

Postpartum Follow-up Care for Pregnant Persons With Opioid Use Disorder and Hepatitis C Virus Infection

Marian Jarlenski, PhD, MPH, Qingwen Chen, MS, Katherine A. Ahrens, PhD, MPH, Lindsay Allen, PhD, MA, Anna E. Austin, PhD, Catherine Chappell, MD, MSc, Julie M. Donohue, PhD, Lindsay Hammerslag, PhD, Paul Lanier, PhD, MSW, Mary Joan McDuffie, MPH, Jeffrey Talbert, PhD, Lu Tang, PhD, and Elizabeth E. Krans, MD, MSc, on behalf of the Medicaid Outcomes Distributed Research Network (MODRN)

Hepatitis C virus (HCV) infections have risen alongside opioid use disorders (OUD) among people of reproductive age.¹⁻³ Enhanced engagement with health care during pregnancy provides an opportunity for persons with OUD to access treatment for

co-occurring disorders such as HCV infection.⁴ Guidelines recommend universal screening for HCV infection in pregnancy, with postpartum treatment for those with chronic HCV infection.⁵ Our objective was to estimate the prevalence of prenatal HCV infection testing and diagnosis and postpartum follow-up care among persons with OUD.

From the Department of Health Policy and Management, University of Pittsburgh School of Public Health, Pittsburgh, Pennsylvania; the Muskie School of Public Service, University of Southern Maine, Portland, Maine; the Department of Health Policy, Management, and Leadership, West Virginia University School of Public Health, Morgantown, West Virginia; the Department of Maternal and Child Health, UNC Gillings School of Global Public Health, Chapel Hill, North Carolina; the Department of Obstetrics, Gynecology and Reproductive Sciences, Magee-Womens Research Institute, University of Pittsburgh, Pittsburgh, Pennsylvania; the Department of Pharmacy Practice and Science, University of Kentucky College of Pharmacy, Lexington, Kentucky; the School of Social Work, University of North Carolina at Chapel Hill, Chapel Hill, North Carolina; the Center for Community Research and Service, University of Delaware, Newark, Delaware; and the Department of Biostatistics, University of Pittsburgh School of Public Health, Pittsburgh, Pennsylvania.

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Corresponding author: Marian Jarlenski, PhD, MPH, Department of Health Policy and Management, University of Pittsburgh, School of Public Health, Pittsburgh, PA; marian.jarlenski@pitt.edu.

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METHODS

We conducted a retrospective cohort study using administrative data from six states in the Medicaid Outcomes Distributed Research Network: Delaware, Kentucky, Maine, North Carolina, Pennsylvania, and West Virginia. The Medicaid Outcomes Distributed Research Network common data model contains a census of enrollment, inpatient and outpatient utilization, and pharmacy records and provides reliable measurement across participating sites.⁶⁻⁸ We included 23,780 people who had a live birth or stillbirth between 2016 and 2019 and a diagnosis of OUD during pregnancy who were followed for 60 days postpartum; a subset of 19,697 (87%) people were followed for 6 months postpartum. This study was approved by site-specific institutional review boards: University of Delaware, University of Kentucky, University of Southern Maine, University of North Carolina at Chapel Hill, University of Pittsburgh, and West Virginia University.

Outcomes included binary measures of HCV antibody or RNA laboratory screening in pregnancy, diagnosis of chronic or acute HCV infection in pregnancy, and *postpartum follow-up for HCV infection*, defined as either an outpatient visit for HCV infection or medication treatment with a direct-acting antiviral.



Table 1. Results From Random-Effects Meta-analysis of Average Predicted Probabilities of Hepatitis C Virus Infection Testing and Diagnosis in Pregnancy and Follow-up in the Postpartum Period Among Women With Opioid Use Disorder in Pregnancy, by Follow-up Period

Postpartum Follow-up Period	Outcome	Pooled Average Predicted Probability*†	95% CI†	90% PI‡	I ² (%)§
60 d	HCV infection testing	70.3	61.5–79.1	52.2–88.4	99.3
	HCV infection diagnosis	30.9	23.8–38.1	16.2–45.7	98.9
	Any follow-up visit or medication	3.2	2.6–3.8	2.1–4.3	76.3
6 mo	HCV infection testing	70.0	60.4–79.5	50.2–89.7	99.2
	HCV infection diagnosis	30.9	23.6–38.2	16.0–45.8	98.7
	Any follow-up visit or medication	5.9	4.9–6.9	4.0–7.8	80.6

PI, prediction interval.

* Pooled results calculated from random-effects meta-analysis pooling state-specific marginal probabilities (derived from state-specific multivariable regression models) and weighting by the inverse of the sum of within-state and between-state variances.

† Average predicted probabilities and related 95% CIs derived using marginal standardization from multivariable regression models adjusting for demographic (age, race and ethnicity, and urban vs rural residence) and clinical (mental health conditions, other nonopioid substance use disorders, medication for opioid use disorder in pregnancy, sequelae of injection drug use, and cirrhosis or other liver disease) characteristics.

‡ The 90% prediction intervals show a range within which predicted probabilities would fall for 90% of states were a different set of states chosen for analysis.

§ The I² statistic quantifies the percentage of variation attributable to cross-state heterogeneity rather than random chance.

Analyses controlled for demographic characteristics including age, race and ethnicity (non-Hispanic Black, non-Hispanic White, Hispanic, or none of these races), and urban compared with rural area of residence.⁹ Race and ethnicity are self-reported during Medicaid enrollment and are included as covariates because they may indicate differences in care due to structural or interpersonal racism.¹⁰ Clinical covariates included indicators for mental health conditions and substance use disorders, medication for OUD in pregnancy, sequelae of injection drug use, liver disease including cirrhosis, and anemia.

For each state, we fit three regression models in which the outcomes were HCV infection testing, HCV infection diagnosis, and postpartum HCV infection follow-up care. From these we derived average predicted probabilities and associated 95% CIs.¹¹ To produce pooled estimates, we conducted a random-effects meta-analysis.¹² We estimated between-state variability¹³ and calculated 90% prediction intervals, which provide a range in which estimates would be expected if we drew data from a different set of states. See Appendices 1–6, available online at <http://links.lww.com/AOG/C664>, for measurement and statistical details.

RESULTS

The pooled average predicted probability of HCV infection testing during pregnancy among people with OUD was 70.3% (95% CI 61.5–79.1), with 90% prediction intervals indicating between-state heterogeneity (52.2–88.4) (Table 1). The average predicted

probability of HCV infection diagnosis during pregnancy was 30.9% (95% CI 23.8–38.1), with 90% prediction intervals indicating heterogeneity (16.2–45.7). At 60 days postpartum, the average predicted probability of receiving any follow-up visit or medication for HCV infection was 3.2% (95% CI 2.6–3.8, 90% prediction interval 2.1–4.3); at 6 months postpartum, the average predicted probability of receiving any follow-up visit or medication for HCV infection was 5.9% (95% CI 4.9–6.9, 90% prediction interval 4.9–7.8).

DISCUSSION

Among Medicaid-enrolled pregnant people with OUD in six states, 70% were tested for HCV infection, 31% were diagnosed with HCV infection, and less than 6% had any follow-up visit or medication treatment within 6 months postpartum. Although HCV screening and diagnosis rates varied across states, postpartum follow-up rates were low. Limitations include reliance on HCV infection diagnosis in medical records rather than detection of HCV RNA in the blood and lack of data on postpartum lactation, which could prevent direct-acting antiviral initiation. Our results suggest a need for improved postpartum HCV infection treatment (or antenatal treatment if safety is established)¹⁴ to realize the public health potential of universal screening in pregnancy.

REFERENCES

1. Haight SC, Ko JY, Tong VT, Bohm MK, Callaghan WM. Opioid use disorder documented at delivery hospitalization -



- United States, 1999-2014. *MMWR Morb Mortal Wkly Rep* 2018;67:845-9. doi: 10.15585/mmwr.mm6731a1
2. Ly KN, Jiles RB, Teshale EH, Foster MA, Pesano RL, Holmberg SD. Hepatitis C virus infection among reproductive-aged women and children in the United States, 2006 to 2014. *Ann Intern Med* 2017;166:775-820. doi: 10.7326/M16-2350
 3. Zibbell JE, Iqbal K, Patel RC, Suryaprasad A, Sanders KJ, Moore-Moravian L, et al. Increases in hepatitis C virus infection related to injection drug use among persons aged ≤ 30 years - Kentucky, Tennessee, Virginia, and West Virginia, 2006-2012. *MMWR Morb Mortal Wkly Rep* 2015;64:453-8.
 4. Krans EE, Patrick SW. Opioid use disorder in pregnancy: health policy and practice in the midst of an epidemic. *Obstet Gynecol* 2016;128:4-10. doi: 10.1097/AOG.0000000000001446
 5. Society for Maternal-Fetal Medicine (SMFM), Dotters-Katz SK, Kuller JA, Hughes BL. Society for Maternal-Fetal Medicine Consult Series #56: hepatitis C in pregnancy—updated guidelines: Replaces Consult Number 43, November 2017. *Am J Obstet Gynecol* 2021;225:B8-18. doi: 10.1016/j.ajog.2021.06.008
 6. Medicaid Outcomes Distributed Research Network (MODRN), Donohue JM, Jarlenski MP, Kim JY, Tang L, Ahrens K, et al. Use of medications for treatment of opioid use disorder among US Medicaid enrollees in 11 states, 2014-2018. *JAMA* 2021; 326:154-64. doi: 10.1001/jama.2021.7374
 7. Adams L, Kennedy S, Allen L, Barnes A, Bias T, Crane D, et al. Innovative solutions for state Medicaid programs to leverage their data, build their analytic capacity, and create evidence-based policy. *EGEMS* 2019;7:41. doi: 10.5334/egems.311
 8. Jarlenski M, Kim JY, Ahrens KA, Allen L, Austin A, Barnes AJ, et al. Healthcare patterns of pregnant women and children affected by OUD in 9 state Medicaid populations. *J Addict Med* 2021;15:406-13. doi: 10.1097/ADM.0000000000000780
 9. U.S. Department of Agriculture. Rural-urban commuting area codes: overview. Accessed Sept 21, 2021. <https://www.ers.usda.gov/data-products/rural-urban-commuting-area-codes/>
 10. Williams DR, Lawrence JA, Davis BA. Racism and health: evidence and needed research. *Annu Rev Public Health* 2019;40:105-25. doi: 10.1146/annurev-publhealth-040218-043750
 11. Muller CJ, MacLehose RF. Estimating predicted probabilities from logistic regression: different methods correspond to different target populations. *Int J Epidemiol* 2014;43:962-70. doi: 10.1093/ije/dyu029
 12. Borenstein M, Hedges LV, Higgins JP, Rothstein HR. A basic introduction to fixed-effect and random-effects models for meta-analysis. *Res Synth Methods* 2010;1:97-111. doi: 10.1002/jrsm.12
 13. IntHout J, Ioannidis JP, Borm GF. The Hartung-Knapp-Sidik-Jonkman method for random effects meta-analysis is straightforward and considerably outperforms the standard DerSimonian-Laird method. *BMC Med Res Methodol* 2014; 14:25. doi: 10.1186/1471-2288-14-25
 14. ClinicalTrials.gov. Sofosbuvir/velpatasvir treatment of chronic hepatitis C during pregnancy (STORC). Accessed February 3, 2022. <https://clinicaltrials.gov/ct2/show/NCT05140941>
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