Original Investigation

Treatment of Young People With Antipsychotic Medications in the United States

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IMPORTANCE Despite concerns about rising treatment of young people with antipsychotic medications, little is known about trends and patterns of their use in the United States.

OBJECTIVE To describe antipsychotic prescription patterns among young people in the United States, focusing on age and sex.

DESIGN, SETTING, AND PARTICIPANTS A retrospective descriptive analysis of antipsychotic prescriptions among patients aged 1 to 24 years was performed with data from calendar years 2006 (n = 765 829), 2008 (n = 858 216), and 2010 (n = 851 874), including a subset from calendar year 2009 with service claims data (n = 53 896). Data were retrieved from the IMS LifeLink LRx Longitudinal Prescription database, which includes approximately 60% of all retail pharmacies in the United States. Denominators were adjusted to generalize estimates to the US population.

MAIN OUTCOMES AND MEASURES The percentage of young people filling 1 or more antipsychotic prescriptions during the study year by sex and age group (younger children, 1-6 years; older children, 7-12 years; adolescents, 13-18 years; and young adults, 19-24 years) was calculated. Among young people with antipsychotic use, percentages with specific clinical psychiatric diagnoses and 1 or more antipsychotic prescriptions from a psychiatrist and from a child and adolescent psychiatrist were also determined.

RESULTS The percentages of young people using antipsychotics in 2006 and 2010, respectively, were 0.14% and 0.11% for younger children, 0.85% and 0.80% for older children, 1.10% and 1.19% for adolescents, and 0.69% and 0.84% for young adults. In 2010, males were more likely than females to use antipsychotics, especially during childhood and adolescence: 0.16% vs 0.06% for younger children, 1.20% vs 0.44% for older children, 1.42% vs 0.95% for adolescents, and 0.88% vs 0.81% for young adults. Among young people treated with antipsychotics in 2010, receiving a prescription from a psychiatrist was less common among younger children (57.9%) than among other age groups (range, 70.4%-77.9%). Approximately 29.3% of younger children treated with antipsychotics in 2010 receiving a child and adolescent psychiatrist. Among young people with claims for mental disorders in 2009 who were treated with antipsychotics, the most common diagnoses were attention-deficit/hyperactivity disorder in younger children (52.5%), older children (60.1%), and adolescents (34.9%) and depression in young adults (34.5%).

CONCLUSIONS AND RELEVANCE Antipsychotic use increased from 2006 to 2010 for adolescents and young adults but not for children aged 12 years or younger. Peak antipsychotic use in adolescence, especially among boys, and clinical diagnosis patterns are consistent with management of developmentally limited impulsive and aggressive behaviors rather than psychotic symptoms.

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Corresponding Author: Mark Olfson, MD, MPH, Department of Psychiatry, College of Physicians and Surgeons, Columbia University, 1051 Riverside Dr, New York, NY (mo49@cumc .columbia.edu). uring the past several years, antipsychotics have gained popularity as treatments for psychiatric disorders in young people.¹ Clinical trials support the efficacy of several antipsychotics for child² and adolescent³ bipolar mania, adolescent schizophrenia,⁴ and irritability associated with autism in adolescents and children as young as 5 (risperidone)⁵ and 6 (aripiprazole)⁶ years. Yet most office visits by children and adolescents that involve antipsychotic treatment do not include one of these clinical diagnoses.⁷⁻⁹ In this setting, the clinical diagnoses associated with antipsychotic treatment vary with patient age. For example, a larger percentage of child (63.0%) than adolescent (33.7%) visits with antipsychotic treatment include disruptive behavior disorder diagnoses, while the reverse is true of bipolar disorder diagnoses (child, 12.2% vs adolescent, 28.8%).⁸

Concerns have been raised regarding the risks of antipsychotic medications to the cardiovascular and metabolic health of young people. Second-generation antipsychotics appear to pose a greater risk of weight gain and dyslipidemia to children than to adults.^{10,11} Within the Medicaid program, growth in the percentage of children and adolescents treated with antipsychotics who also receive other psychotropic medications¹² has further raised concerns regarding adverse effect burden. In the general population, the extent to which young people who are treated with antipsychotics also receive other classes of psychotropic medications is not known.

Concern has particularly focused on the safety of antipsychotic treatment of young children. In preschool-aged children, a paucity of research on psychiatric diagnoses complicates patient selection, the known efficacy of antipsychotics is limited largely to irritability associated with developmental disorders, adverse metabolic and endocrine effects are pronounced,¹³ variation in drug metabolism complicates dosing,¹⁴ and animal studies raise concerns regarding antipsychotic safety on the developing mammalian brain.¹⁵⁻¹⁹ As a result, it is recommended that preschool-aged children presenting for mental health care should receive a comprehensive psychiatric assessment and a trial of a relevant psychosocial intervention before considering psychopharmacologic treatment.²⁰ In this context, the frequency with which preschool-aged children receive antipsychotic prescriptions from nonpsychiatrists or do not receive a trial of psychotherapy suggests potential gaps in care.

The rate of antipsychotic treatment tends to be higher in young adults and adolescents than in children.^{1,8} However, sample size constraints have limited descriptions of antipsychotic treatment to broad age groups.^{1,7-9} Because disruptive behavioral disorders are recorded in a substantial proportion of youth who are treated with antipsychotics²¹ and treatment of these conditions declines in late adolescence,²² an overall decline in antipsychotic treatment during late adolescence would support the hypothesis that diagnoses of disruptive behavioral disorders make important contributions to antipsychotic treatment in this age group.²³

The national prevalence of antipsychotic use by young people is not known. In the Medicaid program and some private insurance plans, antipsychotic use increased among young people during the late 1990s and early 2000s.²¹ On a per-visit basis, office visits by youth that included antipsychotic prescriptions increased rapidly between 1993-1998 and 1999-2004 and then grew more slowly between 1999-2004 and 2005-2009.⁸ A growing awareness of increasing use and safety concerns^{24,25} may have tempered antipsychotic treatment of young people.

We provide, to our knowledge, the first description of patterns and trends in antipsychotic use by young people in the United States. We focus on age-related variations in antipsychotic use, the role of psychiatrists and child and adolescent psychiatrists in prescribing antipsychotics, coprescribed psychotropic medications, and clinical diagnoses.

Methods

Data Sources

From the IMS LifeLink LRx Longitudinal Prescription databases, we obtained data on filled prescriptions for all antipsychotics in 2006, 2008, and 2010, as well as the total population by sex and age covered by these databases who filled 1 or more prescriptions during the year. From the Medical Expenditure Panel Survey we derived the percentages of young people by sex and single year of age who did not fill any prescriptions in 2006, 2008, and 2010.²⁶ We also received data from the IMS LifeLink LRx Longitudinal Prescription database on prescriptions for other psychotropic medications in 2008 but not in 2006 or 2010. The LRx data contain deidentified individual prescriptions from approximately 33 000 retailers. In 2010, IMS LRx data captured 63% of all retail prescriptions in the United States and were nationally representative with respect to age, sex, and insurance.

We also received the 2009 IMS Medical Claims Database, which includes more than 16 million service claims per month from more than 100 000 unique physicians across all 50 states. The 2009 database was merged with pharmacy claims from patients common to the 2009 LRx database. Approximately 5.8% of the 2009 LRx sample of young people treated with antipsychotics was included in the merged file. Because all of the data sets are deidentified, they were exempted from human subjects review by the Yale University and New York State Psychiatric Institute Institutional Review Boards.

Statistical Analysis

We conducted a population-level, retrospective, observational study of antipsychotic use in the United States, focusing on antipsychotic use among young people aged 1 to 24 years (eTable 1 in the Supplement). We adjusted the IMS denominators with Medical Expenditure Panel Survey data on the percentages of young people by sex and single year of age who did not fill any prescriptions in 2006, 2008, and 2010. This adjustment generalized the IMS prevalence estimates of antipsychotic use based on individuals filling 1 or more prescriptions of any medication during the year to the entire US population of young people, including those who did not fill a prescription. The age and sex composition of the IMS population that filled at least 1 prescription of any

	Population Wit	Population With Prescription by Age Category, %					
Year by Sex	1-6 у	7-12 у	13-18 у	19-24 у			
Total							
2006	0.14	0.85	1.10	0.69			
2008	0.16	0.87	1.18	0.75			
2010	0.11	0.80	1.19	0.84			
Males							
2006	0.20	1.28	1.35	0.70			
2008	0.24	1.33	1.43	0.76			
2010	0.16	1.20	1.42	0.88			
Females							
2006	0.08	0.45	0.87	0.68			
2008	0.09	0.47	0.95	0.75			
2010	0.06	0.44	0.95	0.81			

^a Results are based on younger children (1), older children (2), adolescents (3), and young adults (4) receiving antipsychotic prescriptions in the 2006 ($n_1 = 42 459; n_2 = 220 305;$ $n_3 = 305 165; n_4 = 197 900$), 2008 ($n_1 = 50 725; n_2 = 247 111;$ $n_3 = 332 051; n_4 = 228 329$), and 2010 ($n_1 = 36 484; n_2 = 226 914;$ $n_3 = 335 737; n_4 = 252 739$) IMS Health data.

Figure. Percentage of Male and Female Population With Antipsychotic Medication Use by Sex and Age, United States, 2010



kind closely resembled the composition of the corresponding population from the nationally representative Medical Expenditure Panel Survey.

We calculated the percentages of younger children (age, 1-6 years), older children (age, 7-12 years), adolescents (age, 13-18 years), and young adults (age, 19-24 years) who used an antipsychotic overall and by sex for each year (**Table 1**). Rates of any antipsychotic use in 2010 by single year of age were also plotted separately for males and females (**Figure**). Among young people with any antipsychotic use, we determined the percentages with 1 or more antipsychotic prescriptions from a psychiatrist and 1 or more antipsychotic prescriptions from a child and adolescent psychiatrist for each of the 4 age groups in 2006, 2008, and 2010 (**Table 2**). We also determined the percentage of youth in each age group among antipsychotic users who received 1 or more prescriptions for stimulants, antidepressants, mood stabilizers, and benzodiazepines during the same year (**Table 3**).

In the 2009 merged file, we examined the following diagnoses of patients treated with antipsychotics with 1 or more mental disorder diagnoses in any treatment setting: attentiondeficit/hyperactivity disorder (ADHD), disruptive behavior disorders, bipolar disorder, schizophrenia, autism or mental retardation, anxiety, depression, adjustment-related disorders, substance use, and other mental disorder diagnoses. Frequency distributions of these mental disorders and any psychotherapy use (eTable 2 in the Supplement) were examined within the 4 age groups (Table 4).

Results

Overall Use of Antipsychotic Medications

The 2010 LRx database included 36 484 younger children, 226 914 older children, 335 737 adolescents, and 252 739 young adults with 1 or more antipsychotic prescriptions. Accounting for the scope in data coverage, these numbers correspond to approximately 58 000 younger children, 360 000 older children, 530 000 adolescents, and 400 000 young adults with 1 or more antipsychotic prescriptions nationwide.

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Table 2. Percentage of Antipsychotic Users With Any Antipsychotic Prescriptions From a Psychiatrist and From a Child and Adolescent Psychiatrist^a

	Population With Prescription by Year, %			
Age Category by Prescription Source, y	2006	2008	2010	
Psychiatrist				
1-6	61.0	51.2	57.9	
7-12	74.7	71.9	71.9	
13-18	79.7	81.0	77.9	
19-24	71.1	73.4	70.4	
Child and adolescent psychiatrist				
1-6	32.1	31.0	29.3	
7-12	41.9	39.8	39.2	
13-18	40.2	39.0	39.2	
19-24	13.2	13.7	14.2	

^a Results are based on younger children (1), older children (2), adolescents (3), and young adults (4) receiving antipsychotic prescriptions in the 2006 ($n_1 = 42 459$; $n_2 = 220 305$; $n_3 = 305 165$; $n_4 = 197 900$), 2008 ($n_1 = 50 725$; $n_2 = 247 111$; $n_3 = 332 051$; $n_4 = 228 329$), and 2010 ($n_1 = 36 484$; $n_2 = 226 914$; $n_3 = 335 737$; $n_4 = 252 739$) IMS Health data.

Table 3. Stimulant, Antidepressant, Mood Stabilizer, and Benzodiazepine Prescriptions in 2008 Among Young People With Antipsychotic Prescriptions

	Population With Prescription by Age Category, %				
Prescription Medication	1-6 y (n = 50 725)	7-12 y (n = 247 111)	13-18 y (n = 332 051)	19-24 y (n = 228 329)	
Stimulants	58.7	68.7	44.5	17.1	
Antidepressants	20.3	34.0	50.8	59.1	
Mood stabilizers	16.5	24.6	34.9	41.4	
Benzodiazepines	6.4	6.0	11.7	33.5	
Antipsychotics only	27.8	15.0	16.2	18.1	

Table 4. Mental Disorder Diagnoses and Use of Psychotherapy in 2009 Among Young People With Antipsychotic Prescriptions

	Population With Diagnosis by Age Category, %			
Characteristic	1-6 y (n = 925)	7-12 y (n = 5939)	13-18 y (n = 8198)	19-24 y (n = 5353)
Diagnosis ^a				
ADHD	52.5	60.1	34.9	11.3
Autism or mental retardation ^b	23.1	13.8	8.4	5.7
Disruptive behavioral disorders	20.6	15.7	13.0	2.2
Bipolar disorder	8.1	12.7	20.5	26.6
Anxiety	6.9	10.4	13.0	22.9
Depression	2.4	6.1	24.4	34.5
Adjustment-related disorders	1.8	2.6	2.7	2.2
Substance use	0.8	0.3	0.2	0.8
Schizophrenia	0.3	0.2	1.4	7.5
Other mental disorder	23.5	21.0	22.6	18.1
Any psychotherapy	13.5	20.4	24.8	18.8

Abbreviation: ADHD, attention-deficit/hyperactivity disorder. ^a Diagnostic groups are not mutually

exclusive.

^b Includes autism and other pervasive developmental disorders.

There were approximately 270 000 antipsychotic prescriptions dispensed to younger children, 2.14 million to older children, 2.80 million to adolescents, and 1.83 million to young adults nationwide in 2010. Across the 4 age groups, almost all of the antipsychotic prescriptions (96.7%) were for secondgeneration medications.

National Trends in Antipsychotic Use

Antipsychotic use among younger and older children increased between 2006 (younger children, 0.14%; older children, 0.85%) and 2008 (younger children, 0.16%; older children, 0.87%) before declining between 2008 and 2010 (younger children, 0.11%; older children, 0.80%). For adolescents and especially young adults, there was an increase in antipsychotic use between 2006 (adolescents, 1.10%; young adults, 0.69%) and 2010 (adolescents, 1.19%; young adults, 0.84%). The increase in antipsychotic use in the 2 older groups occurred for both males and females (Table 1).

Age and Sex Patterns of Antipsychotic Use

The percentage of persons with any antipsychotic use in 2010 was determined by single year of age for males and females (Figure). The percentage of males with antipsychotic use sharply increased from ages 1 to 10 years, reached a plateau

that continued until age 17 years, rapidly declined to age 19 years, slowly declined to a nadir at age 22 years, and gradually increased until age 24 years. For females, the increase was more gradual and reached a lower peak at age 16 years before declining and following a pattern that resembled the one for males (Figure and eFigures 1 and 2 in the Supplement).

Antipsychotic Prescriptions From Psychiatrists

Most young people treated with antipsychotics in each age group filled 1 or more antipsychotic prescriptions that were written by a psychiatrist (57.9% of young children, 71.9% of older children, 77.9% of adolescents, and 70.4% of young adults in 2010) (Table 2). However, only a minority filled 1 or more antipsychotic prescriptions from a child and adolescent psychiatrist (29.3% of young children, 39.2% of older children, 39.2% of adolescents, and 14.2% of young adults in 2010). Younger children treated with antipsychotics were proportionately the least likely of the 4 age groups to have filled 1 or more antipsychotic prescriptions from a psychiatrist. Within age groups, a similar percentage of males and females treated with antipsychotics filled 1 or more prescriptions from a psychiatrist. There was no consistent temporal trend in the percentage of young people treated with antipsychotics who filled prescriptions from psychiatrists (Table 2).

Other Classes of Psychotropic Medications:

During 2008, most young people who were treated with antipsychotic medications were also treated with other classes of psychotropic medications. Among younger and older children, stimulants were the most commonly prescribed other class of psychotropic medication (younger children, 58.7%; older children, 68.7%). For young adults, antidepressants were the most commonly co-prescribed psychotropic class at 59.1% (Table 3). Among male adolescents, stimulants were the most commonly co-prescribed class, while among female adolescents, antidepressants were the most commonly co-prescribed class.

Clinical Diagnoses and Psychotherapy Use

In the merged 2009 medical claims and LRx sample, most of the younger children (60.0%), older children (56.7%), adolescents (62.0%), and young adults (67.1%) treated with antipsychotics had no outpatient or inpatient claim that included a mental disorder diagnosis. Among antipsychotic-treated children and adolescents with mental disorder claims, the most common diagnosis was ADHD (younger children, 52.5%; older children, 60.1%; adolescents, 34.9%), while depression was the most common diagnosis among young adults (34.5%), followed by bipolar (26.6%) and anxiety disorders (22.9%). Less than one-fourth of the patients in each of the age groups had an outpatient claim for psychotherapy (younger children, 13.5%; older children, 20.4%; adolescents, 24.8%; young adults, 18.8%) (Table 4).

Discussion

In contrast with earlier reports⁷⁻⁹ of increasing antipsychotic treatment of US children through the early 2000s, the percent-

age of children treated with antipsychotics was lower in 2010 than in 2006. Among adolescents and young adults, however, antipsychotic use increased during this period. Consistent with clinical diagnoses suggesting that antipsychotics are primarily used to manage impulsive or aggressive behaviors in children associated with ADHD,²⁷ the highest rate of antipsychotic treatment was in adolescent boys, approximately half of whom also filled prescriptions for stimulants. Young adults treated with antipsychotics were more frequently diagnosed as having depression, bipolar disorder, and anxiety disorder than ADHD.

The single year of age and sex patterns of antipsychotic use offers clues to clinical and social determinants of community antipsychotic prescribing practices. There was a rise in antipsychotic use during childhood that extended into adolescence for females, followed by a plateau and a decline during the transition to early adulthood. This age profile refines earlier reports of steadily increasing rates of antipsychotic use across multiyear developmental categories of childhood to adolescence^{7,28} to early adulthood.²⁹ A peak in antipsychotic use among males aged 11 to 17 years fits with treatment of conduct problems that commonly occur during adolescence.³⁰ In the United States, the rate of arrests for simple assaults peaks at age 16 years before quickly declining.³¹ An important issue remains in determining the extent to which the benefits and safety demonstrated for risperidone in clinical trials of youth with severe conduct disorders and below-average intelligence^{32,33} generalize to other antipsychotics and the broader population of youth with conduct disorders. Much also remains to be learned about the community and clinical prevalence of explosive aggression in children and adolescents.

A decrease in conduct disorders during late adolescence has been linked to suppression of aggression and improved impulse control.³⁴ During late adolescence and early adulthood, neurobiological systems responsible for self-regulation and control undergo a complex maturation. This maturation involves a decrease in prefrontal gray matter related to synaptic pruning, an increase in myelination within the prefrontal cortex, and a proliferation of white matter tracts between cortical and subcortical areas, especially including the prefrontal regions, amygdala, nucleus accumbens, and hippocampus.³⁵ This normal maturation of neurobiological systems may underlie the decrease in antipsychotic treatment prevalence during late adolescence among youth who do not have enduring cognitive impairments and long-term severe behavioral disorders.

Social factors may also play a role in the age-related decrease in antipsychotic treatment in late adolescence. As young people reach the legal age of majority, which is 18 years in most states, they assume greater independence and control of their medical care and may increasingly refuse or self-discontinue use of antipsychotic medications. Because many young people also terminate their formal education during this developmental period, they may also lose a primary point of entry into physical and mental health care. An increase in the incidence of substance disorders during this period³⁶ may further lower the perceived need for mental health care and reduce the likelihood of accessing mental health treatment.

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Male children and adolescents were more likely than their female peers to be treated with antipsychotics. This sex difference is consistent with the known male predominance of ADHD and other disruptive behavior disorders during childhood and adolescence. The narrowing of sex differences in the rate of antipsychotic treatment among young adults may be related to a greater contribution of severe mood disorders to antipsychotic treatment decisions.³⁷ Mood disorders tend to have a later age of onset than does ADHD and to be female predominant.

In accord with a prior analysis of US office-based medical practice,⁸ most young people treated with antipsychotics did not have evidence of a mental disorder diagnosis in the health care claims data. Some primary care physicians may withhold mental disorder diagnoses because they are not reimbursed for these codes³⁸ or to reduce stigma. In other cases, antipsychotics may be used to treat insomnia, agitation, or other symptoms that do not meet criteria for a mental disorder.³⁹

High rates of coprescribing of antipsychotics with other psychotropic classes were observed across all age groups and several medication classes. Stimulants were the most commonly coprescribed psychotropic class during the preadolescent years.¹² Age patterns of coprescribed psychotropic medications reflected corresponding clinical diagnoses. Stimulant prescriptions and ADHD diagnoses peaked among older children, while antidepressant prescriptions and depression diagnoses increased during adolescence.

Maladaptive aggression is a common feature of ADHD. It was present in nearly half of the children in the large National Institute of Mental Health Multimodal Treatment of ADHD trial.⁴⁰ In support of antipsychotic and stimulant combinations, one recent trial of severely aggressive children with ADHD and either oppositional defiant disorder or conduct disorder found that youth who received risperidone augmentation following optimization of stimulants achieved significantly greater reductions in aggressive and disruptive behaviors than did similarly treated children who received placebo as the adjuvant.⁴¹ In our study, among adolescents treated with antipsychotics who had diagnosed psychiatric disorders, the most common diagnoses were ADHD (34.9%), followed by depression (24.4%) and bipolar disorder (20.5%). In the community, the prevalence of adolescents with severe impairment is 8.7% for depression, 4.3% for ADHD, and 2.6% for bipolar disorder.42 These proportions suggest that adolescents with severely impairing ADHD or bipolar disorder may be more likely than those with severely impairing depression to receive antipsychotic treatment.

Some adolescents and young adults treated with antipsychotics were diagnosed with mood or anxiety disorders. Building empirical support for the efficacy of antipsychotics for adolescent bipolar disorder,^{2,3} including US Food and Drug Administration approval of aripiprazole in 2008 for adolescent manic or mixed bipolar disorder, may have contributed to antipsychotic use trends during the study period. Although depressed adults who partially respond to antidepressants sometimes benefit from antipsychotics,⁴³ it is not known whether antipsychotics confer similar benefits to treatmentresistant depression in young people. It is also not known whether anxiolytic properties of antipsychotics that have been observed in some studies of adults⁴⁴ extend to young people with anxiety disorders.

Practice guidelines urge particular caution regarding the use of antipsychotics in young children.²⁰ In preschool-age children with disruptive behavior disorders, consideration of antipsychotic medications should be limited to those who have severe, sustained, and intractable impairment in multiple settings or who pose safety risks. If antipsychotic therapy is initiated, frequent systematic reassessments should be used to minimize treatment duration.²⁰ However, young children treated with antipsychotics commonly receive their prescriptions exclusively from nonpsychiatrist physicians. Given the paucity of high-quality empirical evidence supporting the efficacy and safety of antipsychotic treatment in this age group, these treatment patterns raise potential safety concerns and underscore the importance of improving access for young children with severe mental health problems to high-quality, specialized child and adolescent mental health services.

Despite progress during the past several years in developing effective psychotherapies for disruptive behaviors,⁴⁵ depression,^{46,47} and anxiety⁴⁸ in young people, psychotherapy was provided to only a minority of young people treated with antipsychotics with claims for mental disorder diagnoses. Greater understanding is needed of the factors that impede access to psychosocial interventions for young people with significant psychological distress.

This analysis has several limitations. First, the IMS prescription data capture medicines purchased rather than consumed. Second, no data were available concerning the effectiveness or safety of the antipsychotics. Third, although the population denominator was adjusted for the percentage of the population by age and sex who reported not filling a prescription medication in the study year, it is not possible to estimate the precision of the derived estimates. Fourth, service claims data were available only for a subset of patients. Fifth, the diagnostic data were based on clinical assessments and were not subject to expert validation. Finally, the primary analyses were based on 2010 dispensing patterns; since that time, prescribing practices may have changed.

Conclusions

After several years of increasing rates of antipsychotic treatment of children and adolescents in the United States, the rate of antipsychotic use among children decreased between 2008 and 2010. In view of evidence of widespread antipsychotic prescribing outside of US Food and Drug Administration-labeled indications and concerns regarding the adverse metabolic effects of second-generation antipsychotics,⁴⁹ this decline is a welcome development. Nevertheless, age and sex antipsychotic use patterns suggest that much of the antipsychotic treatment of children and younger adolescents targets agelimited behavioral problems. In older teenagers and young adults, a developmental period of high risk for the onset of psychotic disorders, antipsychotic use increased between 2006 and 2010. Clinical policy makers have opportunities to promote improved quality and safety of antipsychotic medication use in young people through expanded use of quality measures, physician education, telephone- and Internet-based child and adolescent psychiatry consultation models, and improved access to alternative, evidence-based psychosocial treatments.

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Study concept and design: Olfson, Schoenbaum. *Acquisition, analysis, or interpretation of data:* King, Schoenbaum.

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REFERENCES

1. Olfson 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.

2. Findling RL, Nyilas M, Fornes RA, et al. Acute treatment of pediatric bipolar I disorder, manic or mixed episode, with aripiprazole: a randomized, double-blind, placebo-controlled study. *J Clin Psychiatry*. 2009;70(10):1441-1451.

3. Tohen M, Kryzhanovskaya L, Carlson G, et al. Olanzapine versus placebo in the treatment of adolescents with bipolar mania. *Am J Psychiatry*. 2007;164(10):1547-1556.

4. Findling RL, Robb A, Nyilas M, et al. A multiple-center, randomized, double-blind, placebo-controlled study of oral aripiprazole for treatment of adolescents with schizophrenia. *Am J Psychiatry*. 2008;165(11):1432-1441.

5. McCracken JT, McGough J, Shah B, et al; Research Units on Pediatric Psychopharmacology Autism Network. Risperidone in children with autism and serious behavioral problems. *N Engl J Med.* 2002;347(5):314-321.

6. Marcus RN, Owen R, Kamen L, et al. A placebo-controlled, fixed-dose study of aripiprazole in children and adolescents with irritability associated with autistic disorder. J Am Acad Child Adolesc Psychiatry. 2009;48(11):1110-1119.

7. Matone M, Localio R, Huang YS, dosReis S, Feudtner C, Rubin D. The relationship between mental health diagnosis and treatment with second-generation antipsychotics over time: a national study of US Medicaid-enrolled children. *Health Serv Res.* 2012;47(5):1836-1860.

8. Olfson M, Blanco C, Liu SM, Wang S, Correll CU. National trends in the office-based treatment of children, adolescents, and adults with antipsychotics. *Arch Gen Psychiatry*. 2012;69(12):1247-1256.

9. Alexander GC, Gallagher SA, Mascola A, Moloney RM, Stafford RS. Increasing off-label use of antipsychotic medications in the United States, 1995-2008. *Pharmacoepidemiol Drug Saf*. 2011;20 (2):177-184.

10. Tarricone I, Ferrari Gozzi B, Serretti A, Grieco D, Berardi D. Weight gain in antipsychotic-naive patients: a review and meta-analysis. *Psychol Med.* 2010;40(2):187-200.

11. Correll CU, Manu P, Olshanskiy V, Napolitano B, Kane JM, Malhotra AK. Cardiometabolic risk of second-generation antipsychotic medications during first-time use in children and adolescents. *JAMA*. 2009;302(16):1765-1773.

12. Kreider AR, Matone M, Bellonci C, et al. Growth in the concurrent use of antipsychotics with other psychotropic medications in Medicaid-enrolled children. *J Am Acad Child Adolesc Psychiatry*. 2014; 53(9):960-970.e2.

13. De Hert M, Dobbelaere M, Sheridan EM, Cohen D, Correll CU. Metabolic and endocrine adverse effects of second-generation antipsychotics in children and adolescents: a systematic review of randomized, placebo controlled trials and guidelines for clinical practice. *Eur Psychiatry*. 2011; 26(3):144-158.

14. Bartelink IH, Rademaker CMA, Schobben AF, van den Anker JN. Guidelines on paediatric dosing on the basis of developmental physiology and pharmacokinetic considerations. *Clin Pharmacokinet*. 2006;45(11):1077-1097.

15. Costa LG, Steardo L, Cuomo V. Structural effects and neurofunctional sequelae of developmental exposure to psychotherapeutic drugs: experimental and clinical aspects. *Pharmacol Rev.* 2004;56(1): 103-147.

16. Moran-Gates T, Grady C, Shik Park Y, Baldessarini RJ, Tarazi FI. Effects of risperidone on dopamine receptor subtypes in developing rat brain. *Eur Neuropsychopharmacol*. 2007;17(6-7): 448-455.

17. Choi YK, Moran-Gates T, Gardner MP, Tarazi FI. Effects of repeated risperidone exposure on serotonin receptor subtypes in developing rats. *Eur Neuropsychopharmacol*. 2010;20(3):187-194.

 Mandell DJ, Unis A, Sackett GP. Post-drug consequences of chronic atypical antipsychotic drug administration on the ability to adjust behavior based on feedback in young monkeys.
Psychopharmacology (Berl). 2011;215(2):345-352. **19**. Bardgett ME, Franks-Henry JM, Colemire KR, et al. Adult rats treated with risperidone during development are hyperactive. *Exp Clin Psychopharmacol*. 2013;21(3):259-267.

20. Gleason MM, Egger HL, Emslie GJ, et al. Psychopharmacological treatment for very young children: contexts and guidelines. *J Am Acad Child Adolesc Psychiatry*. 2007;46(12):1532-1572.

21. Crystal S, Olfson M, Huang C, Pincus H, Gerhard T. Broadened use of atypical antipsychotics: safety, effectiveness, and policy challenges. *Health Aff* (*Millwood*). 2009;28(5):w770-w781. doi:10.1377 /hlthaff.28.5.w770.

22. Castle L, Aubert RE, Verbrugge RR, Khalid M, Epstein RS. Trends in medication treatment for ADHD. *J Atten Disord*. 2007;10(4):335-342.

23. Penfold RB, Stewart C, Hunkeler EM, et al. Use of antipsychotic medications in pediatric populations: what do the data say [published correction appears in *Curr Psychiatry Rep.* 2014;16(2):432]? *Curr Psychiatry Rep.* 2013;15(12):426.

24. Vitiello B, Correll C, van Zwieten-Boot B, Zuddas A, Parellada M, Arango C. Antipsychotics in children and adolescents: increasing use, evidence for efficacy and safety concerns. *Eur Neuropsychopharmacol*. 2009;19(9):629-635.

25. Correll CU. Antipsychotic use in children and adolescents: minimizing adverse effects to maximize outcomes. *J Am Acad Child Adolesc Psychiatry*. 2008;47(1):9-20.

26. Agency for Healthcare Research and Quality. MEPS-HC panel design and data collection process. http://meps.ahrq.gov/survey_comp/hc_data _collection.jsp. Accessed January 10, 2015.

27. Pathak P, West D, Martin BC, Helm ME, Henderson C. Evidence-based use of second-generation antipsychotics in a state Medicaid pediatric population, 2001-2005. *Psychiatr Serv*. 2010;61(2):123-129.

28. Zito JM, Safer DJ, de Jong-van den Berg LT, et al. A three-country comparison of psychotropic medication prevalence in youth. *Child Adolesc Psychiatry Ment Health*. 2008;2(1):26.

29. Murphy AL, Gardner DM, Cooke C, Kisely S, Hughes J, Kutcher SP. Prescribing trends of antipsychotics in youth receiving income assistance: results from a retrospective population database study. *BMC Psychiatry*. 2013;13:198.

30. Moffitt TE. Adolescence-limited and life-course-persistent antisocial behavior: a developmental taxonomy. *Psychol Rev.* 1993;100 (4):674-701.

31. Snyder HN. Arrest in the United States, 1990-2010. US Department of Justice, Office of Justice Programs, Bureau of Justice Statistics. NJC 239423. http://www.bjs.gov/content/pub/pdf /aus9010.pdf. Published October 2012. Accessed January 12, 2015.

32. Aman MG, De Smedt G, Derivan A, Lyons B, Findling RL; Risperidone Disruptive Behavior Study Group. Double-blind, placebo-controlled study of risperidone for the treatment of disruptive

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behaviors in children with subaverage intelligence. *Am J Psychiatry*. 2002;159(8):1337-1346.

33. Findling RL, Aman MG, Eerdekens M, Derivan A, Lyons B; Risperidone Disruptive Behavior Study Group. Long-term, open-label study of risperidone in children with severe disruptive behaviors and below-average IQ. *Am J Psychiatry*. 2004;161(4): 677-684.

34. Monahan KC, Steinberg L, Cauffman E, Mulvey EP. Trajectories of antisocial behavior and psychosocial maturity from adolescence to young adulthood. *Dev Psychol.* 2009;45(6):1654-1668.

35. Steinberg L. A social neuroscience perspective on adolescent risk-taking. *Dev Rev.* 2008;28(1):78-106.

36. Pedersen CB, Mors O, Bertelsen A, et al. A comprehensive nationwide study of the incidence rate and lifetime risk for treated mental disorders. *JAMA Psychiatry*. 2014;71(5):573-581.

37. Kuehner C. Gender differences in unipolar depression: an update of epidemiological findings and possible explanations. *Acta Psychiatr Scand*. 2003;108(3):163-174.

38. Freeman VG, Rathore SS, Weinfurt KP, Schulman KA, Sulmasy DP. Lying for patients: physician deception of third-party payers. *Arch Intern Med.* 1999;159(19):2263-2270. **39**. Anderson SL, Vande Griend JP. Quetiapine for insomnia: a review of the literature. *Am J Health Syst Pharm.* 2014;71(5):394-402.

40. Jensen PS, Buitelaar J, Pandina GJ, Binder C, Haas M. Management of psychiatric disorders in children and adolescents with atypical antipsychotics: a systematic review of published clinical trials. *Eur Child Adolesc Psychiatry*. 2007;16(2):104-120.

41. Aman MG, Bukstein OG, Gadow KD, et al. What does risperidone add to parent training and stimulant for severe aggression in child attention-deficit/hyperactivity disorder? *J Am Acad Child Adolesc Psychiatry*. 2014;53(1):47-60.e1.

42. Merikangas KR, He JP, Burstein M, et al. Lifetime prevalence of mental disorders in US adolescents: results from the National Comorbidity Survey Replication—Adolescent Supplement (NCS-A). J Am Acad Child Adolesc Psychiatry. 2010; 49(10):980-989.

43. Nelson JC, Papakostas GI. Atypical antipsychotic augmentation in major depressive disorder: a meta-analysis of placebo-controlled randomized trials. *Am J Psychiatry*. 2009;166(9): 980-991.

44. Maher AR, Maglione M, Bagley S, et al. Efficacy and comparative effectiveness of atypical antipsychotic medications for off-label uses in

adults: a systematic review and meta-analysis. *JAMA*. 2011;306(12):1359-1369.

45. Eyberg SM, Nelson MM, Boggs SR. Evidence-based psychosocial treatments for children and adolescents with disruptive behavior. *J Clin Child Adolesc Psychol*. 2008;37(1):215-237.

46. David-Ferdon C, Kaslow NJ. Evidence-based psychosocial treatments for child and adolescent depression. *J Clin Child Adolesc Psychol*. 2008;37 (1):62-104.

47. March J, Silva S, Petrycki S, et al; Treatment for Adolescents With Depression Study (TADS) Team. Fluoxetine, cognitive-behavioral therapy, and their combination for adolescents with depression: Treatment for Adolescents With Depression Study (TADS) randomized controlled trial. *JAMA*. 2004; 292(7):807-820.

48. Silverman WK, Pina AA, Viswesvaran C. Evidence-based psychosocial treatments for phobic and anxiety disorders in children and adolescents. *J Clin Child Adolesc Psychol.* 2008;37(1):105-130.

49. American Diabetes Association; American Psychiatric Association; American Association of Clinical Endocrinologists; North American Association for the Study of Obesity. Consensus development conference on antipsychotic drugs and obesity and diabetes. *Diabetes Care.* 2004;27(2):596-601.