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Children And The Opioid Epidemic: Age-Stratified Exposures And Harms

Using North Carolina Medicaid 2016–18 claims data, we found that approximately one in ten adolescents (10.8 percent) filled at least one opioid prescription per year. Dentists, advanced practice providers, and surgeons were common prescribers of opioids to children. In addition, half of children who experienced opioid-related adverse events had filled opioid prescriptions in the prior six months.

mong adult patients admitted for opioid use disorder treatment, one-third report having their first opioid exposure in childhood, highlighting the importance of addressing early opioid exposures.^{1,2} Children's opioid-related hospitalizations and deaths have doubled or tripled in recent decades.^{3,4} Opioid exposures in childhood are responsible for the majority of drug-related pediatric fatalities and are potentially linked to opioid misuse in adulthood.^{1,5}

Up to 15 percent of children have at least one opioid prescription fill each year, and prescrip-

tions to children have been linked to subsequent opioid-related adverse events.⁶ Disparities in pediatric opioid exposures and opioid-related harms have been reported by age, race, urban/ rural status, and medical complexity.^{3,6-9}

In this study we characterized age-stratified opioid exposures, opioid-related harms, and disparities for North Carolina Medicaid-insured children. As shown in exhibit 1, we found that the yearly prevalence of exposures and harms among children was highest among older adolescents, with one in ten (10.8 percent) adolescents ages 15–17 having at least one opioid prescription fill per year and nearly 280 of every DOI: 10.1377/hlthaff.2020.00724 HEALTH AFFAIRS 39, NO. 10 (2020): -©2020 Project HOPE— The People-to-People Health Foundation, Inc.

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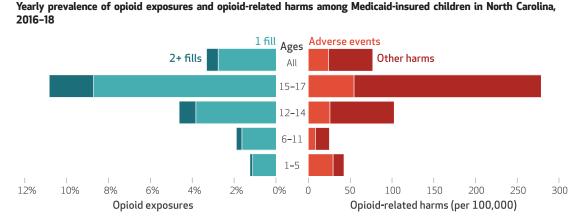
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EXHIBIT 1



SOURCE Authors' analysis of North Carolina Medicaid prescription claims, enrollment, inpatient and outpatient encounter, and provider files for 3,242,196 beneficiaries ages 1–17 enrolled at any point during the period 2016–18. **NOTES** Data from 2016–18 are combined. Yearly prevalence of opioid exposures is presented as percentage; yearly prevalence of harms is presented as rates per 100,000 children. "1 fill" and "2+ fills" are opioid prescription fills per year. Significant differences between age groups were identified for one fill, two or more fills, any fill, opioid-related adverse events (that is, "poisonings by, adverse effects of, or underdosing of opium"), and other opioid-related harms (that is, "opioid abuse," "opioid dependence," or "opioid use, unspecified"; *p* < 0.001) using chi-square tests.

100,000 children of that age experiencing one or more opioid-related harms each year.

Study Data And Methods

We conducted a cross-sectional analysis of 2016– 18 North Carolina Medicaid enrollment and medical/pharmacy claims data. Children ages 1–17 enrolled during the study period were included.

Outpatient opioid prescription fills (exposures) were identified in pharmacy claims data, using National Drug Codes for opioid type and National Provider Identification codes for prescriber type, categorized as physicians by specialty or as dentists or advanced practice providers, which includes nurse practitioners and physician assistants. See the online appendix for prescriber categories.¹⁰

Opioid-related harms were identified from inpatient and outpatient medical claims using International Statistical Classification of Diseases and Related Health Problems, Tenth Revision (ICD-10), codes and were classified as opioidrelated adverse events (for example, T40.0X, "poisoning by, adverse effect of and underdosing of opium") or other opioid-related harms (F11.X, "opioid abuse," "opioid dependence," or "opioid use, unspecified"), as shown in appendix exhibit A1.¹⁰ We included "opioid use, unspecified" under the harms category because most "opioid use, unspecified" subcategories include descriptors of harms (for example, "opioid use, unspecified with intoxication"). Also, documentation of an opioid use diagnosis, although potentially nonspecific, is likely associated with increased risk for opioid-related harm (that is, risk for poisonings or withdrawal) even if opioids are indicated. Enrollment data were used to characterize child age, sex, race, and urban-rural status. Chronic disease status was defined according to complex chronic conditions methodology among children with six months of continuous enrollment.9

The prevalence of opioid exposures and harms for children ages 1–17 was calculated using midyear enrollment as the denominator, presented for 2016–18. For index harms (that is, a harm not preceded by another harm within the previous six months) in children ages 6–17, the most recent opioid prescription fill was identified in the six months before diagnosis; results for children ages 1–5 were not reported because of small sample sizes. The sociodemographic and clinical characteristics of children with opioid exposures and harms were compared using chi-square tests.

This study had several limitations. First, Medicaid pharmacy claims for filled opioid prescrip-

tions likely underestimate children's opioid exposures because they do not capture other sources of exposure such as illicit opioids and household members' prescriptions.¹¹ Second, claims data also lack the clinical information necessary to assess prescription appropriateness or validate the inclusion of diagnoses captured. The inclusion of more ambiguous ICD-10 codes such as "opioid use, unspecified" may overestimate harms. Third, although approximately twothirds of our study population were eligible for the cohort requiring six months of continuous enrollment, results for the total study population (data not shown) were similar in direction and significance to those for the continuous enrollment cohort. Fourth, our Medicaid-specific findings from a single state might not be directly generalizable to all children, although 38 percent of US children are insured by Medicaid.¹² Finally, this cross-sectional analysis could not evaluate the causal association between opioid prescriptions and subsequent opioid-related harms as an adult.

Study Results

PREVALENCE OF EXPOSURES AND HARMS Among 3,242,196 children enrolled in North Carolina Medicaid during 2016–18, each year 3.3 percent had one or more opioid prescription fills, 24.4 per 100,000 experienced adverse events, and 52.2 per 100,000 experienced other opioid-related harms (exhibit 1). Significant differences between age groups were identified for all exposure and harm categories (p < 0.001), with adolescents ages 15-17 experiencing the highest prevalence of opioid prescription fills (10.8 percent), adverse events (54.9 per 100,000), and other opioid-related harms (223.5 per 100,000). Across all ages, the prevalence of one or more prescription fills per year decreased during the study period, as shown in appendix exhibit A2.¹⁰

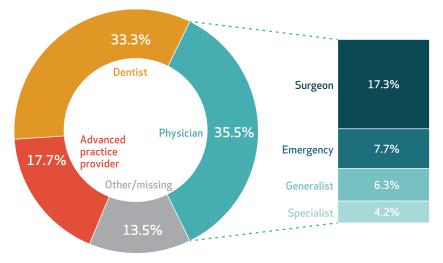
CHARACTERISTICS OF PEDIATRIC EXPOSURES AND HARMS Of 137,710 opioid prescription fills among children during 2016–18, prescribers were most commonly physicians (35.5 percent), dentists (33.3 percent), and advanced practice providers (17.7 percent) (exhibit 2). Exhibit 2 also shows the types of physicians represented; surgeons were the most common type. The most common opioid prescription fill types were hydrocodone (44.9 percent), oxycodone (28.7 percent), and codeine (20.0 percent) (data not shown).

The prevalence of pediatric opioid prescription fills in the cohort of children with six months of continuous enrollment (n = 2,019,211) increased significantly with the number of complex chronic conditions (from 2.7 percent for zero conditions to 15.8 percent for four or more conditions; exhibit 3). The prevalence of opioid prescription fills also was higher for White (3.1 percent versus 2.7 percent Black) and rural-dwelling (3.4 percent versus 3.1 percent suburban and 2.7 percent urban) children. Opioidrelated adverse events were more common in girls (26.2 per 100,000 versus 22.7 per 100,000 boys) and White children (27.5 per 100,000 versus 24.0 per 100,000 Black children), whereas other opioid-related harms were more common in boys (58.1 per 100,000 versus 45.9 per 100,000 girls) and Black (60.2 per 100,000 versus 51.7 per 100,000 White) and urban-dwelling (58.1 per 100,000 versus 49.0 per 100,000 suburban and 38.7 per 100,000 rural) children (exhibit 4).

For children ages 6–17, 48.4 percent of opioid adverse events were preceded by opioid prescription fills within the previous six months—and often within three days—with a higher proportion of Black versus White children having had a recent opioid prescription fill (exhibit 5). Oxycodone (26.6 percent) and hydrocodone (11.3 percent) were the most common opioid types.

EXHIBIT 2

Prescribers of opioids to Medicaid-insured children in North Carolina, 2016-18



SOURCE Authors' analysis of North Carolina Medicaid claims data for 137,710 opioid prescription fills for beneficiaries ages 1–17, 2016–18. **NOTES** The advanced practice provider category includes nurse practitioners and physician assistants. Significant differences between prescriber types and between physician specialties were identified (p < 0.001) using chi-square tests.

EXHIBIT 3

Yearly prevalence of opioid prescription fill exposures among Medicaid-insured children in North Carolina, by age group and sociodemographic and clinical characteristics, 2017–18

		Age, years				
Characteristics	All ages (N = 2,019,211)	1–5 (n = 661,169)	6–11 (n = 790,539)	12-14 (n = 347,255)	15-17 (n = 220,248)	
Sex Male Female	2.9% 3.0	1.3% 0.8	1.7% 1.6	4.5% 4.9	9.3% 11.6	
Race White Black Other	3.1 2.7 3.0	1.0 1.1 1.2	1.9 1.5 1.6	5.2 4.4 4.0	11.6 9.3 9.7	
Urban/rural statu Urban Suburban Rural	s 2.7 3.1 3.4	1.0 0.9 1.3	1.5 1.8 2.0	4.3 5.0 5.4	9.8 11.2 11.5	
Complex chronic 0 1 2 3 4 or more	conditions 2.7 6.5 9.9 12.8 15.8	0.9 3.0 5.5 8.0 11.2	1.5 4.4 9.3 13.9 16.6	4.4 7.7 12.2 16.5 21.1	10.0 14.6 17.6 20.4 30.4	

SOURCE Authors' analysis of North Carolina Medicaid claims data for beneficiaries ages 1-17 with at least six months of continuous enrollment before January 1 of calendar years 2017 and 2018. **NOTES** Calendar year 2016 was not analyzed because of the six-month continuous enrollment criterion. Within each age group, subgroups are significantly different (p < 0.001) using chi-square tests or Cochran-Armitage test (complex chronic conditions categories). Values reflect less than 1 percent missing data for race and urban/rural status.

EXHIBIT 4

		Opioid-related adverse events		Other opioid-related harms	
Characteristics	Total number	Number	Prevalence per 100,000	Number	Prevalence per 100,000
Sex			**		teletet
Male Female	1,654,687 1,587,509	376 416	22.7 26.2	962 729	58.1 45.9
Age, years			skalenkok		sininini
1-5 6-11 12-14 15-17	1,036,623 1,236,172 545,276 424,125	309 108 142 233	29.8 8.7 26.0 54.9	129 198 416 948	12.4 16.0 76.3 223.5
Race			steletok		sininink
White Black Other	1,586,869 1,161,046 489,765	436 279 77	27.5 24.0 15.7	820 699 161	51.7 60.2 32.9
Urban/rural status					telelet
Urban Suburban Rural	2,038,169 393,660 809,442	472 103 216	23.2 26.2 26.7	1,185 193 313	58.1 49.0 38.7

Opioid harms among Medicaid-insured children in North Carolina, by sociodemographic characteristics, 2016-18

SOURCE Authors' analysis of North Carolina Medicaid claims data for beneficiaries ages 1-17, 2016–18. **NOTES** Yearly prevalence is reported per 100,000 patients. For each harm type (adverse event, other opioid-related harms), differences within subgroups were evaluated using chi-square tests. Values reflect less than 1 percent missing data for race and urban/rural status. **p < 0.05 ****p < 0.001

EXHIBIT 5

Opioid prescription fill exposures before index opioid harms among Medicaid-insured children in North Carolina ages 6–17, 2016–18

Characteristics	Opioid-related adverse events (n = 417)	Other opioid- related harms (n = 1,186)
Total with opioid prescription fills in prior 6 months (%)	48.4	9.4
Age, years (%) 6-11 12-14 15-17	61.8 47.0 44.7	7.9 5.8 11.3
Race (%) White Black Other	35.7 59.9 53.4	11.1 7.7 n < 11ª
Most recent opioid type (for those with fill in prior 6 months) (% Oxycodone Hydrocodone) 26.6 11.3	3.3 3.2
Days since last fill (for those with fill in prior 6 months) Median (Quartile 1, quartile 3) Mean (Standard deviation)	3.0 (0.0, 19.0) 20.0 (36.7)	67.0 (5.0, 112.5) 68.5 (58.4)

SOURCE Authors' analysis of North Carolina Medicaid claims data for beneficiaries ages 1–17 with at least one opioid-related harm and six months of continuous Medicaid enrollment before the harm, July 2016–December 2018. **NOTES** Each cell in the age and race rows represents the proportion of children in that subgroup with an opioid prescription fill in the prior six months (for example, among those ages 6–11 with an opioid-related adverse event, 61.8 percent had a prior opioid prescription fill). Children ages 1–5 are not presented because of small sample sizes. ^aDenotes suppressed values resulting from data reporting restrictions.

Discussion

The results of this study further quantify how children have been impacted by opioids and identify disparities by sociodemographic and clinical characteristics. Our findings that more than half of the adverse events reported were preceded by a recent opioid prescription fill suggest a role that the children's own prescriptions may play in subsequent harms. Finally, we identified that children are prescribed opioids by several distinct types of clinicians.

Our findings that opioid exposures and harms disproportionately affect older adolescents compared with younger children are consistent with previous literature.⁶ The increased prevalence of opioid exposures among youth with chronic conditions (for example, cancer and sickle cell anemia) may reflect opioids being appropriately prescribed, as previously reported.⁹

Our findings that Black and urban children were less likely than their counterparts to fill opioid prescriptions or experience adverse events but more likely to experience other opioid-related harms (for example, abuse and dependence) increase the call for future studies to explore racial and geographic opioid-related inequities in children.^{3,7,8,13}

We identified preceding opioid prescription fills in almost 50 percent of adverse events, which may suggest a temporal relationship between children's own prescriptions and subsequent harms. In a 1999–2014 study in Tennessee, 89 percent of chart-adjudicated opioid adverse events were linked to children's prescriptions.⁶ Other adverse events may involve sources of opioids not captured in our data set; exposure to family members' prescriptions, for example, puts adolescents at increased risk for opioid overdose.¹¹ Relative to adverse events, fewer recent prescriptions were identified among youth with other opioid-related harms; these findings are similar to national data that indicate that about a quarter of adolescents ages 12–17 reported the source of their misused opioids as legitimate prescriptions.¹⁴

In addition to dentists, who prescribed approximately one-third of opioids to children,¹⁵ advanced practice providers and surgeons together accounted for another one-third of prescriptions. These three groups often prescribe opioids to children for postprocedural pain.^{15,16} The distinct and separate groups of clinicians who prescribe opioids to children suggest the need for pediatric opioid prescribing guidelines, particularly for postprocedural pain. Professional societies are well positioned to tailor general guidance for their clinicians and patients.¹⁷

Policy Implications

There is an urgent need to prioritize children in federal and state policy and public health measures addressing the opioid epidemic. In addition to pediatric-specific opioid prescribing guidelines, opioid-related harms may be reduced by the dissemination of best practices on opioid safe storage and disposal, naloxone prescribing, and substance use screening and treatment among youth.^{2,18} Stratifying population-level opioid surveillance reports by age and race is needed, given the distinct patterns of exposures and harms in different groups of children. Expected opioid pharmaceutical settlement payouts should support child-centered strategies that are racially and geographically equitable. ■

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