

# A game changer! Long acting Injectables for Opioid Use Disorder Management

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Dr Adam Pastor BA MBBS FAChAM FRACP
Addiction Medicine Specialist and General Physician
Department Addiction Medicine St Vincent's Hospital Melbourne

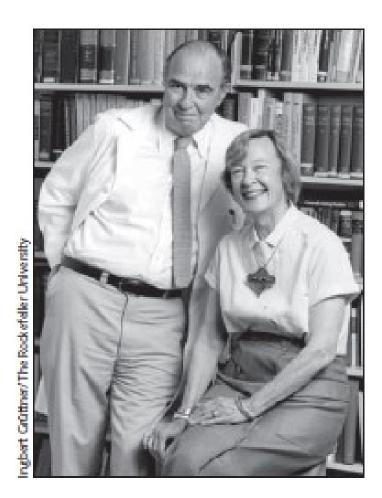
UNDER THE STEWARDSHIP OF MARY AIKENHEAD MINISTRIES



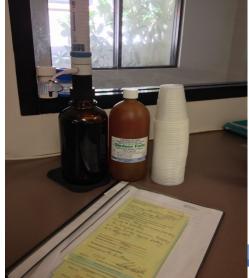
#### Heroin Addiction – A Metabolic Disease

Vincent P. Dole, MD and Marie E. Nyswander, MD, New York

Arch Intern Med-Vol 120, July 1967









#### Nopsad 2021 - Key findings

(National Opioid Pharmacotherapy Statistics Annual Data Collection)



#### **Snapshot day June 2020**

- 53,316 on course of pharmacotherapy for opioid use disorders
- median age 44
- male (67%)
- 3422 individual prescribers
- 71% dosing at local pharmacies
- access in cities > regional > remote



#### **Buvidal**

- Injectable form subcutaneous
- Buprenorphine only
- PBS listed September 2019
- Seven doses available Weekly and Monthly or
  - Weekly 8mg, 16mg, 24mg, 32mg
  - Monthly 64mg, 96mg, 128mg







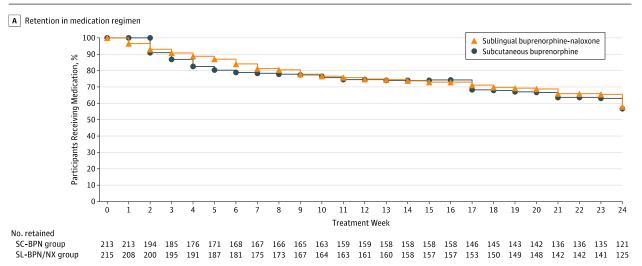








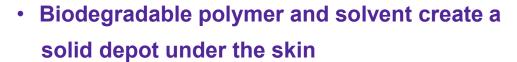
#### Figure 2. Study Outcomes





#### Sublocade

- Injectable form subcutaneous
- Buprenorphine only
- PBS listed May 1st 2020
- Two doses available Monthly administration
  - 300mg in 1.5mL
  - 100mg in 0.5mL



Pre-filled syringe, 19g needle







### **Efficacy**





			Total (N)	Discontinued (n)	Censored (n)	25th percentile days		Log-rank test p value
BL	JP <b>-</b> XR 30	0 mg/300 mg+IDC	196	65	131	104	N/A	<0.0001
BU	P-XR 30	0 mg /100 mg + IDC			129	120	N/A	<0.0001
		Placebo+IDC	99	63	36	26	78	NA
Participants(%) 900 900 900 900 900 900 900 900 900 90		No Contraction of the Contractio	- Landa de la companya de la company					
20		— BUP-XR 300 mg/ — BUP-XR 300 mg/ — Placebo + IDC						
C	0	29	57	85	11	3	141	16
Number at risk		-		Days on stu		-	•	
BUP-XR 300 mg/300 mg + IDC	196	193	170	152	14	4	134	95
BUP-XR 300 mg/100 mg + IDC	194	185	167	153	14	.8	141	80
Placebo + IDC	99	69	57	48	4	4	39	23





#### Effectiveness - Real World Data

Figure 2. Overview of Treatment Satisfaction Questionnaire for Medication (TSQM), Patient Satisfaction-Visual Analog Scale (PS-VAS), Patient Global Impression of Improvement (PGH), Treatment Burden Questionnaire (TBQ), Opioid-Related Behaviors in Treatment (ORBIT), and Substance Like Paccases Scalastor (SUBS) Desuite.

Score, mean (SE)		Treatment difference	Favors	Favors		
	Depot BPN	SL BPN	based on normalized	SL BPN		P
Variable TSQM global satisfa	(n=60)	(n=59)	score (95% CI), %	group	BPN group	Asin
					_	
Week 12	85.2 (2.1)	72.3 (2.2)	12.9 (6.8 to 18.9)			<.00
Wook 24 <sup>a</sup>	82.5 (2.3)	74.3 (2.3)	8.2 (1.7 to 14.6)			.01
Overall	82.4 (1.9)	73.8 (1.9)	8.6 (3.3 to 13.9)		_ <del></del>	.002
TSQM effectiveness					_	
Week 12	83.5 (2.5)	72.2 (2.6)	11.3 (4.2 to 18.5)			.002
Week 24	87.2 (1.9)	75.0 (1.9)	12.2 (6.8 to 17.6)		_ <del>-</del>	<.00
Overall	80.0 (2.0)	74.6 (2.0)	5.4 (-0.2 to 10.9)		-	.06
TSQM convenience						
Week 12	87.7 (2.0)	69.6 (2.0)	18.0 (12.4 to 23.6)			<.00
Wook 24	88.7 (2.2)	69.6 (2.2)	19.1 (12.9 to 25.4)			<.00
Overall	87.4 (1.7)	69.6 (1.7)	17.8 (13.0 to 22.6)			<.00
TSQM side effects						
Week 12	90.9 (2.4)	88.1 (2.4)	2.9 (-3.9 to 9.6)	_	-	.41
Week 24	88.5 (2.6)	90.3 (2.6)	-1.8 (-9.0 to 5.4)	_	<del></del>	.61
Overall	87.49 (2.1)	88.2 (2.1)	-0.3 (-6.2 to 5.6)	_	<del></del>	.91
PS-VAS						
Wook 12	86.9 (2.5)	74.1 (2.5)	12.7 (5.7 to 19.8)			<.00
Week 24	85.4 (2.4)	78.8 (2.4)	6.6 (-0.1 to 13.3)		<u> </u>	.05
Overall	83.9 (2.3)	76.0 (2.3)	7.9 (1.5 to 14.3)			.02
PGI-I score <sup>b</sup>						
Week 12	1.9 (0.1)	3.0 (0.1)	15.9 (10.0 to 21.9)			<.00
Week 24	2.0 (0.2)	2.9 (0.2)	13.0 (6.9 to 19.1)			<.00
Overall	1.9 (0.1)	2.9 (0.1)	14.3 (10.0 to 20.0)			<.00
TBQ global score <sup>b</sup>						
Week 12	13.7 (2.6)	28.3 (2.7)	9.7 (4.8 to 14.7)		<del></del>	<.00
Week 24	13.3 (2.6)	28.6 (2.5)	10.3 (5.5 to 15.0)		<b></b>	<.00
Overall.	13.4 (2.3)	28.3 (2.3)	9.9 (5.7 to 14.2)			<.00
ORBIT total score <sup>b</sup>						
Week 12	1.1 (0.2)	1.2 (0.2)	0.1 (-1.4 to 1.7)	-4	-	.86
Wook 24	0.5 (0.2)	1.2 (0.2)	1.8 (0.5 to 3.0)		-	.006
Overall	0.8 (0.2)	1.2 (0.2)	1.0 (-0.2 to 2.2)		-	.12
SURE total score			,			
Week 12	53.1 (0.8)	53.5 (0.8)	-0.7 (-4.3 to 3.0)	-	_	.70
Week 24	52.8 (1.0)	54.7 (1.0)	-3.0 (-7.4 to 1.5)	_	L	.19
Overall	53.0 (0.8)	54.2 (0.8)	-1.7 (-5.2 to 1.7)		L	.30
	33.2 (0.0)	(0.0)			<del></del>	
			-1		5 10 15 20 25 30	
				Treatmen	t difference based on normalized score (95% CI), %	

Patient-Reported Outcomes of Treatment of Opioid Dependence With Weekly and Monthly Subcutaneous Depot vs Daily Sublingual Buprenorphine A Randomized Clinical Trial

Nicholas Lintzeris, MBBS, PhD; Adrian J. Dunlop, MBBS, PhD; Paul S. Haber, MD, FRACP; Dan I. Lubman, MB ChB, PhD; Robert Graham, MBBS; Sarah Hutchinson, MSc; Shalini Arunogiri, MBBS, PhD; Victoria Hayes, MBBS, MPH; Peter Hjelmström, MD, PhD; Agneta Svedberg, MSc; Stefan Peterson, PhD; Fredrik Tiberg, PhD

	Patients, No. (%)				
Characteristic	Depot buprenorphine group (n = 60)	SL buprenorphine group (n = 59)	Total (N = 119)		
Treatment-emergent AEs	54 (90.0)	49 (83.1)	103 (86.6		
Drug-related ADRs	39 (65.0)	12 (20.3)	51 (42.9)		
SAEs	9 (15.0)	9 (15.3)	18 (15.1)		
Serious drug-related ADRs	1 (1.7)	0	1 (0.8)		
AEs or SAEs leading to withdrawal of trial medication	0	0	0		
Deaths	0	0	0		
Drug overdoses	0	4 (6.8)	4 (3.4)		
AEs occurring in ≥8% of participants					
Injection site					
Pain	11 (18.3)	0	11 (9.2)		
Mass	10 (16.7)	0	10 (8.4)		
Bruising	5 (8.3)	0	5 (4.2)		
Upper respiratory tract infection	7 (11.7)	2 (3.4)	9 (7.6)		
Nausea	5 (8.3)	2 (3.4)	7 (5.9)		
Vomiting	5 (8.3)	2 (3.4)	7 (5.9)		
Toothache	5 (8.3)	1 (1.7)	6 (5.0)		
Arthralgia	5 (8.3)	4 (6.8)	9 (7.6)		



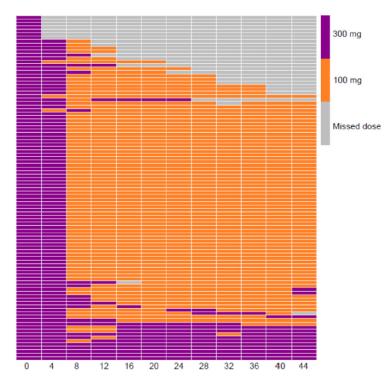


Table 5
Adverse events and events of special interest.

•	
	n
Total number of serious adverse events	26
Total number of incidents where serious adverse events occurred	20
Number of deaths	0
Number of people with a serious adverse event over the treatment period	16
Number of people with a treatment-related adverse event	45
Top 3 treatment-related adverse events	
Withdrawal symptom	17
Injection site pain	11
Injection site itching	14
Number with a serious treatment-related adverse event	0
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 due to faulty syringe	4
Events of special interest	
Pregnancy	2
Buprenorphine overdose	0
Severe hepatic impairment	0
Depot removal	0
Severe precipitated withdrawal	0

Outcomes of a single-arm implementation trial of extended-release subcutaneous buprenorphine depot injections in people with opioid dependence

Michael Farrell <sup>a,\*</sup>, Jeyran Shahbazi <sup>a</sup>, Marianne Byrne <sup>a,b</sup>, Jason Grebely <sup>b</sup>, Nicholas Lintzeris <sup>c,d</sup>, Mark Chambers <sup>a</sup>, Briony Larance <sup>a,e,f</sup>, Robert Ali <sup>a,g</sup>, Suzanne Nielsen <sup>a,h</sup>, Adrian Dunlop <sup>i,j</sup>, Gregory J. Dore <sup>b</sup>, Michael McDonough <sup>k</sup>, Mark Montebello <sup>c,l</sup>, Thomas Nicholas <sup>a</sup>, Rob Weiss <sup>m</sup>, Craig Rodgers <sup>n</sup>, Jon Cook <sup>o</sup>, Louisa Degenhardt <sup>a</sup>, on behalf of the CoLAB study team



### Uptake in Australia (approx.)

November 2021	Victoria	Australia
Buvidal	1300	8200
Sublocade	1200	2600

Why?

**Supervision** 

Cost

**Dispensing** 

**GP** 

Timing?

## ST VINCENT'S HEALTH AUSTRALIA

#### Our experience

- once weekly or monthly contact with a single provider
- limited risk of overdose or diversion
- acceptable to PWID
- assistance of St Vincent's Inclusive Health Fund
- a pilot clinic; low threshold access for people wishing to commence/access LAIB
- Stakeholder feedback including AOD services, specialist and PWID
- no/low cost
- broad referral base including self-referral
- short waiting time
- clear and appropriate screening



#### Pandemic Long Acting Injectable Buprenorphine Clinic

- May 2020 to Jan 2021
- heavily pre-screened; Triage -critical to limit unnecessary attendances
- Level 4 PPE (googles, mask, gown, gloves)
- minimise appointment time, flow through unit was unidirectional
- 'light touch'
- take home naloxone, influenza vaccination
- temperature checks
- SafeScript (real time prescribing) and permit system
- Pathology and Oral Fluid Testing available
- Data collection





Take home naloxone (Nyxoid)

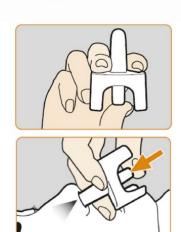
Influenza vaccination

**HCV** screening and treatment - fibroscan

**HBV** screening and referral +/- vaccination

Other AOD and Mental Health







#### Demographics (n=110)

Demographics		Treatment Related	
Mean age	39 years	Heroin Use in last month	41.4%
Age range	20 -74yo	In last week	37.9%
Male	65%	Prescription opioid	12%
Lives Alone	40.2%	Methadone > 50mg	8.2%
Private Accommodation	63.5%	Referral Source	
Worked in last month	12.1%	self	27%
Incarcerated in last 12 months	21.5%	drug court	13%
No prior opioid agonist treatment	10%	PAMS (Pharmacotherapy Advocacy Service)	6%



## Service Delivery

Measure	Number	
Injections	847	
Buvidal	820	519 (63%) monthly formulation 325 (62%) 128mg 112 (22%) 96mg
Sublocade	27	25 (93%) Sublocade 300mg
Take home naloxone (Nyxoid)	30 patients	
Influenza vaccination	22/53 eligible	
HCV testing	81 patients	Treated 4



#### **Outcome Measures**

Measure	Commencement	Follow Up
Retention in Treatment	100%	77%
Heroin Use in last month	41.4%	23%
Heroin Use in last week	37.9%	15.0%
Average # days heroin use	1.6 days/week	0.6 days/week
Employment	10.3%	14.3%
Homeless	12.1%	5.9%



#### Other measures

Psychological Health (Likert Scale 1-10)	5.8	6.3
Physical Health (Likert Scale 1-10)	5.5	6.2
Quality of Life (Likert Scale 1-10)	6.0	6.3

## ST VINCENT'S HEALTH AUSTRALIA

#### Conclusions

- good retention in treatment possible
- new to treatment and prescription opioid users
- LAIB attractive option
- mostly monthly, mostly upper dose ranges
- encouraged uptake during pandemic
- post-pandemic?

#### **New Horizons**



**Commencing directly on LAIB** 

**Pregnancy (and Breastfeeding)** 

Transferring from methadone to LAIB

**Criminal Justice System** 

**Cost and availability** 

Attracting new people to treatment?

Improving/increasing other supports as more regular contact diminishes



#### Acknowledgements

St Vincent's Hospital Inclusive Health Fund

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multidisciplinary staff

- Adam Straub- Doctor
- Helen O'Neill CNC
- Amanda Streitburg pharmacist