Time trends in treated incidence, sociodemographic risk factors and comorbidities: a Finnish nationwide study on anxiety disorders

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Abstract

Background: There has been a lack of research about the time trends and socio-demographic risk factors for children and adolescents who receive treatment for anxiety disorders. This study aimed to fill these gaps in our knowledge by examining a nationwide sample of Finnish children and adolescents diagnosed in specialized healthcare settings.

Methods: This study comprised national register data of all singleton children born in Finland from 1992–2006 who were diagnosed with anxiety disorders from 1998–2012. The changes in time trends in incidence were studied by dividing the study sample into three cohorts by birth years: 1992–1996, 1997–2001 and 2002–2006, who were followed up until the age of 20, 15 and 10 years, respectively. The 22,388 individuals with anxiety disorders were age and gender matched with 76,139 controls from the general population. Logistic regression was used to examine the socio-demographic risk factors and anxiety disorders in the entire sample. Comorbid disorders were examined in the oldest birth cohort (1992–1996 born).

Results: Comparing the 1992–1996 and 2002–2006 cohorts showed that the cumulative incidence of treated anxiety disorders at the age of 10 increased from 0.3 to 1.2% among females and 0.46 to 1.9% among males. Subjects had higher likelihood for being diagnosed with an anxiety disorder if their mothers had low maternal socio-economic status class at birth (OR 1.53, 95% CI 1.45–1.61) compared to higher SES class, and marital status was single at the time of birth (OR 2.02, 95% CI 1.87–2.17) compared to married or in a relationship. They had lower risk of anxiety disorders diagnosis if born in rural (OR 0.82, 95% CI 0.79–0.86) or semi-urban areas (OR 0.79, 95% CI 0.76–0.82) when compared to urban residence. There was a wide range of psychiatric comorbidities, and unipolar depression was the most common (31.2%).

Conclusion: Anxiety disorders diagnosed by specialized Finnish services increased from 1998–2012 in both genders. This could indicate a real increase in overall anxiety disorders or an increase in treatment seeking. The findings on maternal socioeconomic status and single parenting improve the recognition of the environmental risk factors for anxiety disorders among children and adolescents.
Background
Anxiety disorders are very common in childhood and adolescence and global estimates suggest they affect 6.5% of individuals under 19 years of age [1]. The prevalence varies for different anxiety disorders and so does the usual age at onset. Child onset disorders usually include separation anxiety, specific phobias and selective mutism, whereas social phobia, agoraphobia, panic disorder and generalized anxiety disorder are more likely to start in adolescence [2]. Anxiety disorders cause significant functional impairment at home, at school and with friends [3]. They can manifest as somatic symptoms [3, 4] and increase the risk of several other psychiatric disorders, which lead to further deterioration in the prognosis [5, 6]. When they are left untreated, anxiety disorders tend to become chronic [7] and may even lead to disabilities [8]. Even with their widespread prevalence, the treatment rates for anxiety disorders are low [9]. These issues highlight how important it is to recognise the risk factors for anxiety disorders, so that children and adolescents can be diagnosed and treated as early as possible.

Previous time trend studies for anxiety disorders have showed a phenomenon of increased health care visits [10–14] especially by girls [10, 11, 14]. However, these studies had limitations with regard to sample sizes, definition of diagnoses and study periods. In addition, a number only compared two cohorts and this introduced the risk that any changes could have been down to natural fluctuations, rather than specific factors. Time trend studies provide information for health care planning and identify which factors need to be further investigated when changes are observed. These can include different cultural and lifestyle aspects and sociodemographic structures.

The aetiology of anxiety disorders comprises both genetic and environmental risk factors [2, 15]. The existing studies on environmental risk factors, such as sociodemographic risk factors, are mostly cross-sectional or small cohort studies, and only few studies have followed a prospective design. There have been a few larger register-based studies that have focused on socio-demographic factors, such as parental unemployment [16], welfare benefits [17], low maternal socio-economic status (SES) [18], family breakdowns and single parents [16–18]. These factors have all been associated with a higher risk for anxiety disorders. In addition, two Danish registry studies found that the risk for diagnosed anxiety disorders was higher among those living in urban areas [16, 19]. However, all the previous studies have had either very narrow inclusion criteria for anxiety disorders [18, 19] or a wide selection of diagnoses, which included obsessive compulsive disorder and post-traumatic stress disorder [16, 17]. These are not considered anxiety disorders in the Diagnostic and Statistical Manual of Mental Disorders, 5th Edition, as their aetiology is different [20]. Moreover, all these studies had relatively small sample sizes compared to the present study.

Finland has well-established national registers, which contain valuable psychiatric epidemiological data [21–23]. The aim of our study was to provide a comprehensive description of the anxiety disorders observed among a large sample of children and adolescents being cared for by specialized Finnish mental health services. We used three birth cohorts to study the treated incidence, cumulative incidence and time trends and examined the socio-demographic factors and psychiatric comorbidities associated with diagnosed anxiety disorders.

Materials and methods
This nationwide register-based case-control study is a part of Finnish Prenatal Study of Anxiety Disorders. It included information on all singleton children born in Finland between 1 January 1992 and 31 December 2006 and followed up until 31 December 2016. The cases included individuals who were diagnosed with an anxiety disorder between 1 January 1998 and 31 December 2012. The study used a number of Finnish nationwide registers that provided comprehensive information on various datasets.

National registers
The Finnish registers used in this study were the Care Register for Health Care recognized also as Finnish Hospital Discharge Register, the Central Population Register and the Finnish Medical Birth Register. Detailed descriptions of these registers have previously been published [22, 23] and are briefly explained here.

The Care Register includes information on all visits to specialized health services in Finland. From 1969 to 1998, this database contained information from all inpatient wards in somatic and psychiatric hospitals, local health care centres, private hospitals, prison hospitals and military wards. Since 1998, it has also included outpatient diagnoses. This study used the Care Register to identify cases diagnosed with anxiety disorders by specialized mental health services and to collect any comorbid diagnoses that were recorded during the study period. During our study period the diagnoses were coded using the
International Classification of Diseases, Tenth Revision (ICD-10). The Care Register is a continuation of the Discharge Register, and its inclusiveness and accuracy have been assessed to be satisfactory to very good [24].

The Population Register provides information on all Finnish citizens and permanent residents and it is maintained by the Population Register Centre and local register offices. It includes information such as the individual’s name, address, data of birth and death, family members and any immigration or emigration status. In this study the Population Register was used to collect information on two risk factors for the cases and matched controls, namely the place of birth and whether someone lived in an urban or non-urban area.

The Birth Register was established in 1987 and it contains standardized data on the prenatal and perinatal periods of infants who are born alive and stillborn. In this study the Register was used to collect information on the mother’s socio-economic and marital status at the time of her child’s birth. The database has previously been described [25].

The linkage of information between these registers was possible using the personal identity code issued for each Finnish resident at birth or when they move to the country. The approval for the utilization and linkage of the register data was provided by the Data Protection Ombudsman and National Institute for Health and Welfare, Finland, and the ethical approval was obtained from the Ethics Committee of the Hospital District of Southwest Finland. This study is solely based on register data hence obtaining informed consent was not required by the ethics committee. This study was done in accordance to the research guidelines and regulations of National Institute for Health and Welfare, Finland.

Study participants
Cases and controls
This study comprises of a nested case-control sample. It used national register data of all singleton children born in Finland in 1992–2006. There were 22,388 cases identified from the Care Register, who were diagnosed with anxiety disorders from 1998 to 2012. The sample was further divided into three birth cohorts, 1992–1996 (13,806 cases), 1997–2001 (6453 cases) and 2002–2006 born individuals (2129 cases). These individuals were followed up until the age of 20, 15 and 10 years, respectively.

Anxiety disorders were defined using the ICD-10 classifications of: agoraphobia (F40.0, F40.00, F40.01), social phobia (F40.1, F93.2), specific phobias (F40.2, F40.8, F40.9, F93.1), panic disorder (F41.0, F41.00, F41.01, F41.08, F41.09), generalized anxiety disorder (F41.1, F93.80), separation anxiety (F93.0), selective mutism (F94.0) and other, nonspecific anxiety disorders (F41.2, F41.3, F41.8, F41.9, F93.3, F93.89, F93.9). This study includes cases that were diagnosed with any anxiety disorders at least once when they were at least six years old. We excluded obsessive compulsive disorder and stress-related disorders and cases diagnosed with moderate or severe intellectual disability (ICD-10: F72 and F73).

The cases were matched with controls from the Population Register by their date of birth (± 30 days), and gender. The controls were still living in Finland when their matched cases were first diagnosed with an anxiety disorder. The controls had not been diagnosed with any anxiety disorder or moderate to severe intellectual disability (ICD-10 F72, F73) during the study period. A total of 76,139 controls were included in the analyses.

Demographic variables
Maternal SES and marital status at birth were obtained from the Birth Register. Maternal SES was divided into the five maternal occupation categories used by Gissler et al. [26]. These were upper white collar (including experts and managers), lower white collar (including clerical workers), blue collar (including manual workers), others (including entrepreneurs, students, people who were unemployed and stay-at-home parents) and missing data. This classification is also the national Finnish classification for occupations and socio-economic groups. The region of birth and urbanicity were obtained from the Population Register and defined as Southern, Eastern, Northern and Western Finland. These areas were further broken down by urban, semi-urban or rural, based on the definitions from Statistics Finland. Finland is one of the mostly sparsely populated countries in Europe. According to Statistics Finland, the average population density is 18 inhabitants per square kilometres, varying from 0.2 in some areas in the north to almost 3000 in the capital city. In semi-urban areas the inhabitant rate varies from 6 to 62 [27].

Comorbidities
The comorbid psychiatric diagnoses were retrieved from the Care Register for the oldest cohort born in 1992–1996 in order to have longer follow-up period. Diagnoses at 6–16 years of age were included to make the sample uniform. The comorbidities were defined using the ICD-10 classification of mental disorders (F10-F99), excluding anxiety disorders (Table 1).

Statistical analyses
We studied the time trends of the treated incidence and cumulative incidence of diagnosed anxiety disorders, by birth year: 1992–1996, 1997–2001 and 2002–2006. First, we analysed the changes in incidence rates among diagnosed anxiety disorder cases. The population at risk were
children living in Finland at the end of 2012 and were obtained from Population Register. The yearly incidence of diagnosed anxiety disorders per 100 people at risk was calculated by gender. Second, the yearly cumulative incidence was calculated with odds ratios (OR) and 95% confidence intervals (95% CI). The cumulative incidence was one minus the estimated survival function, which was calculated using the Kaplan-Meier estimate. Third, both the yearly incidence and cumulative incidence were calculated separately for the three birth cohorts, at 20, 15 and 10 years of age, respectively. The statistical difference between the treated incidence rates for males and females was calculated using the log-rank test. In addition, we examined if there were any cases that received anxiety disorder diagnoses at the age before six and was not followed up after age six (n = 88).

We used nested case-control design to examine the possible association between sociodemographic factors and anxiety disorders in the entire sample. Conditional logistic regression was used to examine the unadjusted associations in the total sample and by gender. The ORs and 95% CIs were calculated for each variable and the outcome was an anxiety disorder diagnosis. Then adjustments were made for significant covariates, p-value <0.05 was considered to indicate statistical significance. We examined the randomness of the missing values among the risk factors and used multiple imputation method as missing values were non-random. Twenty imputed data sets were produced and logistic regression model was fitted with each dataset. These results were pooled to obtain the final results. The frequencies for the anxiety disorder subgroups in the total population and comorbid diagnoses in the oldest cohort, born in 1992–1996, were calculated. Chi-square test was done to see the differences in the prevalence of comorbid diagnoses between genders. The statistical analyses including multiple imputation were performed using SAS statistical software, version 9.4. (SAS Institute Inc., Cary, NC, USA).

Results

Characteristics of the study population

The characteristics of the study population is presented in Table 2. The total sample comprised of 98,527 children and adolescents. Of these, 22,388 were cases diagnosed with anxiety disorders, of which 56.4% were female. The mean age at diagnosis was 11.3 years (standard deviation (SD) 2.9, age range 6–20 years). The mean age was 11.6 years (SD 3.1) for females and 10.9 years (SD 2.7) for males. For the three birth cohorts, 1992–1996, 1997–2001 and 2002–2006, the mean ages for the females and males were 15.5 and 14.4 (SD 2.5, 3.3, range 6–20 years), 11.6 and 10.5 (SD 2.5, 2.2, range 6–15 years) and 7.7 and 7.8 (SD 1.3, 1.2, range 6–10 years), respectively.

Yearly incidence of diagnosed anxiety disorders by age and gender

The age-specific changes in diagnosed anxiety disorders were examined by analyzing the yearly incidence by both age and gender (Fig. 1). The incidence showed a notable increase among females from 12.5–17 years of age, whereas the increase in incidence was smaller and practically stable in males from 6–20 years of age. Males had more anxiety disorders at 6–12 years of age than females (p < 0.001) and for females it was from 13–20 years of age than males (p < 0.001).

Cumulative incidence of diagnosed anxiety disorders by birth cohort and gender

Figures 2 and 3 show the cumulative incidences of diagnosed anxiety disorders per 100 live births by birth cohort and gender. The first birth cohort was followed

Table 1 ICD-10 diagnostic codes for comorbid psychiatric diagnosis

| Subgroup                                | ICD-10 Code(s)               |
|-----------------------------------------|------------------------------|
| Intellectual disability                 | F70–F77                      |
| Developmental delay and disorders of childhood | F78–F81                  |
| Speech disorders                        | F82                          |
| Learning disabilities                   | F83                          |
| Febrile convulsions                     | F84                          |
| Attention deficit disorders             | F85                          |
| Sensory integration disorder            | F86                          |
| Speech and language disorders           | F87                          |
| Communication disorders                 | F88                          |
| Social communication deficits           | F89                          |
| Social anxiety                          | F90                          |
| Conduct and oppositional disorders     | F91, F92                     |
| Conduct disorder                        | F93                          |
| Attention deficit disorder              | F94.1, F94.2                 |
| Tic disorders                           | F95                          |
| Sleep disorders                         | F51                          |
| Sleep-wake disorders                    | F51.3, F51.4                 |
| Eating disorders                        | F50                          |
| Personality disorders                   | F60–F63                      |
| Impulse disorders                       | F63                          |
| Autism spectrum disorders               | F84                          |
| Schizophrenia spectrum disorders        | F20–F25, F28, F29            |
| Bipolar disorders                       | F30, F31                     |
| Unipolar depressive disorders           | F32, F34                     |
| Phobic disorder                         | F42                          |
| Obsessive-compulsive disorders          | F43                          |
| Anxiety disorders                       | F45                          |

Table 2 Gender distribution and mean age of anxiety disorder diagnosis in the whole sample and in the three birth cohorts

| Cohorts             | Total n of cases | Female n (%) | Male n (%) |
|---------------------|------------------|--------------|------------|
|                     | Mean age at first diagnosis | Mean age (SD) | Mean age (SD) |
| Entire sample 1992–2006 (6–20 years) | 22,388 | 12,628 (56.4%) | 9760 (43.6%) |
| Birth cohort 1992–1996 (6–20 years) | 13,806 | 8793 (63.7%) | 5013 (36.3%) |
| Birth cohort 1997–2001 (6–15 years) | 6453 | 3043 (47.2%) | 3410 (52.8%) |
| Birth cohort 2002–2006 (6–10 years) | 2129 | 792 (37.2%) | 1337 (62.8%) |
up until the age of 20 years and the other cohorts were followed until 15 and 10 years of age, respectively. Individuals born in 1992–1996, and diagnosed with anxiety disorders by the age of 20, had cumulative incidences of 5.7% (95% CI 5.60–5.82) by the age of 20 and 0.4% (95% CI 0.36–0.41) by the age of 10. The incidence had increased in the 1997–2001 and 2002–2006 groups by the age of 10, with cumulative incidences of 0.9% (95% CI 0.91–0.99) and 1.5% (95% CI 1.46–1.62), respectively. Similar trends were noticed in both genders. The cumulative incidence at the age of 10 increased when the first and last, 1992–1996 and 2002–2006, cohorts were compared. They increased from 0.3% (95% CI 0.28–0.34) to 1.18% (95% CI 1.09–1.29) among females and 0.46%, (95% CI 0.43–0.49) to 1.9% (95% CI 1.76–2.00) among males.

Demographic factors and the risk of anxiety disorders
Table 3 shows the sociodemographic characteristics of the study sample. Of the 98,527 individuals, 5348 were missing data on maternal SES, 431 on urbanicity and 8151 on marital status. When the upper white-collar SES class was used as the reference category, children born to the mothers of blue collar (OR 1.53, 95% CI 1.45–1.61) and lower white-collar (OR 1.18, 95% CI 1.12–1.24) SES classes were more likely to get diagnosed with anxiety disorders. These results were obtained after adjusting for the other sociodemographic covariates, gender and time of birth; p < 0.0001. Another significant finding observed in the adjusted analysis was that children born to single mothers had higher likelihood of anxiety disorder diagnoses (OR 2.02, 95% CI 1.87–2.17) p < 0.0001, when compared to the reference category of children born to mothers who were married or in a relationship. The likelihood of being diagnosed with an anxiety disorder was lower in Western Finland, (OR 0.68, 95% CI 0.66–0.71), Northern Finland (OR 0.59, 95% CI 0.56–0.62) and Eastern Finland (OR 0.67, 95% CI 0.63–0.71); (p < 0.0001 for each) when Southern Finland was used as the reference category. The odds of being diagnosed with an anxiety disorder was also lower in rural, (OR 0.82, 95% CI 0.79–0.86) and semi-urban, (OR 0.79, 95% CI 0.76–0.82) (both p < 0.0001) when urban residence was used as the reference category.

![Fig. 1 Yearly incidence of diagnosed anxiety disorders by age and gender per 100 people](image-url)
Gender distribution of anxiety disorder subgroups among total cases

Most of the diagnoses were described as unspecified anxiety disorders, with separate data for females (76.6%) and males (74.1%). Other commonly diagnosed anxiety disorders were specific phobias (female 10.0%, male 13.6%), panic disorders (female 9.8%, male 5.1%), generalized anxiety disorders (female 9.1%, male 10.2%), social phobias (female 9.1%, male 8.1%) and agoraphobia (female 1.6%, male 1.2%).

Gender differences in comorbid diagnosis in the 1992–1996 birth cohort

We restricted the distribution and comorbidity analysis to the oldest subgroup, which were born between 1992 and 1996 and this showed that 13,806 individuals had been diagnosed with anxiety disorders (Fig. 4). The most common comorbidities among females were unipolar depressive disorders (35.9%), stress related and dissociative disorders (11.3%), conduct and oppositional disorders (9.1%), eating disorders (8.9%) and learning and coordination disorders (7.5%). Among males they were unipolar depressive disorders (22.8%), conduct and oppositional disorders (18.9%), learning and coordination disorders (18.9%) and ADHD (13.1%). Significant gender differences were observed in several comorbidities, ADHD (female 2.4%, male 13.1%), ASD (female 2.3%, male 8.1%), tic disorders (female 0.4%, male 2.3%), learning and coordination disorders (female 7.5%, male 18.9%), and eating disorders (female 8.9%, male 1.6%); p < 0.0001.

Discussion

This nationwide birth cohort study of 22,388 children and adolescents, and matched controls, provides comprehensive data on the incidence, comorbidities and socio-demographic risk factors associated with diagnoses of anxiety disorders.

There were three key findings. First, the treated incidence of anxiety disorders increased with age, with rapid increases in females from 12.5 years, but more stable increases in males. Our data confirmed previous research that males had more anxiety disorders diagnosed in childhood, but the incidence was higher in females from adolescence onwards [28, 29]. Previous studies have discussed whether girls were more vulnerable to internalizing disorders in adolescence because of psychological, environmental, genetic, or biological factors, such as puberty, hormones and immune system regulation.
### Table 3  Demographic factors and risk of anxiety disorders

|                                  | Cases          | Controls       | Unadjusted       | Adjusteda                  |
|----------------------------------|----------------|----------------|------------------|---------------------------|
|                                  | N   | %   | N   | %   | OR  | 95% CI | p-value | OR  | 95% CI | p-value |
| **Maternal SES Class**           |     |     |     |     |     |       |         |     |       |         |
| Upper white-collar workers       | 2707 | 12.1 | 11,660 | 16.1 | Ref |       |         |     |       |         |
| Lower white-collar workers       | 8972 | 42.8 | 34,040 | 47.1 | 1.13 | 1.07–1.18 | <0.0001 | 1.18 | 1.12–1.24 | <0.0001 |
| Blue collar workers              | 4722 | 22.6 | 14,009 | 19.4 | 1.44 | 1.37–1.52 | <0.0001 | 1.53 | 1.45–1.61 | <0.0001 |
| Other                            | 4541 | 21.7 | 12,528 | 17.3 | 1.55 | 1.47–1.64 | <0.0001 | 1.65 | 1.56–1.74 | <0.0001 |
| **Region of birth**              |     |     |     |     |     |       |         |     |       |         |
| Southern Finland                 | 11,331 | 50.6 | 30,192 | 39.7 | Ref |       |         |     |       |         |
| Western Finland                  | 6743 | 30.1 | 27,006 | 35.5 | 0.66 | 0.64–0.69 | <0.0001 | 0.68 | 0.66–0.71 | <0.0001 |
| Northern Finland                 | 2322 | 10.4 | 10,788 | 14.2 | 0.57 | 0.55–0.60 | <0.0001 | 0.59 | 0.56–0.62 | <0.0001 |
| Eastern Finland                  | 1992 | 8.9  | 8153  | 10.7 | 0.65 | 0.61–0.68 | <0.0001 | 0.67 | 0.63–0.71 | <0.0001 |
| **Urbanicity**                   |     |     |     |     |     |       |         |     |       |         |
| Urban                            | 15,161 | 68.0 | 46,199 | 60.9 | Ref |       |         |     |       |         |
| Semi-urban                       | 3249 | 14.6 | 13,175 | 17.4 | 0.75 | 0.72–0.78 | <0.0001 | 0.79 | 0.76–0.82 | <0.0001 |
| Rural                            | 3881 | 17.4 | 16,431 | 21.7 | 0.72 | 0.69–0.75 | <0.0001 | 0.82 | 0.79–0.86 | <0.0001 |
| **Maternal marital status**      |     |     |     |     |     |       |         |     |       |         |
| Married/ in a relationship       | 18,681 | 93.8 | 68,457 | 97.1 | Ref |       |         |     |       |         |
| Single                           | 1224 | 6.1  | 2014  | 2.9  | 2.28 | 2.11–2.45 | <0.0001 | 2.02 | 1.87–2.17 | <0.0001 |

* The multivariate model included maternal SES class, region of birth, urbanicity, maternal marital status, gender and birth year

Ref Reference category, OR Odds ratio, CI Confidence interval
Girls appear to perceive more stress, especially from school environments and self-imposed demands [32, 33]. The timing of puberty [34] and body image concerns [35] might predispose girls to anxiety symptoms. Genders show different emotional responses to stress [36] and in how they identify and react to symptoms [37, 38], how openly they talk about their feelings [39] and how they seek help [38, 40]. This may be why girls are more likely to be treated for anxiety disorders. Another possibility is that the assessment methods tend to favour the symptoms experienced by girls [30], or that early school-based interventions for externalizing behaviour have improved more than those for internalizing symptoms, which are more common among girls [31], and hence girls could be directed more likely into specialized services. In our sample the girls’ incidence rate decreased after the age of 17. This could be due to the early occurrence of anxiety disorders, special vulnerability of girls in puberty or unspecific anxiety being part of the human development during teenage, especially among girls [2, 29, 30]. Additionally, the decrease in the incidence rates at older ages could be explained by the Finnish health care system; the treatment of anxiety disorders in adults is more concentrated in the primary health care.

We found that the cumulative treated incidence rates had increased with time among both genders, in line with previous studies [11, 13, 14]. Previous time trend studies have suggested that this increase is possibly due to increase in health care visits especially by girls [10–14]. The increased resources in mental health services in Finland have helped expanding need-based psychosocial support, treatment and rehabilitation, raising mental health awareness and change in attitude towards psychiatry in general population [11]. It is also possible that the incidence rates in our sample have increased due to prior diagnosis and the rates could be comparable at older ages. A study from UK found the increases in the incidence rates in primary care to be highest among those aged 10–16 years, not among 6 to 9-year-olds [14]. In Finland, emotional symptoms increased in female adolescents from 1988 to 2014 [41] and they fluctuated in eight-year-old children from 1989 to 2013, but with no significant increases [42]. However, service use increased in these community-based samples [41–43], especially among children with significant problems [43, 44].

Secondly, we assessed the associations between socio-demographic factors and anxiety disorders in a nested case-control setting. The risk factors appeared to be equal for both genders which includes low maternal socio-economic status, single mothers, urban living and being born in Southern Finland. Previous reports also found that low maternal SES increased the odds of anxiety disorders being diagnosed in children when the highest SES class was used as the reference [17, 45, 46]. In general, low SES has been linked with various somatic, cognitive and socioemotional problems in children [47,
learning the importance of assessing comorbidities when treating anxiety disorders. It is also important to avoid unintended over diagnosis [69], as this can have a significant impact on the severity and duration of symptoms [6].

Our study had some limitations that needs to be considered when interpreting the findings. First, the anxiety disorders subjects from six years of age onwards were identified from Finnish Hospital Discharge Register diagnosed by specialized mental health services. Individuals with less severe symptoms may not receive a diagnosis, end up in specialized services or not even seek for treatment. Additionally, parental factors contribute in the help-seeking of minors. Hence, information on these children and adolescents is not available in register. Our rate estimates should be interpreted as treated incidence rates and cannot be regarded as population-based incidence rates. Second, the register data did not enable addressing the differences in the incidence rates between the genders. Third, this study analyzed data on children, adolescents, and young adults and hence the results cannot be interpreted beyond the follow-up age. Since the socio-demographic data were recorded at the time of birth, we were unaware of any changes later in life. Finally, most of the anxiety disorder diagnoses were unspecified, therefore we could not study risk factors and comorbidities associated with anxiety disorder subgroups at this stage.

Conclusion
We believe that this study covered the largest sample of children and adolescents treated by specialized mental health services for anxiety disorders to date. It remains unknown whether the increased treatment rates found in our study reflect a real increase in overall anxiety, merely increased treatment seeking or treatment seeking at earlier ages. Future studies are needed to improve our understanding of the specific reasons behind these increases. The demand for specialized mental health services for children and adolescents is increasing, which also indicates that the currently available primary care services might not be sufficient. The findings on low maternal SES and single parenting increasing the risk for anxiety disorders emphasizes the impact of environmental factors.

Abbreviations
ADHD: Attention Deficit Hyperactivity Disorder; ASD: Autism Spectrum Disorder; CI: Confidence Interval; CRHC: Care Register for Health Care; DSM-5: Diagnostic and Statistical Manual of Mental Disorder, 5th Edition; ICD-9: International classification of Disease, Ninth revision; ICD-10: International
classification of Disease, Tenth revision; OCD: Obsessive compulsive disorder; OR: Odds ratio; SD: Standard deviation; SES: Socioeconomic status.

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Authors’ contributions
PK and TS shared the authorship by designing the research plan, statistical plan as well as sharing the work in writing the manuscript. TL, DG, KK contributed with the manuscript planning, interpretation of results and revision. AS1 mentored the process as senior researcher and principle investigator. AS2 contributed with the statistical analysis. All the authors read and approved the final manuscript.

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Availability of data and materials
The data that support the findings of this study are available from Finnish Social and Health Data Permit Authority- FINDATA, but restrictions apply to the availability of these data, which were used under license for the current study, and so are not publicly available. Data are however available from the authors upon reasonable request and with permission of FINDATA.

Declarations

Ethics approval and consent to participate
The approval for the utilization and linkage of the register data was provided by the Data Protection Ombudsman and National Institute for Health and Welfare, Finland. The ethical approval was obtained from the Ethics Committee, Hospital District of Southwest Finland. This study is solely based on register data hence obtaining informed consent was not required by the Ethics committee, Hospital District of Southwest Finland.

Consent for publication
Not applicable.

Competing interests
The authors declare that they have no competing interest.

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References
1. Polanczyk GV, Salum GA, Sugaya LS, Caye A, Rohde LA. Annual research review: A meta-analysis of the worldwide prevalence of mental disorders in children and adolescents. J Child Psychol Psychiatry Allied Discip. 2015;56(3):345–65. https://doi.org/10.1111/jcpp.12391.
2. Beesdo-Baum K, Knappe S. Developmental Epidemiology of Anxiety Disorders. Child Adolesc Psychiatr Clin N Am. 2012;21(3):457–78. https://doi.org/10.1016/j.chc.2012.05.001.
3. Essau CA, Conrod J, Petermann F. Frequency, comorbidity, and psychosocial impairment of anxiety disorders in German adolescents. J Anxiety Disorder. 2000;14(3):263–79. https://doi.org/10.1016/s0887-6185(99)00239-0.
4. Shannon RA, Berggren MD, Matthews A. Frequent Visitors: Somatization in School-Age Children and Implications for School Nurses. J Sch Nurs. 2010;26(3):169–82. https://doi.org/10.1177/1059840509356777.
5. Kaufman J, Chaney D. Comorbidity of mood and anxiety disorders. Depress Anxiety. 2000;12 Suppl 1:69-76. https://doi.org/10.1002/1520-6394(2000)12:1+<69:AID-DA9>3.0.CO;2-K.
6. Lamers F, van Opppen P, Corjiss HC, Smit JH, Spinheoven P, van Balkom AJLM, et al. Comorbidity patterns of anxiety and depressive disorders in a large cohort study: the Netherlands Study of Depression and Anxiety (NEDSA). J Clin Psychiatry. 2011;72(3):341–8. https://doi.org/10.4088/JCP.10m0176blu.
7. Pine DS, Cohen P, Gurlay D, Brook J, Ma Y. The risk for early-adulthood anxiety and depressive disorders in adolescents with anxiety and depressive disorders. Arch Gen Psychiatry. 1998;55(1):36–64. https://doi.org/10.1001/archpsyc.55.1.56.
8. Ahola K, Vrtanen M, Honkonen T, Isometsa E, Aromaa A, Lonnqvist J. Common mental disorders and subsequent work disability: A population-based Health 2000 Study. J Affect Disord. 2011;134. https://doi.org/10.1016/j.jad.2011.05.028.
9. Alonso J, Liu Z, Evans-Lacklo S, Sadikova E, Sampson N, Chatterji S, et al. WHO World Mental Health Survey Collaborators. Treatment gap for anxiety disorders is global: Results of the World Mental Health Surveys in 21 countries. Depress Anxiety. 2018;35(3):195–208. https://doi.org/10.1002/da.22711.
10. Ask H, Handal M, Hauge LJ, Reichborn-Kjennerud T, Skurtvet S. Incidence of diagnosed pediatric anxiety disorders and use of prescription drugs: a nationwide registry study. Eur Child Adolesc Psychiatry. 2020;29(8):1063–73. https://doi.org/10.1007/s00127-019-01419-0.
11. Gyllenberg D, Marttila M, Sund R, Jokiranta-Olkoniemi E, Sourander A, Gissler M, et al. Temporal changes in the incidence of treated psychiatric and neurodevelopmental disorders during adolescence: an analysis of two national Finnish birth cohorts. Lancet Psychiatry. 2018;5(3):227–36. https://doi.org/10.1016/S2215-0066(18)30038-5.
12. John A, Marchant AL, McGregor JJ, Tan JOA, Hutchings HA, Kovess V, et al. Recent trends in the incidence of anxiety and prescription of antidepressants and hypnotics in children and young people: An e-cohort study. J Affect Disord. 2015;183:134–41. https://doi.org/10.1016/j.jad.2015.05.002.
13. Olsson M, Blanco C, Wang S, Laje G, Correll CU. National trends in the mental health care of children, adolescents, and adults by office-based physicians. JAMA Psychiatry. 2014;71(1):81–90. https://doi.org/10.1001/jamapsychiatry.2013.3074.
14. Cybulski L, Ashcroft DM, Carr MJ, et al. Temporal trends in annual incidence rates for psychiatric disorders and self-harm among children and adolescents in the UK, 2003–2018. BMC Psychiatry. 2021;21:229. https://doi.org/10.1186/s12888-021-03235-w.
15. John A, Marchant AL, McGregor JJ, Tan JOA, Hutchings HA, Kovess V, et al. Recent trends in the incidence of anxiety and prescription of antidepressants and hypnotics in children and young people: An e-cohort study. J Affect Disord. 2015;183:134–41. https://doi.org/10.1016/j.jad.2015.05.002.
16. Hyland P, Shevlin M, Eikliir A, Christoffersen M, Murphy J. Social, familial and psychological risk factors for mood and anxiety disorders in childhood and early adulthood: a birth cohort study using the Danish Registry System. Soc Psychiatry Psychiatr Epidemiol. 2016;51(3):331–8. https://doi.org/10.1007/s00127-016-1171-1.
17. Guhl M, Emerson SD, Mahdavi D, Gadamermann AM. Associations of Birth Factors and Socio-Economic Status with Indicators of Early Emotionality and Risk of Anxious Attachment Behavior. J Affect Disord. 2020;267:105061. https://doi.org/10.1016/j.jad.2020.105061.
18. Koskelo M, Chudal R, Luntamo T, Suominen A, Steinhausen H-C, Sourander A. The impact of parental psychopathology and sociodemographic factors in selective mutism - a nationwide population-based study. BMC Psychiatry. 2020;20(1):221. https://doi.org/10.1186/s12888-020-02637-6.
19. Helenius D, Munk-Jørgensen P, Steinhausen HC. Family load estimates and risk factors of anxiety disorders in a nationwide three generation
59. Peen J, Schoevers RA, Beekman AT, Dekker J. The current status of urban-rural differences in psychiatric disorders. Acta Psychiatr Scand. 2010;121(2):84–93. https://doi.org/10.1111/j.1600-0447.2009.01438.x.

60. Penkalla AM, Kohler S. Urbanicity and mental health in Europe: A systematic review. Eur J Ment Heal. 2014;9(2):163–77.

61. Verheij RA. Explaining urban-rural variations in health: A review of interactions between individual and environment. Soc Sci Med. 1996;42(6):923–35. https://doi.org/10.1016/0277-9536(95)00190-5.

62. Dekker J, Peen J, Koelen J, Smit F, Schoevers R. Psychiatric disorders and urbanization in Germany. BMC Public Health. 2008;8(1):17. https://doi.org/10.1186/1471-2458-8-17.

63. Jones AR, Cook TM, Wang J. Rural–urban differences in stigma against depression and agreement with health professionals about treatment. J Affect Disord. 2011;134(1):145–50. https://doi.org/10.1016/j.jad.2011.05.013.

64. Paananen R, Ristikari T, Merikukka M, Gissler M. Social determinants of mental health: a Finnish nationwide follow-up study on mental disorders. J Epidemiol Community Health. 2013;67(12):1025–31. https://doi.org/10.1136/jech-2013-202768.

65. Paananen R, Santalahti P, Merikukka M, Rämo A, Wahlbeck K, Gissler M. Socioeconomic and regional aspects in the use of specialized psychiatric care—a Finnish nationwide follow-up study. Eur J Pub Health. 2013;23(3):372–7. https://doi.org/10.1093/europub/cks147.

66. Essau CA, Lewinsohn PM, Lim JX, Ho M, Ho R, Rohde P. Incidence, recurrence and comorbidity of anxiety disorders in four major developmental stages. J Affect Disord. 2018;228(December 2017):248–53. https://doi.org/10.1016/j.jad.2017.12.014.

67. Wittchen HU, Kessler RC, Pfister H, Hofler M, Lieb R. Why do people with anxiety disorders become depressed? A prospective-longitudinal community study. Acta Psychiatr Scand Suppl. 2000;102(406):14–23.

68. Margari L, Buttiglione M, Craig F, Cristella A, de Giambattista C, Matera E, et al. Neuropsychopathological comorbidities in learning disorders. BMC Neurol. 2013;13(13):198. https://doi.org/10.1186/1471-2377-13-198.

69. Merten EC, Cwik JC, Margraf J, Schneider S. Overdiagnosis of mental disorders in children and adolescents (in developed countries). Child Adolesc Psychiatry Ment Health. 2017;11(1):5. https://doi.org/10.1186/s13034-016-0140-5.

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