

MOTHER'S COUNTRY OF BIRTH AND PSYCHOTROPIC DRUG USE IN SWEDISH ADOLESCENTS: A LIFE-COURSE APPROACH

Journal:	BMJ Open
Manuscript ID:	bmjopen-2012-001260
Article Type:	Research
Date Submitted by the Author:	13-Apr-2012
Complete List of Authors:	van Leeuwen, Willemijn; Lund University, Faculty of Medicine, Social Epidemiology Nilsson, Sofia; Lund University, Faculty of Medicine, Social Epidemiology Merlo, Juan; Lund University, Faculty of Medicine, Social Epidemiology
Primary Subject Heading :	Mental health
Secondary Subject Heading:	Epidemiology, Public health, Sociology, Health services research, Pharmacology and therapeutics
Keywords:	EPIDEMIOLOGY, Quality in health care < HEALTH SERVICES ADMINISTRATION & MANAGEMENT, PUBLIC HEALTH, SOCIAL MEDICINE, Community child health < PAEDIATRICS, Child & adolescent psychiatry < PSYCHIATRY

SCHOLARONE™ Manuscripts

MOTHER'S COUNTRY OF BIRTH AND PSYCHOTROPIC DRUG USE IN SWEDISH ADOLESCENTS: A LIFE-COURSE APPROACH

Corresponding author

Juan Merlo

MD, PhD, Tenured Professor

Unit for Social Epidemiology, Skåne University Hospital

Jan Waldenströms gata 35

SE-20502 Malmö, Sweden

E-mail: juan.merlo@med.lu.se

Tel. +46 40391329

Authors

Willemijn Van Leeuwen^{1,2}

Sofia Nilsson¹

Juan Merlo¹

Affiliations

Running title:

Psychotropic drug use in Swedish adolescents

Keywords

Psychotropic drugs, adolescent, second-generation immigrants, needs assessment, bias (epidemiology)

Word count:

Abstracts: 246

Manuscript: 2916

¹ Unit for Social Epidemiology, Faculty of Medicine, Lund University, Malmö, Sweden

² Dept. of Public Health, Academic Medical Center (AMC) University of Amsterdam, Netherlands

ABSTRACT

Objectives

In addition to medical needs, psychotropic drug use in adolescence might be conditioned by the cultural context of the family. This knowledge is relevant for both detecting inequities in health care, and identifying information bias in epidemiological studies using psychotropic drugs as a proxy for impaired psychological health. Therefore, we investigated whether, independent of needs, the socioeconomic characteristics of the mother's country of birth are associated with psychotropic drug use in Swedish-born adolescents.

Methods

By linking the Swedish Medical Birth Registry to other national registers, we identified all 324510 singletons born between 1988 and 1990 who were alive and residing in Sweden until the age of 18 (years 2006 to 2008). We measured their psychotropic drug use in the year they turned 18. Applying a life-course approach, we included both the mother's and the children's characteristics throughout pregnancy, delivery, infancy, childhood and adolescence when calculating a risk score to adjust for needs. We classified the mother's country of birth according to the gross national income (GNI) per capita of each country.

Results

Overall, the lower the income of the mother's birth country, the lower the probability of psychotropic drug use among children. When adjusting for needs, the association became even stronger.

Discussion

Besides medical needs, use of psychotropic drugs by second-generation immigrants seems conditioned by the socioeconomic characteristics of the mothers' countries of birth. The threat of information bias must be considered when psychotropic drug use is employed as a proxy for impaired psychological health.

INTRODUCTION

In Sweden, as in many other countries, the healthcare system aims to allocate resources on equal terms and according to needs rather than by gender, socioeconomic position or country of birth [1].

Psychotropic drugs are potent therapy for treating impaired psychological disorders in childhood and adolescence [2]. Despite the many knowledge gaps concerning their safety and efficacy [3], the use of these drugs in adolescents has increased considerably during recent decades [4-6]. In addition, these drugs are used as a proxy for impaired psychological health in adolescents and young adults in register-based epidemiological studies [7-13]. Psychotropic drug use in adolescents should be conditioned by needs. However, the concept of 'need' is ambiguous, and care-seeking and care-giving are influenced by other factors in addition to disease [14, 15]. There are also complex relationships between symptoms, expectations, social factors and an individual's conception of disease [16, 17]. Furthermore, the concepts of disease, illness and need are culturally defined, to a great extent, and are influenced by societal and contextual factors acting over and above individual characteristics [18]. Therefore, psychotropic drug use may be conditioned by an individual's country of birth or interaction with Swedish society [19]. Patterns of healthcare utilisation may also be transmitted across generations. Nevertheless, there are many knowledge gaps concerning the determinants of psychotropic drug use in immigrants. To the best of our knowledge, this information is almost absent for second-generation immigrants. Nevertheless, this information is relevant not only for detecting inequities in health care but also for identifying information bias in epidemiological studies that use psychotropic drugs as a proxy for impaired psychological health.

In this context, we investigated psychotropic drug use in 18 year old adolescents born in

Sweden. We aimed to compare adolescents born to immigrant women (i.e., second-generation immigrants) with those born to native mothers. Our hypothesis was that any difference would disappear or at least decrease after adjusting for needs. We appraised needs using a risk score [20] for psychotropic drug use, including information on the health and socioeconomic status of the mothers and the children during pregnancy and delivery as well as throughout the children's infancy, childhood and adolescence [21]. We studied every 18-year-old adolescent in Sweden during the years 2006-2008.

POPULATION AND METHODS

Study population

The Swedish Medical Birth Registry is linked to several other national databases, including the National Inpatient Registry, the National Mortality Registry, the Swedish Drug Prescription Register, the Register of the Total Population, and the Swedish 1990 population census, using a unique personal identification number. These registries are administered by the Swedish authorities (i.e., The National Board of Health and Welfare and Statistics Sweden). To ensure the anonymity of the subjects, these authorities encrypted the original personal identification numbers of the individuals before delivering the data to us. The study was approved by the Regional Ethics Review Board in southern Sweden.

Using the Medical Birth Register, we identified all singleton children born in Sweden between 1988 and 1990 who were alive and residing in Sweden from birth until the year in which the child turned 18 (2006-2008) and whose mother's country of birth could be identified. The final study population represented 94% (324510/347148) of all children born in 1988, 1989 and 1990. Figure 1 shows a flow diagram of the selection criteria used to define the study population.

Assessment of the outcome variable

The outcome variable was the use of psychotropic drugs (i.e., dispensation at the pharmacy) according to the Drug Prescription Registry. We obtained this information in the calendar year that the child turned 18. Psychotropic drugs were defined as all drugs with Anatomic Therapeutic Chemical (ATC) classification system [22] codes starting with N5 or N6 (see Table 1). We dichotomised the outcome variable into use or non-use of psychotropic drugs.

Assessment of child's characteristics

We categorised *birth year* into 1988, 1989 or 1990, using 1988 as the reference group. *Apgar score* was measured on a scale from 1-10 at the first, fifth and 10^{th} minute after birth, where a score of 10 indicates a delivery without distress for the infant. We created dummy variables where an *Apgar score* ≤ 5 was classified as 'low', and an *Apgar score* >5 was classified as 'not low'. The latter was used as the referent. We recoded the missing values at 5 or 10 minutes as 'not low Apgar' because it can be assumed that the subsequent *Apgar scores* were high and therefore not measured if the Apgar score after one minute was high [23]. This was the case for 37% of the children.

For every hospital discharge, a main diagnosis and up to seven secondary diagnoses are recorded in the Patient Register and coded according to the International Classification of Diseases (ICD). We defined a child admitted to a hospital with a diagnosis coded 290 to 319 or F00 to F90 (ICD versions 9 and 10, respectively) as *severe psychiatric morbidity*. Children never admitted to a hospital with any of these diagnoses were established as the reference group in the comparisons.

For every child, we calculated the total number of days spent in the hospital from birth until their 15th birthday. We distinguished between children with *extensive hospitalisation* (i.e.,

children spending 30 days or more in the hospital) and children with less than 30 days in the hospital or who were never hospitalised, using the latter group as the referent. We combined *history of psychiatric diagnosis* and *extensive hospitalisation* of the children into four categories, as indicated in table 2.

Assessment of mother's characteristics

We categorised the mothers' *country of birth* by two different types of criteria: socioeconomic and geographic. Both categorisations are listed in table 2. In our analyses, we used the socioeconomic criterion, which categorises the mother's country of birth by means of the World Bank classification of country economies. Countries were grouped according to their gross national income (GNI) per capita using the World Bank Atlas method into low-, lower middle-, upper middle-, and high-income countries. Sweden was included in the high-income category. However, we separated Sweden from the group of high-income countries and used Sweden as the referent in the comparisons because we conducted the study in the context of Swedish culture.

We classified *mother's age at delivery* into six groups (i.e., <20 years, 20-24 years, 25-29 29... >39 years). We considered mothers from the age group 25-29 the reference group in the comparisons.

We classified *maternal smoking during pregnancy* into four categories: 'no smoking', 'light smokers' (1-9 cigarettes per day), 'heavy smokers' (>9 cigarettes per day) and 'missing values'. The missing values ranged from 7.6% among the Swedes to 11.8% among mothers from low-income countries. We considered the non-smoking group the reference category. The information on smoking was self-reported and recorded when the mother attended her first scheduled antenatal care visit.

We identified all mothers with a diagnosis of *pre-eclampsia* (ICD-9 code 642) and considered mothers who did not suffer from pre-eclampsia the reference group in the comparisons.

We defined mother's *educational achievement* as the highest completed level of education in 1990. This variable was categorised into four groups: 'elementary school or lower' (9 years), 'lower secondary school' (12 years total), 'higher educational achievement' (>12 years) and 'missing information'. We used the 'higher educational achievement' category as the reference group in the comparisons.

Information regarding whether parents were *living together* was based on self-reported status when the mother was first seen for antenatal care. We considered mothers living together with the father of the child the reference group.

We identified mothers with *severe psychiatric morbidity* and with *extensive hospitalisation* in a manner similar to that used for children (see above). In addition, we combined a history of psychiatric diagnosis with extensive hospitalisation in the same way as we did for the children (see table 2).

Statistical Analyses

To estimate needs, we applied a logistic regression and constructed an equation that predicted the probability (i.e., the risk score) [20] of using psychotropic drugs around the age of 18 in the unexposed group (i.e., children with mothers born in Sweden). This equation included the maternal and child's characteristics mentioned above.

These variables were chosen using both previous knowledge and an explorative approach. When two variables showed multicollinearity, we selected the variable that provided a better goodness of fit by means of a chi-square test (e.g., mother's education compared with household income).

Subsequently, we applied the risk score equation to the entire population (i.e., children with mothers born in Sweden *and* children with immigrant mothers).

The equation of the risk score is presented in Table 1. The individual value of the risk score (RS) (i.e., predicted probability) was obtained with the following formula:

RS =
$$[e^{logit} / (1 + e^{logit})] * 100.$$

In the analyses, we categorised the risk score into four groups by quartiles and used the group with the lowest risk score as the reference group in the comparisons.

Using logistic regression, we developed two consecutive models to estimate the relative differences (i.e., odds ratios) in the use of psychotropic drugs. In the first model, we investigated the bare association between mother's country of birth and the use of psychotropic drugs in adolescence. In the second model, we included the risk score described above to adjust for possible dissimilarities in needs between the different maternal countries of birth. We analysed boys and girls separately in the models because there are cultural differences in gender roles, and gender may influence healthcare and psychotropic drug use [24-26]. We used SPSS version 18 for all analyses.

RESULTS

Table 1 indicates the regression coefficients of the variables used in the logistic regression to obtain the risk score (i.e., predicted probability) for psychotropic drug use.

> **Table 1.** Equation for estimation of the risk score (RS) used in the analyses to approximate the need for psychotropic drugs. The equation was calculated in the population of children born to Swedish mothers and subsequently applied to all the children. The RS corresponds to the predicted probability of using psychotropic drugs as a function of the variables included in the model.

```
Logit (Use of psychotropic drugs) =
-3.467 = intercept
+0.180 * 'smoking 1-9 cigarettes per day'
+0.291 * 'smoking more than 9 cigarettes per day'
+0.057 * 'no information on smoking'
+0.126 * 'pre-eclampsia'
+1.742 * 'no information on age'
+0.219 * 'age under 20'
+0.071 * 'age 20 to 24'
+0.001 * 'age 30 to 34'
+0.094 * 'age 35 to 39'
+0.204 * 'age over 39'
+0.057 * 'low Apgar after 1 min'
+0.192 * 'low Apgar after 1 and 5 min'
+0.324 * 'low Apgar after 10 min'
+0.087 * 'no information on Appar score'
+0.140 * 'no information on family situation'
+0.221 * 'not living together'
+0.152 * 'birth year 1989'
+0.238 * 'birth year 1990'
+0.164 * 'low educational level'
+0.028 * 'middle educational level'
+0.258 * 'no information on educational level'
+0.259 * 'mother extensive hospitalisation and no psychiatric diagnosis'
+0.530 * 'mother psychiatric diagnosis and no extensive hospitalisation'
+0.605 * 'mother extensive hospitalisation and psychiatric diagnoses'
+0.566 * 'child extensive hospitalisation and no psychiatric diagnosis'
+1.539 * 'child psychiatric diagnosis and no extensive hospitalisation'
+2.479 * 'child extensive hospitalisation and psychiatric diagnoses'
```

Table 2 shows the characteristics of the adolescents at different life stages and classified according to the GNI categories of the mother's country of birth. This table indicates that adolescents born to mothers from low-income countries used fewer psychotropic drugs than do adolescents with mothers born in Sweden. However, adolescents with mothers from low-income countries were hospitalised more frequently and had more psychiatric diagnoses than were adolescents with Swedish mothers.

Table 2. Maternal country of birth classified according to World Bank Gross National Income (GNI) by child's characteristics. Numbers are percentages unless otherwise indicated.

	Sweden	HIC	UMIC	LMIC	LIC
Number of children	288 626	15 905	9572	7972	2435
Number of cirilden	140 235	7743	4656	3883	1174
Use of psychotropics at age 18 (ATC code)					
 Psychotropics unspecified (N5+N6) 	4.7	5.0	3.9	3.4	2.2
 Antipsychotics (N5A) 	0.4	0.5	0.5	0.5	0.2
Anxiolytics (N5B)	1.5	1.6	1.4	1.3	0.7
 Hypnotics and sedatives (N5C) 	1.2	1.5	1.0	1.2	0.4
 Antidepressants (N6A) 	2.8	2.9	2.0	1.6	0.9
 Psychostimulants (N6B) 	0.8	0.7	0.5	0.4	0.3
Child's characteristics during perinatal period					
Birth year					
- 1988	32.1	33.2	31.2	28.7	28.1
- 1989	33.1	32.3	32.8	32.2	32.4
- 1990	34.8	34.5	35.9	39.1	39.4
Apgar score					
 Low Apgar score 	94.1	94.4	93.6	93.8	91.7
 Low Apgar score (<6) after 1 min only 	3.0	2.4	3.3	3.4	4.1
 Low Apgar score (<6) after 1 and 5 min. 	0.7	0.7	0.9	0.7	1.2
 Low Apgar score (<6) after 10 min 	0.4	0.5	0.4	0.5	0.8
 Apgar scores missing 	1.9	2.0	1.8	1.6	2.2
Child's characteristics until age 15					
Hospitalised >30 days	2.1	2.0	2.7	2.2	3.0
Psychiatric diagnosis according to Patient Registry	1.6	1.8	1.9	1.8	2.5
Hospitalised >30 days (A) and psychiatric	1.0	1.0	1.7	1.0	2.3
diagnosis (B) combined					
- (A) no and (B) no	96.7	96.5	95.9	96.3	94.9
- (A) yes and (B) no	1.8	1.7	2.3	1.9	2.5
- (A) no and (B) yes	1.2	1.4	1.4	1.5	2.1
- (A) yes and (B) yes	0.4	0.4	0.5	0.3	0.5

HIC: high-income country, UMIC: upper middle-income country, LMIC: lower middle-income country, LIC: low-income country.

For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

Table 3 is similar to Table 1 but provides information on maternal characteristics. Low-income countries are those countries outside of Europe. Compared with Swedish mothers, mothers from low-income countries have lower income, lower educational achievement and are less likely to live with the father of their children. Furthermore, a higher percentage of mothers from low-income countries have been hospitalised for more than 30 days, and mothers from low-income countries have a much lower smoking prevalence.

Table 3. Maternal country of birth classified according to World Bank Gross National Income (GNI) by maternal characteristics. Numbers are percentages unless otherwise indicated.

Maternal characteristics during perinatal period	Sweden	HIC	UMIC	LMIC	LIC
Geographical region of birth					
- Sweden	100				
Nordic country outside Sweden	100	79.4			
Rest of Europe		14.6	63.4	27.6	
Rest of EuropeRest of the world		6.0	36.6	72.4	100
Years spent in Sweden at delivery		0.0	20.0	,	100
- >18	98.9	49.8	9.5	13.2	7.6
- 6-18	0.6	32.5	35.3	20.7	30.9
- 0-5	0.4	15.9	38.1	46.5	44.4
 No information 	0.1	1.9	17.2	19.6	17.1
Age at delivery in years					
No information	0.0	0.2	8.3	8.4	8.3
- <20	2.8	2.7	5.3	3.6	2.9
- 20-24	24.3	19.4	24.9	22.5	22.8
- 25-29	36.9	33.5	26.6	32	35.7
- 30-34	24.9	27.6	21.3	22.5	22.2
- 35-39	9.3	13.2	11.3	9.2	7.3
- >39	1.8	3.3	2.4	1.8	0.9
Equalised disposable household income in 1990					
- Low	31.0	34.9	62.1	56.7	57.2
- Middle	33.1	30.4	19.3	20	22.9
- High	33.4	31.6	13.5	14.8	12.8
 No information 	2.5	3.1	5.1	8.4	7.1
Highest completed educational level in 1990					
 Primary school (9 years) or less 	17.2	25.2	41.2	36.8	46.9
 Lower secondary (12 years) 	54.9	47.7	32.2	33.6	24.6
 Higher education (15 years) 	25.1	22.1	14.6	13.4	9.9
 No information 	2.9	5.0	12.1	16.2	18.6
Living with child's father					
- Yes	90.6	88.1	85.5	87	82.2
- No	5.2	6.9	6.9	5.4	9.8
 No information 	4.2	5.0	7.6	7.6	8.0
Smoking during pregnancy					
- No	68.0	62.4	69.3	75.5	83.1
- 1-9 cigarettes per day	15.0	16.3	_12.5	8.6	4.1
- >9 cigarettes per day	9.3	12.6	6.4	4.7	1.0
 No information 	7.6	8.7	11.8	11.2	11.8
Pre-eclampsia	2.1	2.0	1.5	1.2	2.4
Maternal characteristics until child aged 15					
Death	0.6	1.0	0.8	0.6	0.9
Hospitalised >30 days	7.0	8.1	7.2	6.9	11.0
Psychiatric diagnosis according to Patient Registry	1.4	2.1	1.5	1.4	1.3
Hospitalised >30 days (A) and psychiatric	1.4	2.1	1.5	1	1.5
diagnosis (B) combined					
- (A) no and (B) no	91.2	89.1	90.0	90.2	86.4
- (A) yes and (B) no	4.9	5.1	5.2	4.9	8.4
- (A) no and (B) yes	1.9	2.8	2.7	2.9	2.5
- (A) yes and (B) yes	2.1	3.0	2.1	2.0	2.6
HIC: high-income country LIMIC: upper middle-income					

HIC: high-income country, UMIC: upper middle-income country, LMIC: lower middle-income country, LIC: low-income country.

Table 4 shows the odds ratios and 95% confidence intervals for psychotropic drug use in adolescence as a function of the GNI categories of maternal country of birth stratified by gender. Overall, the lower the GNI of the mother's country of birth, the lower the probability of use of psychotropic drugs among second-generation immigrants compared with adolescents born to Swedish-born mothers. In the unadjusted analysis (Table 3, model 1), this association was more evident for girls than for boys. Nevertheless, after adjustment for the risk score (Table 3, model 2), the association was accentuated and evident in both sexes. The lowest risk for psychotropic drug use was found in girls born to mothers from low-income countries.

Table 4. Association between maternal birth country and psychotropic drug use at age 18, unadjusted (model 1) and adjusted (model 2) for risk score. * Values are odds ratios (ORs) and 95% confidence interval (CIs).

	Boys Mode		16	Mode	el 2		Girls Mode			Mode	el 2	
	OR	95%	6 CI	OR	95%	6 CI	OR	95%	6 CI	OR	95%	% CI
Maternal country of birth												
– Sweden	ref			ref			ref			ref		
 High-income country 	1.08	(0.96)	1.21)	1.01	(0.90)	1.13)	1.08	(0.99)	1.19)	1.03	(0.94)	1.13)
 Upper middle-income country 	0.94	(0.80)	1.10)	0.80	(0.69)	0.94)	0.74	(0.64)	0.85)	0.66	(0.57)	0.76)
 Lower middle-income country 	0.90	(0.75)	1.07)	0.78	(0.66)	0.94)	0.59	(0.50)	0.70)	0.53	(0.45)	0.63)
 Low-income country 	0.69	(0.48)	0.98)	0.57	(0.40)	0.82)	0.32	(0.21)	0.48)	0.28	(0.19)	0.42)
Risk score for psychotropic drug use*												
1st quartile				ref						ref		
 2nd quartile 				1.17	(1.01)	1.29)				1.10	(1.04)	1.21)
 3rd quartile 				1.45	(1.32)	1.59)				1.31	(1.13)	1.31)
4th quartile				2.55	(2.34	2.77)				1.81	(1.81	2.01)

^{*}Variables included in the risk score: Perinatal period (Apgar score, child's birth year, maternal smoking during pregnancy, maternal age, maternal educational level, family situation); Infancy, Childhood and Adolescence (extensive hospitalisation child, psychiatric diagnosis child, extensive hospitalisation mother, psychiatric diagnosis mother).

DISCUSSION

Our analyses indicate that Swedish adolescents born to immigrant women have a lower use of psychotropic drugs in adolescence than do the offspring of native Swedish women. This observation was evident in both sexes but was especially pronounced in girls. Moreover, the relative use of psychotropic drugs decreased linearly with the income (i.e., GNI per capita) of the mother's country of birth and was lowest for girls with mothers born in low-income countries.

Several mechanisms could explain our findings. Adolescents born to immigrant women might have better psychological health and therefore lower needs for psychotropic drug treatment than the offspring of native Swedish women. If this were true, adjusting for needs would make any difference disappear or at least decrease. To create a risk score, we adopted a life-course approach [27] and observed that a number of maternal and offspring characteristics during pregnancy, delivery, infancy, childhood and adolescence predicted psychotropic drug use during adolescence. Nevertheless, after adjusting for needs using the risk score, the influence of mother's country of birth (i.e., lower use of psychotropic drugs among adolescents with mothers from low-income countries) remained independent and even increased. This finding suggests that differences in psychotropic drug use may reflect divergences in access to healthcare resources rather than lower needs. However, we cannot exclude the possibility that we missed other relevant factors, such as childhood adversities, which affect psychological health later in life (e.g., neglect and verbal or physical abuse) [28]. Because we were restricted to the available information, our adjustment for needs is inherently incomplete.

One explanation for the observed underutilisation of psychotropic drugs by second-generation immigrants may be that the cultural context in which one grows up influences healthcare-

seeking behaviour. Examples of these cultural influences are different conceptions of disease and taboos [29]. Another factor affecting the lower use of psychotropic drugs by second-generation immigrants may be so-called health illiteracy (i.e., unfamiliarity with the country's healthcare system) [30].

These cultural and health illiteracy-related aspects may explain the underutilisation of health care by mothers. However, the children in our study were *second*-generation immigrants who grew up and attended school in Sweden. Thus, they had the opportunity to become familiar with the Swedish system. Nevertheless, we think it is likely that the effects of cultural influences and health illiteracy on mothers' healthcare-seeking behaviours are transmitted to the next generation because it is often mothers who regulate healthcare contacts for children.

Interestingly, our results suggesting a relative underutilisation of psychotropic drugs in 18-year-old second-generation immigrants contrast with previous findings that observed a higher rate of psychotropic drug use in first-generation immigrants [31]. In addition, a study in the Netherlands [32] found that in comparison with the native population, first- and second-generation Turkish and Moroccan immigrants had an increased rate of antidepressant and antipsychotic drug prescriptions and a decreased rate of ADHD medication and lithium prescriptions. However, in our earlier investigations in the county of Scania, we observed that overall utilisation of psychiatric healthcare resources in 18- to 80-year-old immigrants was considerably less than expected according to self-reported needs [30].

As expected, we found a high correlation between time spent in Sweden before delivery and mothers' country of birth. We observed the lowest percentage of mothers with more than 18 years in Sweden in the group of mothers from low-income countries. Because spending fewer years in Sweden reduces the opportunity for acculturation, this variable may have contributed to the relative underutilisation of psychotropic drugs in this group.

In the present analysis, we only considered prescribed drugs. Therefore, the results may be biased by the sale of over-the-counter (OTC) drugs. In Sweden, the only available OTC psychotropic drug is Propiomazine. Nevertheless, we believe it is improbable that our results would be affected by the inclusion of Propiomazine in the analyses because this assumption would imply that second-generation immigrants use this OTC drug as a substitute for all other psychotropic drugs.

One strength of our study is that we had information from registries covering the entire Swedish population. Furthermore, because giving birth at home is very unusual in Sweden, nearly all births are registered in the Medical Birth Register. The information in the Medical Birth Register, National Patient Register and National Cause of Death Register is mandatorily reported by law, and the quality of the registries is regularly evaluated by the National Board of Health and Welfare and Statistics Sweden.

Measurement of psychotropic drug use using administrative registries reflects both access to healthcare and the presence of psychological disorders. From the perspective of community medicine, the identification of an imbalance between needs and utilisation of psychotropic drugs raises questions about equity in access to health care resources. However, our findings also identify the existence of information bias when using a register-based measurement of psychotropic drug use as a proxy for impaired psychological health if second-generation immigrants are included in the analyses.

We performed analyses investigating specific groups of psychotropic drugs (results available on request), but the results were similar to those found when analysing the entire medication group. Moreover, because the therapeutic profiles of the studied medication groups overlap each other (i.e., different psychiatric disorders can be treated with the same drug group), we cannot create a direct link between specific medications and diagnoses.

The suitability of treating young people with psychotropic drugs is polemic. Certainly, there are risks when these drugs are used more often than necessary [33, 34]. However, underuse may engender avoidable suffering [35]. It is not clear whether adolescents in Sweden and other high-income countries are prescribed these drugs in excess (i.e., overutilisation) [36].

If underutilisation of psychotropic drugs by adolescent second-generation immigrants occurs, policy makers in the Swedish healthcare system should take this situation into account. Swedish healthcare policy is based on the principle of equity that supports the distribution of healthcare resources on equal terms and according to needs, regardless of any other determinants. This aspect is particularly relevant because one of the main goals stated by the Swedish National Institute of Public Health is 'secure and favourable conditions during childhood and adolescence' and, specifically, 'improved mental health amongst children and young people' (39). Paraphrasing Wittkampf [32], we need further research to clarify whether patients of different ethnic backgrounds with the same symptoms receive similar diagnoses and adequate treatment.

COMPETING INTERESTS

None declared

FUNDING

This work was supported by the Swedish Council for Working Life and Social Research (FAS) [Dnr 2010-0402, PI Juan Merlo], the Swedish Research Council (VR) [Dnr K2011-69X-15377-07-6, PI Juan Merlo] and the Centre for Economic Demography at Lund University.

STATEMENT

"The Corresponding Author has the right to grant on behalf of all authors and does grant on behalf of all authors an exclusive licence (or non-exclusive for UK Crown Employees) on a worldwide basis to the BMJ Publishing Group Ltd, and its Licensees to permit this article (if accepted) to be published in The Journal of Epidemiology and Community Health and any other BMJPGL products and to exploit all subsidiary rights, as set out in our licence."

CONTRIBUTORSHIP STATEMENT

All authors WvL and JM hade the original idea of the study. JM, WvL and SN participated in the design, analysis, interpretation of data, and drafting of the article. All authors approved the final version to be published. JM is the guarantor of the article.

REFERENCES

- Health on equal terms--national goals for public health. *Scand J Public Health* 2001; **Suppl 57**:1-68.
- Practice parameter on the use of psychotropic medication in children and adolescents. *J Am Acad Child Psy* 2009;**48**:961-73.
- Vitiello B. An international perspective on pediatric psychopharmacology. *Int Rev Psychiatry* 2008;**20**:121-6.
- 4 Zito JM, Safer DJ, dosReis S, *et al.* Trends in the prescribing of psychotropic medications to preschoolers. *JAMA*: the journal of the American Medical Association 2000;**283**:1025-30.
- 5 Zito JM, Safer DJ, DosReis S, *et al.* Rising prevalence of antidepressants among US youths. *Pediatrics* 2002;**109**:721-7.
- 6 Thomas CP, Conrad P, Casler R, *et al.* Trends in the use of psychotropic medications among adolescents, 1994 to 2001. *Psychiatr Serv* 2006;**57**:63-9.
- Sondergard L, Kvist K, Andersen PK, *et al.* Do antidepressants precipitate youth suicide?: a nationwide pharmacoepidemiological study. *Eur Child Adoles Psy* 2006;**15**:232-40.
- 8 Ekblad M, Gissler M, Lehtonen L, *et al.* Relation of prenatal smoking exposure and use of psychotropic medication up to young adulthood. *American journal of epidemiology* 2011;**174**:681-90.
- 9 Gyllenberg D, Sourander A. Psychotropic drug and polypharmacy use among adolescents and young adults: Findings from the Finnish 1981 Nationwide Birth Cohort Study. *Nord J Psychiat* 2012.
- 10 Lindblad F, Ringback Weitoft G, Hjern A. Maternal and paternal psychopathology increases risk of offspring ADHD equally. *Epidemiol Psychiatr Sci* 2011;**20**:367-72.
- Lindstrom K, Lindblad F, Hjern A. Preterm birth and attention-deficit/hyperactivity disorder in schoolchildren. *Pediatrics* 2011;**127**:858-65.
- 12 Lindblad F, Hjern A. ADHD after fetal exposure to maternal smoking. *Nicotine & tobacco research : official journal of the Society for Research on Nicotine and Tobacco* 2010;**12**:408-15.
- 13 Hjern A, Weitoft GR, Lindblad F. Social adversity predicts ADHD-medication in school children--a national cohort study. *Acta paediatrica* 2010;**99**:920-4.
- 14 Campbell SM, Roland MO. Why do people consult the doctor? *Family practice* 1996:**13**:75-83.

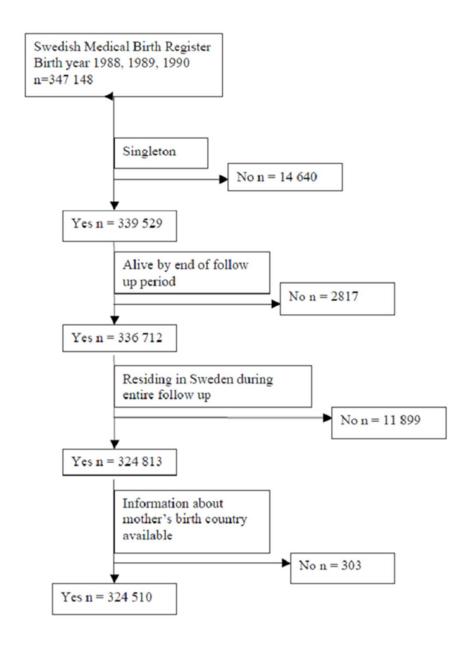
- Norcross WA, Ramirez C, LA P. The influence of women on the health care-seeking behavior of men. *J Fam Pract 1996 Nov;43(5):475-480* 1996;**45**:475-80.
- Hopton J, Hogg R, McKee I. Patients' accounts of calling the doctor out of hours. *BMJ* 1996;**313**:991-4.
- 17 Fylkesnes K. Determinants of health care utilization--visits and referrals. *Scand J Soc Med* 1993;**21**:40-50.
- 18 Adamson J, Ben-Shlomo Y, Chaturvedi N, *et al.* Ethnicity, socio-economic position and gender--do they affect reported health-care seeking behaviour? *Soc Sci Med* 2003;**57**:895-904.
- 19 Lay B, Nordt C, Roessler W. Mental hospital admission rates of immigrants in Switzerland. *Social Psychiatry and Psychiatric Epidemiology* 2007;**42**:229-36.
- 20 Arbogast PG, Ray WA. Use of disease risk scores in pharmacoepidemiologic studies. *Statistical Methods in Medical Research* 2009;**18**:67-80.
- 21 Ben-Shlomo Y, Kuh D. A life course approach to chronic disease epidemiology: conceptual models, empirical challenges and interdisciplinary perspectives. *Int J Epidemiol* 2002;**31**:285-93.
- WHO. ATC -Structure and principles. WHO Collaborating Centre for Drug Statistics Methodology 2011.
- 23 EpC. The Swedish Medical Birth Register -A summary of content and quality. 2003.
- 24 Cafferata GL, Meyers SM. Pathways to Psychotropic-Drugs Understanding the Basis of Gender Differences. *Medical Care* 1990;**28**:285-300.
- 25 Benyamini Y, Blumstein T, Boyko V, *et al.* Cultural and educational disparities in the use of primary and preventive health care services among midlife women in Israel. *Womens Health Issues* 2008;**18**:257-66.
- 26 Ayonrinde O. Importance of cultural sensitivity in therapeutic transactions Considerations for healthcare providers. *Disease Management & Health Outcomes* 2003;**11**:233-48.
- 27 Ben-Shlomo Y, Kuh D. A life course approach to chronic disease epidemiology: conceptual models, empirical challenges and interdisciplinary perspectives. *International journal of epidemiology* 2002;**31**:285-93.
- 28 Lanius RA, Vermetten E, Pain C. *The impact of early life trauma on health and disease : the hidden epidemic.* Cambridge, UK; New York: Cambridge University Press 2010.
- 29 Lindert J, Schouler-Ocak A, Heinz A, *et al.* Mental health, health care utilisation of migrants in Europe. *European Psychiatry* 2008;**23**:S14-S20.
- 30 Merlo J. Fördelning av psykisk ohälsa och sjukvårdsresurser i Malmö, region Skåne. In: Östman M, ed. *Migration och psykisk ohälsa*. Malmö: Malmö högskola; Hälsa och samhalle 2008:45-94.

- 31 Hjern A. High use of sedatives and hypnotics in ethnic minorities in Sweden. EthnHealth 2001;**6**:5-11.
- 32 Wittkampf LC, Smeets HM, Knol MJ, et al. Differences in psychotropic drug prescriptions among ethnic groups in the Netherlands. Social psychiatry and psychiatric epidemiology 2010;45:819-26.
- 33 Zakriski Aea. Justifiable psychopharmacology or overzealous prescription? Examining parental reports of lifetime prescription histories of psychiatrically hospitalised children. Child and Adolescent Mental Health 2005;10:7.
- 34 Jureidini JN, Doecke CJ, Mansfield PR, et al. Efficacy and safety of antidepressants for children and adolescents. BMJ 2004;328:879-83.
- 35 Gentile S. Efficacy of antidepressant medications in children and adolescents with obsessive-compulsive disorder: a systematic appraisal. J Clin Psychopharmacol **:31**:625-32.
- 36 Victorri-Vigneau C, Basset G, Jolliet P. How a novel programme for increasing awareness of health professionals resulted in a 14% decrease in patients using excessive doses of psychotropic drugs in western France. European Journal of Clinical Pharmacology 2006;62:311-6.

LEGEND TO THE FIGURE

Figure 1. Information flow showing the number of individuals remaining after the application of the exclusion criteria





Information flow showing the number of individuals remaining after the application of the exclusion criteria 143x200mm (96 x 96 DPI)



MOTHER'S COUNTRY OF BIRTH AND PRESCRIPTION OF PSYCHOTROPIC MEDICATION IN SWEDISH ADOLESCENTS: A LIFE COURSE APPROACH

Journal:	BMJ Open
Manuscript ID:	bmjopen-2012-001260.R1
Article Type:	Research
Date Submitted by the Author:	18-Jul-2012
Complete List of Authors:	van Leeuwen, Willemijn; Lund University, Faculty of Medicine, Social Epidemiology Nilsson, Sofia; Lund University, Faculty of Medicine, Social Epidemiology Merlo, Juan; Lund University, Faculty of Medicine, Social Epidemiology
Primary Subject Heading :	Mental health
Secondary Subject Heading:	Epidemiology, Public health, Sociology, Health services research, Pharmacology and therapeutics
Keywords:	EPIDEMIOLOGY, Quality in health care < HEALTH SERVICES ADMINISTRATION & MANAGEMENT, PUBLIC HEALTH, SOCIAL MEDICINE, Community child health < PAEDIATRICS, Child & adolescent psychiatry < PSYCHIATRY

SCHOLARONE™ Manuscripts

MOTHER'S COUNTRY OF BIRTH AND PRESCRIPTION OF PSYCHOTROPIC MEDICATION IN SWEDISH ADOLESCENTS: A LIFE COURSE APPROACH

Corresponding author

Juan Merlo

MD, PhD, Tenured Professor

Unit for Social Epidemiology, Skåne University Hospital

Jan Waldenströms gata 35

SE-20502 Malmö, Sweden

E-mail: juan.merlo@med.lu.se

Tel. +46 40391329

Authors

Willemijn Van Leeuwen^{1,2} Sofia Nilsson¹ Juan Merlo¹

Affiliations

Running title:

Psychotropic drug use in Swedish adolescents

Keywords

Psychotropic drugs, adolescent, second-generation immigrants, needs assessment, bias (epidemiology)

Word count:

Abstracts: 247

Manuscript: 2916

¹ Unit for Social Epidemiology, Faculty of Medicine, Lund University, Malmö, Sweden

² Dept. of Public Health, Academic Medical Center (AMC) University of Amsterdam, Netherlands

Page 2 of 23

ABSTRACT

Objectives

 Besides medical needs, psychotropic medication use in adolescence might be conditioned by the cultural context of the family. This knowledge is relevant for both detecting inequities in health care, and identifying information bias in epidemiological studies using psychotropic medication as a proxy for impaired psychological health. Therefore, we investigated whether, independent of needs, the socioeconomic characteristics of the mother's country of birth are associated with psychotropic medication use in Swedish-born adolescents.

Methods

By linking the Swedish Medical Birth Registry to other national registers, we identified all 324510 singletons born between 1988 and 1990 and who were alive and residing in Sweden until the age of 18 (years 2006 to 2008). We measured their use of psychotropic medication in the year they turned 18. Applying a life-course approach, we included both the mother's and the children's characteristics throughout pregnancy, delivery, infancy, childhood and adolescence when calculating a risk score to adjust for needs. We classified the mother's country of birth according to the gross national income (GNI) per capita of each country.

Results

Overall, the lower the income of the mother's birth country, the lower the probability of psychotropic medication use among children. When adjusting for needs, the association became even stronger.

Discussion

Besides medical needs, use of psychotropic medication by descendants of immigrants seems conditioned by the socioeconomic characteristics of the mothers' countries of birth. The threat of information bias must be considered if psychotropic medication is used a proxy for impaired psychological health in descendants of immigrants.

INTRODUCTION

There are many knowledge gaps concerning the determinants of psychotropic drug use in immigrants and, to the best of our knowledge, this information is almost absent for descendants of immigrants. In Sweden, the healthcare system aims to allocate resources on equal terms and according to needs rather than by gender, socioeconomic position or country of birth^{1 2}. Therefore, from a public health perspective, it is important to regularly examine the distribution of health care resources in relation to health care needs across different socioeconomic groups, gender and country of births.

Psychotropic medication are potent therapy for treating impaired psychological disorders in childhood and adolescence³. Despite the many uncertainties concerning their safety and efficacy⁴, the use of this medication in adolescents has increased considerably during recent decades⁵⁻⁷. In addition, psychotropic medication is used as a proxy for impaired psychological health in adolescents and young adults in register-based epidemiological studies⁸⁻¹⁴.

Psychotropic drug use in adolescents should be conditioned by needs. However, the concept of 'need' is ambiguous, and care-seeking and care-giving are influenced by other factors in addition to disease ¹⁵ ¹⁶. There are also complex relationships between symptoms, expectations, social factors and an individual's conception of disease ¹⁷ ¹⁸. Furthermore, the concepts of disease, illness and need are culturally defined to a great extent, and are influenced by societal and contextual factors acting over and above individual characteristics ¹⁹. Therefore, psychotropic drug use may be conditioned by an individual's country of birth per se or in interaction with Swedish society ²⁰. Patterns of healthcare utilisation may also be transmitted across generations. This information is relevant not only for detecting inequities in health care but also for identifying information bias in epidemiological studies that use psychotropic medication as a proxy for impaired

psychological health.

In this context, we investigated psychotropic medication use in 18 year old adolescents born in Sweden. We aimed to compare adolescents born to immigrant women (i.e., descendants of immigrants) with those born to native mothers. Our hypothesis was that any difference would disappear, or at least decrease, after adjusting for needs. We appraised needs using a risk score²¹ for psychotropic drug use including information on the health and socioeconomic status of the mothers and the children during pregnancy and delivery as well as throughout the children's infancy, childhood and adolescence²². We studied every 18-year-old adolescent in Sweden during the years 2006-2008.

POPULATION AND METHODS

Study population

The Swedish Medical Birth Registry is linked to several other national databases, including the National Inpatient Registry, the National Mortality Registry, the Swedish Drug Prescription Register, the Register of the Total Population, and the Swedish 1990 population census, using a unique personal identification number. These registries are administered by the Swedish authorities (i.e., The National Board of Health and Welfare and Statistics Sweden). To ensure the anonymity of the subjects, these authorities encrypted the original personal identification numbers of the individuals before delivering the data to us. The database was approved by the Regional Ethics Review Board in southern Sweden.

Using the Medical Birth Register, we identified all singleton children born in Sweden between 1988 and 1990 who were alive and residing in Sweden from birth until the year in which the child turned 18 (2006-2008) and whose mother's country of birth could be identified. The final study population represented 94% (324510/347148) of all children born

in 1988, 1989 and 1990. Figure 1 shows a flow diagram of the selection criteria used to define the study population.

Assessment of the outcome variable

The outcome variable was the use of psychotropic medication (i.e., dispensation at the pharmacy) according to the Drug Prescription Registry. We obtained this information in the calendar year that the child turned 18. Psychotropic medication were defined as all drugs with Anatomic Therapeutic Chemical (ATC) classification system²³ codes starting with N5 or N6 (see Table 1). We dichotomised the outcome variable into use or non-use of psychotropic medication.

Assessment of child's characteristics

We categorised *birth year* into 1988, 1989 or 1990, using 1988 as the reference group. *Apgar score* was measured on a scale from 1-10 at the first, fifth and 10^{th} minute after birth, where a score of 10 indicates a delivery without distress for the infant. We created dummy variables where an Apgar score ≤ 5 was classified as 'low', and an Apgar score ≥ 5 was classified as 'not low'. The latter was used as the referent. We recoded the missing values at 5 or 10 minutes as 'not low Apgar' because it can be assumed that the subsequent Apgar scores were high and therefore not measured if the Apgar score after one minute was high 24 . This was the case for 37% of the children.

For every hospital discharge, a main diagnosis and up to seven secondary diagnoses are recorded in the Patient Register and coded according to the International Classification of Diseases (ICD). We defined a child admitted to a hospital with a diagnosis coded 290 to 319 or F00 to F90 (ICD versions 9 and 10, respectively) as *severe psychiatric morbidity*. Children never admitted to a hospital with any of these diagnoses were established as the reference

group in the comparisons.

For every child, we calculated the total number of days spent in the hospital from birth until their 15th birthday. We distinguished between children with *extensive hospitalisation* (i.e., children spending 30 days or more in the hospital) and children with less than 30 days in the hospital or who were never hospitalised, using the latter group as the referent. We combined *history of psychiatric diagnosis* and extensive hospitalisation of the children into four categories, as indicated in table 2.

Assessment of mother's characteristics

We categorised the mothers' *country of birth* by two different types of criteria: socioeconomic and geographic. Both categorisations are listed in table 2. In our analyses, we used the socioeconomic criterion, which categorises the mother's country of birth by means of the World Bank classification of country economies. Using this classification we assume that countries in the same group resemble each other in access to material and welfare resources and medical resources in particular. This procedure seems less prone to stigmatize individuals than when using their demographic (e.g., immigrant vs. Swedish-born), or ethnic (e.g., Caucasian, Asian, Black) characteristics or their geographic origin (e.g., Northern European, African).

Countries were grouped according to their gross national income (GNI) per capita using the World Bank Atlas method into low-, lower middle-, upper middle-, and high-income countries. Sweden was included in the high-income category. However, we separated Sweden from the group of high-income countries and used Sweden as the referent in the comparisons because we conducted the study in the context of Swedish culture.

We classified mother's age at delivery into six groups (i.e., <20 years, 20-24 years, 25-29

 29... >39 years). We considered mothers from the age group 25-29 the reference group in the comparisons.

We classified *maternal smoking during pregnancy* into four categories: 'no smoking', 'light smokers' (1-9 cigarettes per day), 'heavy smokers' (>9 cigarettes per day) and 'missing values'. The missing values ranged from 7.6% among the Swedes to 11.8% among mothers from low-income countries. We considered the non-smoking group the reference category. The information on smoking was self-reported and recorded when the mother attended her first scheduled antenatal care visit.

We identified all mothers with a diagnosis of *pre-eclampsia* (ICD-9 code 642) and considered mothers who did not suffer from pre-eclampsia the reference group in the comparisons.

We defined mother's *educational achievement* as the highest completed level of education in 1990. This variable was categorised into four groups: 'elementary school or lower' (9 years), 'lower secondary school' (12 years total), 'higher educational achievement' (>12 years) and 'missing information'. We used the 'higher educational achievement' category as the reference group in the comparisons.

Information regarding whether parents were *living together* was based on self-reported status when the mother was first seen for antenatal care. We considered mothers living together with the father of the child the reference group.

We identified mothers with *severe psychiatric morbidity* and with *extensive hospitalisation* in a manner similar to that used for children (see above). In addition, we combined a history of psychiatric diagnosis with extensive hospitalisation in the same way as we did for the children (see table 2).

Statistical Analyses

 To estimate needs, we applied a logistic regression and constructed an equation that predicted the probability (i.e., the risk score) ²¹ of using psychotropic medications around the age of 18 in the unexposed group (i.e., children with mothers born in Sweden). This equation included the maternal and child's characteristics mentioned above.

These variables were chosen using both previous knowledge and an explorative approach. When two variables showed multicollinearity, we selected the variable that provided a better goodness of fit by means of a chi-square test (e.g., mother's education compared with household income).

Subsequently, we applied the risk score equation to the entire population (i.e., children with mothers born in Sweden *and* children with immigrant mothers).

The equation of the risk score is presented in Table 1. The individual value of the risk score (RS) (i.e., predicted probability) was obtained with the following formula:

$$RS = [e^{logit} / (1 + e^{logit})] * 100.$$

In the analyses, we categorised the risk score into four groups by quartiles and used the group with the lowest risk score as the reference group in the comparisons.

Using logistic regression, we developed two consecutive models to estimate the relative differences (i.e., odds ratios) in the use of psychotropic medications. In the first model, we investigated the bare association between mother's country of birth and the use of psychotropic medication in adolescence. In the second model, we included the risk score described above to adjust for possible dissimilarities in needs between the different maternal countries of birth. We analysed boys and girls separately in the models because there are cultural differences in gender roles, and gender may influence healthcare and psychotropic drug use²⁵⁻²⁷. We used SPSS version 18 for all analyses.

RESULTS

Table 1 indicates the regression coefficients of the variables used in the logistic regression to obtain the risk score (i.e., predicted probability) for psychotropic drug use.

Table 1. Equation for estimation of the risk score (RS) used in the analyses to approximate the need for psychotropic drugs. The equation was calculated in the population of children born to Swedish mothers and subsequently applied to all the children. The RS corresponds to the predicted probability of using psychotropic drugs as a function of the variables included in the model.

```
Logit (Use of psychotropic drugs) =
-3.467 = intercept
+0.180 * 'smoking 1-9 cigarettes per day'
+0.291 * 'smoking more than 9 cigarettes per day'
+0.057 * 'no information on smoking'
+0.126 * 'pre-eclampsia'
+1.742 * 'no information on age'
+0.219 * 'age under 20'
+0.071 * 'age 20 to 24'
+0.001 * 'age 30 to 34'
+0.094 * 'age 35 to 39'
+0.204 * 'age over 39'
+0.057 * 'low Apgar after 1 min'
+0.192 * 'low Apgar after 1 and 5 min'
+0.324 * 'low Apgar after 10 min'
+0.087 * 'no information on Apgar score'
+0.140 * 'no information on family situation'
+0.221 * 'not living together'
+0.152 * 'birth year 1989'
+0.238 * 'birth year 1990'
+0.164 * 'low educational level'
+0.028 * 'middle educational level'
+0.258 * 'no information on educational level'
+0.259 * 'mother extensive hospitalisation and no psychiatric diagnosis'
+0.530 * 'mother psychiatric diagnosis and no extensive hospitalisation'
+0.605 * 'mother extensive hospitalisation and psychiatric diagnoses'
+0.566 * 'child extensive hospitalisation and no psychiatric diagnosis'
+1.539 * 'child psychiatric diagnosis and no extensive hospitalisation'
+2.479 * 'child extensive hospitalisation and psychiatric diagnoses'
```

Table 2 shows the characteristics of the adolescents at different life stages and classified according to the GNI categories of the mother's country of birth. This table indicates that adolescents born to mothers from low-income countries used less psychotropic medication than do adolescents with mothers born in Sweden. However, adolescents with mothers from low-income countries were more frequently hospitalized, and also more often hospitalized with a psychiatric diagnosis, than adolescents with Swedish mothers.

Table 2. Child's characteristics by maternal country of birth classified according to World Bank Gross National Income (GNI) by. Numbers are percentages unless otherwise indicated.

	Sweden	HIC	UMIC	LMIC	LIC
N. 1 (171)	288 626	15 905	9572	7972	2435
Number of children Number of girls	140 235	7743	4656	3883	1174
Use of newshatuenic medication at age 19 (ATC and a)					
Use of psychotropic medication at age 18 (ATC code) - Psychotropic medication unspecified (N5+N6)	4.7	5.0	3.9	3.4	2.2
- Antipsychotics (N5A)	0.4	0.5	0.5	0.5	0.2
- Anxiolytics (N5B)	1.5	1.6	1.4	1.3	0.2
Anxiorytics (N5D)Hypnotics and sedatives (N5C)	1.2	1.5	1.0	1.2	0.7
- Antidepressants (N6A)	2.8	2.9	2.0	1.6	0.4
- Psychostimulants (N6B)	0.8	0.7	0.5	0.4	0.3
Child's characteristics during perinatal period					
Birth year					
- 1988	32.1	33.2	31.2	28.7	28.1
- 1989	33.1	32.3	32.8	32.2	32.4
- 1990	34.8	34.5	35.9	39.1	39.4
Apgar score					
 Low Apgar score 	94.1	94.4	93.6	93.8	91.7
 Low Apgar score (<6) after 1 min only 	3.0	2.4	3.3	3.4	4.1
 Low Apgar score (<6) after 1 and 5 min. 	0.7	0.7	0.9	0.7	1.2
 Low Apgar score (<6) after 10 min 	0.4	0.5	0.4	0.5	0.8
 Apgar scores missing 	1.9	2.0	1.8	1.6	2.2
Child's characteristics until age 15					
Hospitalised >30 days	2.1	2.0	2.7	2.2	3.0
Psychiatric diagnosis according to Patient Registry	1.6	1.8	1.9	1.8	2.5
Hospitalised >30 days (A) and psychiatric					
diagnosis (B) combined					
- (A) no and (B) no	96.7	96.5	95.9	96.3	94.9
- (A) yes and (B) no	1.8	1.7	2.3	1.9	2.5
- (A) no and (B) yes	1.2	1.4	1.4	1.5	2.1
- (A) yes and (B) yes	0.4	0.4	0.5	0.3	0.5

HIC: high-income country, UMIC: upper middle-income country, LMIC: lower middle-income country, LIC: low-income country.

Table 3 is similar to Table 1 but provides information on maternal characteristics. Low-income countries are those countries outside of Europe. Compared with Swedish mothers, mothers from low-income countries have lower income, lower educational achievement and are less likely to live with the father of their children. Furthermore, a higher percentage of mothers from low-income countries have been hospitalised for more than 30 days, and mothers from low-income countries have a much lower smoking prevalence.

Table 3. Maternal characteristics by country of birth classified according to World Bank Gross National Income (GNI) by. Numbers are percentages

Ma	ternal characteristics during perinatal period	Sweden	ніс	UMIC	LMIC	LIC
ıvıa	ternar characteristics during permatar period	Sweden	me	UIVIIC	LIVIIC	LIC
Geo	graphical region of birth					
_	Sweden	100				
_	Nordic country outside Sweden		79.4			
_	Rest of Europe		14.6	63.4	27.6	
_	Rest of the world		6.0	36.6	72.4	100
Yea	rs spent in Sweden at delivery					
_	>18	98.9	49.8	9.5	13.2	7.6
_	6-18	0.6	32.5	35.3	20.7	30.9
_	0-5	0.4	15.9	38.1	46.5	44.4
_	No information	0.1	1.9	17.2	19.6	17.1
Age	at delivery in years					
_	No information	0.0	0.2	8.3	8.4	8.3
_	<20	2.8	2.7	5.3	3.6	2.9
_	20-24	24.3	19.4	24.9	22.5	22.8
_	25-29	36.9	33.5	26.6	32	35.7
_	30-34	24.9	27.6	21.3	22.5	22.2
_	35-39	9.3	13.2	11.3	9.2	7.3
_	>39	1.8	3.3	2.4	1.8	0.9
Hig	hest completed educational level in 1990					
_	Primary school (9 years) or less	17.2	25.2	41.2	36.8	46.9
_	Lower secondary (12 years)	54.9	47.7	32.2	33.6	24.6
_	Higher education (15 years)	25.1	22.1	14.6	13.4	9.9
_	No information	2.9	5.0	12.1	16.2	18.6
Livi	ing with child's father					
_	Yes	90.6	88.1	85.5	87	82.2
_	No	5.2	6.9	6.9	5.4	9.8
_	No information	4.2	5.0	7.6	7.6	8.0
Smo	oking during pregnancy					
_	No	68.0	62.4	69.3	75.5	83.1
_	1-9 cigarettes per day	15.0	16.3	12.5	8.6	4.1
_	>9 cigarettes per day	9.3	12.6	6.4	4.7	1.0
_	No information	7.6	8.7	11.8	11.2	11.8
Pre-	eclampsia	2.1	2.0	1.5	1.2	2.4

Maternal characteristics until child aged 15					
Death	0.6	1.0	0.8	0.6	0.9
Hospitalised >30 days	7.0	8.1	7.2	6.9	11.0
Psychiatric diagnosis according to Patient Registry	1.4	2.1	1.5	1.4	1.3
Hospitalised >30 days (A) and psychiatric					
diagnosis (B) combined					
- (A) no and (B) no	91.2	89.1	90.0	90.2	86.4
- (A) yes and (B) no	4.9	5.1	5.2	4.9	8.4
- (A) no and (B) yes	1.9	2.8	2.7	2.9	2.5
- (A) yes and (B) yes	2.1	3.0	2.1	2.0	2.6

HIC: high-income country, UMIC: upper middle-income country, LMIC: lower middle-income country, LIC: low-income country.

Table 4 shows the odds ratios and 95% confidence intervals for psychotropic medication use in adolescence as a function of the GNI categories of maternal country of birth stratified by gender. Overall, the lower the GNI of the mother's country of birth, the lower the probability of use of psychotropic medication among descendants of immigrants compared with adolescents born to Swedish-born mothers. In the unadjusted analysis (Table 3, model 1), this association was more evident for girls than for boys. Nevertheless, after adjustment for the risk score (Table 3, model 2), the association was accentuated and evident in both sexes. The lowest risk for psychotropic drug use was found in girls born to mothers from low-income countries.

Table 4. Association between maternal country of birth and psychotropic medication use at age 18, unadjusted (model 1) and adjusted (model 2) for the risk score for psychotropic medication use*. Values are odds ratios (ORs) and 95% confidence interval (CIs).

	Boys						Girls					
	Model 1			Model 2		Model 1			Model 2			
Maternal country of birth	OR	OR 95% CI		OR	95% CI		OR	95% CI		OR	95% CI	
- Sweden	Ref			Ref			Ref			Ref		
High-income country	1.08	(0.96)	1.21)	1.01	(0.90)	1.13)	1.08	(0.99)	1.19)	1.03	(0.94	1.13)
 Upper middle-income country 	0.94	(0.80)	1.10)	0.80	(0.69)	0.94)	0.74	(0.64)	0.85)	0.66	(0.57)	0.76)
 Lower middle-income country 	0.90	(0.75)	1.07)	0.78	(0.66)	0.94)	0.59	(0.50)	0.70)	0.53	(0.45)	0.63)
 Low-income country 	0.69	(0.48)	0.98)	0.57	(0.40)	0.82)	0.32	(0.21)	0.48)	0.28	(0.19)	0.42)
Risk score for psychotropic drug use*												
1st quartile				Ref						Ref		
 2nd quartile 				1.17	(1.01	1.29)				1.10	(1.04)	1.21)
 3rd quartile 				1.45	(1.32	1.59)				1.31	(1.13)	1.31)
 4th quartile 				2.55	(2.34)	2.77)				1.81	(1.81)	2.01)

^{*}Variables included in the risk score for psychotropic drug use: Perinatal period (Apgar score, child's birth year, maternal smoking during pregnancy, maternal age, maternal educational level, family situation); Infancy, Childhood and Adolescence (extensive hospitalisation child, psychiatric diagnosis child, extensive hospitalisation mother, psychiatric diagnosis mother).

DISCUSSION

Our analyses indicate that Swedish adolescents born to immigrant women have a lower use of psychotropic medication in adolescence than do the offspring of native Swedish women. This observation was evident in both sexes but was especially pronounced in girls. Moreover, the relative use of psychotropic medication decreased linearly with the income (i.e., GNI per capita) of the mother's country of birth and was lowest for girls with mothers born in low-income countries.

Several mechanisms could explain our findings. Adolescents born to immigrant women might have better psychological health and therefore lower needs for psychotropic drug treatment than the offspring of native Swedish women. If this were true, adjusting for needs would make any difference disappear or at least decrease. To create a risk score, we adopted a life-course approach²⁸ and observed that a number of maternal and offspring characteristics during pregnancy, delivery, infancy, childhood and adolescence predicted psychotropic drug use during adolescence. Nevertheless, after adjusting for needs using the risk score, the influence of mother's country of birth (i.e., lower use of psychotropic medication among adolescents with mothers from low-income countries) remained independent and even increased. This finding suggests that differences in psychotropic drug use may reflect divergences in access to healthcare resources rather than lower needs. However, we cannot exclude the possibility that we missed other relevant factors that condition needs, such as childhood adversities, which affect psychological health later in life (e.g., neglect and verbal or physical abuse)²⁹. Because we were restricted to the available information, our adjustment for needs is inherently incomplete.

Since the National Prescription Register contains only prescriptions that were handled by the pharmacies, information on medication prescribed but never picked up at the pharmacy is not

 available. Therefore, the observed differences in psychotropic drug use might reflect disparities in primary compliance rather than in prescription behavior (i.e., medication could be prescribed but the individual never pick it up from the pharmacy). In Sweden, at the time of our study, the total yearly cost that a particular individual needed to pay for prescribed medications was 4300 SEK (i.e., about 500 EU). Beyond this ceiling, medication was free of charge³⁰. Therefore, individuals with a low socioeconomic position may be less prone to pick up prescribed medication. We adjusted for socioeconomic factors in the risk score for psychotropic drug use but this procedure might be insufficient.

One explanation for the observed underutilisation of psychotropic medication by descendants of immigrants may be that the cultural context in which one grows up influences healthcare-seeking behaviour. Examples of these cultural influences are different conceptions of disease and taboos³¹. Another factor affecting the lower use of psychotropic medication by descendants of immigrants may be so-called health illiteracy (i.e., unfamiliarity with the country's healthcare system)³². These cultural and health illiteracy-related aspects may explain the underutilisation of health care by mothers. However, the children in our study grew up and attended school in Sweden. Thus, they had the opportunity to become familiar with the Swedish system. Nevertheless, it is likely that the effects of cultural influences and health illiteracy on mothers' healthcare-seeking behaviours are transmitted to the next generation because it is often mothers who regulate healthcare contacts for children.

Another circumstance behind the observed patterns of use of medication could be the communication between the patient and the health care provider. In this sense, insufficient intercultural competence among the health care staff might affect the interpretation of the patient's symptoms and hence, the prescription of medication.

The process of incorporating the host culture into an individual's behavior, such as for example health care behavior, is known as acculturation³³. From this perspective the number

of years spent in Sweden might condition the degree of acculturation. Indeed, we found a high correlation between time spent in Sweden before delivery and mothers' country of birth, the lowest percentage of mothers with more than 18 years in Sweden being the group of mothers from low-income countries.

Interestingly, our results suggesting a relative underutilisation of psychotropic medication in 18-year-old descendants of immigrants contrast with previous findings that observed a higher rate of psychotropic drug use in first-generation immigrants³⁴. In addition, a study in the Netherlands³⁵ found that in comparison with the native population, first- and descendants of Turkish and Moroccan immigrants had an increased rate of antidepressant and antipsychotic drug prescriptions and a decreased rate of ADHD medication and lithium prescriptions. However, in our earlier investigations in the county of Scania, we observed that overall utilisation of psychiatric healthcare resources in 18 to 80-year-old immigrants was considerably less than expected according to self-reported needs ³².

In the present analysis we only considered prescribed medication. Therefore, the results may be biased by the sale of over-the-counter (OTC) medication. In Sweden, the only available OTC psychotropic drug is Propiomazine. Nevertheless, we believe it is improbable that our results would be affected by the inclusion of Propiomazine in the analyses because this assumption would imply that descendants of immigrants use this OTC drug as a substitute for all other psychotropic medication.

Strength of our study is that we had information from registries covering the entire Swedish population. Furthermore, 98.6 % of all births are registered in the Swedish Medical Birth Registry. Therefore, we do not believe that giving birth at home is an important source of bias in our estimations. It could be possible that some mothers give birth at home because they are illegally residing in Sweden or because the Swedish health care is very different to their

original traditions as is the case for Somali women³⁶. However, we do not have information on the country of origin of the 1.4% of the women giving birth at home in Sweden.

Furthermore, all women in Sweden with a Personal Identification Number (PIN) have the right to attend antenatal care and the majority of them do indeed use this service. However, a minor percentage do not attend all visits that they have the right to, and immigrated women tend to come later in the pregnancy then native Swedish women³⁷. Also, even if information was quite complete overall, data on some variables was more frequently missing in children with an immigrant mother than in children with a native mother which may originate information bias. Besides, since smoking is a self-reported variable, there might be a bias due to linguistic barriers or cultural differences in the attitude towards reporting smoking.

The information recorded in the Medical Birth Register, the National Patient Register and the National Cause of Death Register is mandatorily reported by law, and the quality of the registries is regularly evaluated by the National Board of Health and Welfare and Statistics Sweden.

Measurement of psychotropic drug use using administrative registries reflects both access to healthcare and the presence of psychological disorders. From the perspective of community medicine, the identification of an imbalance between needs and utilisation of psychotropic medication raises questions about equity in access to health care resources. However, our findings also identify the existence of information bias when using a register-based measurement of psychotropic drug use as a proxy for impaired psychological health if descendants of immigrants are included in the analyses.

We performed analyses investigating specific groups of psychotropic medication (results available on request), but the results were similar to those found when analysing the entire medication group. Moreover, because the therapeutic profiles of the studied medication

groups overlap each other (i.e., different psychiatric disorders can be treated with the same drug group), we cannot create a direct link between specific medications and diagnoses.

The suitability of treating young people with psychotropic medication is polemic. Side effects can be severe and, certainly, there are risks when these medications are used in children and adolescents^{38 39}. Also, it is not clear whether adolescents in Sweden and other high-income countries are prescribed these medication in excess (i.e., overutilization)⁴⁰. However, underuse may engender avoidable suffering⁴¹ and therefore it is important to ensure that all people in need are granted access to medication.

If underutilisation of psychotropic medication by adolescent descendants of immigrants occurs, policy makers in the Swedish healthcare system should take this situation into account. Swedish healthcare policy is based on the principle of equity that supports the distribution of healthcare resources on equal terms and according to needs, regardless of any other determinants. This aspect is particularly relevant because one of the main goals stated by the Swedish National Institute of Public Health is achieve 'secure and favourable conditions during childhood and adolescence' and, specifically, 'improved mental health amongst children and young people' (39). Paraphrasing Wittkampf³⁵, we need further research to clarify whether patients of different ethnic backgrounds with the same symptoms receive similar diagnoses and adequate treatment.

COMPETING INTERESTS

None declared

FUNDING

This work was supported by the Swedish Council for Working Life and Social Research (FAS) [Dnr 2010-0402, PI Juan Merlo], the Swedish Research Council (VR) [Dnr K2011-69X-15377-07-6, PI Juan Merlo] and the Centre for Economic Demography at Lund University.

Contributorship Statement

 WvL and JM hade the original idea of the study. JM, WvL and SN participated in the design, analysis, interpretation of data, and drafting of the article. All authors approved the final version to be published. JM is the guarantor of the article.

Data Sharing Statement

The original data used in this study was obtained from the Swedish Board of Health and Welfare and from Statistics Sweden. The authors are not allowed to share the database without permission of those Swedish authorities.

REFERENCES

- 1. Health on equal terms--national goals for public health. *Scand J Public Health* 2001;Suppl 57:1-68.
- 2. Wadman C, Boström G, Karlsson AS. Health On Equal Terms? Results from the 2006 Swedish National Public Health Survey. *STATENS FOLKHÄLSOINSTITUT A 2008:1, Stockholm, Sweden* 2008.
- 3. Practice parameter on the use of psychotropic medication in children and adolescents. *J Am Acad Child Psy* 2009;48(9):961-73.
- 4. Vitiello B. An international perspective on pediatric psychopharmacology. *Int Rev Psychiatry* 2008;20(2):121-6.
- 5. Zito JM, Safer DJ, dosReis S, et al. Trends in the prescribing of psychotropic medications to preschoolers. *JAMA*: the journal of the American Medical Association 2000;283(8):1025-30.
- 6. Zito JM, Safer DJ, DosReis S, et al. Rising prevalence of antidepressants among US youths. *Pediatrics* 2002;109(5):721-7.
- 7. Thomas CP, Conrad P, Casler R, et al. Trends in the use of psychotropic medications among adolescents, 1994 to 2001. *Psychiatr Serv* 2006;57(1):63-9.
- 8. Sondergard L, Kvist K, Andersen PK, et al. Do antidepressants precipitate youth suicide?: a nationwide pharmacoepidemiological study. *Eur Child Adoles Psy* 2006;15(4):232-40.
- 9. Ekblad M, Gissler M, Lehtonen L, et al. Relation of prenatal smoking exposure and use of psychotropic medication up to young adulthood. *American journal of epidemiology* 2011;174(6):681-90.
- 10. Gyllenberg D, Sourander A. Psychotropic drug and polypharmacy use among adolescents and young adults: Findings from the Finnish 1981 Nationwide Birth Cohort Study. *Nord J Psychiat* 2012.
- 11. Lindblad F, Ringback Weitoft G, et al. Maternal and paternal psychopathology increases risk of offspring ADHD equally. *Epidemiol Psychiatr Sci* 2011;20(4):367-72.
- 12. Lindstrom K, Lindblad F, Hjern A. Preterm birth and attention-deficit/hyperactivity disorder in schoolchildren. *Pediatrics* 2011;127(5):858-65.
- 13. Lindblad F, Hjern A. ADHD after fetal exposure to maternal smoking. *Nicotine & tobacco research : official journal of the Society for Research on Nicotine and Tobacco* 2010;12(4):408-15.
- 14. Hjern A, Weitoft GR, Lindblad F. Social adversity predicts ADHD-medication in school children--a national cohort study. *Acta paediatrica* 2010;99(6):920-4.

- 15. Campbell SM, Roland MO. Why do people consult the doctor? *Family practice* 1996;13(1):75-83.
- 16. Norcross WA, Ramirez C, LA P. The influence of women on the health care-seeking behavior of men. *J Fam Pract 1996 Nov;43(5):475-480* 1996;45(5):475-80.
- 17. Hopton J, Hogg R, McKee I. Patients' accounts of calling the doctor out of hours. *BMJ* 1996;313:991-4.
- 18. Fylkesnes K. Determinants of health care utilization--visits and referrals. *Scand J Soc Med* 1993;21(1):40-50.
- 19. Adamson J, Ben-Shlomo Y, Chaturvedi N, et al. Ethnicity, socio-economic position and gender--do they affect reported health-care seeking behaviour? *Soc Sci Med* 2003;57(5):895-904.
- 20. Lay B, Nordt C, Roessler W. Mental hospital admission rates of immigrants in Switzerland. *Social Psychiatry and Psychiatric Epidemiology* 2007;42(3):229-36.
- 21. Arbogast PG, Ray WA. Use of disease risk scores in pharmacoepidemiologic studies. *Statistical Methods in Medical Research* 2009;18(1):67-80.
- 22. Ben-Shlomo Y, Kuh D. A life course approach to chronic disease epidemiology: conceptual models, empirical challenges and interdisciplinary perspectives. *Int J Epidemiol* 2002;31(2):285-93.
- 23. WHO. ATC -Structure and principles: WHO Collaborating Centre for Drug Statistics Methodology, 2011.
- 24. EpC. The Swedish Medical Birth Register -A summary of content and quality, 2003.
- 25. Cafferata GL, Meyers SM. Pathways to Psychotropic-Drugs Understanding the Basis of Gender Differences. *Medical Care* 1990;28(4):285-300.
- 26. Benyamini Y, Blumstein T, Boyko V, et al. Cultural and educational disparities in the use of primary and preventive health care services among midlife women in Israel. *Womens Health Issues* 2008;18(4):257-66.
- 27. Ayonrinde O. Importance of cultural sensitivity in therapeutic transactions Considerations for healthcare providers. *Disease Management & Health Outcomes* 2003;11(4):233-48.
- 28. Ben-Shlomo Y, Kuh D. A life course approach to chronic disease epidemiology: conceptual models, empirical challenges and interdisciplinary perspectives. *International journal of epidemiology* 2002;31(2):285-93.
- 29. Lanius RA, Vermetten E, Pain C. *The impact of early life trauma on health and disease : the hidden epidemic.* Cambridge, UK; New York: Cambridge University Press, 2010.
- 30. Socialdepartementet. Uppdaterade högkostnadsskydd öppen hälso- och sjukvård samt läkemedel. Ds 2011:23. *Departementsserien (Ds)* 27 juni 2011.
- 31. Lindert J, Schouler-Ocak A, Heinz A, Priebe S. Mental health, health care utilisation of migrants in Europe. *European Psychiatry* 2008;23:S14-S20.
- 32. Merlo J. Fördelning av psykisk ohälsa och sjukvårdsresurser i Malmö, region Skåne. In: Östman M, editor. *Migration och psykisk ohälsa*. Malmö: Malmö högskola; Hälsa och samhalle, 2008:45-94.
- 33. Thomson MD, Hoffman-Goetz L. Defining and measuring acculturation: a systematic review of public health studies with Hispanic populations in the United States. *Soc Sci Med* 2009;69(7):983-91.
- 34. Hjern A. High use of sedatives and hypnotics in ethnic minorities in Sweden. *Ethn.Health* 2001;6(1):5-11.
- 35. Wittkampf LC, Smeets HM, Knol MJ, et al. Differences in psychotropic drug prescriptions among ethnic groups in the Netherlands. *Social psychiatry and psychiatric epidemiology* 2010;45(8):819-26.

36. Essen B, Johnsdotter S, Hovelius B, et al. Qualitative study of pregnancy and childbirth experiences in Somalian women resident in Sweden. *BJOG* 2000;107(12):1507-12.

- 37. Ny P, Dykes AK, Molin J, et al. Utilisation of antenatal care by country of birth in a multi-ethnic population: a four-year community-based study in Malmo, Sweden. *Acta obstetricia et gynecologica Scandinavica* 2007;86(7):805-13.
- 38. Zakriski Aea. Justifiable psychopharmacology or overzealous prescription? Examining parental reports of lifetime prescription histories of psychiatrically hospitalised children. *Child and Adolescent Mental Health* 2005;10(1):7.
- 39. Jureidini JN, Doecke CJ, Mansfield PR, et al. Efficacy and safety of antidepressants for children and adolescents. *BMJ* 2004;328(7444):879-83.
- 40. Victorri-Vigneau C, Basset G, Jolliet P. How a novel programme for increasing awareness of health professionals resulted in a 14% decrease in patients using excessive doses of psychotropic drugs in western France. *European Journal of Clinical Pharmacology* 2006;62(4):311-16.
- 41. Gentile S. Efficacy of antidepressant medications in children and adolescents with obsessive-compulsive disorder: a systematic appraisal. *J Clin Psychopharmacol* 2011;31(5):625-32.

LEGEND TO THE FIGURE

Figure 1. Flow diagram indicating the number of individuals remaining in the study sample after the application of the inclusion criteria

