



Review article

Strengths and weaknesses of existing data sources to support research to address the opioids crisis

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ABSTRACT

Better opioid prescribing practices, promoting effective opioid use disorder treatment, improving naloxone access, and enhancing public health surveillance are strategies central to reducing opioid-related morbidity and mortality. Successfully advancing and evaluating these strategies requires leveraging and linking existing secondary data sources.

We conducted a scoping study in Fall 2017 at RAND, including a literature search (updated in December 2018) complemented by semi-structured interviews with policymakers and researchers, to identify data sources and linking strategies commonly used in opioid studies, describe data source strengths and limitations, and highlight opportunities to use data to address high-priority public health research questions.

We identified 306 articles, published between 2005 and 2018, that conducted secondary analyses of existing data to examine one or more public health strategies. Multiple secondary data sources, available at national, state, and local levels, support such research, with substantial breadth in data availability, data contents, and the data's ability to support multi-level analyses over time. Interviewees identified opportunities to expand existing capabilities through systematic enhancements, including greater support to states for creating and facilitating data use, as well as key data challenges, such as data availability lags and difficulties matching individual-level data over time or across datasets.

Multiple secondary data sources exist that can be used to examine the impact of public health approaches to addressing the opioid crisis. Greater data access, improved usability for research purposes, and data element standardization can enhance their value, as can improved data availability timeliness and better data comparability across jurisdictions.

1. Introduction

The United States is suffering its most serious drug-related public health crisis in a generation (Kolodny et al., 2015). Prescription opioid-related mortality rates increased by nearly 400% between 2000 and 2014; this period has also seen substantial increases in prevalence of opioid use disorder and rates of opioid-related hospitalizations (Dart et al., 2015; Han et al., 2015; Jones et al., 2015; Rudd et al., 2016; Tedesco et al., 2017). Heroin overdose deaths have more than quadrupled since 2010, and of the more than 47,000 opioid overdose deaths in 2017, nearly one-third involved heroin and over half involved

synthetic opioids (e.g., fentanyl) (Scholl et al., 2018). Multiple factors have contributed to the rise in opioid-related morbidity and mortality, and reducing the social and public health costs of opioid harms requires a multi-pronged approach (Cicero et al., 2015; Cicero et al., 2014; Kolodny et al., 2015; Lasser, 2017; Webster et al., 2011). To this end, the Department of Health and Human Services (HHS)¹ has identified five key strategies to combat the opioid crisis: 1) advancing better pain management practices; 2) improving addiction prevention, treatment, and recovery services; 3) promoting use of overdose reversing drugs; 4) strengthening data for better public health surveillance; and 5) supporting better research across the first four strategies (Price, 2017; U.S.

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¹ Selected abbreviations, see Appendix 2 for full abbreviations list. EHR = Electronic health record; EMS = Emergency medical services; HHS = Department of Health and Human Services; OEND = Overdose education and naloxone distribution; PDMP = Prescription drug monitoring program.

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Health and Human Services, xxxx).

Advancing these strategies often relies on analyses of non-clinical secondary data, yet researchers may be unaware of many available existing data sources (Sherman et al., 2016). Organized by the first four HHS strategies (Commission on Evidence-Based Policymaking, 2017), this review seeks to address this issue through identifying commonly used secondary data sources, the types of outcomes they are used to examine, their strengths and limitations, and promising data-linkage opportunities to support better research. Using a mixed-methods approach combining qualitative interviews with a scoping study to identify commonly used secondary data sources and data linkage strategies that could support better research, this article complements existing reviews of available data sources and metrics for studying prescription opioid use (Cochran et al., 2015; Schmidt et al., 2014; Secora et al., 2014).

2. Methods

We employed a multi-phase approach to synthesize information from the literature, opioid research experts, and policymakers as part of this HHS-funded study. We first conducted a scoping study, consistent with established methods (Arkskey and O'Malley, 2005; Levac et al., 2010), to identify commonly used data sources and data linking strategies in existing opioid research, focused on the United States context. The scoping study was complemented by semi-structured interviews with policymakers and opioid services and policy researchers to identify existing data source strengths and limitations, innovative uses of data and data linkages, and opportunities to use such data to address high-priority research questions. The RAND Institutional Review Board determined the project exempt.

To identify data sources, we searched for literature published between 2005 and 2017 through databases including PubMed, OVID, CINAHL, and PsycINFO using terms such as “opioid,” “buprenorphine,” “methadone,” and “naloxone,” as well as terms specific to opioid policy interventions such as “prescription monitoring program,” “pill mill,” and “Good Samaritan.” We used similar terms to conduct an internet search for relevant non-peer reviewed reports or presentations, and we reviewed additional articles and reports cited in key documents. We extracted information related to each document’s content, including research objective, outcome measures, and key variables and identified specific data sources, geographic coverage, time period, and data linkages in documents using empirical data. Data linkages were defined as any analysis combining data from multiple sources to study the same individual, policy, or geographic area.

The scoping study was complemented by 30-minute semi-structured interviews with sixteen opioid policy researchers and federal program officials conducted in August and September of 2017 (see Appendix for interview guide). Interviewees were selected by HHS officials to obtain a diverse set of perspectives. Discussions were tailored to the interviewees’ expertise and designed to gather insights on existing dataset strengths, limitations, and promising opportunities for dataset linkage. Research team members used detailed interview notes to identify common themes related to current dataset uses as well as potential opportunities to address key policymaker questions. In the twelve months subsequent to the interviews, the scoping study was updated to capture more recent literature published through December 2018, with particular attention to research questions, datasets, or data linkages previously identified as gaps by interviewees.

3. Review

The scoping study identified 446 articles and reports; 306 (68.6%) involved discussion or analyses of existing datasets; the remainder involved primary data collection or did not use empirical data (e.g., editorials, reviews). Existing datasets were wide-ranging but categorized generally as national surveys, electronic health records (EHR) and

claims, mortality records, prescription drug monitoring program (PDMP) data, contextual or policy data, and other national, state, or local data sources (e.g., national poison control center data, state arrest records). Interviewees discussed barriers or challenges in accessing datasets, their experiences linking datasets, and how datasets could be used to answer key research questions.

In Sections 3.1 through 3.4, we provide further detail on commonly used data sources, organized by HHS strategy. In the tables, we provide information on commonly used data sources and specific data elements, strengths and limitations for the different types of data, as well as data linking strategies for each HHS area. We subsequently highlight common topics arising during semi-structured interviews.

3.1. Advancing better pain management practices

An estimated 20% of non-cancer outpatients with pain receive opioid analgesics (Daubresse et al., 2013), chronic use of which increases risk of opioid use disorder (Boscarino et al., 2010; Chou et al., 2014) and opioid-related harms (Chou et al., 2015; Substance Abuse and Mental Health Services Administration, 2013). Researchers have commonly sought to identify the relationship between prescribing policy interventions, opioid analgesic prescribing and distribution, opioid-related overdose, and state- or community-level contextual factors (Table 1). In this section, we review measures, data sources, and linkages commonly used in this research, and we summarize common themes in this area from the interviews.

The most common studies of opioid prescribing interventions examine the impact of PDMPs on opioid analgesic prescribing and opioid-related overdose. Data regarding PDMP policies (Dave et al., 2017; Rutkow et al., 2015) commonly comes from the National Alliance for Model State Drug Laws (NAMSDL) or Prescription Drug Abuse Policy System (PDAPS) (Baehren et al., 2010; Bao et al., 2016; Buchmueller and Carey, 2018; Chang et al., 2016; Delcher et al., 2015; Gilson et al., 2012; Green et al., 2013; Li et al., 2014; Lin et al., 2017; Moyo et al., 2017; Pardo, 2017; Patrick et al., 2016; Paulozzi and Stier, 2010; Rasubala et al., 2015; Ringwalt et al., 2015a; Rutkow et al., 2015; Wen et al., 2017; Yarbrough, 2017), with additional information about PDMP components obtained from Temple University’s Policy Surveillance Program or Brandeis’ PDMP Training Technical Assistance Center (Buchmueller and Carey, 2018; Dave et al., 2017; Pardo, 2017; Patrick et al., 2016; Rasubala et al., 2015; Weiner et al., 2017a; Wen et al., 2017). Case studies of opioid prescribing guidelines or directives generally rely on data from site-specific implementation (Bujold et al., 2012; Chen et al., 2016; del Portal et al., 2016; Hansen et al., 2017; Johnson et al., 2011; Von Korff et al., 2016; Westanmo et al., 2015). Studies of other state prescribing regulations such as ID laws, continuing education requirements, doctor shopping laws, and physician exam requirements use CDC Public Health Law Program or original review of legal documents (Barber et al., 2017; Dave et al., 2017; Davis and Carr, 2016; Kuo et al., 2016; Popovici et al., 2017). Finally, studies evaluating the effects of Florida’s pill mill laws use information on the policy’s implementation (Chang et al., 2016; Kennedy-Hendricks et al., 2016; Rutkow et al., 2015).

Research examining opioid analgesic prescription characteristics, prescribing behavior, and dispensing patterns (Table 1) commonly uses prescription information from commercial (Cepeda et al., 2012, 2013a; Cepeda et al., 2013b; Chang et al., 2016; Dowell et al., 2016; Guy et al., 2017; Laroche et al., 2016; Liu et al., 2013; Qureshi et al., 2015; Rutkow et al., 2015; Schnell and Currie, 2018) and Medicaid pharmacy claims (Braden et al., 2010; Cochran et al., 2017; Garg et al., 2017; Hartung et al., 2017; Kim et al., 2016; Liu et al., 2013; Mack et al., 2015; Ray et al., 2016; Roberts et al., 2016; Turner and Liang, 2015; Wen et al., 2017; Yang et al., 2015), Medicare Part D Prescription Drug Event data (Buchmueller and Carey, 2018; Gellad et al., 2017; Hernandez et al., 2018; Kuo et al., 2016; Moyo et al., 2017; Willy et al., 2014; Yarbrough, 2017), Veterans Health Administration (VHA) data

Table 1
Secondary Data Sources to Support Research toward Advancing Better Pain Management Practices.

Data Elements (by Topic)	Sources	Strengths and Limitations
Policy data		
Prescribing interventions	<ul style="list-style-type: none"> ● PDAPS ● NAMSDL* ● CDC Public Health Law Program* 	<p>Strengths: + Can be linked with outcome data to examine state policy impact</p> <p>Limitations: - Some data not provided in analyzable format - May not fully capture heterogeneity in state laws - Some policy information not available historically for longitudinal analysis</p>
<ul style="list-style-type: none"> ● Pain clinic laws ● Education requirements ● Prescribing limits 		
EHR and claims data		
Opioid prescribing and distribution	Commercial claims	<p>Strengths: + Multi-payer and may include cash payments</p> <p>Limitations: - Not set up to track people long-term given insurance coverage transitions - Limited information on patient diagnoses or healthcare utilization - Difficult to link to outcomes (e.g., mortality)</p>
<ul style="list-style-type: none"> ● Opioid analgesic prescribing ● Prescription characteristics (opioid type, dose, days' supply, MED) ● Other prescriptions ● Payment 	<ul style="list-style-type: none"> ● Healthcare ● Marketscan ● IQVIA 	
Opioid-related overdose	Federal claims	<p>Strengths: + Can link hospital and pharmacy claims + Can look at Rx histories of patients who go to a hospital/ED for overdose</p> <p>Limitations: - Provides information on one population (Medicare or Medicaid enrollees) - Not set up to track people long-term given insurance coverage transitions - Cannot measure opioid mortality as provides date but not cause of death</p>
<ul style="list-style-type: none"> ● Diagnostic codes for nonfatal overdose 	<ul style="list-style-type: none"> ● Medicare data ● National or state Medicaid datasets 	
Detection of opioid misuse & morbidity		
<ul style="list-style-type: none"> ● Inpatient stays and ED visits ● Diagnoses and procedures ● Costs 		
	VHA data warehouse	<p>Strengths: + VHA data warehouse enables linkages across datasets + Has been linked to NDI</p> <p>Limitations: - Limited accessibility</p>
	HCUP (national and state inpatient and emergency department databases)	<p>Strengths: + Large collection of longitudinal data, nation-wide and state-level; free portal access to opioid-related data + State data is mapped to a standardized format</p> <p>Limitations: - Not all states participate in the databases - Costs to obtain full datasets</p>
Prescription drug monitoring data		
Opioid prescribing and distribution	<ul style="list-style-type: none"> ● State PDMPs ● PBSS ● ARCOS 	<p>Strengths: + Comprehensive data on distribution (ARCOS) or prescribing (PDMP) + PDMPs used to develop measures for patient/prescriber risk behaviors</p> <p>Limitations: - Access barriers - ARCOS not available in computable formats (i.e., in PDF form) - State capacity issues may limit ability to link PDMP data with other datasets - PDMP systems may lack unique IDs or have ID entry errors, creating issues in identifying individual-level matches</p>
<ul style="list-style-type: none"> ● Prescription name/type ● Prescription dose, days' supply, MED ● Prescriber ● Payment 		
Mortality data		
Opioid-related overdose	<ul style="list-style-type: none"> ● NDI ● NVSS MCOD ● CDC WONDER* ● State vital records 	<p>Strengths: + National data with information on opioid overdose mortality + CDC WONDER is readily downloadable and publicly available</p> <p>Limitations: - Lags in data availability - Variation in quality of reporting detail on drug involvement</p>
<ul style="list-style-type: none"> ● Cause of death ● Drugs involved in death ● Demographics 		
Contextual data		
Contextual factors ■ Unemployment rate ■ Physician density ■ Demographics	<ul style="list-style-type: none"> ● BEA*; CPS* ● BLS*; ACS* ● AHRF* ● CMS* 	<p>Strengths: + Allows analyses to control for state or county factors related to opioid analgesic use or opioid analgesic prescribing</p> <p>Limitations: - Lags in data availability</p>

* Publicly available at no cost.

(Barber et al., 2017; Bohnert et al., 2011; Edlund et al., 2007; Miller et al., 2015; Olivia et al., 2017; Park et al., 2015; Zedler et al., 2014), and PDMP data (Baehren et al., 2010; Becker et al., 2017; Delcher et al., 2015; Deyo et al., 2017; Dowell et al., 2016; Gilson et al., 2012; Gwira Baumbblatt et al., 2014; Hall et al., 2008; Katz et al., 2010; Kreiner et al., 2017; Mercado et al., 2018; Rasubala et al., 2015; Ringwalt et al., 2015a; Roberts et al., 2016). PDMP studies usually entail single-state analyses, although the Prescription Behavior Surveillance System (PBSS), which compiles PDMP data from multiple states (Paulozzi et al., 2015), has allowed for multi-state comparisons of opioid misuse indicators. Several studies have examined state-level opioid analgesic distribution using the Automation of Reports and Consolidated Orders System (ARCOS) (Alpert et al., 2016; Brady et al., 2014; Paulozzi and Stier, 2010; Reisman et al., 2009).

To examine the relationship between opioid analgesic use and overdose, studies use person-level mortality records from the National

Death Index (NDI) (Bohnert et al., 2016; Bohnert et al., 2011; Park et al., 2015) or state death certificate data (Dasgupta et al., 2016; Dunn et al., 2010; Garg et al., 2017; Gwira Baumbblatt et al., 2014; Hall et al., 2008; Hirsch et al., 2014; Mercado et al., 2018; Ray et al., 2016) and opioid-related toxicity or overdose event measures from Medicare (Buchmueller and Carey, 2018; Kuo et al., 2016), commercial claims (Braden et al., 2010; Laroche et al., 2016; Turner and Liang, 2015), Medicaid (Cochran et al., 2017; Yang et al., 2015), and VHA databases (Miller et al., 2015; Zedler et al., 2014). Other research examines aggregate state- or county-level rates of fatal opioid overdose using state death certificate data (Kennedy-Hendricks et al., 2016), the National Vital Statistics System Multiple Cause of Death (NVSS MCOD) microdata (Alpert et al., 2016; Dowell et al., 2016; Li et al., 2014), and CDC WONDER (Compton et al., 2016; Gomes et al., 2018; Pardo, 2017; Patrick et al., 2016; Rigg et al., 2018).

To evaluate contextual factors related to opioid prescribing or

Table 2
Secondary Data Sources to Support Research on Improving Prevention, Treatment, and Recovery Services.

Data elements (by Topic)	Sources	Strengths and limitations
EHR and claims data		
Opioid misuse or use disorders	Commercial claims	Strengths: + Prescription data can capture the population treated with buprenorphine
<ul style="list-style-type: none"> • Opioid use disorder diagnosis • Opioid-related inpatient stays and ED visits 	<ul style="list-style-type: none"> • IQVIA • Marketscan • Symphony Health 	Limitations: - Limited information on patient diagnoses or other healthcare utilization
Treatment demand & utilization		- Requires triangulating with other sources to fully assess treatment need
<ul style="list-style-type: none"> • Buprenorphine prescriptions • Payment • Monthly prescriber patient census 	National or state Medicaid datasets	Strengths: + Can link hospital and pharmacy claims
Individual-level risk factors		+ Single-state analyses have linked to death data
<ul style="list-style-type: none"> • Other Rx use or healthcare utilization • Socio-demographics; comorbidities 	VHA data warehouse	Limitations: - Only provides information on Medicaid enrollees
		- Misses those receiving other publicly funded substance abuse treatment
	HCUP (national and state inpatient and emergency department databases)	Strengths: + Facilitates linkage to treatment facility-level variables
		+ Has been linked to NDI
		Limitations: - Limited accessibility and specific population
		Strengths: + Large collection of longitudinal data, nation-wide and state-level; free portal access to opioid-related data
		+ State data is mapped to a standardized format
		Limitations: - Not all states participate in the databases
		- Costs to obtain full datasets
National surveys		
Opioid misuse or use disorders	Household surveys	Strengths: + National data with rich information on substance use & mental health
<ul style="list-style-type: none"> • Nonmedical use of opioids • Opioid use disorder symptoms 	<ul style="list-style-type: none"> • NSDUH* • NESARC 	+ NSDUH 2015 redesign asks about any pain reliever use (not only misuse)
Treatment demand & utilization		Limitations: - Does not ask about medications used for treatment or treatment retention
<ul style="list-style-type: none"> • Opioid use disorder treatment • Source of payment 		- Screens for use disorder symptoms, but does not ask about formal diagnosis
Individual-level risk factors		- Sample may miss high-risk populations (e.g., homeless, arrestees)
<ul style="list-style-type: none"> • Mental health, substance use • Socio-demographics 		- State identifiers restricted
Treatment demand & utilization	Treatment facility surveys	Strengths: + National data on admissions to treatment & public-sector specialty care
<ul style="list-style-type: none"> • # treatment admissions • # patients receiving methadone in OTPs (N-SSATS) • Referral source 	<ul style="list-style-type: none"> • TEDS* • N-SSATS* 	+ TEDS has patient demographic data
Treatment supply & capacity (N-SSATS only)		+ Up to 3 drugs of abuse listed (differentiate heroin & opioid analgesics)
<ul style="list-style-type: none"> • Treatment facility characteristics • Estimated operating capacity 		+ N-SSATS includes both public and private facilities
		Limitations: - TEDS only includes agonist treatments; cannot differentiate MAT types
		- Limited information on payment
		- Quality control issues with TEDS, as states may not consistently report on similar patients or have consistent procedures to assess data quality
		- TEDS data do not include private for-profit treatment facilities
Mortality data		
Opioid-related overdose	<ul style="list-style-type: none"> • NDI • NVSS MCODE • CDC WONDER* • State vital records 	Strengths: + National data with information on opioid overdose mortality
<ul style="list-style-type: none"> • Cause of death • Drugs involved in death • Demographics 		+ CDC WONDER is readily downloadable and publicly available
Other national data sources		Limitations: - Lags in data availability
Treatment supply & capacity	Provider censuses	- Variation in quality of reporting detail on drug involvement
<ul style="list-style-type: none"> • Waivered physicians • Patient caps • Physician address, ZIP 	<ul style="list-style-type: none"> • SAMHSA database* • DEA ACSA 	Strengths: + Measures supply/capacity of waivered physicians at geographic detail
Policy data		+ Can link to AMA Physician Masterfile
Treatment policies	<ul style="list-style-type: none"> • RAND/NCSL • ASAM 	Limitations: - Costs to obtain DEA ACSA
<ul style="list-style-type: none"> • Medicaid coverage information • Formulary placement • Copays, prior authorization, etc. 		- SAMHSA publicly available data captures around 55% of physicians
Contextual data		
Contextual factors	<ul style="list-style-type: none"> • BEA* • AHRF* 	Strengths: + Can be linked to outcomes to examine effects of state policies
<ul style="list-style-type: none"> • Physician density • Hospital beds per capita • State or county economic factors 		Limitations: - Collected through retrospective surveys, thus potentially inaccurate
		- Data is missing for some states
		Strengths: + Can control for state or county factors related to healthcare access or treatment need
		Limitations: - Lags in data availability

* Publicly available at no cost.

opioid-related harms, studies commonly include state- or county-level measures of the unemployment rate and income per capita from the Bureau of Economic Analysis (BEA) (Dave et al., 2017), Bureau of Labor Statistics (BLS) (Patrick et al., 2016), or American Community Survey (ACS) (Guy et al., 2017; Schnell and Currie, 2018; Yarbrough, 2017); information on physician density and demographics from the Area Health Resource Files (AHRF) (Dave et al., 2017; Guy et al., 2017); and rates of health insurance coverage from the Current Population Survey (CPS) or Centers for Medicare & Medicaid Services (CMS) (Guy et al., 2017; Wen et al., 2017; Yarbrough, 2017).

To examine how policies or community factors influence pain management practices, studies link state policy data and state- or county-level contextual factors to data on opioid prescribing (Brady et al., 2014; Buchmueller and Carey, 2018; Haffajee et al., 2018; Kuo et al., 2016; Moyo et al., 2017; Wen et al., 2017; Yarbrough, 2017) or overdose mortality records (Dowell et al., 2016; Li et al., 2014; Pardo, 2017; Patrick et al., 2016). Research examining potentially inappropriate prescribing generally links opioid prescription data with opioid overdose data at the person level. These include studies linking PDMP data with Medicaid claims (Hartung et al., 2017; Kim et al.,

2016), hospital discharges (Baehren et al., 2010; Deyo et al., 2017), death certificates or toxicology reports (Albert et al., 2011; Deyo et al., 2017; Gwira Baublatt et al., 2014; Mercado et al., 2018), or data capturing state medical board actions (Kreiner et al., 2017); analyses of multiple linked VHA databases (Bohnert et al., 2011; Gellad et al., 2017; Westanmo et al., 2015); and research linking Medicaid claims with state vital records, hospital discharge data, or the NDI (Garg et al., 2017; Massachusetts Department of Public Health, 2016; Olfson et al., 2018; Ray et al., 2016).

3.1.1. Common interview themes

The insufficient understanding of factors influencing opioid analgesic use and subsequent outcomes was a common theme, with interviewees noting a paucity of empirical research examining how changes in opioid prescribing guidelines, pain reimbursement policies, or clinician education protocols influence treatment of pain and subsequent risk for opioid misuse, addiction, and overdose. While recent studies have examined the impact of opioid prescribing guidelines within a single state (Gillette et al., 2018; Tenney et al., 2019; Weiner et al., 2017b), the absence of systematically collected information on how guidelines are being implemented across states and over time complicates identification of the policy features that are effective. Interviewees also stressed the need for additional research examining longer-term effectiveness of opioid and non-opioid analgesic interventions for chronic pain given questions about the comparative effectiveness of opioid analgesics in managing some types of chronic pain (Krebs et al., 2018; Krebs et al., 2010).

Additional common themes were the need for analyses of provider- or hospital-level opioid prescribing patterns to identify factors underlying provider- or practice-level variation in risky or inappropriate prescribing, and the need for longitudinal patient-level analyses with sufficient temporal coverage to examine the pathways and sequences of events associated with adverse outcomes following opioid analgesic prescribing. Interviewees also frequently observed that all-payer claims databases, such as that developed by Massachusetts (Massachusetts Department of Public Health, 2017), may facilitate important longitudinal analyses that unlike Medicaid and commercial claims can track individuals as they transition across different types of insurance or across plans within a given insurance type.

3.2. Improving addiction prevention, treatment, and recovery services

Despite considerable improvement in the availability of medication-assisted treatment (Volkow et al., 2014) substantial gaps between opioid use disorder treatment need and capacity persist (Feder et al., 2017b; Dick et al., 2015; Hadland et al., 2017; Jones et al., 2015; Morgan et al., 2018; Saloner and Karthikeyan, 2015). In this section, we provide information about measures, data sources, and data linkages commonly used to study prevalence of opioid misuse or use disorders, treatment demand and utilization, supply and capacity of treatment, treatment policies, and contextual factors associated with treatment need and access (Table 2), and we summarize common themes in this area from the interviews.

Self-reported measures of opioid misuse or opioid use disorder symptoms come from national household surveys such as the National Survey on Drug Use and Health (NSDUH) and National Epidemiologic Survey on Alcohol and Related Conditions (NESARC) (Becker et al., 2008; Compton et al., 2016; Martins et al., 2012; McCabe et al., 2008; Rigg and Monnat, 2015; Secora et al., 2014). The NSDUH's information on self-reported receipt of and need for opioid use disorder treatment has also informed research on treatment need and utilization trends (Becker et al., 2008; Feder et al., 2017a; Han et al., 2015; Jones, 2017; Jones et al., 2015; Saloner and Karthikeyan, 2015; Wu et al., 2016). Commercial and Medicaid claims data (Braden et al., 2010; Edlund et al., 2014; Liu et al., 2013; Ray et al., 2016; Turner and Liang, 2015), Veterans Health Administration data (Bohnert et al., 2011; Edlund

et al., 2007), inpatient and emergency department databases (Guy et al., 2018; Tedesco et al., 2017), and electronic health records (Boscarino et al., 2010; Carrell et al., 2015; PCOR, 2018) are also used to estimate rates of potential opioid misuse or opioid use disorders. These data sources are also commonly used to examine person-level sociodemographic and clinical risk factors associated with development of opioid use disorder (Becker et al., 2008; Bohnert et al., 2011; Braden et al., 2010; Compton et al., 2016; Edlund et al., 2014; Edlund et al., 2007; Martins et al., 2012; McCabe et al., 2008; Ray et al., 2016; Rigg and Monnat, 2015; Secora et al., 2014; Turner and Liang, 2015).

Opioid use disorder treatment rates have often been studied using the National Survey of Substance Abuse Treatment Services Data (N-SSATS) and the Treatment Episodes Data Set (TEDS) (Ducharme and Abraham, 2008; Feder et al., 2017b; Jones et al., 2015; Martin et al., 2015; Saloner et al., 2016). Analyses of treatment trajectories, variation in buprenorphine utilization, quality of care and patient adherence to buprenorphine, as well as buprenorphine providers' patient censuses (Stein et al., 2016), instead generally use commercial or Medicaid claims (Baxter et al., 2015; Gordon et al., 2015; Lo-Ciganic et al., 2016; Morgan et al., 2018; Saloner et al., 2017; Stein et al., 2012; Stein et al., 2016; Turner et al., 2013; Turner et al., 2015).

Research describing national trends and geographic variation in treatment supply and capacity often uses SAMHSA's Buprenorphine Waiver Notification System (Dick et al., 2015; Stein et al., 2015a; Stein et al., 2015b) or the DEA's Active Controlled Substances Act Registrants Database (ACSA) (Andrilla et al., 2019; Knudsen, 2015; Rosenblatt et al., 2015) to examine the supply of buprenorphine waived physicians, while studies assessing the capacity of opioid treatment programs or availability of various types of medication-assisted treatment use N-SSATS state- or county-level data (Dick et al., 2015; Ducharme and Abraham, 2008; Jones et al., 2018; Jones et al., 2015; Stein et al., 2015b).

Studies of state Medicaid policies' effects on treatment access and utilization of methadone and buprenorphine commonly use policy information from the RAND/National Conference of State Legislatures (RAND/NCSL) Survey (Burns et al., 2016; Stein et al., 2015a) or the American Society of Addiction Medicine (ASAM) survey of Medicaid programs (Rinaldo and Rinaldo, 2013; Saloner et al., 2016), while research examining state- or county-level factors related to treatment supply or demand often use BEA or AHRF measures of the unemployment rate and income per capita (Dick et al., 2015; Knudsen, 2015; Stein et al., 2015a); and AHRF information on physician density, percent of adults uninsured, hospital beds per capita, and urbanicity (Dick et al., 2015; Stein et al., 2015a; Stein et al., 2015b).

To examine state and community-level factors associated with treatment utilization or supply, studies often link policy and contextual data sources at the state or county level to outcome data on the location of buprenorphine waived physicians or buprenorphine use (IMS Institute for Healthcare Informatics, 2016; Knudsen, 2015; Saloner et al., 2016; Stein et al., 2015a; Stein et al., 2012). Others link aggregate measures of treatment need with measures of treatment capacity to identify areas with treatment shortages (Dick et al., 2015; Jones et al., 2015).

3.2.1. Common interview themes

Interviewees frequently noted that most existing data sources do not contain information on block grant funded treatment, thereby providing only a partial picture of treatment utilization, and limiting accurate identification of treatment shortage areas. Interviewees also observed that current analyses of treatment patterns (i.e., patient or provider trajectories) are commonly unable to track individuals across insurance coverage transitions. Interviewees stressed the need to better understand the effects of opioid use disorder treatment quality on outcomes, studies for which EHRs can complement claims data (Campbell et al., 2019; Garnick et al., 2012; Haddad et al., 2015). Finally, interviewees highlighted the need for further study of opioid use

Table 3
Secondary Data Sources to Support Research Promoting Use of Overdose-Reversing Drugs.

Data Elements (by Topic)	Sources	Strengths and Limitations
Policy data		
Naloxone policies	<ul style="list-style-type: none"> ● PDAPS ● NAMSDDL* ● NCSL* 	Strengths: + Can be linked with data on opioid outcomes to examine state policy impact Limitations: - May not capture state variation in nominally identical naloxone policies - Data on EMS protocols not readily available - Some data not provided in readily analyzable format
Mortality data		
Opioid overdose mortality	<ul style="list-style-type: none"> ● CDC WONDER* ● NVSS MCODE 	Strengths: + National data with information on opioid overdose mortality + CDC WONDER is readily downloadable and publicly available Limitations: - Lags in data availability - Variation in quality of reporting detail on drug involvement due to differences across states in rigor of medical examiner/coroner procedures
<ul style="list-style-type: none"> ● Opioid analgesic, heroin, or synthetic overdose deaths ● Age, gender, race/ethnicity ● State or county 		
EHR and claims data		
Naloxone distribution	Pharmacy claims <ul style="list-style-type: none"> ● IQVIA ● Symphony Health VHA data warehouse	Strengths: + Measures pharmacy distribution of naloxone Limitations: - Only captures the distribution of naloxone via pharmacy channel; does not capture purchase and distribution via state or community programs Strengths: + Rich information on patient characteristics + Able to examine naloxone refills and renewals Limitations: - Limited accessibility
<ul style="list-style-type: none"> ● Naloxone prescriptions ● Prescriber specialty ● Patient age and gender ● Naloxone formulation 		
Other national and local sources		
Naloxone distribution	OEND Program Data <ul style="list-style-type: none"> ● MA OOP Pilot Program ● Harm Reduction Coalition 	Strengths: + Fills in some gaps regarding naloxone distributed via state or local programs Limitations: - Data collection on OEND programs not standardized - National data not systematically collected, updated, or made publicly available
<ul style="list-style-type: none"> ● # persons trained ● # naloxone kits provided ● # overdose reversals 		
Other national sources		
Naloxone distribution	EMS data <ul style="list-style-type: none"> ● NEMSIS* 	Strengths: + Naloxone administration is a fairly high-quality variable + Can do small area analysis Limitations: - Not a registry of patients receiving care - Data quality differs across agencies/states - Some elements restricted; contains no diagnosis information - Barriers to linking
<ul style="list-style-type: none"> ● EMS naloxone administration ● EMS provider level ● 911 call info ● Information on incident and transport 		
Contextual data		
Contextual factors	<ul style="list-style-type: none"> ● CPS* ● BLS* ● US Census* ● PDAPS ● NAMSDDL* 	Strengths: + Can control for state or county factors associated with opioid mortality Limitations: - Lags in data availability - Policy data often not available in readily analyzable format
<ul style="list-style-type: none"> ● Other opioid-related policies ● State or county-level demographics, socioeconomics 		

* Publicly available at no cost.

disorder treatment among justice-involved individuals (Acevedo et al., 2015; Garnick et al., 2014; Krawczyk et al., 2017), likely requiring linked substance abuse treatment and arrest or incarceration databases.

3.3. Promoting use of overdose-reversing drugs

Overdose-reversing drugs, such as naloxone, play a critical role in opioid overdose prevention (Boyer, 2012; Davis and Carr, 2015; van Dorp et al., 2007). In this section, we describe measures, data sources, and data linkages used to describe policies to promote naloxone distribution and use, and to evaluate how naloxone policies or programs relate to naloxone distribution, opioid overdose mortality, and contextual factors.

Information on state naloxone policies regarding use by community bystanders, emergency medical services (EMS) personnel, and other first responders is generally drawn from original reviews of legal databases (Brodrick et al., 2016; Burris et al., 2017; Davis and Carr, 2015; Davis et al., 2014a), with some groups, such as PDAPS, compiling data on the timing and provisions of certain laws into a single source (Table 3).

Studies of community-based overdose education and naloxone distribution (OEND) programs (Clark et al., 2014; Giglio et al., 2015; Haegerich et al., 2014; Kerensky and Walley, 2017; Mueller et al., 2015) commonly rely on surveys of OEND program participants, including reported overdose reversals, number of naloxone administrations, number of naloxone kits distributed, and overdose response, collected by OEND programs (Bennett et al., 2011; Doe-Simkins et al., 2014; Enteen et al., 2010; Jones et al., 2014b; Oliva et al., 2016; Walley et al., 2013a; Walley et al., 2013b; Wheeler et al., 2012; Wheeler et al.,

2015). National data on the locations of OEND programs has been compiled by the Harm Reduction Council, but the data are not publicly available (Lambdin et al., 2018a; Lambdin et al., 2018b). Fewer studies have examined retail pharmacy naloxone dispensing using pharmacy claims (e.g., Symphony Health, IQVIA) (Freeman et al., 2018; Jones et al., 2016; Xu et al., 2018) or EMS naloxone administration using National EMS Information System (NEMSIS) data to examine trends and geographic variation in naloxone distribution (Cash et al., 2018; Faul et al., 2015; Faul et al., 2017). Another set of studies evaluated naloxone prescribing through the VHA OEND program (Bounthavong et al., 2017; Oliva et al., 2017).

To examine how state naloxone policies or local OEND programs influence mortality, multi-state analyses generally use state-level data on opioid overdose mortality from the NVSS MCODE microdata or CDC WONDER (Frank and Pollack, 2017; Pardo, 2017; Rees et al., 2017; Wheeler et al., 2015), while single-state analyses more commonly use state- or county-level measures collected from state death certificates (Albert et al., 2011; Burrell et al., 2017; Maxwell et al., 2006; Walley et al., 2013b).

Studies of state naloxone policies' effects on opioid overdose generally merge state-level opioid overdose mortality data with information on state naloxone policies (Pardo, 2017; Rees et al., 2017); other community-level contextual factors, such as unemployment rates or per capita income from the CPS or US Census (Pardo, 2017; Rees et al., 2017; Walley et al., 2013b); and information about other state opioid policies (e.g., pain clinic laws) from PDAPS, the Policy Surveillance Program, or NAMSDDL (Pardo, 2017; Rees et al., 2017). Studies of the impact of OEND programs instead often use multiple complementary datasets, including parallel analyses of trends in emergency department

visits, fatal accident poisonings, and outpatient-dispensed controlled substances (Albert et al., 2011; Walley et al., 2013b). Sub-county level studies using linked data are rare. One study linked police naloxone use to EMS data to assess the proportion of cases in which EMS administered additional naloxone doses (Fisher et al., 2016), while another single-county study mapped naloxone-carrying pharmacies with overdose death data at the ZIP Code level (Burrell et al., 2017).

3.3.1. Common interview themes

Interviewees frequently noted that more systematic collection of data on naloxone distribution outside of outpatient pharmacy channels would further understanding of naloxone access barriers and inform effective approaches for distribution and use. Interviewees also discussed how determining optimal naloxone dosing, particularly in the context of more widespread use of synthetic opioids (Frank and Pollack, 2017), would benefit from better data about naloxone reversals and the surrounding circumstances. Several interviewees noted the potential value of EMS data (Table 3), but observed that variation in EMS data quality and completeness across agencies and regulatory barriers precluding individual level linkages currently limit its value, as analyses of EMS naloxone administration and subsequent patient outcomes are often confined to a single jurisdiction (Belz et al., 2006; Knowlton et al., 2013; Levine et al., 2016; Ray et al., 2018). Many interviewees also noted the potential value of longitudinal studies linking data on persons receiving naloxone with claims data, which would enable researchers to follow individuals through the health care system.

3.4. Strengthening data for better public Health surveillance

The rapid evolution of opioid use and markets has generated efforts to improve data collection and surveillance tools to monitor medical and non-medical opioid use. In this section, we describe measures, data sources, and linkages used to study opioid surveillance topics not discussed extensively in the sections above, including detection of misuse, product-specific use and emerging trends, toxico-surveillance, and illicit markets (Table 4), and we summarize common themes in this area from the interviews.

State PDMP data systems, now present to some degree in all 50 states, are increasingly being used to develop risk indicators for inappropriate prescriber behavior (Kreiner et al., 2017; Porucznik et al., 2014; Ringwalt et al., 2015b) and to detect inappropriate or problematic patterns in opioid analgesic prescribing, dispensing, and use (Katz et al., 2010; O'Kane et al., 2016; U.S. Department of Health and Human Services and Behavioral Health Coordinating Committee, 2013). EHR data is also used to improve surveillance of problematic opioid use and opioid-related harms (Olivia et al., 2017), occasionally using natural language processing to text mine clinicians' notes (Canan et al., 2017; Carrell et al., 2015).

Proprietary databases, such as RADARS and NAVIPPRO, are also being used for near-real-time surveillance of opioid use. RADARS consists of several programs that collect and compile data on product-specific drug diversion and nonfatal overdose, opioid use and treatment, and street drug prices (Bau et al., 2016; Butler et al., 2013; Cassidy et al., 2014; Cepeda et al., 2017; Cicero et al., 2007; Coplan et al., 2016; Dart et al., 2015; Davis et al., 2014b; Inciardi et al., 2009; Secora et al., 2014). NAVIPPRO collects and compiles information on product-specific opioid use, initiation, route of administration, and source of opioids from two proprietary systems and several publicly available data sources (Butler et al., 2018; Butler et al., 2008; Butler et al., 2013; Cepeda et al., 2017; Coplan et al., 2016; Secora et al., 2014). Non-traditional data resources such as Twitter, web forum postings, Google trends, and cryptomarket forums on the Dark Web are also drawing attention as means to bolster public health surveillance, better understand opioid misuse and prescription drug diversion (Anderson et al., 2017; Chan et al., 2015; Katsuki et al., 2015), forecast state-level mortality or nonfatal overdose (Parker et al., 2017; Young

et al., 2018), and assess emerging trends in new psychoactive substances (Van Hout and Hearne, 2017).

RADARS data on diversion has been used to examine illicit pharmaceutical opioid markets (Coplan et al., 2016; Dart et al., 2015; Inciardi et al., 2009), and NSDUH (Inciardi et al., 2009; Jones et al., 2014a) and NAVIPPRO (Cassidy et al., 2014) includes information on self-reported sources of prescription opioids for nonmedical use. While national data on drug seizures, drug testing, and illicit drug prices that could be used to examine trends and geographic variation in illicit opioid markets exist in the National Forensic Laboratory Information System (NFLIS) or System to Retrieve Information from Drug Evidence (STRIDE) (National Academies of Sciences Engineering and Medicine, 2017; Secora et al., 2014), we identified few empirical analyses using these measures (Rosenblum et al., 2014; Stein et al., 2015a), and found local or state law enforcement databases to be more common sources of drug seizures and arrest data (Bujold et al., 2012; Piper et al., 2016; Ray et al., 2017).

While mortality microdata help monitor drug overdose mortality and polysubstance involvement in fatal overdose (Jalal et al., 2018; Kandel et al., 2017), concerns about its use for public health surveillance have been raised due to state variation in procedures used by medical examiners and coroners to record manner of death and specific drugs involved in overdoses (Davis et al., 2014b; Lucyk and Nelson, 2017; Ruhm, 2017, 2018; Warner et al., 2013). Alternative data sources that have been used to examine trends, geographic "hot spots," and product-specific characteristics for opioid-related overdose include Drug Abuse Warning Network (DAWN) emergency department data (Bau et al., 2016; Secora et al., 2017), opioid-related toxic exposures through RADARS or the National Poison Data System (NPDS) (Bau et al., 2016; Coplan et al., 2016; Coplan et al., 2013; Davis et al., 2014b; Mowry et al., 2016), detailed and timely information on fatal and nonfatal overdose through the Enhanced State Opioid Overdose Surveillance (ESOOS)/State Unintentional Drug Overdose Reporting System (SUDORS) (Mattson et al., 2018; Seth et al., 2018; Vivolo-Kantor et al., 2018), and information about opioid-related overdose from state hospital discharge databases (Cerdeira et al., 2017) or emergency department syndromic surveillance systems (Albert et al., 2011; Daly et al., 2017; Tomassoni et al., 2017). While containing less detailed information on specific products involved in overdose, the Healthcare Cost and Utilization Project (H-CUP) suite of inpatient and emergency department databases have also been used to assess temporal and geographic variation in nonfatal opioid-related overdose (Guy et al., 2018; Sakhuja et al., 2017; Tedesco et al., 2017; Unick et al., 2014; Unick and Ciccarone, 2017).

Much of the effort toward bettering data for public health surveillance involves state strategies to facilitate linkages of multiple data sources (Albert et al., 2011; Bau et al., 2016; Cepeda et al., 2017; Coplan et al., 2016; Davis et al., 2014b; Inciardi et al., 2009), across multiple state agencies. For example, with Chapter 55 of the Acts of 2015, Massachusetts' Department of Public Health developed a data warehouse providing person-level linkages across ten datasets managed by five state agencies, including the state all-payer claims database; state PDMP; death certificate records and toxicology results; substance abuse treatment information; hospital, emergency department, and outpatient records; criminal justice incarceration and treatment records; and emergency medical service data (Massachusetts Department of Public Health, 2017). Maryland also is advancing efforts to link person-level data from the PDMP, drug use and alcohol treatment admissions, hospital admissions, fatalities, and criminal justice data (Lyons and Madison, 2017; Saloner, 2016).

3.4.1. Common interview themes

Interviewees highlighted the need for surveillance efforts to consider the opioid crisis as a dynamic system with multiple agents and networks of interacting individuals and agencies (Burke, 2016; Wakeland et al., 2015), involving both licit and illicit markets. Linking

Table 4
Secondary Data Sources to Support Strengthening Data for Better Public Health Surveillance.

Data Elements (by Topic)	Sources	Strengths and Limitations
Prescription drug monitoring data		
Detection of opioid misuse <ul style="list-style-type: none"> • Prescription name/type • Prescription dose • Prescriber • Payment 	<ul style="list-style-type: none"> • State PDMP • PBSS 	<p>Strengths: + Comprehensive data on prescribing (i.e., multi-payer) + Can be used to develop measures around patient, prescriber, and pharmacist risky behaviors</p> <p>Limitations: - Access barriers - State capacity issues may limit ability to link PDMP data with other datasets</p>
Mortality data		
Opioid-related overdose <ul style="list-style-type: none"> • Cause of death • Drugs involved in death • Demographics 	<ul style="list-style-type: none"> • NDI • NVSS MCODE • CDC WONDER* • State vital records 	<p>Strengths: + National data with information on opioid overdose mortality + CDC WONDER is readily downloadable and publicly available</p> <p>Limitations: - Lags in data availability - Variation in quality of reporting detail on drug involvement</p>
Other national sources		
Detection of opioid misuse <ul style="list-style-type: none"> • Inpatient stays and ED visits • Nonfatal overdose • Opioid use disorder • Diagnoses and procedures 	<ul style="list-style-type: none"> • HCUP (national and state inpatient and emergency department databases) 	<p>Strengths: + Large collection of longitudinal data, nation-wide and state-level + State data is mapped to a standardized format</p> <p>Limitations: - Not all states participate in the three state-level databases - Costs to obtain full datasets</p>
	Enhanced state opioid overdose surveillance	<p>Strengths: + Very rich detail integrated from ED hospital billing, EMS, and syndromic surveillance data + Timely data availability and comparability across jurisdictions</p> <p>Limitations: - Not currently available for all states</p>
Toxico-surveillance <ul style="list-style-type: none"> • Opioid-related poison center calls • Exposure type (e.g., intentional abuse exposures) 	Poison Control <ul style="list-style-type: none"> • NPDS 	<p>Strengths: + Product and drug specific information</p> <p>Limitations: - Must be requested and purchased - Lags in availability vary by poison center</p>
Product-specific use & trends <ul style="list-style-type: none"> • Opioid use/initiation • Route of administration 	Proprietary surveillance <ul style="list-style-type: none"> • RADARS • NAVIPPRO 	<p>Strengths: + Multifaceted data collection including product and drug specific information + Can identify exposure among high-risk groups (e.g., pregnant women) + RADARS has information on product street prices</p> <p>Limitations: - Not nationally representative - Possible sampling biases - Costs to obtain</p>
Toxico-surveillance <ul style="list-style-type: none"> • Nonfatal opioid overdose 		
Illicit opioid markets <ul style="list-style-type: none"> • Source of opioids 		<p>Limitations: - Not nationally representative - Possible sampling biases - Costs to obtain</p>
Toxico-surveillance <ul style="list-style-type: none"> • Opioid-related ED visits • Substance with composition and formulation-specific differentiation 	ED surveillance <ul style="list-style-type: none"> • DAWN* 	<p>Strengths: + Nationally representative and generalizable + Mortality data available for a subset of states</p> <p>Limitations: - Discontinued in 2011 - Possible sampling and information biases</p>
Illicit opioid markets <ul style="list-style-type: none"> • Drug category; drug chemistry • Prevalence and location of emerging drugs • Street price (STRIDE) 	Drug seizure or testing data <ul style="list-style-type: none"> • NFLIS • STRIDE 	<p>Strengths: + Data on illicit drug supply, prices, and purity + Seizure data often available with less lag time + Useful in constructing models of the likely course of the epidemic</p> <p>Limitations: - Access barriers (particularly for sub-state data) - Some drugs seizures are not analyzed by participating laboratories</p>
Other state and local sources		
Illicit opioid markets <ul style="list-style-type: none"> • Criminal history • Drug-related offenses and arrests • Demographics 	Drug arrest data from state or local criminal justice agencies	<p>Strengths: + Could be used to examine network patterns of co-arrests + If linked with other data, can assess systematic histories leading to arrest</p> <p>Limitations: - Often not available in electronic form that is usable - Difficulties in obtaining data use permissions</p>
Detection of opioid misuse <ul style="list-style-type: none"> • Opioid-related inpatient stays and ED visits • Diagnoses and procedures • Costs 	HCUP (State Inpatient and State Emergency Department Databases)	<p>Strengths: + Large collection of state-level longitudinal data + State data is mapped to a standardized format</p> <p>Limitations: - Not all states participate - Costs to obtain full datasets</p>
	State inpatient, ED, mortality, or syndromic surveillance sources	<p>Strengths: + Often available with less time lag than national sources + May be linkable to variety of state data sources</p> <p>Limitations: - Access and cost barriers vary across sources - State-specific so challenges for cross-state comparison</p>
National surveys		
Illicit opioid markets <ul style="list-style-type: none"> • Self-reported drug use • Urinalysis test results • Substance abuse treatment history • Drug acquisition and payment 	Arrestee Survey <ul style="list-style-type: none"> • ADAM* 	<p>Strengths: + Captures a high-risk population with uniform data collection across sites</p> <p>Limitations: - Limited to few sites collecting data and male arrestees only - No longer fully operational - Certain data elements are restricted access</p>

* Publicly available at no cost.

opioid prescribing or dispensing data with data about illicit opioid users and illicit drug markets, such as that available in the recently scaled back Arrestee Drug Abuse Monitoring System (ADAM; Table 4), could be used to systematically examine individuals' histories associated with arrests, indicators of diversion, or movement between heroin and opioid analgesic markets. Interviewees also commonly discussed the need for more rapid data collection and analyses of other data sources, such as nonfatal overdose or drug seizure data, that can complement mortality data (Ruhm, 2017; Warner et al., 2013) and allow timelier understanding of emerging trends and facilitate more appropriately tailored interventions (Houry, 2017). Rhode Island's Opioid Overdose Reporting System (McCormick et al., 2017) and North Carolina's Disease Event Tracking and Epidemiologic Collection Tool are examples of state efforts toward near-real time collection and analysis of statewide nonfatal overdose data (Ising et al., 2016). Many interviewees also mentioned other novel efforts to leverage novel data sources (e.g., social media, the Dark Web) combined with machine learning techniques to identify risks and emerging trends (Brownstein et al., 2009; Kalyanam et al., 2017; Kalyanam and Mackey, 2017), as well as the potential benefits of linking claims or PDMP data with social services data (e.g., child welfare data) to augment ecological analyses (Ghertner et al., 2018; Orsi et al., 2018; Quast, 2018; Quast et al., 2019; Quast et al., 2018) and better understand the consequences of opioid misuse and opioid use disorder treatment on child welfare outcomes.

4. Discussion

Many efforts to inform strategies to combat the opioid crisis rely on analyses of secondary data. To further these efforts, this study is intended to enhance researcher awareness regarding the many existing data sources that can be used to address key HHS strategies, identify ways in which data sources can be used together to address questions more effectively than is possible with a single data source, and highlight existing data source strengths and limitations, innovative uses of data and data linkages, and opportunities to use such data to address high-priority research questions.

We identified a broad range of available data resources that researchers are using to examine a range of issues related to the opioid crisis, as well as many of the combinations of data sources being used by researchers to examine how the community and policy context relates to opioid-related outcomes. The value and availability of HHS support for data collection, aggregation and dissemination in addressing the opioid crisis is highlighted by the frequency with which researchers are using federal data sources, including surveys, claims data, policy data, and data from the census and other federal agencies. Such federal investments, and the consideration of future investments to enhance the quality and availability of data, such as linking mortality data to federal claims data, supporting the development of and access to all-payer claims databases, and encouraging the integration of criminal justice and public health datasets, are highlighted by our findings as critical steps to enhance the quality of future opioid-related research.

Our discussions with experts also emphasized a range of actions that do not require a substantial investment but appear likely to enhance the quality, availability, and usability of existing data. These include establishing standards for determining opioid-related cause of death, making overdose data available in a timelier manner, and ensuring available data is provided in formats that facilitate incorporation into analytic software. Even in the short time period since our interviews took place, some progress has been made to fill the identified gaps in research. Researchers have increasingly leveraged information from state APCDs – linked or as a standalone data source – to understand the intersection of patient conditions, opioid use, non-opioid therapies, and opioid-related harms; and to better estimate state-level population prevalence of opioid use disorder (Barocas et al., 2018; Bartels et al., 2018; Laroche et al., 2018; Malon et al., 2018; Whedon et al., 2018). Recent funding for the Enhanced State Opioid Overdose Surveillance

(ESOOS) system has allowed for the collection of more timely and comprehensive data on fatal overdoses from over 30 states; however, to our knowledge, these data have not yet been made widely available to researchers for use beyond in the creation of reports by state health departments and the CDC (Goldschmidt et al., 2018; Mattson et al., 2018; O'Donnell et al., 2018; Schilke et al., 2019; Vivolo-Kantor et al., 2018). Making such data available to a broader array of researchers, and facilitating their linkage with other data sources, such as those with information on prescription drug use or criminal justice history, is one potential opportunity that could greatly enhance the value of these existing data sources.

5. Limitations

There are a number of limitations of this work that merit discussion. There is a tremendous amount of research being done related to the opioid crisis, with new papers being published in high quality journals weekly. Furthermore, the scoping study should not be considered a structured systematic literature review, thus there are studies and data sources not captured in this document and many of the key questions identified are ones that we expect investigators are already examining. Furthermore, we recognize that categorizing data sources and research questions by HHS strategy is somewhat arbitrary, and that the most influential research often crosses these categories.

Finally, while this review has taken an expansive perspective to highlight the breadth of potential resources available to researchers studying opioid policy, a deeper dive into any one area may yield further insights and challenges. The opioid crisis is complex, and there is a need to better understand the expected time course of a given policy's effect, determine the role of heterogeneous policy implementation in differentially influencing outcomes, understand how the adoption of multiple policies may interact to enhance or diminish any given policy's impact, and determine how a variety of important outcomes may be impacted by policy even if not the intended target of the intervention. Existing ecological research has highlighted the need to monitor multiple datasets simultaneously, and further research that can leverage individual-level record linkages and longitudinal information on individual outcomes will enhance our understanding of ecological associations in order to guide more informed policy design.

5.1. Conclusions

Given the human and societal toll of the opioid crisis, efforts to create and make available improved data assets to support more informed efforts to address the opioid crisis are a public health imperative. Overall, there are a variety of areas in which resources and time may be invested to enhance use and linkage of existing secondary data sources for opioid research. A tremendous amount of work is being done at the federal, state, and local levels to combat the opioids crisis. There has also been a substantial increase in research that has improved our understanding of the complex and multi-dimensional nature of the opioid crisis, as well as advanced the evidence base regarding the effectiveness of opioid policies and initiatives toward reducing opioid-related harms. While significant resources for the use and analysis of secondary data exist, not all are being optimized. This work serves to enhance awareness of existing data resources relevant to opioid research, describe the scope of research leveraging these datasets, and highlight some key research gaps, data limitations, and data linkage needs that future research can address to further efforts to combat the opioids crisis.

Declaration of Competing Interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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Author contributions

Author RS managed the literature search, assisted with data extraction for all documents included, conducted interviews with key informants, and wrote the first draft of the manuscript. Authors CAK and EAT co-wrote the draft, checked the full text of identified documents and conducted data extraction activities, and conducted interviews with key informants. Author BDS oversaw environmental scan activities, conducted interviews with key informants, and contributed to writing the manuscript. SL and SRS helped formulate the research questions and contributed to drafting the manuscript.

Appendix A. Supplementary data

Supplementary data to this article can be found online at <https://doi.org/10.1016/j.pmedr.2019.101015>.

References

- Acevedo, A., Garnick, D.W., Dunigan, R., Horgan, C.M., Ritter, G.A., Lee, M.T., Panas, L., Campbell, K., Haberman, K., et al., 2015. Performance measures and racial/ethnic disparities in the treatment of substance use disorders. *J. Stud. Alcohol Drugs* 76, 57–67.
- Albert, S., Brason 2nd, F.W., Sanford, C.K., Dasgupta, N., Graham, J., Lovette, B., 2011. Project Lazarus: community-based overdose prevention in rural North Carolina. *Pain Med.* 12 (Suppl 2), S77–S85.
- Alpert, A., Powell, D., Pacula, R.L., 2016. Supply-Side Drug Policy in the Presence of Substitutes: Evidence from the Introduction of Abuse-Deterrent Opioids. RAND Corporation, Santa Monica, CA.
- Anderson, L.S., Bell, H.G., Gilbert, M., Davidson, J.E., Winter, C., Barratt, M.J., Win, B., Painter, J.L., Menone, C., et al., 2017. Using social listening data to monitor misuse and nonmedical use of bupropion: a content analysis. *JMIR Public Health Surveill.* 3 e6.
- Andrilla, C.H.A., Moore, T.E., Patterson, D.G., Larson, E.H., 2019. Geographic distribution of providers with a DEA waiver to prescribe buprenorphine for the treatment of opioid use disorder: a 5-year update. *J. Rural Health* 35, 108–112.
- Arkskey, H., O'Malley, L., 2005. Scoping studies: towards a methodological framework. *Int. J. Soc. Res. Methodol.* 8, 19–32.
- Baehren, D.F., Marco, C.A., Droz, D.E., Sinha, S., Callan, E.M., Akpunonu, P., 2010. A statewide prescription monitoring program affects emergency department prescribing behaviors. *Ann. Emergency Med.* 56 (19–23), e1–e3.
- Bao, Y., Pan, Y., Taylor, A., Radakrishnan, S., Luo, F., Pincus, H.A., Schackman, B.R., 2016. Prescription drug monitoring programs are associated with sustained reductions in opioid prescribing by physicians. *Health Aff. (Millwood)* 35, 1045–1051.
- Barber, C., Gagnon, D., Fonda, J., Cho, K., Hermos, J., Miller, M., 2017. Assessing the impact of prescribing directives on opioid prescribing practices among Veterans Health Administration providers. *Pharmacoepidemiol. Drug Saf.* 26, 40–46.
- Barocas, J.A., White, L.F., Wang, J., Walley, A.Y., LaRochelle, M.R., Bernson, D., Land, T., Morgan, J.R., Samet, J.H., et al., 2018. Estimated prevalence of opioid use disorder in Massachusetts, 2011–2015: a capture-recapture analysis. *Am. J. Public Health* 108, 1675–1681.
- Bartels, K., Fernandez-Bustamante, A., McWilliams, S.K., Hopfer, C.J., Mikulich-Gilbertson, S.K., 2018. Long-term opioid use after inpatient surgery: a retrospective cohort study. *Drug Alcohol Depend.* 187, 61–65.
- Bau, G., Bucher Bartelson, B., Severtson, S.G., Green, J., Dart, R.C., 2016. Comparison of Population Rates between the Drug Abuse Warning Network (DAWN) and the RADARS System Poison Center Program.
- Baxter, J.D., Clark, R.E., Samnaliev, M., Aweh, G., O'Connell, E., 2015. Adherence to buprenorphine treatment guidelines in a medicaid program. *Subst. Abuse* 36, 174–182.
- Becker, W.C., Fiellin, D.A., Merrill, J.O., Schulman, B., Finkelstein, R., Olsen, Y., Busch, S.H., 2008. Opioid use disorder in the United States: Insurance status and treatment access. *Drug Alcohol Depend.* 94, 207–213.
- Becker, W.C., Fenton, B.T., Brandt, C.A., Doyle, E.L., Francis, J., Goulet, J.L., Moore, B.A., Torrice, V., Kerns, R.D., et al., 2017. Multiple sources of prescription payment and risky opioid therapy among veterans. *Med. Care* (55 Suppl 7 Suppl 1), S33–S36.
- Belz, D., Lieb, J., Rea, T., Eisenberg, M.S., 2006. Naloxone use in a tiered-response emergency medical services system. *Prehosp. Emergency Care* 10, 468–471.
- Bennett, A.S., Bell, A., Tomedi, L., Hulsey, E.G., Kral, A.H., 2011. Characteristics of an overdose prevention, response, and naloxone distribution program in Pittsburgh and Allegheny County, Pennsylvania. *J. Urban Health* 88, 1020–1030.
- Bohnert, A.S., Valenstein, M., Bair, M.J., Ganoczy, D., McCarthy, J.F., Ilgen, M.A., Blow, F.C., 2011. Association between opioid prescribing patterns and opioid overdose-related deaths. *JAMA* 305, 1315–1321.
- Bohnert, A.S., Logan, J.E., Ganoczy, D., Dowell, D., 2016. A detailed exploration into the association of prescribed opioid dosage and overdose deaths among patients with chronic pain. *Med. Care* 54, 435–441.
- Boscarino, J.A., Rukstalis, M., Hoffman, S.N., Han, J.J., Erlich, P.M., Gerhard, G.S., Stewart, W.F., 2010. Risk factors for drug dependence among out-patients on opioid therapy in a large US health-care system. *Addiction* 105, 1776–1782.
- Bounthavong, M., Harvey, M.A., Wells, D.L., Popish, S.J., Himstreet, J., Oliva, E.M., Kay, C.L., Lau, M.K., Randeria-Noor, P.P., et al., 2017. Trends in naloxone prescriptions prescribed after implementation of a National Academic Detailing Service in the Veterans Health Administration: a preliminary analysis. *J. Am. Pharm. Assoc.* 2003 (57), S68–S72.
- Boyer, E.W., 2012. Management of opioid analgesic overdose. *N. Engl. J. Med.* 367, 146–155.
- Braden, J.B., Russo, J., Fan, M.Y., Edlund, M.J., Martin, B.C., DeVries, A., Sullivan, M.D., 2010. Emergency department visits among recipients of chronic opioid therapy. *Arch. Intern. Med.* 170, 1425–1432.
- Brady, J.E., Wunsch, H., DiMaggio, C., Lang, B.H., Giglio, J., Li, G., 2014. Prescription drug monitoring and dispensing of prescription opioids. *Public Health Rep.* 129, 139–147.
- Brodrick, J.E., Brodrick, C.K., Adinoff, B., 2016. Legal regimes surrounding naloxone access: considerations for prescribers. *Am. J. Drug Alcohol Abuse* 42, 117–128.
- Brownstein, J.S., Freifeld, C.C., Madoff, L.C., 2009. Digital disease detection: harnessing the Web for public health surveillance. *N. Engl. J. Med.* 360 (2153–2155), 57.
- Buchmueller, T.C., Carey, C., 2018. The effect of prescription drug monitoring programs on opioid utilization in medicare. *Am. Econ. J.: Econ. Policy* 10, 77–112.
- Bujold, E., Huff, J., Staton, E.W., Pace, W.D., 2012. Improving use of narcotics for non-malignant chronic pain: a lesson from Community Care of North Carolina. *J. Opioid Manage.* 8, 363–367.
- Burke, D.S., 2016. Forecasting the opioid epidemic. *Science* 354, 529.
- Burns, R.M., Pacula, R.L., Bauhoff, S., Gordon, A.J., Hendrikson, H., Leslie, D.L., Stein, B.D., 2016. Policies related to opioid agonist therapy for opioid use disorders: the evolution of state policies from 2004 to 2013. *Subst. Abuse* 37, 63–69.
- Burrell, A., Ethun, L., Fawcett, J.A., Rickard-Aasen, S., Williams, K., Kearney, S.M., Pringle, J.L., 2017. The pharmacist's role in overdose: using mapping technologies to analyze naloxone and pharmacy distribution. *J. Am. Pharm. Assoc.* 2003, 57 S73–S77 e1.
- Burris, S., Johnson, S., Ibrahim, J., Platt, E., Allen, L., 2017. State-level opioid antagonist access laws: the emergence of three distinct strategies, 2001–2015. *Drug Alcohol Depend.* 171 e29.
- Butler, S.F., Budman, S.H., Licari, A., Cassidy, T.A., Lioy, K., Dickinson, J., Brownstein, J.S., Benneyan, J.C., Green, T.C., et al., 2008. National addictions vigilance intervention and prevention program (NAVIPPRO): a real-time, product-specific, public health surveillance system for monitoring prescription drug abuse. *Pharmacoepidemiol. Drug Saf.* 17, 1142–1154.
- Butler, S.F., Cassidy, T.A., Chilcoat, H., Black, R.A., Landau, C., Budman, S.H., Coplan, P.M., 2013. Abuse rates and routes of administration of reformulated extended-release oxycodone: initial findings from a sentinel surveillance sample of individuals assessed for substance abuse treatment. *J. Pain* 14, 351–358.
- Butler, S.F., Black, R.A., Fleming, A.B., 2018. Relative abuse of crush-resistant prescription opioid tablets via alternative oral modes of administration. *Pain Med.* 19, 1613–1627.
- Campbell, C.I., Weisner, C., Binswanger, I.A., Lapham, G.T., Ahmedani, B.K., Yarborough, B.J.H., Haller, I.V., Altschuler, A., Hechter, R.C., et al., 2019. Predictors of healthcare effectiveness data and information set (HEDIS) treatment initiation and engagement among patients with opioid use disorder across 7 health systems. *Subst. Abuse* 1–7.
- Canan, C., Polinski, J.M., Alexander, G.C., Kowal, M.K., Brennan, T.A., Shrank, W.H., 2017. Automatable algorithms to identify nonmedical opioid use using electronic data: a systematic review. *J. Am. Med. Inform. Assoc.* 24, 1204–1210.
- Carrell, D.S., Cronkite, D., Palmer, R.E., Saunders, K., Gross, D.E., Masters, E.T., Hyland, T.R., Von Korff, M., 2015. Using natural language processing to identify problem usage of prescription opioids. *Int. J. Med. Inf.* 84, 1057–1064.
- Cash, R.E., Kinsman, J., Crowe, R.P., Rivard, M.K., Faul, M., Panchal, A.R., 2018. Naloxone administration frequency during emergency medical service events - United States, 2012–2016. *MMWR Morb. Mortal. Wkly. Rep.* 67, 850–853.
- Cassidy, T.A., DasMahapatra, P., Black, R.A., Wieman, M.S., Butler, S.F., 2014. Changes in prevalence of prescription opioid abuse after introduction of an abuse-deterrent opioid formulation. *Pain Med.* 15, 440–451.
- Cepeda, M.S., Fife, D., Chow, W., Mastrogianni, G., Henderson, S.C., 2012. Assessing opioid shopping behaviour: a large cohort study from a medication dispensing database in the US. *Drug Saf.* 35, 325–334.
- Cepeda, M.S., Fife, D., Chow, W., Mastrogianni, G., Henderson, S.C., 2013a. Opioid

- shopping behavior: how often, how soon, which drugs, and what payment method. *J. Clin. Pharmacol.* 53, 112–117.
- Cepeda, M.S., Fife, D., Vo, L., Mastrogianni, G., Yuan, Y., 2013b. Comparison of opioid doctor shopping for tapentadol and oxycodone: a cohort study. *J. Pain* 14, 158–164.
- Cepeda, M.S., Coplan, P.M., Kopper, N.W., Maziere, J.Y., Wedin, G.P., Wallace, L.E., 2017. ER/LA opioid analgesics REMS: overview of ongoing assessments of its progress and its impact on health outcomes. *Pain Med.* 18, 78–85.
- Cerda, M., Gaidus, A., Keyes, K.M., Ponicki, W., Martins, S., Galea, S., Gruenewald, P., 2017. Prescription opioid poisoning across urban and rural areas: identifying vulnerable groups and geographic areas. *Addiction* 112, 103–112.
- Chan, B., Lopez, C., Sarkar, U., 2015. The Canary in the coal mine tweets: social media reveals public perceptions of non-medical use of opioids. *PLoS One* 10 e0135072.
- Chang, H.Y., Lyapustina, T., Rutkow, L., Daubresse, M., Richey, M., Faul, M., Stuart, E.A., Alexander, G.C., 2016. Impact of prescription drug monitoring programs and pill mill laws on high-risk opioid prescribers: a comparative interrupted time series analysis. *Drug Alcohol Depend.* 165, 1–8.
- Chen, J.H., Hom, J., Richman, I., Asch, S.M., Podchyska, T., Johansen, N.A., 2016. Effect of opioid prescribing guidelines in primary care. *Medicine (Baltimore)* 95 e4760.
- Chou, R., Deyo, R.A., Devine, B., Hansen, R., Sullivan, S., Jarvik, J., Blazina, I., Dana, T., Bougatsos, C., et al., 2014. The Effectiveness and Risks of Long-Term Opioid Treatment of Chronic Pain; Evidence Report/Technology Assessment Number 218. Agency for Health Care Research and Quality, Rockville MD.
- Chou, R., Turner, J.A., Devine, E.B., Hansen, R.N., Sullivan, S.D., Blazina, I., Dana, T., Bougatsos, C., Deyo, R.A., 2015. The effectiveness and risks of long-term opioid therapy for chronic pain: a systematic review for a National Institutes of Health Pathways to Prevention Workshop. *Ann. Intern. Med.* 162, 276–286.
- Cicero, T.J., Dart, R.C., Inciardi, J.A., Woody, G.E., Schnoll, S., Munoz, A., 2007. The development of a comprehensive risk-management program for prescription opioid analgesics: researched abuse, diversion and addiction-related surveillance (RADARS). *Pain Med.* 8, 157–170.
- Cicero, T.J., Ellis, M.S., Surratt, H.L., Kurtz, S.P., 2014. The changing face of heroin use in the United States: a retrospective analysis of the past 50 years. *JAMA Psychiatry* 71, 821–826.
- Cicero, T.J., Ellis, M.S., Harney, J., 2015. Shifting patterns of prescription opioid and heroin abuse in the United States. *N. Engl. J. Med.* 373, 1789–1790.
- Clark, A.K., Wilder, C.M., Winstanley, E.L., 2014. A systematic review of community opioid overdose prevention and naloxone distribution programs. *J. Addict. Med.* 8, 153–163.
- Cochran, G., Woo, B., Lo-Ciganic, W.H., Gordon, A.J., Donohue, J.M., Gellad, W.F., 2015. Defining nonmedical use of prescription opioids within health care claims: a systematic review. *Subst. Abuse* 36, 192–202.
- Cochran, G., Gordon, A.J., Lo-Ciganic, W.H., Gellad, W.F., Frazier, W., Lobo, C., Chang, C.H., Zheng, P., Donohue, J.M., 2017. An examination of claims-based predictors of overdose from a large Medicaid program. *Med. Care* 55, 291–298.
- Commission on Evidence-Based Policymaking, 2017. The Promise of Evidence-based Policymaking. Commission on Evidence-Based Policymaking, Washington, DC.
- Compton, W.M., Jones, C.M., Baldwin, G.T., 2016. Relationship between nonmedical prescription-opioid use and heroin use. *N. Engl. J. Med.* 374, 154–163.
- Coplan, P.M., Kale, H., Sandstrom, L., Landau, C., Chilcoat, H.D., 2013. Changes in oxycodone and heroin exposures in the National Poison Data System after introduction of extended-release oxycodone with abuse-deterrent characteristics. *Pharmacoepidemiol. Drug Saf.* 22, 1274–1282.
- Coplan, P.M., Chilcoat, H.D., Butler, S.F., Sellers, E.M., Kadakia, A., Harikrishnan, V., Haddox, J.D., Dart, R.C., 2016. The effect of an abuse-deterrent opioid formulation (OxyContin) on opioid abuse-related outcomes in the postmarketing setting. *Clin. Pharmacol. Ther.* 100, 275–286.
- Daly, E.R., Dufault, K., Swenson, D.J., Lakevicius, P., Metcalf, E., Chan, B.P., 2017. Use of emergency department data to monitor and respond to an increase in opioid overdoses in New Hampshire, 2011–2015. *Public Health Rep.* 132, 735–795.
- Dart, R.C., Surratt, H.L., Cicero, T.J., Parrino, M.W., Severtson, S.G., Bucher-Bartelson, B., Green, J.L., 2015. Trends in opioid analgesic abuse and mortality in the United States. *N. Engl. J. Med.* 372, 241–248.
- Dasgupta, N., Funk, M.J., Proescholdbell, S., Hirsch, A., Ribisl, K.M., Marshall, S., 2016. Cohort study of the impact of high-dose opioid analgesics on overdose mortality. *Pain Med.* 17, 85–98.
- Daubresse, M., Chang, H.Y., Yu, Y., Viswanathan, S., Shah, N.D., Stafford, R.S., Kruszewski, S.P., Alexander, G.C., 2013. Ambulatory diagnosis and treatment of nonmalignant pain in the United States, 2000–2010. *Med. Care* 51, 870–878.
- Dave, D., Grecu, A., Saffer, H., 2017. Mandatory access prescription drug monitoring programs and prescription drug abuse. NBER Working Paper Series Working Paper 23537.
- Davis, C.S., Carr, D., 2015. Legal changes to increase access to naloxone for opioid overdose reversal in the United States. *Drug Alcohol Depend.* 157, 112–120.
- Davis, C.S., Carr, D., 2016. Physician continuing education to reduce opioid misuse, abuse, and overdose: many opportunities, few requirements. *Drug Alcohol Depend.* 163, 100–107.
- Davis, J.M., Severtson, S.G., Bucher-Bartelson, B., Dart, R.C., 2014b. Using poison center exposure calls to predict prescription opioid abuse and misuse-related emergency department visits. *Pharmacoepidemiol. Drug Saf.* 23, 18–25.
- Davis, C.S., Southwell, J.K., Niehaus, V.R., Walley, A.Y., Dailey, M.W., 2014a. Emergency medical services naloxone access: a national systematic legal review. *Acad. Emergency Med.* 21, 1173–1177.
- del Portal, D.A., Healy, M.E., Satz, W.A., McNamara, R.M., 2016. Impact of an opioid prescribing guideline in the acute care setting. *J. Emergency Med.* 50, 21–27.
- Delcher, C., Wagenaar, A.C., Goldberger, B.A., Cook, R.L., Maldonado-Molina, M.M., 2015. Abrupt decline in oxycodone-caused mortality after implementation of Florida's Prescription Drug Monitoring Program. *Drug Alcohol Depend.* 150, 63–68.
- Deyo, R.A., Hallvik, S.E., Hildebran, C., Marino, M., Dexter, E., Irvine, J.M., O'Kane, N., Van Otterloo, J., Wright, D.A., et al., 2017. Association between initial opioid prescribing patterns and subsequent long-term use among opioid-naive patients: a statewide retrospective cohort study. *J. Gen. Intern. Med.* 32, 21–27.
- Dick, A.W., Pacula, R.L., Gordon, A.J., Sorbero, M., Burns, R.M., Leslie, D.L., Stein, B.D., 2015. Growth in buprenorphine waivers for physicians increased potential access to opioid agonist treatment, 2002–11. *Health Aff.* 34, 1028–1034.
- Doe-Simkins, M., Quinn, E., Xuan, Z., Sorensen-Alawad, A., Hackman, H., Ozonoff, A., Walley, A.Y., 2014. Overdose rescues by trained and untrained participants and change in opioid use among substance-using participants in overdose education and naloxone distribution programs: a retrospective cohort study. *BMC Public Health* 14, 297.
- Dowell, D., Zhang, K., Noonan, R.K., Hockenberry, J.M., 2016. Mandatory provider review and pain clinic laws reduce the amounts of opioids prescribed and overdose death rates. *Health Aff. (Millwood)* 35, 1876–1883.
- Ducharme, L., Abraham, A., 2008. State policy influence on the early diffusion of buprenorphine in community treatment programs. *Subst. Abuse Treat. Prev. Policy* 3, 17–27.
- Dunn, K.M., Saunders, K.W., Rutter, C.M., Banta-Green, C.J., Merrill, J.O., Sullivan, M.D., Weisner, C.M., Silverberg, M.J., Campbell, C.I., et al., 2010. Opioid prescriptions for chronic pain and overdose: a cohort study. *Ann. Intern. Med.* 152, 85–92.
- Edlund, M.J., Steffick, D., Hudson, T., Harris, K.M., Sullivan, M., 2007. Risk factors for clinically recognized opioid abuse and dependence among veterans using opioids for chronic non-cancer pain. *Pain* 129, 355–362.
- Edlund, M.J., Martin, B.C., Russo, J.E., DeVries, A., Braden, J.B., Sullivan, M.D., 2014. The role of opioid prescription in incident opioid abuse and dependence among individuals with chronic noncancer pain: the role of opioid prescription. *Clin. J. Pain* 30, 557–564.
- Enteen, L., Bauer, J., McLean, R., Wheeler, E., Hurliaux, E., Kral, A.H., Bamberger, J.D., 2010. Overdose prevention and naloxone prescription for opioid users in San Francisco. *J. Urban Health* 87, 931–941.
- Faul, M., Dailey, M.W., Sugerma, D.E., Sasser, S.M., Levy, B., Paulozzi, L.J., 2015. Disparity in naloxone administration by emergency medical service providers and the burden of drug overdose in US rural communities. *Am. J. Public Health* 105 (Suppl 3), e26–e32.
- Faul, M., Lurie, P., Kinsman, J.M., Dailey, M.W., Crabaugh, C., Sasser, S.M., 2017. Multiple naloxone administrations among emergency medical service providers is increasing. *Prehosp. Emergency Care* 1–8.
- Feder, K.A., Krawczyk, N., Saloner, B., 2017b. Medication-assisted treatment for adolescents in specialty treatment for opioid use disorder. *J. Adolesc. Health* 60, 747–750.
- Feder, K., Mojtabai, R., Krawczyk, N., Young, A., Kealhofer, M., Tormohlen, K., Crum, R., 2017a. Trends in insurance coverage and treatment among persons with opioid use disorders following the Affordable Care Act. *Drug Alcohol Depend.* 179, 271–274.
- Fisher, R., O'Donnell, D., Ray, B., Rusyniak, D., 2016. Police officers can safely and effectively administer intranasal naloxone. *Prehosp. Emergency Care* 20, 675–680.
- Frank, R.G., Pollack, H.A., 2017. Addressing the fentanyl threat to public health. *N. Engl. J. Med.* 376, 605–607.
- Freeman, P.R., Hankosky, E.R., Lofwall, M.R., Talbert, J.C., 2018. The changing landscape of naloxone availability in the United States, 2011–2017. *Drug Alcohol Depend.* 191, 361–364.
- Garg, R.K., Fulton-Kehoe, D., Franklin, G.M., 2017. Patterns of opioid use and risk of opioid overdose death among Medicaid patients. *Med. Care* 55, 661–668.
- Garnick, D.W., Horgan, C.M., Acevedo, A., McCorry, F., Weisner, C., 2012. Performance measures for substance use disorders: what research is needed? *Addict. Sci. Clin. Pract.* 7, 18.
- Garnick, D.W., Horgan, C.M., Acevedo, A., Lee, M.T., Panas, L., Ritter, G.A., Dunigan, R., Bidorin, A., Campbell, K., et al., 2014. Criminal justice outcomes after engagement in outpatient substance abuse treatment. *J. Subst. Abuse Treat.* 46, 295–305.
- Gellad, W.F., Zhao, X., Thorpe, C.T., Thorpe, J.M., Sileanu, F.E., Cashy, J.P., Mor, M., Hale, J.A., Radomski, T., et al., 2017. Overlapping buprenorphine, opioid, and benzodiazepine prescriptions among veterans dually enrolled in Department of Veterans Affairs and Medicare Part D. *Subst. Abuse* 38, 22–25.
- Ghertner, R., Waters, A., Radel, L., Crouse, G., 2018. The role of substance use in child welfare caseloads. *Child Youth Serv. Rev.* 90, 83–93.
- Giglio, R.E., Li, G., DiMaggio, C.J., 2015. Effectiveness of bystander naloxone administration and overdose education programs: a meta-analysis. *Inj. Epidemiol.* 2, 10.
- Gillette, C., Bush, M.A., Rogers, K.M.L., Mospan, G., Nealy, K., DeGeeter, M., Robinson, A.M., 2018. Association between the North Carolina Medical Board opioid guideline update and opioid prescriptions in Medicare Part D beneficiaries. *J. Opioid Manage.* 14, 239–243.
- Gilson, A.M., Fishman, S.M., Wilsey, B.L., Casamalhuapa, C., Baxi, H., 2012. Time series analysis of California's prescription monitoring program: impact on prescribing and multiple provider episodes. *J. Pain* 13, 103–111.
- Goldschmidt, A., Koziol, J., McCormick, M., Viner-Brown, S., 2018. State unintentional drug overdose reporting surveillance: opioid overdose deaths and characteristics in Rhode Island. *Rhode Island Med. J.* 101, 25–30.
- Gomes, T., Tadrous, M., Mamdani, M.M., Paterson, J.M., Juurlink, D.N., 2018. The burden of opioid-related mortality in the United States. *JAMA Network Open* 1 e180217.
- Gordon, A.J., Lo-Ciganic, W.H., Cochran, G., Gellad, W.F., Cathers, T., Kelley, D., Donohue, J.M., 2015. Patterns and quality of buprenorphine opioid agonist treatment in a large Medicaid program. *J. Addict. Med.* 9, 470–477.
- Green, T.C., Mann, M.R., Bowman, S.E., Zaller, N., Soto, X., Gadea, J., Cordy, C., Kelly, P., Friedmann, P.D., 2013. How does use of a prescription monitoring program change pharmacy practice? *J. Am. Pharm. Assoc.* 2003 (53), 273–281.

- Guy Jr., G.P., Zhang, K., Bohm, M.K., Losby, J., Lewis, B., Young, R., Murphy, L.B., Dowell, D., 2017. Vital signs: changes in opioid prescribing in the United States, 2006–2015. *MMWR Morb. Mortal. Wkly. Rep.* 66, 697–704.
- Guy Jr., G.P., Pasalic, E., Zhang, K., 2018. Emergency department visits involving opioid overdoses, U.S., 2010–2014. *Am. J. Prev. Med.* 54, e37–e39.
- Gwira Baumblatt, J.A., Wiedeman, C., Dunn, J.R., Schaffner, W., Paulozzi, L.J., Jones, T.F., 2014. High-risk use by patients prescribed opioids for pain and its role in overdose deaths. *JAMA Intern. Med.* 174, 796–801.
- Haddad, M.S., Zelenev, A., Altice, F.L., 2015. Buprenorphine maintenance treatment retention improves nationally recommended preventive primary care screenings when integrated into urban federally qualified health centers. *J. Urban Health* 92, 193–213.
- Hadland, S.E., Wharam, J.F., Schuster, M.A., Zhang, F., Samet, J.H., Larochelle, M.R., 2017. Trends in receipt of buprenorphine and naltrexone for opioid use disorder among adolescents and young adults, 2001–2014. *JAMA Pediatr.* 171, 747–755.
- Haegerich, T.M., Paulozzi, L.J., Manns, B.J., Jones, C.M., 2014. What we know, and don't know, about the impact of state policy and systems-level interventions on prescription drug overdose. *Drug Alcohol Depend.* 145, 34–47.
- Haffajee, R.L., Mello, M.M., Zhang, F., Zaslavsky, A.M., Larochelle, M.R., Wharam, J.F., 2018. Four states with robust prescription drug monitoring programs reduced opioid dosages. *Health Aff. (Millwood)* 37, 964–974.
- Hall, A.J., Logan, J.E., Toblin, R.L., Kaplan, J.A., Kraner, J.C., Bixler, D., Crosby, A.E., Paulozzi, L.J., 2008. Patterns of abuse among unintentional pharmaceutical overdose fatalities. *JAMA* 300, 2613–2620.
- Han, B., Compton, W.M., Jones, C.M., Cai, R., 2015. Nonmedical prescription opioid use and use disorders among adults aged 18 through 64 years in the United States, 2003–2013. *JAMA* 314, 1468–1478.
- Hansen, R.N., Walker, R.L., Shortreed, S.M., Dublin, S., Saunders, K., Ludman, E.J., Von Korff, M., 2017. Impact of an opioid risk reduction initiative on motor vehicle crash risk among chronic opioid therapy patients. *Pharmacoepidemiol. Drug Saf.* 26, 47–55.
- Hartung, D.M., Ahmed, S.M., Middleton, L., Van Otterloo, J., Zhang, K., Keast, S., Kim, H., Johnston, K., Deyo, R.A., 2017. Using prescription monitoring program data to characterize out-of-pocket payments for opioid prescriptions in a state Medicaid program. *Pharmacoepidemiol. Drug Saf.* 26, 1053–1060.
- Hernandez, I., He, M., Brooks, M.M., Zhang, Y., 2018. Exposure-response association between concurrent opioid and benzodiazepine use and risk of opioid-related overdose in medicare part D beneficiaries. *JAMA Network Open* 1, e180919.
- Hirsch, A., Proescholdbell, S.K., Bronson, W., Dasgupta, N., 2014. Prescription histories and dose strengths associated with overdose deaths. *Pain Med.* 15, 1187–1195.
- Houry, D., 2017. Testimony from Debra Houry, M.D. on Fentanyl: The Next Wave of the Opioid Crisis before Committee on Energy and Commerce on March 21.
- IMS Institute for Healthcare Informatics, 2016. Use of Opioid Recovery Medications: Recent Evidence on State Level Buprenorphine Use and Payment Types, Parsippany, NJ.
- Inciardi, J.A., Surratt, H.L., Cicero, T.J., Kurtz, S.P., Martin, S.S., Parrino, M.W., 2009. The “black box” of prescription drug diversion. *J. Addict. Dis.* 28, 332–347.
- Ising, A., Proescholdbell, S., Harmon, K.J., Sachdeva, N., Marshall, S.W., Waller, A.E., 2016. Use of syndromic surveillance data to monitor poisonings and drug overdoses in state and local public health agencies. *Inj. Prev.* 22 (Suppl 1), i43–i49.
- Jalal, H., Buchanich, J.M., Roberts, M.S., Balmert, L.C., Zhang, K., Burke, D.S., 2018. Changing dynamics of the drug overdose epidemic in the United States from 1979 through 2016. *Science* 361.
- Johnson, E.M., Porucznik, C.A., Anderson, J.W., Rolfs, R.T., 2011. State-level strategies for reducing prescription drug overdose deaths: Utah's prescription safety program. *Pain Med.* 12 (Suppl 2), S66–S72.
- Jones, C.M., 2017. The paradox of decreasing nonmedical opioid analgesic use and increasing abuse or dependence: an assessment of demographic and substance use trends, United States, 2003–2014. *Addict. Behav.* 65, 229–235.
- Jones, A., Bonermann, B., Sharp, A., Millett, G., 2018. Where multiple modes of medication-assisted treatment are available. *Health Aff. Blog.*
- Jones, C.M., Paulozzi, L.J., Mack, K.A., 2014a. Sources of prescription opioid pain relievers by frequency of past-year nonmedical use United States, 2008–2011. *JAMA Intern. Med.* 174, 802–803.
- Jones, C.M., Campopiano, M., Baldwin, G., McCance-Katz, E., 2015. National and state treatment need and capacity for opioid agonist medication-assisted treatment. *Am. J. Public Health* 105, e55–e63.
- Jones, C.M., Lurie, P.G., Compton, W.M., 2016. Increase in naloxone prescriptions dispensed in US Retail pharmacies since 2013. *Am. J. Public Health* 106, 689–690.
- Jones, J.D., Roux, P., Stancliff, S., Matthews, W., Comer, S.D., 2014b. Brief overdose education can significantly increase accurate recognition of opioid overdose among heroin users. *Int. J. Drug Policy* 25, 166–170.
- Kalyanam, J., Katsuki, T., Lanckriet, R.G., Mackey, T.K., 2017. Exploring trends of non-medical use of prescription drugs and polydrug abuse in the Twittersphere using unsupervised machine learning. *Addict. Behav.* 65, 289–295.
- Kalyanam, J., Mackey, T.K., 2017. A review of digital surveillance methods and approaches to combat prescription drug abuse. *Curr. Addict. Rep.*
- Kandel, D.B., Hu, M.C., Griesler, P., Wall, M., 2017. Increases from 2002 to 2015 in prescription opioid overdose deaths in combination with other substances. *Drug Alcohol Depend.* 178, 501–511.
- Katsuki, T., Mackey, T.K., Cuomo, R., 2015. Establishing a link between prescription drug abuse and illicit online pharmacies: analysis of Twitter data. *J. Med. Internet Res.* 17, e280.
- Katz, N., Panas, L., Kim, M., Audet, A.D., Bilansky, A., Eadie, J., Kreiner, P., Paillard, F.C., Thomas, C., et al., 2010. Usefulness of prescription monitoring programs for surveillance: analysis of Schedule II opioid prescription data in Massachusetts, 1996–2006. *Pharmacoepidemiol. Drug Saf.* 19, 115–123.
- Kennedy-Hendricks, A., Richey, M., McGinty, E.E., Stuart, E.A., Barry, C.L., Webster, D.W., 2016. Opioid overdose deaths and Florida's crackdown on pill mills. *Am. J. Public Health* 106, 291–297.
- Kerensky, T., Walley, A.Y., 2017. Opioid overdose prevention and naloxone rescue kits: what we know and what we don't know. *Addict. Sci. Clin. Pract.* 12, 4.
- Kim, H., Hartung, D.M., Jacob, R.L., McCarty, D., McConnell, K.J., 2016. The concentration of opioid prescriptions by providers and among patients in the Oregon Medicaid program. *Psychiatr. Serv.* 67, 397–404.
- Knowlton, A., Weir, B.W., Hazzard, F., Olsen, Y., McWilliams, J., Fields, J., Gaasch, W., 2013. EMS runs for suspected opioid overdose: implications for surveillance and prevention. *Prehosp. Emergency Care* 17, 317–329.
- Knudsen, H.K., 2015. The supply of physicians waived to prescribe buprenorphine for opioid use disorders in the United States: a state-level analysis. *J. Stud. Alcohol Drugs* 76, 644–654.
- Kolodny, A., Courtwright, D.T., Hwang, C.S., Kreiner, P., Eadie, J.L., Clark, T.W., Alexander, G.C., 2015. The prescription opioid and heroin crisis: a public health approach to an epidemic of addiction. *Annu. Rev. Public Health* 36, 559–574.
- Krawczyk, N., Picher, C.E., Feder, K.A., Saloner, B., 2017. Only one in twenty justice-referred adults in specialty treatment for opioid use receive methadone or buprenorphine. *Health Aff. (Millwood)* 36, 2046–2053.
- Krebs, E.E., Lurie, J.D., Fanciullo, G., Tosteson, T.D., Blood, E.A., Carey, T.S., Weinstein, J.N., 2010. Predictors of long-term opioid use among patients with painful lumbar spine conditions. *J. Pain* 11, 44–52.
- Krebs, E.E., Gravelly, A., Nugent, S., Jensen, A.C., DeRonne, B., Goldsmith, E.S., Kroenke, K., Bair, M.J., Noorbaloochi, S., 2018. Effect of opioid vs nonopioid medications on pain-related function in patients with chronic back pain or hip or knee osteoarthritis pain: the SPACE randomized clinical trial. *JAMA* 319, 872–882.
- Kreiner, P.W., Strickler, G.K., Undurraga, E.A., Torres, M.E., Nikitin, R.V., Rogers, A., 2017. Validation of prescriber risk indicators obtained from prescription drug monitoring program data. *Drug Alcohol Depend.* 173 (Suppl 1), S31–S38.
- Kuo, Y.F., Raji, M.A., Chen, N.W., Hasan, H., Goodwin, J.S., 2016. Trends in opioid prescriptions among part D medicare recipients from 2007 to 2012. *Am. J. Med.* 129 (221), e21–e30.
- Lambdin, B.H., Davis, C.S., Wheeler, E., Tueller, S., Kral, A.H., 2018a. Naloxone laws facilitate the establishment of overdose education and naloxone distribution programs in the United States. *Drug Alcohol Depend.* 188, 370–376.
- Lambdin, B.H., Zibbell, J., Wheeler, E., Kral, A.H., 2018b. Identifying gaps in the implementation of naloxone programs for laypersons in the United States. *Int. J. Drug Policy* 52, 52–55.
- Larochelle, M.R., Liebschutz, J.M., Zhang, F., Ross-Degnan, D., Wharam, J.F., 2016. Opioid prescribing after nonfatal overdose and association with repeated overdose: a cohort study. *Ann. Intern. Med.* 164, 1–9.
- Larochelle, M.R., Bernson, D., Land, T., Stopka, T.J., Wang, N., Xuan, Z., Bagley, S.M., Liebschutz, J.M., Walley, A.Y., 2018. Medication for opioid use disorder after non-fatal opioid overdose and association with mortality: a cohort study. *Ann. Intern. Med.* 169, 137–145.
- Lasser, K.E., 2017. Prescription opioid use among U.S. adults: our brave new world. *Ann. Intern. Med.* 167, 351–352.
- Levac, D., Colquhoun, H., O'Brien, K.K., 2010. Scoping studies: advancing the methodology. *Implement. Sci.* 5, 69.
- Levine, M., Sanko, S., Eckstein, M., 2016. Assessing the risk of prehospital administration of naloxone with subsequent refusal of care. *Prehosp. Emergency Care* 20, 566–569.
- Li, G., Brady, J.E., Lang, B.H., Giglio, J., Wunsch, H., DiMaggio, C., 2014. Prescription drug monitoring and drug overdose mortality. *Inj. Epidemiol.* 1, 9.
- Lin, D.H., Lucas, E., Murimi, I.B., Jackson, K., Baier, M., Frattaroli, S., Gielen, A.C., Moyo, P., Simoni-Wastila, L., et al., 2017. Physician attitudes and experiences with Maryland's prescription drug monitoring program (PDMP). *Addiction* 112, 311–319.
- Liu, Y., Logan, J.E., Paulozzi, L.J., Zhang, K., Jones, C.M., 2013. Potential misuse and inappropriate prescription practices involving opioid analgesics. *Am. J. Manage. Care* 19, 648–665.
- Lo-Ciganic, W.H., Gellad, W.F., Gordon, A.J., Cochran, G., Zemaits, M.A., Cathers, T., Kelley, D., Donohue, J.M., 2016. Association between trajectories of buprenorphine treatment and emergency department and in-patient utilization. *Addiction* 111, 892–902.
- Lucyk, S.N., Nelson, L.S., 2017. Toxicosurveillance in the US opioid epidemic. *Int. J. Drug Policy* 46, 168–171.
- Lyons, B., Madison, K., 2017. Predictive Risk Evaluation to Combat Overdose Grant (PRECOG). Behavioral Health Administration.
- Mack, K.A., Zhang, K., Paulozzi, L., Jones, C., 2015. Prescription practices involving opioid analgesics among Americans with Medicaid, 2010. *J. Health Care Poor Underserved* 26, 182–198.
- Malon, J., Shah, P., Koh, W.Y., Cattabriga, G., Li, E., Cao, L., 2018. Characterizing the demographics of chronic pain patients in the state of Maine using the Maine all payer claims database. *BMC Public Health* 18, 810.
- Martin, C.E., Longinaker, N., Terplan, M., 2015. Recent trends in treatment admissions for prescription opioid abuse during pregnancy. *J. Subst. Abuse Treat.* 48, 37–42.
- Martins, S.S., Fenton, M.C., Keyes, K.M., Blanco, C., Zhu, H., Storr, C.L., 2012. Mood and anxiety disorders and their association with non-medical prescription opioid use and prescription opioid-use disorder: longitudinal evidence from the National Epidemiologic Study on Alcohol and Related Conditions. *Psychol. Med.* 42, 1261–1272.
- Massachusetts Department of Public Health, 2016. An assessment of opioid-related deaths in Massachusetts (2013–2014).
- Massachusetts Department of Public Health, 2017. An assessment of fatal and nonfatal opioid overdoses in Massachusetts (2011–2015).
- Mattson, C.L., O'Donnell, J., Kariisa, M., Seth, P., Scholl, L., Gladden, R.M., 2018.

- Opportunities to prevent overdose deaths involving prescription and illicit opioids, 11 States, July 2016–June 2017. *MMWR Morb. Mortal. Wkly. Rep.* 67, 945–951.
- Maxwell, S., Bigg, D., Stanczykiewicz, K., Carlberg-Racich, S., 2006. Prescribing naloxone to actively injecting heroin users: a program to reduce heroin overdose deaths. *J. Addict. Dis.* 25, 89–96.
- McCabe, S.E., Cranford, J.A., West, B.T., 2008. Trends in prescription drug abuse and dependence, co-occurrence with other substance use disorders, and treatment utilization: results from two national surveys. *Addict. Behav.* 33, 1297–1305.
- McCormick, M., Koziol, J., Sanchez, K., 2017. Development and use of a new opioid overdose surveillance system, 2016. *Subst. Abuse* 1091, 71–76.
- Mercado, M.C., Sumner, S.A., Spelke, M.B., Bohm, M.K., Sugerma, D.E., Stanley, C., 2018. Increase in drug overdose deaths involving Fentanyl-Rhode Island, January 2012–March 2014. *Pain Med.* 19, 511–523.
- Miller, M., Barber, C.W., Leatherman, S., Fonda, J., Hermos, J.A., Cho, K., Gagnon, D.R., 2015. Prescription opioid duration of action and the risk of unintentional overdose among patients receiving opioid therapy. *JAMA Intern. Med.* 175, 608–615.
- Morgan, J.R., Schackman, B.R., Leff, J.A., Linas, B.P., Walley, A.Y., 2018. Injectable naltrexone, oral naltrexone, and buprenorphine utilization and discontinuation among individuals treated for opioid use disorder in a United States commercially insured population. *J. Subst. Abuse Treat.* 90–96.
- Mowry, J.B., Spyker, D.A., Brooks, D.E., Zimmerman, A., Schauben, J.L., 2016. 2015 annual report of the American Association of Poison Control Centers' National Poison Data System (NPDS): 33rd annual report. *Clin. Toxicol. (Phila)* 54, 924–1109.
- Moyo, P., Simoni-Wastila, L., Griffin, B.A., Onukwugha, E., Harrington, D., Alexander, G.C., Palumbo, F., 2017. Impact of prescription drug monitoring programs (PDMPs) on opioid utilization among Medicare beneficiaries in 10 U.S. states. *Addiction* 112, 1784–1796.
- Mueller, S.R., Walley, A.Y., Calcaterra, S.L., Glanz, J.M., Binswanger, I.A., 2015. A review of opioid overdose prevention and naloxone prescribing: implications for translating community programming into clinical practice. *Subst. Abuse* 36, 240–253.
- National Academies of Sciences Engineering and Medicine, 2017. *Pain Management and the Opioid Epidemic: Balancing Societal and Individual Benefits and Risks of Prescription Opioid Use*. National Academies Press, Washington, DC.
- O'Donnell, J., Gladden, R.M., Mattson, C.L., Kariisa, M., 2018. Notes from the field: overdose deaths with carfentanyl and other fentanyl analogs detected - 10 States, July 2016–June 2017. *MMWR Morb. Mortal. Wkly. Rep.* 67, 767–768.
- O'Kane, N., Hallvik, S.E., Marino, M., Van Otterloo, J., Hildebran, C., Leichting, G., Deyo, R.A., 2016. Preparing a prescription drug monitoring program data set for research purposes. *Pharmacoepidemiol. Drug Saf.* 25, 993–997.
- Olfson, M., Wall, M., Wang, S., Crystal, S., Blanco, C., 2018. Service use preceding opioid-related fatality. *Am. J. Psychiatry* 175, 538–544.
- Oliva, E.M., Nevedal, A., Lewis, E.T., McCaa, M.D., Cochran, M.F., Konicki, P.E., Davis, C.S., Wilder, C., 2016. Patient perspectives on an opioid overdose education and naloxone distribution program in the U.S. Department of Veterans Affairs. *Subst. Abuse* 37, 118–126.
- Oliva, E.M., Christopher, M.L.D., Wells, D., Bounthavong, M., Harvey, M., Himstreet, J., Emmendorfer, T., Valentino, M., Franchi, M., et al., 2017. Opioid overdose education and naloxone distribution: development of the Veterans Health Administration's national program. *J. Am. Pharm. Assoc.* 2003, 57.
- Olivia, E., Bowe, T., Tavakoli, S., Martins, S., Lewis, E., Paik, M., Wiechers, I., Henderson, P., Harvey, M.A., et al., 2017. Development and applications of the Veterans Health Administration's Stratification Tool for Opioid Risk Mitigation (STORM) to improve opioid safety and prevent overdose and suicide. *Psychol. Serv.* 14, 34–49.
- Orsi, R., Yuma-Guerrero, P., Sergi, K., Pena, A.A., Shillington, A.M., 2018. Drug overdose and child maltreatment across the United States' rural-urban continuum. *Child Abuse Negl.* 86, 358–367.
- Pardo, B., 2017. Do more robust prescription drug monitoring programs reduce prescription opioid overdose? *Addiction* 112 (10), 1773–1783.
- Park, T.W., Saitz, R., Ganoczy, D., Ilgen, M.A., Bohnert, A.S., 2015. Benzodiazepine prescribing patterns and deaths from drug overdose among US veterans receiving opioid analgesics: case-cohort study. *BMJ* 350 h2698.
- Parker, J., Cuthbertson, C., Loveridge, S., Skidmore, M., Dyar, W., 2017. Forecasting state-level premature deaths from alcohol, drugs, and suicides using Google Trends data. *J. Affect. Disord.* 213, 9–15.
- Patrick, S.W., Fry, C.E., Jones, T.F., Buntin, M.B., 2016. Implementation of prescription drug monitoring programs associated with reductions in opioid-related death rates. *Health Aff. (Millwood)* 35, 1324–1332.
- Paulozzi, L.J., Stier, D.D., 2010. Prescription drug laws, drug overdoses, and drug sales in New York and Pennsylvania. *J. Public Health Policy* 31, 422–432.
- Paulozzi, L.J., Strickler, G.K., Kreiner, P.W., Koris, C.M., Centers for Disease, C., Prevention, 2015. Controlled substance prescribing patterns: prescription behavior surveillance system, eight states, 2013. *MMWR Surveill. Summ.* 64, 1–14.
- PCOR, 2018. *pcornet Partner Networks*.
- Piper, B.J., Desrosiers, C.E., Lipovsky, J.W., Rodney, M.A., Baker, R.P., McCall, K.L., Nichols, S.D., Martin, S.L., 2016. Use and misuse of opioids in maine: results from pharmacists, the prescription monitoring, and the diversion alert programs. *J. Stud. Alcohol Drugs* 77, 556–565.
- Popović, I., Maclean, J.C., Hijazi, B., Radakrishnan, S., 2017. The effect of state laws designed to prevent nonmedical prescription opioid use on overdose deaths and treatment. *Health Econ.* 27, 294–305.
- Porucznik, C.A., Johnson, E.M., Rolfs, R.T., Sauer, B.C., 2014. Specialty of prescribers associated with prescription opioid fatalities in Utah, 2002–2010. *Pain Med.* 15, 73–78.
- Price, T., 2017. Secretary Price Announces HHS Strategy for Fighting Opioid Crisis. Health and Human Services, National Rx Drug Abuse and Heroin Summit, Atlanta, GA.
- Quast, T., 2018. State-level variation in the relationship between child removals and opioid prescriptions. *Child Abuse Negl.* 86, 306–313.
- Quast, T., Storch, E.A., Yampolskaya, S., 2018. Opioid prescription rates and child removals: evidence from florida. *Health Aff. (Millwood)* 37, 134–139.
- Quast, T., Bright, M.A., Delcher, C., 2019. The relationship between foster care entries and high-dose opioid prescribing in California. *Addict. Behav.* 93, 52–58.
- Qureshi, N., Wesolowicz, L.A., Liu, C.M., Tungol Lin, A., 2015. Effectiveness of a retrospective drug utilization review on potentially unsafe opioid and central nervous system combination therapy. *J. Manage Care Spec. Pharm.* 21, 938–944.
- Rasubala, L., Pernapati, L., Velasquez, X., Burk, J., Ren, Y.F., 2015. Impact of a mandatory prescription drug monitoring program on prescription of opioid analgesics by dentists. *PLoS One* 10 e0135957.
- Ray, W.A., Chung, C.P., Murray, K.T., Hall, K., Stein, C.M., 2016. Prescription of long-acting opioids and mortality in patients with chronic noncancer pain. *JAMA* 315, 2415–2423.
- Ray, B.R., Lowder, E.M., Kivisto, A.J., Phalen, P., Gil, H., 2018. EMS naloxone administration as non-fatal opioid overdose surveillance: 6-year outcomes in Marion County, Indiana. *Addiction* 113, 2271–2279.
- Ray, B., Quinet, K., Dickinson, T., Watson, D.P., Ballew, A., 2017. Examining fatal opioid overdoses in Marion County, Indiana. *J. Urban Health: Bull. N.Y. Acad. Med.* 94, 301–310.
- Rees, D., Sabia, J., Argys, L., Latshaw, J., Dave, D., 2017. With a little help from my friends: The effects of Naloxone access and good Samaritan laws on opioid-related deaths. *NBER Working Paper Series NBER Working Paper* 23171.
- Reisman, R.M., Shenoy, P.J., Atherly, A.J., Flowers, C.R., 2009. Prescription opioid usage and abuse relationships: an evaluation of state prescription drug monitoring program efficacy. *Subst. Abuse* 3, 41–51.
- Rigg, K.K., Monnat, S.M., 2015. Urban vs. rural differences in prescription opioid misuse among adults in the United States: informing region specific drug policies and interventions. *Int. J. Drug Policy* 26, 484–491.
- Rigg, K.K., Monnat, S.M., Chavez, M.N., 2018. Opioid-related mortality in rural America: geographic heterogeneity and intervention strategies. *Int. J. Drug Policy* 57, 119–129.
- Rees, D., Sabia, J., Argys, L., Latshaw, J., Dave, D., 2017. With a little help from my friends: The effects of Naloxone access and good Samaritan laws on opioid-related deaths. *NBER Working Paper Series NBER Working Paper* 23171.
- Ringwalt, C., Garretson, M., Alexandridis, A., 2015a. The effects of North Carolina's prescription drug monitoring program on the prescribing behaviors of the state's providers. *J. Prim. Prev.* 36, 131–137.
- Ringwalt, C., Schiro, S., Shanahan, M., Proescholdbell, S., Meder, H., Austin, A., Sachdeva, N., 2015b. The use of a prescription drug monitoring program to develop algorithms to identify providers with unusual prescribing practices for controlled substances. *J. Prim. Prev.* 36, 287–299.
- Roberts, A.W., Farley, J.F., Holmes, G.M., Oramasionwu, C.U., Ringwalt, C., Sleath, B., Skinner, A.C., 2016. Controlled substance lock-in programs: examining an unintended consequence of a prescription drug abuse policy. *Health Aff. (Millwood)* 35, 1884–1892.
- Rosenblatt, R.A., Andrilla, C.H., Catlin, M., Larson, E.H., 2015. Geographic and specialty distribution of US physicians trained to treat opioid use disorder. *Ann. Fam. Med.* 13, 23–26.
- Rosenblum, D., Unick, G.J., Ciccarone, D., 2014. The entry of Colombian-sourced heroin into the US market: the relationship between competition, price, and purity. *Int. J. Drug Policy* 25, 88–95.
- Rudd, R.A., Seth, P., David, F., Scholl, L., 2016. Increases in drug and opioid-involved overdose deaths - United States, 2010–2015. *MMWR Morb. Mortal. Wkly. Rep.* 65, 1445–1452.
- Ruhm, C.J., 2017. Geographic variation in opioid and heroin involved drug poisoning mortality rates. *Am. J. Prev. Med.* 53, 745–753.
- Ruhm, C.J., 2018. Corrected US opioid-involved drug poisoning deaths and mortality rates, 1999–2015. *Addiction* 113, 1339–1344.
- Rutkow, L., Chang, H.Y., Daubresse, M., Webster, D.W., Stuart, E.A., Alexander, G.C., 2015. Effect of Florida's prescription drug monitoring program and pill mill laws on opioid prescribing and use. *JAMA Intern Med* 175, 1642–1649.
- Sakhuja, A., Sztajnkrzyer, M., Vallabhajosyula, S., Cheungpasitpon, W., Patch 3rd, R., Jentzer, J., 2017. National trends and outcomes of cardiac arrest in opioid overdose. *Resuscitation* 121, 84–89.
- Saloner, B., Karthikeyan, S., 2015. Changes in substance abuse treatment use among individuals with opioid use disorders in the United States, 2004–2013. *JAMA* 314, 1515–1517.
- Saloner, B., Stoller, K.B., Barry, C.L., 2016. Medicaid coverage for methadone maintenance and use of opioid agonist therapy in specialty addiction treatment. *Psychiatr. Serv.* 67, 676–679.
- Saloner, B., Daubresse, M., Caleb Alexander, G., 2017. Patterns of buprenorphine-naloxone treatment for opioid use disorder in a multistate population. *Med. Care* 55, 669–676.
- Saloner, B., 2016. Panel Paper: Using Data Science to Identify Individuals at High Risk of Opioid Overdose: A Multiyear Data Linkage Project in Maryland, Association for Public Policy Analysis and Management 38th Annual Fall Research Conference: The Role of Research in Making Government More Effective, Washington, DC.
- Schilke, R., Card, K., Jiang, J., Sturns, J., McCoy, S., Colston, L., 2019. Validating syndromic data for opioid overdose surveillance in Florida. *Online J. Public Health Inf. Schmidt, T., Zimam, A., Nielsen, A., Wakeland, W., 2014. Data sources regarding the nonmedical use of pharmaceutical opioids in the United States. Rev. Health Care* 5, 33–50.
- Schnell, M., Currie, J., 2018. Addressing the opioid epidemic: is there a role for physician education? *Am. J. Health Econ.* 4, 383–410.

- Scholl, L., Seth, P., Kariisa, M., Wilson, N., Baldwin, G., 2018. Drug and opioid-involved overdose deaths - United States, 2013–2017. *MMWR Morb. Mortal. Wkly. Rep.* 67, 1419–1427.
- Secora, A.M., Dormitzer, C.M., Staffa, J.A., Dal Pan, G.J., 2014. Measures to quantify the abuse of prescription opioids: a review of data sources and metrics. *Pharmacoepidemiol. Drug Saf.* 23, 1227–1237.
- Secora, A., Trinidad, J.P., Zhang, R., Gill, R., Dal Pan, G., 2017. Drug availability adjustments in population-based studies of prescription opioid abuse. *Pharmacoepidemiol. Drug Saf.* 26, 180–191.
- Seth, P., Scholl, L., Rudd, R.A., Bacon, S., 2018. Overdose deaths involving opioids, cocaine, and psychostimulants - United States, 2015–2016. *MMWR Morb. Mortal. Wkly. Rep.* 67, 349–358.
- Sherman, R.E., Anderson, S.A., Dal Pan, G.J., Gray, G.W., Gross, T., Hunter, N.L., LaVange, L., Marinac-Dabic, D., Marks, P.W., et al., 2016. Real-world evidence - what is it and what can it tell us? *N. Engl. J. Med.* 375, 2293–2297.
- Stein, B.D., Gordon, A.J., Sorbero, M., Dick, A.W., Schuster, J., Farmer, C., 2012. The impact of buprenorphine on treatment of opioid dependence in a Medicaid population: Recent service utilization trends in the use of buprenorphine and methadone. *Drug Alcohol Depend.* 123, 72–78.
- Stein, B.D., Gordon, A.J., Dick, A.W., Burns, R.M., Pacula, R.L., Farmer, C.M., Leslie, D.L., Sorbero, M., 2015a. Supply of buprenorphine waived physicians: the influence of state policies. *J. Subst. Abuse Treat.* 48, 104–111.
- Stein, B.D., Pacula, R.L., Gordon, A.J., Burns, R.M., Leslie, D.L., Sorbero, M.J., Bauhoff, S., Mandell, T.W., Dick, A.W., 2015b. Where is buprenorphine dispensed to treat opioid use disorders? The role of private offices, opioid treatment programs, and substance abuse treatment facilities in urban and rural counties. *Milbank Q.* 93, 561–583.
- Stein, B.D., Sorbero, M., Dick, A.W., Pacula, R.L., Burns, R.M., Gordon, A.J., 2016. Physician capacity to treat opioid use disorder with buprenorphine-assisted treatment. *JAMA* 316, 1211–1212.
- Substance Abuse and Mental Health Services Administration, 2013. Drug Abuse Warning Network, 2011: National Estimates of Drug-Related Emergency Department Visits. HHS Publication No. (SMA) 13-4760, DAWN Series D-39. Substance Abuse and Mental Health Administration, Rockville, MD.
- Tedesco, D., Asch, S.M., Curtin, C., Hah, J., McDonald, K.M., Fantini, M.P., Hernandez-Boussard, T., 2017. Opioid abuse and poisoning: trends in inpatient and emergency department discharges. *Health Aff. (Millwood)* 36, 1748–1753.
- Tenney, L., McKenzie, L.M., Matus, B., Mueller, K., Newman, L.S., 2019. Effect of an opioid management program for Colorado workers' compensation providers on adherence to treatment guidelines for chronic pain. *Am. J. Ind. Med.* 62, 21–29.
- Tomassoni, A.J., Hawk, K.F., Jubanyik, K., Noguee, D.P., Durant, T., Lynch, K.L., Patel, R., Din, D., Ulrich, A., et al., 2017. Multiple fentanyl overdoses - New Haven, Connecticut, June 23, 2016. *MMWR Morb. Mortal. Wkly. Rep.* 66, 107–111.
- Turner, L., Kruszewski, S., Mojtai, R., Webster, D.W., Nesbit, S., Stafford, R., Alexander, G., 2013. Trends in buprenorphine and methadone sales and utilization in the United States, 1997–2012. *Value Health* 16, A56.
- Turner, L., Kruszewski, S.P., Alexander, G.C., 2015. Trends in the use of buprenorphine by office-based physicians in the United States, 2003–2013. *Am. J. Addict.* 24, 24–29.
- Turner, B.J., Liang, Y., 2015. Drug overdose in a retrospective cohort with non-cancer pain treated with opioids, antidepressants, and/or sedative-hypnotics: interactions with mental health disorders. *J. Gen. Intern. Med.* 30, 1081–1096.
- U.S. Department of Health and Human Services, Behavioral Health Coordinating Committee, 2013. Addressing prescription drug abuse in the United States: current activities and future opportunities.
- U.S. Health and Human Services, Help, Resources, and Information: National Opioid Crisis.
- Unick, G.J., Ciccarone, D., 2017. US regional and demographic differences in prescription opioid and heroin-related overdose hospitalizations. *Int. J. Drug Policy* 46, 112–119.
- Unick, G., Rosenblum, D., Mars, S., Ciccarone, D., 2014. The relationship between US heroin market dynamics and heroin-related overdose, 1992–2008. *Addiction* 109, 1889–1898.
- van Dorp, E., Yassen, A., Dahan, A., 2007. Naloxone treatment in opioid addiction: the risks and benefits. *Expert Opin. Drug Saf.* 6, 125–132.
- Van Hout, M.C., Hearne, E., 2017. New psychoactive substances (NPS) on cryptomarket for: an exploratory study of characteristics of forum activity between NPS buyers and vendors. *Int. J. Drug Policy* 40, 102–110.
- Vivolo-Kantor, A.M., Seth, P., Gladden, R.M., Mattson, C.L., Baldwin, G.T., Kite-Powell, A., Coletta, M.A., 2018. Vital signs: trends in emergency department visits for suspected opioid overdoses - United States, July 2016–September 2017. *MMWR Morb. Mortal. Wkly. Rep.* 67, 279–285.
- Volkow, N.D., Frieden, T.R., Hyde, P.S., Cha, S.S., 2014. Medication-assisted therapies-tackling the opioid-overdose epidemic. *N. Engl. J. Med.* 370, 2063–2066.
- Von Korff, M., Dublin, S., Walker, R.L., Parchman, M., Shortreed, S.M., Hansen, R.N., Saunders, K., 2016. The impact of opioid risk reduction initiatives on high-dose opioid prescribing for patients on chronic opioid therapy. *J. Pain* 17, 101–110.
- Wakeland, W., Nielsen, A., Geissert, P., 2015. Dynamic model of nonmedical opioid use trajectories and potential policy interventions. *Am. J. Drug Alcohol Abuse* 41, 508–518.
- Walley, A.Y., Doe-Simkins, M., Quinn, E., Pierce, C., Xuan, Z., Ozonoff, A., 2013a. Opioid overdose prevention with intranasal naloxone among people who take methadone. *J. Subst. Abuse Treat.* 44, 241–247.
- Walley, A.Y., Xuan, Z., Hackman, H.H., Quinn, E., Doe-Simkins, M., Sorensen-Alawad, A., Ruiz, S., Ozonoff, A., 2013b. Opioid overdose rates and implementation of overdose education and nasal naloxone distribution in Massachusetts: interrupted time series analysis. *BMJ* 346, f174.
- Warner, M., Paulozzi, L., Nolte, K., Davis, G., Nelson, L.S., 2013. State variation in certifying manner of death and drugs involved in drug intoxication deaths. *Acad. Forensic Pathol.* 3, 231–237.
- Webster, L.R., Cochella, S., Dasgupta, N., Fakata, K.L., Fine, P.G., Fishman, S.M., Grey, T., Johnson, E.M., Lee, L.K., et al., 2011. An analysis of the root causes for opioid-related overdose deaths in the United States. *Pain Med.* 12 (Suppl 2), S26–S35.
- Weiner, S.G., Baker, O., Poon, S.J., Rodgers, A.F., Garner, C., Nelson, L.S., Schuur, J.D., 2017b. The effect of opioid prescribing guidelines on prescriptions by emergency physicians in Ohio. *Ann. Emerg. Med.* 70 (799–808) e1.
- Weiner, J., Bao, Y., Meisel, Z., 2017. Prescription Drug Monitoring Programs: Evolution and Evidence, *CHERISH Issue Brief*.
- Wen, H., Schackman, B.R., Aden, B., Bao, Y., 2017. States with prescription drug monitoring mandates saw a reduction in opioids prescribed to Medicaid enrollees. *Health Aff. (Millwood)* 36, 733–741.
- Westanno, A., Marshall, P., Jones, E., Burns, K., Krebs, E.E., 2015. Opioid dose reduction in a VA health care system-implementation of a primary care population-level initiative. *Pain Med.* 16, 1019–1026.
- Whedon, J.M., Toler, A.W.J., Goehl, J.M., Kazal, L.A., 2018. Association between utilization of chiropractic services for treatment of low-back pain and use of prescription opioids. *J. Altern. Complement. Med.* 24, 552–556.
- Wheeler, E., Davidson, P.J., Jones, T., Irwin, K., 2012. Community based opioid overdose prevention programs providing naloxone — United States, 2010. *MMWR Morb. Mortal. Wkly. Rep.* 61, 101–105.
- Wheeler, E., Jones, T.S., Gilbert, M.K., Davidson, P.J., 2015. Opioid overdose prevention programs providing naloxone to laypersons - United States, 2014. *MMWR Morb. Mortal. Wkly. Rep.* 64, 631–635.
- Willy, M.E., Graham, D.J., Racoosin, J.A., Gill, R., Kropp, G.F., Young, J., Choi, J., MaCurdy, T.E., et al., 2014. Candidate metrics for evaluating the impact of prescriber education on the safe use of extended-release/long-acting (ER/LA) opioid analgesics. *Pain Med.* 15, 1558–1568.
- Wu, L.T., Zhu, H., Swartz, M.S., 2016. Treatment utilization among persons with opioid use disorder in the United States. *Drug Alcohol Depend.* 169, 117–127.
- Xu, J., Davis, C.S., Cruz, M., Lurie, P., 2018. State naloxone access laws are associated with an increase in the number of naloxone prescriptions dispensed in retail pharmacies. *Drug Alcohol Depend.* 189, 37–41.
- Yang, Z., Wilsey, B., Bohm, M., Weyrich, M., Roy, K., Ritley, D., Jones, C., Melnikow, J., 2015. Defining risk of prescription opioid overdose: Pharmacy shopping and overlapping prescriptions among long-term opioid users in Medicaid. *J. Pain* 16, 445–453.
- Yarbrough, C.R., 2017. Prescription drug monitoring programs produce a limited impact on painkiller prescribing in Medicare part D. *Health Serv. Res.* 53, 671–689.
- Young, S.D., Zheng, K., Chu, L.F., Humphreys, K., 2018. Internet searches for opioids predict future emergency department heroin admissions. *Drug Alcohol Depend.* 190, 166–169.
- Zedler, B., Xie, L., Wang, L., Joyce, A., Vick, C., Kariburyo, F., Rajan, P., Baser, O., Murrelle, L., 2014. Risk factors for serious prescription opioid-related toxicity or overdose among Veterans Health Administration patients. *Pain Med.* 15, 1911–1929.