25 760	(13.4)
1999-2003	228 184	(20.4)	35 751	(18.5)

2004-2008	219 962	(19.7)	47 239	(24.5)
2009-2013	210 392	(18.8)	66 985	(34.7)
Maternal age at birth, mean (\pm SD)	29.0	(5.1)	29.4	(5.3)
Marital status, n (%)				
Married or cohabitant	1 025 052	(91.8)	177 543	(92.0)
Not married or cohabitant	91 902	(8.2)	15 349	(8.0)
Parity, n (%)				
0	459 884	(41.2)	79 055	(41.0)
1	403 764	(36.1)	63 271	(32.8)
2	187 040	(16.7)	29 426	(15.3)
3	48 427	(4.3)	11 917	(6.2)
4	17 839	(1.6)	9223	(4.8)
Educational level, n (%)				
No education	43	(<1)	4329	(3.0)
Primary school	240 915	(21.6)	45 764	(31.7)
Secondary school	428 907	(38.5)	39 168	(27.1)
University/college	444 869	(39.9)	55 086	(38.2)
Educational level, missing n (%)	2220	(0.2)	48 545	(25.2)
Income level, n (%)				
<25 percentile	251 088	(24.5)	32 762	(29.2)
25-50 percentile	261 566	(25.6)	22 334	(19.9)
50-75 percentile	256 578	(25.1)	27 372	(24.4)
\geq 75 percentile	254 058	(24.8)	29 844	(26.6)
Income level, missing n (%)	93 664	(8.4)	80 580	(41.8)
Health region in Norway n (%)				
South/East	562 873	(50.4)	130 005	(67.5)
West	262 865	(23.5)	34 094	(17.7)
Middle	168 260	(15.1)	17 047	(8.8)
North	122 207	(10.9)	11 490	(6.0)
Health region in Norway, missing n (%)	749	(<1)	256	(<1)
Length of residence (year), mean (\pm SD)			6.0	(6.0)
Gestational diabetes mellitus n (%)				

No	1 107 556	(99.2)	188 670	(97.8)
Yes	9398	(0.8)	4222	(2.2)

Abbreviations: SD, standard deviation

* Non-immigrant women: Norwegian-born women with two Norwegian-born parents

† Immigrant women: Foreign-born women with two foreign-born parents.

Table 2. Crude and adjusted odds ratios (ORs) of gestational diabetes mellitus (GDM) in immigrant women by the seven global burden of disease super regions of maternal birth and by length of residence in Norway

	N	n GDM	%	Crude OR	(95% CI)	Adjusted OR*	(95% CI)*
Maternal birth region							
Pregnancies to non-immigrant women [†]	1 116 954	9398	0.8	1.00	Reference	1.00	Reference
Pregnancies to immigrant women							
Central Europe, Eastern Europe; and Central Asia	39 530	470	1.2	1.42	(1.29, 1.56)	0.96	(0.87, 1.06)
High income countries	43 804	437	1.0	1.19	(1.08, 1.31)	0.97	(0.88, 1.07)
Latin America and Caribbean	5096	112	2.2	2.65	(2.19, 3.20)	1.63	(1.35, 1.97)
North Africa and Middle East	31 077	793	2.6	3.09	(2.87, 3.32)	2.28	(2.11, 2.47)
South Asia	16 506	730	4.4	5.45	(5.05, 5.89)	5.51	(5.07, 5.98)
Southeast Asia, East Asia, and Oceania	32 629	1080	3.3	4.03	(3.78, 4.30)	3.00	(2.80, 3.21)
Sub-Saharan Africa	24 250	600	2.5	2.99	(2.75, 3.25)	1.87	(1.71, 2.06)
Length of residence (years)							
Pregnancies to non-immigrant women [†]	1 116 954	9 398	0.8	1.00	Reference	1.00	Reference
Pregnancies to immigrant women							
<2	46 401	609	1.3	1.57	(1.44, 1.70)	1.25	(1.14, 1.36)
2-3	39 778	739	1.9	2.23	(2.07, 2.41)	1.66	(1.54, 1.8)
4-5	29 086	625	2.2	2.59	(2.38, 2.81)	1.84	(1.69, 2.01)

6-7	21 162	501	2.4	2.86	(2.61, 3.13)	1.99	(1.81, 2.19)
8-9	15 176	423	2.8	3.38	(3.06, 3.73)	2.23	(2.02, 2.47)
>9	39 925	1314	3.3	4.01	(3.78, 4.25)	2.55	(2.39, 2.71)
Per category change				1.28	(1.27, 1.29)	1.18	(1.17, 1.19)

Abbreviations: CI, confidence intervals; OR, odds ratio

*Adjusted for year of birth, region of birth, maternal age at delivery, marital status, parity, education and income.

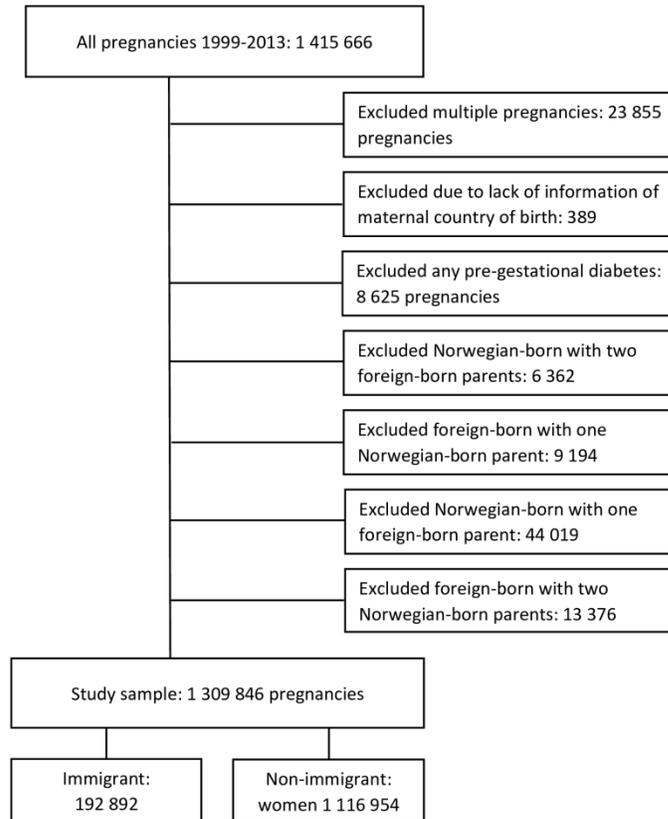
†Reference group: Norwegian-born non-immigrant women

Figure Legends

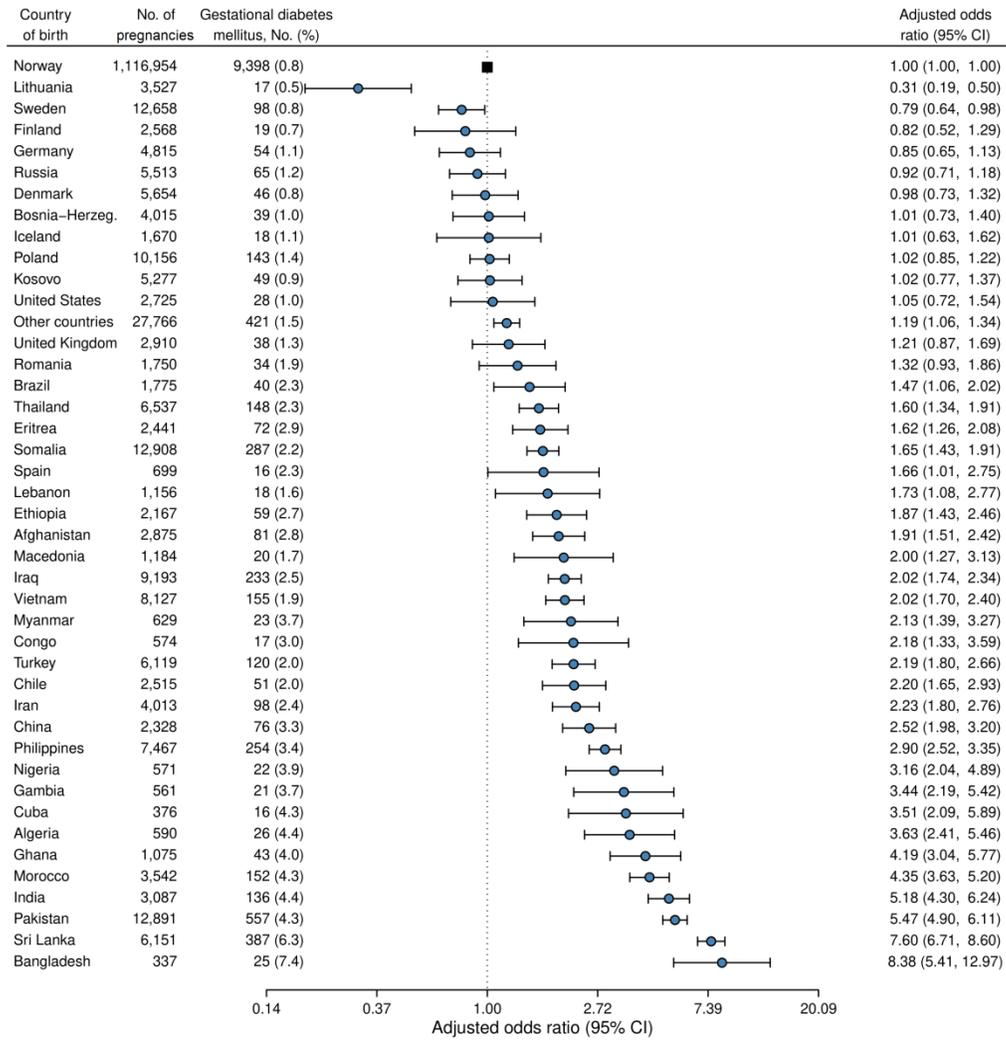
Figure 1. Flow chart of the exclusion process. The total number of excluded pregnancies to foreign-born women was 22 570. Of these, 77% (17 398/22 570) pregnancies were to foreign-born women from Sweden, United States, South Korea, Denmark, Great Britain, Germany, Colombia, and Canada, India, and the Netherlands.

Figure 2. Prevalence and adjusted odds ratios (adjusted for year of birth, maternal age at birth, parity, health region in Norway, marital status at birth, education, and income) of gestational diabetes mellitus (GDM) in women by maternal country of birth compared with non-immigrant women

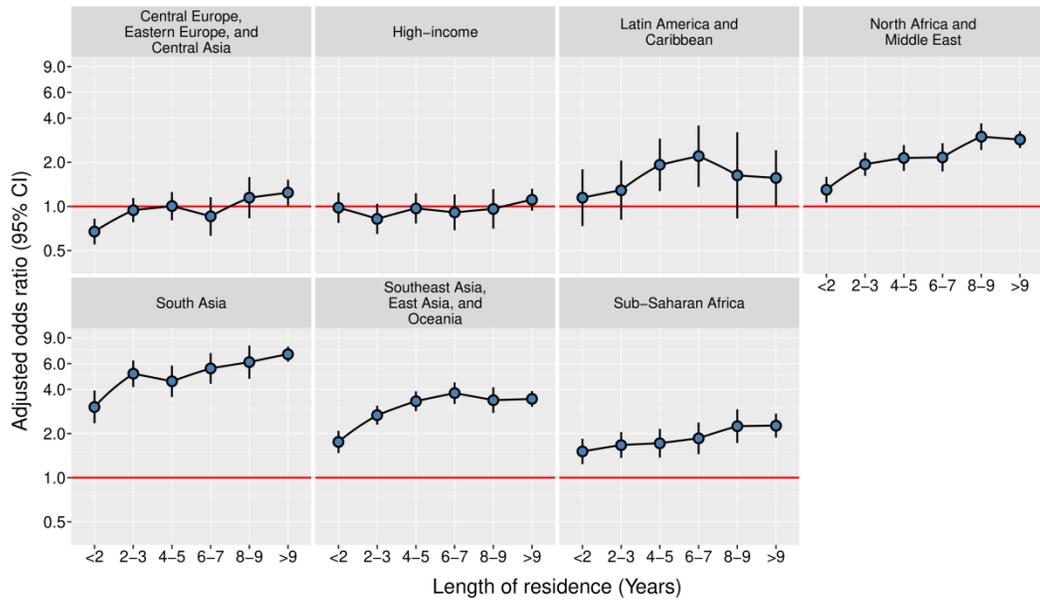
Figure 3. Adjusted odds ratios of gestational diabetes mellitus (GDM) in immigrant women by length of residence in Norway, compared to non-immigrant women (adjusted for year of birth, maternal age at birth, parity, health region in Norway, marital status at birth, education, and income). Stratified analyses by regions of maternal birth according to the seven global burden of disease super regions



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