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Introduction

- Opioid Use Disorder (OUD) is a chronic, treatable condition associated with increased rates of morbidity and mortality, including overdose.¹
- Multiple effective medications for OUD (MOUD) are approved by the Food and Drug Administration: oral methadone, sublingual or transmucosal buprenorphine and buprenorphine/naloxone (BUP), subcutaneous extended-release buprenorphine (BUP-XR), and intramuscular extended-release naltrexone (NTX-XR).^{1,2}
- Lack of MOUD <u>initiation</u>: 17% of the 4.8 million people aged ≥12 years in the United States who were classified as having an OUD in 2023–2024 received MOUD.³
- Data shows that patients with OUD who remain on MOUD for >6 months have fewer overdoses and less serious opioid-related acute healthcare use compared with those who had shorter treatment duration or no treatment.⁴
- Lack of MOUD <u>retention</u>: Most patients (51–64%) discontinue treatment by 6 months.^{5,6}
- This study seeks to fill the gap in evidence regarding the real-world adherence to BUP-XR and its association with patient outcomes and costs.

Objectives



To evaluate patient characteristics, treatment patterns, healthcare resource utilization (HCRU), and costs following initiation of BUP-XR among commercially-insured patients with OUD, overall and stratified by level of adherence to BUP-XR treatment.

Methods

Retrospective Claims Analysis

- Administrative claims data from Merative™ MarketScan® Commercial and Medicare Databases
 Index selection period: March 1, 2019—December 31, 2022; the index date was the first claim for BUP-XR (i.e., SUBLOCADE®) during the index selection period.
- Study period: March 1, 2018-December 31, 2023
- Patients were required to have continuous enrollment with medical, pharmacy, and behavioral health benefits 12 months prior to index (baseline period) and 12 months after and including index (follow-up period).
- Patients with BUP-XR utilization 12 months prior to index or those with NTX-XR utilization 30 days prior to index were excluded.
- Patient profiles including demographics, clinical characteristics, and medication use were assessed at baseline. HCRU and cost were assessed during the follow-up period.
- Patients were categorized into sub-cohorts by proportion of days covered (PDC) to BUP-XR during the first 6 months of the follow-up period.

Results

Baseline Demographics and Comorbidities

- The 663 BUP-XR patients identified were on average 37.5±11.4 years old, 67.4% male, and 84.8% resided in urban areas (**Table 1**).
- Common baseline comorbidities and concomitant medications are presented in **Table 1**.

MOUD Adherence

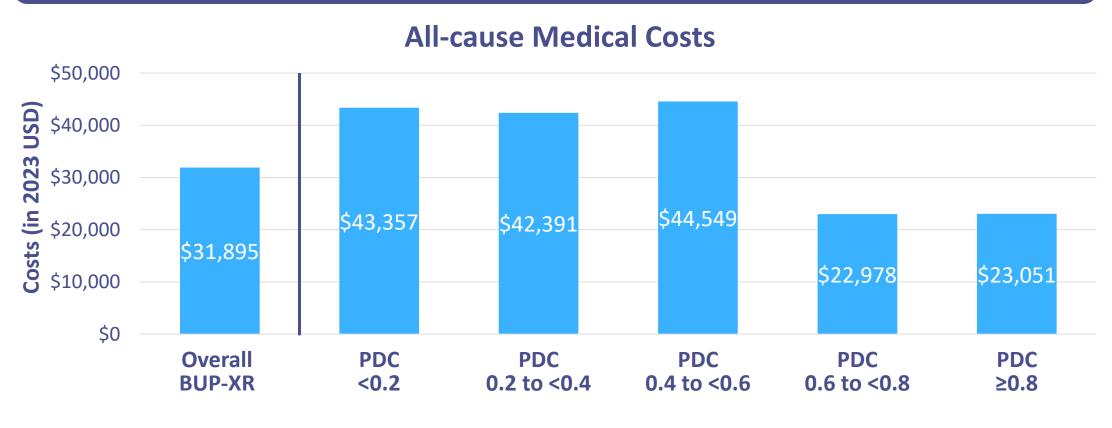
- <u>BUP-XR Adherence</u>: Average BUP-XR PDC during the first 6 months of the follow-up period was 0.6, with 42.4% (n=281) of patients with high adherence (PDC≥0.8) and 16.3% (n=108) with low adherence (PDC<0.2) (**Table 2**).
- MOUD Overall Adherence: Patients with low BUP-XR adherence had an average PDC of 0.5 for all MOUD (BUP-XR plus all other MOUD) during the first 6 months of the follow-up period (**Table 2**).

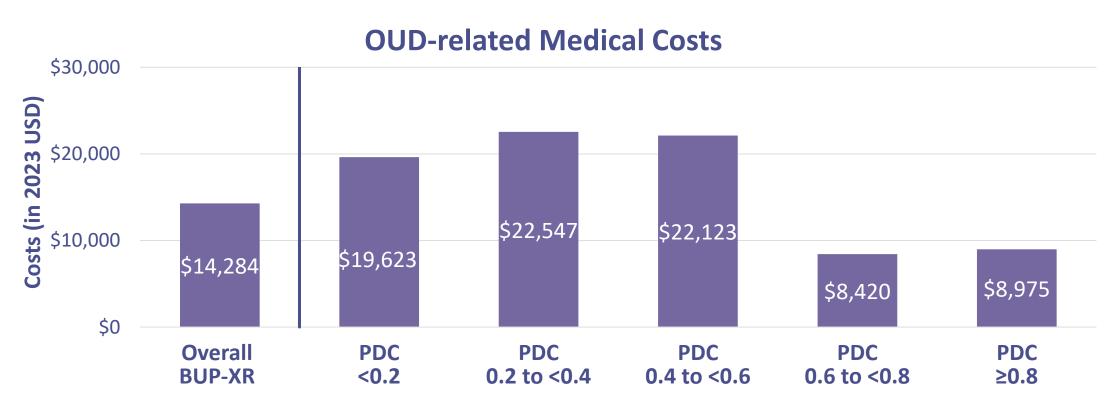
Table 1. Demogra	ohics and	Baselin	e Clinical	Characte	ristics	
	Overall	PDC	PDC	PDC	PDC	PDC
	BUP-XR	<0.2	0.2 to <0.4	0.4 to <0.6	0.6 to < 0.8	≥0.8
	(N=663)	(n=108)	(n=99)	(n=82)	(n=93)	(n=281)
Demographics (on index date)						
Age, mean (SD)	37.5 (11.4)	37.0 (12.3)	34.6 (10.8)	35.1 (11.6)	37.8 (12.0)	39.3 (10.6)
Sex, %						
Male	67.4	65.7	64.7	68.3	75.3	66.2
Female	32.6	34.3	35.4	31.7	24.7	33.8
Population density, %						
Urban	84.8	83.3	78.8	93.9	90.3	82.9
Rural	14.5	16.7	19.2	6.1	8.6	16.4
Unknown	0.8	0.0	2.0	0.0	1.1	0.7
Insurance plan type, %						
Comprehensive/Indemnity	2.6	3.7	3.0	2.4	0.0	2.9
EPO/PPO	56.3	55.6	50.5	54.9	64.5	56.2
POS/POS with capitation	10.7	9.3	12.1	11.0	4.3	12.8
HMO	11.6	9.3	16.2	12.2	16.1	9.3
CDHP/HDHP	17.7	18.5	17.2	19.5	14.0	18.2
Other/Unknown	1.2	3.7	1.0	0.0	1.1	0.7
Baseline clinical characteristics (in 12-n	nonth pre-ir	ndex period	plus index d	late)		
Charlson Comorbidity Index, mean (SD)	0.4 (0.8)	0.6 (1.3)	0.3 (0.6)	0.3 (0.8)	0.2 (0.6)	0.3 (0.8)
Select clinical condition, %						
OUD	88.5	87.0	86.9	93.9	91.4	87.2
AUD	22.2	25.9	27.3	19.5	19.4	20.6
SUD (other than OUD/AUD)	52.8	60.2	55.6	51.2	49.5	50.5
Depression/bipolar disorder	47.1	50.9	50.5	47.6	47.3	44.1
Generalized anxiety disorder	52.2	55.6	60.6	53.7	47.3	49.1
Chronic pain conditions	39.5	42.6	37.4	31.7	44.1	39.9
Concomitant medications, %						
Narcotic pain medications	19.9	23.2	24.2	15.9	23.7	17.1
Benzodiazepines	29.4	38.9	35.4	25.6	36.6	22.4
Sedative/hypnotics	37.1	37.0	41.4	51.2	39.8	30.6
Antidepressants	64.6	57.4	73.7	67.1	63.4	63.7
Antipsychotics	29.6	38.0	33.3	31.7	36.6	22.1

Table 2. Study Outcomes											
	Overall	PDC	PDC	PDC	PDC	PDC					
	BUP-XR	<0.2	0.2 to <0.4	0.4 to <0.6	0.6 to < 0.8	≥0.8					
	(N=663)	(n=108)	(n=99)	(n=82)	(n=93)	(n=281)					
PDC during 6-month post-index period, mean (SD)											
BUP-XR	0.6 (0.3)	0.2 (0.0)	0.3 (0.0)	0.5 (0.0)	0.7 (0.0)	0.9 (0.1)					
All MOUD ¹	0.8 (0.3)	0.5 (0.3)	0.6 (0.3)	0.6 (0.2)	0.8 (0.1)	0.9 (0.1)					
All-cause healthcare utilization during 12-month post-index period, %											
Admission (excluding for detoxification)	7.2	6.5	8.1	7.3	9.7	6.4					
ED visit (excluding for detoxification)	32.0	38.9	39.4	36.6	35.5	24.2					
Detoxification ²	16.0	20.4	27.3	19.5	15.1	9.6					
Recurrence ³ of OUD symptoms	12.1	16.7	23.2	13.4	12.9	5.7					
Days to recurrence, mean (SD)	130 (105)	78 (85)	141 (110)	94 (75)	139 (97)	192 (114)					

- ¹Included BUP, BUP-XR, and NTX-XR
- ²Identified by claims for inpatient and outpatient service (including ED visit) with procedure codes for detoxification.
- ³Defined by claims for inpatient admission, ED visit, or detoxification with a diagnosis for current opioid use/abuse/dependence or opioid overdose in the primary position for inpatient claims or any position for outpatient claims.

Figure 1. Non-MOUD Costs During 12-month Post-index Period





OUD-related costs were identified through medical claims with a diagnosis for current opioid use/abuse/dependence or opioid overdose in the primary position for inpatient claims or any position for outpatient claims.

Healthcare Resource Utilization and Costs

- Compared to BUP-XR patients with low adherence (**Table 2**), patients with high adherence had lower proportions of:
- All-cause ED visits excluding detoxification (24.2% vs. 38.9%)
- All-cause detoxification visits (9.6% vs. 20.4%)
- Evidence of OUD symptom recurrence (i.e., return to use) (5.7% vs. 16.7%)
- After excluding MOUD-related pharmacy costs, patients with high adherence had lower all-cause costs (\$23,051 vs. \$43,357) and OUD-related costs (\$8,975 vs. \$19,623) compared to patients with low adherence (**Figure 1**).

Conclusions

- Patient characteristics, healthcare utilization, and costs differ between patients with high (PDC≥0.8) vs. low (PDC<0.2) adherence to BUP-XR.
- Patients with low adherence are more likely to experience complex medical histories including comorbid substance use and psychiatric disorders.
- A benefit is observed among patients with high adherence, including fewer ED and detoxification visits, a lower rate of OUD symptom recurrence (i.e., return to use) and lower non-MOUD costs.
- Additional studies evaluating the impact of BUP-XR adherence on real-world outcomes are warranted.

Limitations

- Administrative claims data are subject to limitations in data coding and data entry errors (e.g., the use of MOUD may be under-reported in claims if patients receive the medications during inpatient hospitalization or pay for them out of pocket).
- Study results may not be generalizable to populations with other types of health coverage (e.g., Medicaid or Tricare) or the uninsured.
- The study period occurred during the COVID-19 pandemic when access restrictions to care and disruption of the healthcare system could influence findings.
- Study results have not been adjusted for differences in baseline characteristics between adherence groups.
- Although PDC≥0.8 is commonly used in claims-based studies as the level indicative of medication adherence, this threshold may not capture nuances of real-world treatment behaviors for a long-acting medication.

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Abbreviations

AUD: alcohol use disorder; BUP: sublingual or transmucosal buprenorphine and buprenorphine/naloxone; BUP-XR: buprenorphine extended-release; CDHP: consumer-driven health plan; ED: emergency department; EPO: exclusive provider organization; HCRU: healthcare resource utilization and costs; HDHP: high deductible health plan; HMO: health maintenance organization; MOUD: medications for opioid use disorder; NTX-XR: intramuscular extended-release naltrexone; PDC: proportion of days covered; POS: point of service; PPO: preferred provider organization;

OUD: opioid use disorder; SD: standard deviation; SUD: substance use disorder.

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Disclosures

This study was funded by Indivior Inc. Authors MJ and ATT are full-time employees of Merative which was contracted by Indivior Inc. to conduct this work. Author WM is full-time employee of Indivior Inc. Authors CF and MLT were full-time employees of Indivior Inc. at the time the study was conducted.