Estimated Savings From International Reference Pricing for Prescription Drugs

High prescription drug prices have important implications for health care spending,¹ patient financial burden,² and adherence.³ Prices for brand-name drugs in particular are higher in the US compared with other high-income coun-



Supplemental content

tries, most of which regulate drug prices. However, inconsistent availability of data on

net prices (ie, prices after rebates and other discounts) complicates international comparisons of drug prices. A 2021 study found that US prices for brand-name drugs were 344% of those in other high-income countries at manufacturer ("list") prices, but the difference was smaller (230%) after an adjustment to approximate lower US net prices.⁴

The Elijah E. Cummings Lower Drug Costs Now Act, HR 3, would allow the US Secretary of Health and Human Services to negotiate prices with drug manufacturers on behalf of Medicare and private insurers, up to a cap of 120% of prices in 6 countries (Australia, Canada, France, Germany, Japan, and the UK). Negotiation would apply to all insulins and at least 25 other single-source, brand-name drugs selected by the secretary in the first year and 50 in the second year. The Congressional Budget Office estimated that this application of international reference pricing, a price control tool used by many other countries, would save \$456 billion for Medicare alone over 10 years. ⁵

We estimated what 2020 national US savings would have been at HR 3 maximum international prices rather than US manufacturer and net prices for insulins and 50 top brandname drugs by sales.

Methods | We linked 2020 SSR Health product-level US net sales data to 2020 IQVIA MIDAS product-level estimates of national volume and sales at manufacturer prices for the US and

all 6 HR 3 countries. We identified insulins and the top 50 single-source brand-name products in terms of US net sales. We calculated US net prices by dividing product US net sales from SSR Health by US volume from MIDAS. We calculated what payments to drug manufacturers would have been if 2020 US volumes for study products were bought at US manufacturer prices (from MIDAS), US net prices (as described above), and international prices (defined as 120% of volume-weighted mean MIDAS manufacturer prices across HR 3 countries). We compared 2020 US spending for study products at different prices, first overall and then by therapeutic class (from SSR Health), to assess whether US net-to-international reductions differ by class. We analyzed data using Stata, version 16. See the Supplement for additional details.

Results | International reference pricing would have lowered 2020 US spending on study products by 52.3% or \$83.5 billion, from \$159.9 billion at US net prices to \$76.3 billion (Figure 1A). For comparison, US spending at manufacturer prices was \$240.1 billion. US net-to-international discounts were relatively larger and more tightly distributed (median [IQR] of 52.0% [24.8%]) compared with US net-to-manufacturer discounts (median [IQR] of 21.5% [48.0%]; Wilcoxon signed rank P < .001) (Figure 2).

US net-to-international discounts ranged from 44.4% to 57.3% across therapeutic classes (Figure 1B). Spending would have been 53.7% lower at international rather than US net prices for oncology drugs, for which US manufacturer-to-net discounts are relatively small. Spending would have been 44.4% lower at international prices for insulins, for which US manufacturer-to-net discounts are substantial.

Discussion | A national US net-to-international discount of 52% is lower than the estimate of the Congressional Budget Office of 68% for Medicare. ⁶ The current analysis does not

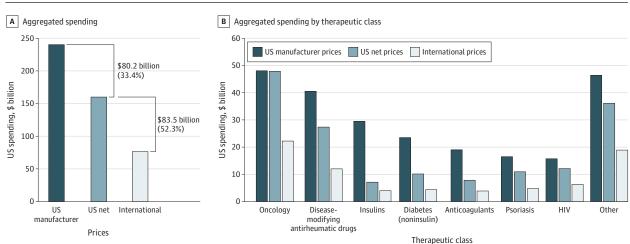


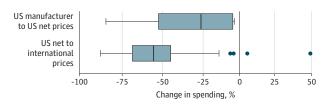
Figure 1. Aggregated Spending on Prescription Drugs at US Manufacturer, US Net, and International Prices

Analysis of SSR Health US net sales and IQVIA MIDAS sales at manufacturer prices and volume data for the US and 6 HR 3 countries. The analysis is limited to insulins (10 SSR Health products) and 50 select single-source, brand-name

drugs (including 2 SSR Health products assigned to the anticoagulant class, 4 noninsulin diabetes drugs, 4 HIV drugs, 4 psoriasis drugs, 5 disease-modifying antirheumatic drugs, 15 oncology drugs, and 16 other drugs).

1744

Figure 2. Product-Level Percent Changes in Spending at 2020 US Volume and Different Prices



The analysis was limited to 10 insulin products and 50 other select single-source, brand-name drugs. Vertical lines through the solid boxes represent medians, the widths of the solid boxes mark the IQR, the whiskers represent adjacent values, which are defined as the largest and smallest data points within 1.5 times the IQR, and dots indicate points that fall beyond the whiskers.

reflect several components of HR 3, including price negotiation, and Medicare vs national net prices may differ. The 52% discount is likely conservative due to data limitations, including use of net prices to manufacturers rather than to payers (including supply chain markups) and the absence of net price data for HR 3 countries. Furthermore, actual spending under HR 3 may be lower if Medicaid and other payers already face subinternational prices for some drugs. Although international reference pricing would yield considerable savings, other important considerations around the design and implementation of drug price regulation include incentives for research and development, industry launch and pricing strategies, and increasing utilization in response to lower prices.

Andrew W. Mulcahy, MPP, PhD Daniel Schwam, MA Preethi Rao, PhD Stephanie Rennane, PhD Kanaka Shetty, MD, MS

Author Affiliations: The RAND Corporation, Arlington, Virginia (Mulcahy, Schwam, Rao, Rennane); The RAND Corporation, Santa Monica, California (Shetty).

Corresponding Author: Andrew W. Mulcahy, MPP, PhD, 1200 S Hayes St, Arlington, VA 22202 (amulcahy@rand.org).

Accepted for Publication: July 26, 2021.

Published Online: September 10, 2021. doi:10.1001/jama.2021.13338

Author Contributions: Dr Mulcahy had full access to all of the data in the study and takes responsibility for the integrity of the data and the accuracy of the data analysis.

Concept and design: Mulcahy, Rennane, Shetty.

 $\label{lem:acquisition} \textit{Acquisition, analysis, or interpretation of data:} \ \textbf{All authors.}$

 ${\it Drafting~of~the~manuscript:}~{\it Mulcahy}.$

Critical revision of the manuscript for important intellectual content: All authors. Statistical analysis: Mulcahy.

Obtained funding: Mulcahy, Rennane.

Administrative, technical, or material support: Schwam, Rao, Rennane, Shetty. Supervision: Mulcahy.

Conflict of Interest Disclosures: None reported.

Funding/Support: This study was funded by Arnold Ventures.

Role of the Funder/Sponsor: The funder had no role in the design and conduct of the study; collection, management, analysis, and interpretation of the data; preparation, review, or approval of the manuscript; and decision to submit the manuscript for publication.

- 1. Keehan SP, Cuckler GA, Poisal JA, et al. National Health Expenditure projections, 2019-28: expected rebound in prices drives rising spending growth. *Health Aff (Millwood)*. 2020;39(4):704-714. doi:10.1377/hlthaff.2020.
- 2. Yang EJ, Galan E, Thombley R, et al. Changes in drug list prices and amounts paid by patients and insurers. *JAMA Netw Open*. 2020;3(12):e2028510-e2028510. doi:10.1001/jamanetworkopen.2020.28510
- 3. Kirzinger A, Lopes L, Wu B, Brodie M. KFF tracking poll February 2019: prescription drugs. *KFF Health*. March 1, 2019. Accessed May 17, 2021. https://www.kff.org/health-costs/poll-finding/kff-health-tracking-poll-february-2019-prescription-drugs/
- 4. Mulcahy A, Whaley C, Tebeka M, Schwam D, Edenfield N, Becerra-Ornelas A. International Prescription Drug Price Comparisons: Current Empirical Estimates and Comparisons With Previous Studies. RAND Corp; 2021. Accessed May 17, 2021. https://www.rand.org/pubs/research_reports/RR2956.html
- 5. Budgetary Effects of H.R. 3, the Elijah E. Cummings Lower Drug Costs Now Act. Congressional Budget Office; 2019. Accessed May 17, 2021. https://www.cbo.gov/publication/55936
- **6**. Adams C, Herrnstadt H. CBO's model of drug price negotiations under the Elijah E. Cummings Lower Drug Costs Now Act. Congressional Budget Office; February 2021. Accessed May 17, 2021. https://www.cbo.gov/publication/56905

Rates of Prenatal Cannabis Use Among Pregnant Women Before and During the COVID-19 Pandemic

Cannabis use among pregnant women is common and has increased in recent years in the US, from an estimated 3.4% in 2002 to 7.0% in 2017. Pregnant women report using cannabis to relieve stress and anxiety, and prenatal

+

Supplemental content

cannabis use may have risen during the COVID-19 pandemic as pregnant women

faced general and pregnancy-specific COVID-related stressors (eg, social isolation, financial and psychosocial distress, increased burden of childcare, changes in prenatal care, and concerns about heightened risks of COVID-19).^{3,4}

Considered an essential business in California, cannabis retailers remained open during the pandemic with record sales in 2020.⁵ We used data from Kaiser Permanente Northern California (KPNC), a large integrated health care delivery system with universal screening for prenatal cannabis use to test the hypothesis that rates of prenatal cannabis use increased during the COVID-19 pandemic.

Methods | The sample comprised all KPNC pregnant women screened for prenatal cannabis use via a universal urine toxicology test from January 1, 2019, through December 31, 2020, during standard prenatal care (at ≈8 weeks' gestation). The institutional review board of KPNC approved this study and waived the need for informed consent.

We computed monthly rates of prenatal cannabis use standardized to the year 2020 age and race and ethnicity distribution. We fit interrupted time-series (ITS) models to monthly rate data using negative binomial regression, adjusted for age (<25, 25 to <35, ≥35 years) and self-reported race and ethnicity (Asian/Pacific Islander, Black, Hispanic, non-Hispanic White, or other or unknown), which were included because of the known age and race and ethnicity differences in the prevalence of prenatal cannabis use. The prepandemic period was defined as urine toxicology tests conducted from January 2019 to March 2020 and the

JAMA November 2, 2021 Volume 326, Number 17