# Psychiatric comorbidities in children with conduct disorder: a descriptive analysis of real-world data

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## To the editor:

Two recent advisories from the US surgeon general have underscored the unprecedented public health crisis in youth mental health and emphasised the need for "timely data collection and research to identify and respond to youth mental health needs more rapidly".<sup>1</sup> Increased rates of suicidal behaviour, depression and anxiety symptoms and substance use, particularly in adolescents and young adults, are part of an emerging landscape of high volumes of emergency room visits and hospitalisations for behavioural problems.<sup>2 3</sup> Conduct disorder (CD) has emerged as an increasingly common presenting concern in adolescents and young adults receiving emergency and inpatient psychiatric care.4-6 Yet, CD remains among the least studied of paediatric psychiatric disorders, even though it is thought to be associated with a myriad of devastating social, legal and psychiatric consequences that may persist into adulthood and is frequently comorbid with other psychiatric disorders.<sup>7</sup> Although recent studies estimate that CD affects approximately 2%-3% of the school-aged paediatric population worldwide,<sup>8</sup> ambiguity remains about its association with other psychiatric conditions. CD is posited to be associated with increased rates of anxiety, depression, substance use and attention-deficit hyperactivity disorder (ADHD), as well as other externalising disorders, with a multisite analysis in Europe examining nearly 800 children with CD and found that more than one-third likely suffered from current ADHD symptoms and over three-fourths suffered from oppositional defiant disorder symptoms.<sup>9</sup> Yet, estimates of comorbidity based on clinical data collected in settings that reflect actual psychiatric care, particularly in the USA, are lacking. Furthermore, many of the estimates of psychiatric comorbidity in adolescents and

young adults, defined as ages 12–25, with CD, were conducted >10 years ago, preceding the ongoing paediatric mental health crisis in the USA.

Untreated co-occurring psychiatric conditions in adolescents and young adults with behavioural disorders like CD can exacerbate behavioural dysregulation and increase the likelihood of criminal legal system involvement.<sup>10-12</sup> Against this backdrop, there is an urgent need for up-to-date and populationlevel estimates of the psychiatric comorbidities in young people with CD. To our knowledge, there are no observational studies that provide essential descriptive statistics on the prevalence of psychiatric comorbidities in young people with CD. Recent advances in the analysis of real-world clinical evidence, defined as routinely collected healthcare data derived from sources outside typical clinical research settings (ie, electronic health records (EHRs), insurance claims and billing data, and government registries), may hold promise in helping us understand the prevalence of co-occurring psychiatric conditions in people with CD. To address these gaps in our understanding of CD and its co-occurring psychiatric conditions, this brief report used real-world EHR spanning over 113 million people worldwide, with the greatest representation in the USA, to estimate the rates of psychiatric comorbidity in adolescents and young adults, defined as ages 12-25 years, with CD.

### METHODS

This descriptive analysis used the TriNetX data, which is a federated research network capturing deidentified real-time EHRs. This dataset includes comprehensive data related to clinical diagnoses, procedures and treatments, prescription medications and

laboratory data, sourced from over 119 million people (including adolescents and young adults, defined as ages 12–25 years) across 80 healthcare organisations (HCOs) in multiple countries). The TriNetX data include data from inpatient and outpatient (ie, specialty, primary care) settings. Due to the nature of the protected health information in the TriNetX data, detailed geographic and institutional information is not provided. TriNetX uses privacy-preserving record linkage (via cryptographic tokens) to enrich clinical data from the EHR of HCOs with closed claims data, and hospital and government mortality data. These data were determined not to be human subjects research by the Institutional Review Board, as our data were deidentified. We followed the Strengthening the Reporting of Observational Studies in Epidemiology and the Reporting of Studies Conducted Observational Routinely Collected Using Health Data Statement for Pharmacoepidemiology reporting guidelines.

Our sample consisted of people between the ages of 12 and 25 years who were in the TriNetX databases on and before the date of 1 October 2023. We selected the lower threshold of 12 years to focus our sample beginning from early adolescence to an upper threshold of 25 years to represent a commonly used upper limit of young adulthood. We obtained a cohort of people with an International Classification of Diseases-10 Clinical Modification (ICD-10-CM) diagnosis of CD (F91). Given that the specificity of ICD codes is typically improved with multiple claims, we required at least two diagnoses of CD to mitigate misclassification bias.<sup>13</sup> The presented data were gueried on 22 November 2023 and again on 28 January 2024. Among all persons with CD, we extracted descriptive characteristics such as age at the time of index diagnosis (the first time they received the diagnosis), sex (assigned male at birth, assigned female at birth), race/ethnicity and clinical characteristics, namely co-occurring psychiatric disorders such as ADHD, mood and anxiety disorders, psychotic disorders, personality disorders and substance use disorders. To evaluate the types of treatments individuals were receiving, we used linked pharmacy records to extract data on prescriptions (filled in the 2 years preceding and including the date of the index/first CD diagnosis) for antipsychotics, selective serotonin reuptake inhibitors (SSRIs), stimulants for ADHD (ie, amphetamine salts, methylphenidate, lisdexamfetamine) and mood stabilisers such as lithium, lamotrigine, carbamazepine and valproate, using a combination of RxNorm and Anatomical Therapeutic Chemical codes. We subsequently conducted descriptive analyses using the TriNetX data to query the prevalence of co-occurring psychiatric disorders in the 2 years preceding and including the date of the index/ first diagnosis (observation window shown in online supplemental efigure), as well as demographic characteristics of people who received a diagnosis of CD. Overall, 92.9% (66 376/71419) of the sample had greater than 2 years of data. For HCOs that returned data coded by ICD-9, TriNetX mapped ICD-9 codes to ICD-10-CM codes using General Equivalence Mapping algorithms. Diagnostic codes are provided in online supplemental etable.

## RESULTS

As shown in table 1, a total of 71 419 persons between the ages of 12 and 25 who had at least two diagnoses of CD were included in this study. The mean (standard deviation (SD)) age at the time of index CD diagnosis was 14.7 (2.7) years. 64.8% were assigned male at birth. 15 303 (21.4%) were non-Hispanic black and 41 248 (57.8%) non-Hispanic white. In the retrieved cohort of CD individuals, more than 90% of them were in the USA.

Psychiatric comorbidities were common with more than half of persons exhibiting co-occurring disorders. Overall, 41608 (58.3%) had a diagnosis of ADHD in the 2 years preceding the diagnosis of CD. 32.6% (n=23256) received psychostimulant scripts for the treatment of ADHD in the 2 years preceding CD diagnosis. Specifically, 7633 (10.7%) received amphetamine salts, 7861 (11.0%) received dextroamphetamine, 12220 (17.1%) received methylphenidate, 3651 (5.1%) received dexmethylphenidate, and 5738 (8.0%) received lisdexamfetamine.

Furthermore, 33246 (46.6%) had a diagnosis of a mood disorder (ie, major depressive disorder, bipolar disorder), 6981 (9.8%) had post-traumatic stress disorder, 7341 (10.3%) had a generalised anxiety disorder, 3074 (4.3%) had schizophrenia-spectrum disorders, 3334 (4.7%) had a diagnosis of a personality disorder), and 4998 (7.0%) had a diagnosis of an impulse control disorder.

Notably, approximately as many individuals received antipsychotics (n=21359, 29.9%) as SSRIs (n=20468, 28.7%). Finally, 12.5% of individuals had diagnoses for autism spectrum disorders (n=8943) and 14.0% for substance use disorders (n=9963), with cannabis use disorder overwhelmingly being the most common substance use problem (n=6356, 8.9%) among people in the cohort who had a diagnosis of substance use disorder.

## DISCUSSION

Overall, this analysis uses real-time administrative claims data for over 100 million enrollees, spanning public and private insurance (as well as uninsured people receiving medical care who are captured in EHR), making this the largest and most comprehensive characterisation of psychiatric comorbidities in people with CD to date. Comprehensive real-world data on the prevalence of co-occurring psychiatric disorders in adolescents and young adults, ages 12-25 years, with CD, is much needed amid the current paediatric mental health crisis in the US. People with CD often receive treatment in acutecare settings such as emergency rooms, where crowding, short patient stays, clinician task interruption and lack of longitudinal patient-physician relationships can hinder screening and intervention.<sup>14</sup> In a recent national survey, paediatric emergency medicine physicians (including both trainees and faculty) in the USA reported feeling underprepared in evaluating and treating children with unmet social needs.<sup>14</sup> Against this backdrop, our analysis, to the best of our knowledge, is among the first papers to use real-world data to characterise the prevalence of co-occurring psychiatric disorders in adolescents and young adults with CD, providing

 Table 1
 Descriptive characteristics of people with conduct disorder

	Individuals diagnosed with conduct disorder	
	n=71419 Percentage	
Male (vs female)	46082	64.5%
Racial and ethnic characteristics		
Non-Hispanic white	41248	57.8%
Non-Hispanic black	15303	21.4%
Hispanic	6965	9.8%
Unknown ethnicity	12978	18.2%
Unknown race	9993	14.0%
Asian	1049	1.5%
Other race	2994	4.2%
Native Hawaiian or Pacific Islander	438	0.6%
Native American	394	0.6%
Age at time of index (first) diagnosis, in years (SD)	14.7	2.7
Psychiatric diagnoses		
Attention-deficit hyperactivity disorder	41 608	58.3%
Generalised anxiety disorder	7341	10.3%
Post-traumatic stress disorder	6981	9.8%
Adjustment disorder	7957	11.1%
Obsessive-compulsive disorder	2682	3.8%
Mood (affective) disorders (major depressive or bipolar disorder)	33246	46.6%
Schizophrenia-spectrum disorders	3074	4.3%
Personality disorders	3334	4.7%
Impulse control disorders	4998	7.0%
Autism spectrum disorders	8943	12.5%
Intellectual disabilities	5846	8.2%
Any substance use disorder (alcohol, stimulant, cannabis, sedative)	9963	14.0%
Cannabis use disorder	6356	8.9%
Psychotropic medications		
Antipsychotics	21359	29.9%
Risperidone	8088	11.3%
Aripiprazole	7551	10.6%
Quetiapine	4191	5.9%
Olanzapine	3760	5.3%
Haloperidol	3543	5.0%
Ziprasidone	1579	2.2%
Lurasidone	666	0.9%
Psychostimulants	23256	32.6%
Amphetamine stimulants	12046	16.9%
Amphetamine Salts	7633	10.7%
Dextroamphetamine	7861	11.0%
Methylphenidate	12220	17.1%
Dexmethylphenidate	3651	5.1%
Lisdexamfetamine	5738	8.0%
Antidepressants and mood stabilisers		

Continued

## Table 1 Continued

	Individuals diagnosed with conduct disorder	
	n=71419	Percentage
ny selective serotonin reuptake inhibitors	20468	28.7%
thium	613	0.9%
amotrigine	2640	3.7%
arbamazepine	529	0.7%
alproate	3592	5.0%

up-to-date data on the epidemiology of CD to optimise treatment for this critically underserved population Our findings reveal a high prevalence of co-occurring psychiatric conditions, particularly ADHD and affective disorders. A sizeable minority of patients received diagnoses for addictive disorders, with cannabis use disorder being the most prevalent of substance use disorders. Even though almost 50% of the sample was documented to have a diagnosis of ADHD, only 33% received psychostimulants for ADHD. Approximately as many adolescents and young adults with CD received antipsychotics as psychostimulants in our cohort.

This study is strengthened by its use of real-world clinical evidence, allowing us to study the epidemiology of CD using larger and more inclusive populations than what is often accomplished in clinical trials and cohort studies, in line with the US surgeon general's call for "integrated, real-time data infrastructure for understanding youth mental health trends".<sup>1</sup> Yet, there are key limitations. First, we cannot rule out misclassification of diagnoses and medication consumption in the administrative data, especially since pharmacy claims may not reflect the actual consumption of medication. Second, while the TriNetX data is multinational, the overwhelming majority of the data comes from the USA; as the Diagnostic and Statistical Manual of Mental Disorders (DSM) is typically used to diagnose CD in the USA, our data may not necessarily be generalisable to countries that use ICD systems. Third, by nature of its deidentified real-time data, the TriNetX accessible to us lacks refinement capabilities (ie, data on the type of clinician that gave the diagnosis of CD), as seen in other administrative databases such as Market-Scan and CMS Medicaid data, which limits our ability to conduct in-depth causal inference work beyond descriptive data. While CD diagnoses are typically made using DSM criteria, our TriNetX interface only permits access to ICD codes. Because TriNetX data stem from a conglomerate of EHR data, people who do not have access to healthcare (ie, people diagnosed with CD who are incarcerated and are not receiving consistent healthcare) are under-represented.

Finally, against the backdrop of mass incarceration and structural racism in the US mental health systems, it is important to emphasise concern surrounding how the diagnosis of CD may increase the likelihood

of criminal legal system involvement.<sup>10-12</sup> The USA has one of the highest rates of incarceration in the world, with the growth in incarceration rates occurring disproportionately in black Americans, many of whom suffer from trauma and have unmet mental health needs.<sup>15</sup> Studies have shown that adolescents who are involved in the criminal justice system have significant substance use disorders and mental health needs that are often untreated.<sup>12</sup> Data from the present study are likely to help efforts, such as the Juvenile Justice Behavioral Health Services Cascade,<sup>12</sup> to address the mental health needs of justice-involved children and provide trauma-informed alternatives to incarceration. Future research is urgently needed to elucidate racial and ethnic disparities in CD diagnoses and downstream mental health outcomes, as well as comprehensively assess the extent to which co-occurring psychiatric disorders in adolescents and young adults with CD, ages 12-25 years, are being adequately treated.

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