

National Drug & Alcohol Research Centre

The Difference is Research



Community studies of Long-Acting Buprenorphine injections for opioid dependence



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Disclosures

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Depot buprenorphine injections

Rationale:

- Less need for frequent attendance at clinic / pharmacy
- Less inconvenience and costs for clients and service providers
- Better adherence to medication (no missed doses) and better treatment outcomes
- Less diversion of medication
- Reduced capacity for patient interaction with the medication
- Available PBS-listed medications:
 - 1) Sublocade (Indivior) monthly SC injection by HCP
 - 2) Buvidal (Camurus) weekly or monthly SC injection by HCP





Objectives of CoLAB study

- To **document** and **evaluate the implementation** of monthly BPN injections in different community treatment settings (barriers and facilitators).
- To **examine key client outcomes,** including impacts on treatment engagement, retention and patient satisfaction.

Primary Outcome:

• Proportion of participants retained in treatment at 48 weeks following initiation of monthly depot buprenorphine injections. Treatment retention is defined as remaining on active depot buprenorphine medication at 48 weeks





Secondary Objectives

- 1. To evaluate opioid craving, withdrawal, opioid and other substance use
- 2. To evaluate utilisation of buprenorphine medication during the study, including Sublocade dose variation, adherence with dosing schedule (e.g. timing of doses, missed doses) and dose supplementation (additional sublingual buprenorphine)
- 3. To evaluate treatment safety and tolerability
- 4. To describe patient-reported changes to health and social well-being
- 5. To evaluate demographic, drug use and treatment factors associated with treatment outcomes (e.g. retention)
- 6. To evaluate patient-reported experience of treatment
- 7. To examine treatment retention at 24 weeks
- 8. To document the cost of treatment at different settings



Study design – Main Study





Study design – Extension phase









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CoLAB network – 7 sites



Participants



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Baseline characteristics:

Age and Gender

- Age, mean(SD): 44 (9)
- Female: 28

Background and Education

- Born in Australia: 88%
- Main source of income pension or benefit: 70%
- Completed year 10 education or more: 69

Present Living Condition

- · Boarding house: 6
- Privately owned house or flat: 22
- Rented house or flat : 51
- Other: 21





Baseline characteristics:



- Pharmaceutical opioid: 41%
- Other : 1%





Baseline characteristics: OAT treatment history

First Treatment Episode

- Age : 34 years (mean)
- Methadone : 41%
- Buprenorphine : 59%

Lifetime duration of OAT treatment in years (median)

- Length : 2.3
- Methadone : 0.5
- Buprenorphine : 3

OAT treatment in the past three month

- Methadone: 7
- Buprenorphine-naloxone : 99
- Buprenorphine : 4



Baseline characteristics



- No opioid withdrawal reported : 90
- Mild withdrawal reported : 10
- Significant withdrawal reported : 0

Other baseline characteristics:

- Non-fatal overdose in the past year : 14
- Opioid Craving VAS, median : 0
- Subjective Opiate Withdrawal Scale, median : 0
- Moderately-severely depressed (PHQ-9), n (%): 42



Baseline characteristics: substance use

| Past month substance use, n | |
|-----------------------------|----|
| Any illicit drug use | 54 |
| Opioids | 28 |
| Injected any drug | 28 |
| Heroin | 20 |
| Other Opioids | 11 |
| Amphetamine Type Substances | 25 |
| Cocaine | 4 |
| Benzodiazepines | 33 |
| Alcohol | 50 |
| Cannabis | 35 |
| Daily tobacco smoking | 73 |





Treatment retention and adherence



Predictors of treatment drop out: unadjusted

Bivariate analyses shows **heroin use** and **injecting drug use in the past month** are statistically significant predictors of treatment drop out

| | | Unadjusted Models | | |
|--|------|-------------------|---------|--|
| Independent variables | HR | 95% CI | P-value | |
| Age | 0.96 | 0.91 - 1.00 | 0.053 | |
| Gender | 1.52 | 0.67 - 3.44 | 0.314 | |
| Injecting drug use in the past month | 3.46 | 1.58 - 7.59 | 0.002 | |
| Heroin use in the past month | 3.22 | 1.44 - 7.18 | 0.004 | |
| Other non-prescribed opioids in the past month | 0.31 | 0.04 - 2.27 | 0.249 | |
| Amphetamine use in the past month | 1.83 | 0.81 - 4.14 | 0.148 | |
| Length of time in OAT treatment prior to study (years) | 0.89 | 0.77 - 1.03 | 0.127 | |

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Predictors of treatment drop out: adjusted

- Adjusting for age, gender and length of OAT treatment, injecting drug use in the past month is again a significant predictors of treatment drop out
- Adjusting for age, gender and substance use (including heroin, other non-prescribed opioids and amphetamine); heroin use in the past month becomes a significant predictor of treatment drop out

| | | Adjusted Mode | 1 | | Adjusted Mode | 12 |
|--|--------------|----------------------------|----------------|--------------|----------------------------|----------------|
| Independent variables | HR | 95% CI | P-value | HR | 95%CI | P-value |
| Age Gender | 0.95 2.00 | 0.91 - 0.99 0.84 - 4.75 | 0.032 0.115 | 0.97 1.52 | 0.92 - 1.00 0.62 - 3.69 | 0.122 0.357 |
| Injecting drug use in the past month | 4.10 | 1.81 - 9.32 | 0.001 | | 0.0 | 0 0 |
| Heroin use in the past month | | 0.0 | | 2.93 | 1.19 - 7.16 | 0.019 |
| Other non-prescribed opioids in the past month | | | | 0.28 | 0.04 - 2.16 | 0.223 |
| Amphetamine use in the past month | | | | 1.32 | 0.49 - 3.53 | 0.577 |
| Length of time in OAT treatment prior to study (years) | 0.92 | 0.80 - 1.06 | 0.267 | 0.94 | 0.81 - 1.08 | 0.388 |

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Drug use among people receiving depot buprenorphine



Opioid use



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Employment and depression







Quality of Life, pain and treatment satisfaction





Sublingual buprenorphine top up prescription

Sublingual buprenorphine was primarily prescribed during the first three month of the study

| Percentage of participants who received sublingual top up at each interval | |
|--|--------|
| Week 4 - 12 | 17 |
| Week 16 - 24 | 3 |
| Week 28 - 36 | 1 |
| Week 40 - 48 | 1 |
| mean duration of sublingual buprenorphine top-up (days) mean dose of sublingual buprenorphine top-up (mg) | 5 7 |
| Main reasons for sublingual prescription | |
| Patient request | 2 |
| Patient reporting withdrawal symptoms | 16 |
| Investigator decision | 4 |





Adverse events

| Number of people with a treatment-related adverse events | 45 | |
|---|----------|--|
| Number with a serious treatment-related adverse event | 0 | |
| | | |
| Top 3 treatment-related adverse events Withdrawal symptom | 17 | |
| Top 3 treatment-related adverse events Withdrawal symptom Injection site pain | 17 11 | |

| Top 10 Adverse Events (number of events) | |
|--|----|
| Withdrawal symptom | 44 |
| Injection site pain | 16 |
| Injection site itching | 14 |
| Headache | 11 |
| Injection site lump | 9 |
| Constipation | 8 |
| Lethargy | 7 |
| Nausea | 7 |
| Injection site redness | 6 |
| Product leakage | 4 |



Conclusion

- This study showed:
 - Successful implementation of buprenorphine depot treatment
 - High treatment retention rate
 - High treatment adherence
 - Reduced substance use
 - Reduced depression
 - Improved quality of life
 - Improved treatment satisfaction
- However further longer term observational studies with larger cohorts are required in order to gain deeper understanding on the benefits of buprenorphine depot medications.





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