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Presentation to the New Mexico State Legislative Health and Human Services Committee November 8, 2018

Oral Agonist Treatment (OAT)

- main treatment since 1960s
- methadone, buprenorphine
- for some patients, OAT works
 - decreases craving
 - decreases illicit injections
- but not effective for all
 - low retention rates
 - continued illicit drug use even during Rx

After 50 years of OAT treatments

- effective for some but sub-optimal
- not successful for everyone
- highest risk of overdose death comes right after coming off OAT
- opioid dependence remains a major public health and social problem

The Burning Clinical Question

How should we treat people who have not benefited from these existing treatments?

Try these existing treatments yet again?

Try something new?



The Choices

Try these existing treatments yet again?

Try something new?



Randomized Controlled Trial

NAOMI

NORTH AMERICAN OPIATE MEDICATION INITIATIVE

Injectable Diacetylmorphine (DAM)



Optimized Oral methadone

Randomized Controlled Trial

for chronic injection opioid users who previously did not benefit from available therapies and remain outside the addiction treatment system

Theory of Injectable Opioid Treatment

- many drug users remain out of treatment
- if we could attract them into treatment
 - stop endless "grind" of crime and sex work
 - stabilize chaotic lives and improve health
 - avoid HIV, HCV, other diseases, and death
 - provide counselling
 - provide other treatments if desired



Enrolment

Profile of the Participants

Characteristics	Total (n=251) % or mean	
Age	39.7	
Female Gender	38.6%	
First Nation	23.9%	
Precarious housing	72.9%	
Unemployed over the past 3 years	70.9%	
Public Assistance or Welfare	76.1%	
Illegal Sources of Income	67.3%	
Sex work	17.5%	
Ever convicted in life	81.7%	
Illegal activities, days in the prior 30	15	
Money spent on drugs, prior month	\$1500	
Number of prior Methadone Treatments	3	
Number of previous treatments	7	
Overdoses in life	4.1	



RR = 1.62; p<0.001; 95% CI= 1.35-1.95

RR = 1.40; p=0.004; 95% CI= 1.11-1.77

"Street" Heroin Use

NORTH AMERICAN OPIATE MEDICATION INITIATIVE



The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Diacetylmorphine versus Methadone for the Treatment of Opioid Addiction

Eugenia Oviedo-Joekes, Ph.D., Suzanne Brissette, M.D., David C. Marsh, M.D., Pierre Lauzon, M.D., Daphne Guh, M.Sc., Aslam Anis, Ph.D., and Martin T. Schechter, M.D., Ph.D.

N ENGLJ MED 361;8 NEJM.ORG AUGUST 20, 2009



The Hydromorphone Surprise



Street heroin use



Type of opioid the patient thought s/he received during treatment	DAM (n=115)	HDM (n=25)	Total (n=140)
Heroin definitely – n (%)	53 (46.1)	8 (32.0)	61 (43.6)
Heroin possibly – n (%)	18 (15.7)	8 (32.0)	26 (18.6)
Not sure – n (%)	22 (19.1)	4 (16.0)	26 (18.6)
Dilaudid [®] possibly – n (%)	9 (7.8)	3 (12.0)	12 (8.6)
Dilaudid [®] definitely– n (%)	3 (2.6)	0 (0.0)	3 (2.1)
No data* – n (%)	10 (8.7)	2 (8.0)	12 (8.5)

(*) One participant switched to oral methadone without receiving any HDM dosage; 2 participants did not complete the treatment retention questionnaire after leaving the clinic; 9 participants completed the retention questionnaire before the evaluation of the double bind was introduced.

SALOMÉ

Study to Assess Longer-term Opioid Medication Effectiveness

SALOME: Total Street Acquired Opioid Use



JAMA Psychiatry

Research

Original Investigation

Hydromorphone Compared With Diacetylmorphine for Long-term Opioid Dependence A Randomized Clinical Trial

Eugenia Oviedo-Joekes, PhD; Daphne Guh, MSc; Suzanne Brissette, MD; Kirsten Marchand, BSc; Scott MacDonald, MD; Kurt Lock, BA; Scott Harrison, MA; Amin Janmohamed, MSc; Aslam H. Anis, PhD; Michael Krausz, MD; David C. Marsh, MD; Martin T. Schechter, MD

Questions about Injectable Opioid Treatment

1. Can this be done safely?

Vancouver Injection Room



Safety



- addiction medicine specialists
- nursing
- safety procedures
 - pre and post-injection assessments
 - time and identifier bar codes
 - pharmaceutical GMP
 - emergency protocols





NORTH AMERICAN OPIATE MEDICATION INITIATIVE



- 109,171 treatment injections
- common expected side effects:
 - drowsiness
 - local histamine reaction (itchiness, pins and needles)
- only 13 episodes of serious over-sedation
 - 1 per 8,400 injections
 - treated on site with O₂ and naloxone
 - no hospitalizations; no lasting effects
 - often associated with benzodiazepines

Questions about Injectable Opioid Treatment

- 1. Can this be done safely?
- 2. Won't drug users want ever higher doses?



Day since Treatment Initiation

Average Daily Dose of DAM

Questions about Injectable Opioid Treatment

- 1. Can this be done safely?
- 2. Won't drug users want ever higher doses?
- 3. How can we possibly afford this?

Cost Considerations

- Dutch findings:
 - program costs more
 - decreased crime, enforcement, medical costs
 - overall savings: 12,793 euros (p.p. per year)
- NAOMI Cost-effectiveness Evaluation
 - about \$23 per day for DAM treatment
 - dominant strategy better outcomes, lower cost
- SALOME Cost-effectiveness Evaluation
 - about \$23 per day for DAM treatment
 - dominant strategy better outcomes, lower cost

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Cost utility analysis of co-prescribed heroin compared with methadone maintenance treatment in heroin addicts in two randomised trials

Marcel G W Dijkgraaf, Bart P van der Zanden, Corianne A J M de Borgie, Peter Blanken, Jan M van Ree, Wim van den Brink

CMAJ

Research

Cost-effectiveness of diacetylmorphine versus methadone for chronic opioid dependence refractory to treatment

Bohdan Nosyk PhD, Daphne P. Guh MSc, Nicholas J. Bansback PhD, Eugenia Oviedo-Joekes PhD, Suzanne Brissette MD, David C. Marsh MD, Evan Meikleham MSc, Martin T. Schechter MD PhD, Aslam H. Anis PhD

ADDICTION





doi:10.1111/add.14171

Cost-effectiveness of hydromorphone for severe opioid use disorder: findings from the SALOME randomized clinical trial

Nick Bansback^{1,2}, Daphne Guh¹, Eugenia Oviedo-Joekes^{1,2}, Suzanne Brissette³, Scott Harrison⁴, Amin Janmohamed¹, Michael Krausz^{1,5}, Scott MacDonald⁴, David C. Marsh⁶, Martin T. Schechter^{1,2} & Aslam H. Anis^{1,2}

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Thank you for your attention



Heroin

The Harms of Opioids

Harm

euphoria/sedation withdrawal constipation flushing overdose and death viral infections bacterial infections violence illegal activity social disintegration



Harm	The Drug?	The Circumstances?
euphoria/sedation	X	
withdrawal	X	
constipation	X	
flushing	X	
overdose and death		X
viral infections		X
bacterial infections		X
violence		X
illegal activity		X
social disintegration		X





<u>Societal Additives</u> crime, prison, violence, disorder, disintegration, hospitalization, infection, death..... <u>Physical Additives</u> fentanyl, starch, icing sugar, puddle water.....

- Remove the Societal Additives
- Remove the Physical Additives

What would be left?

Heroin







 $C_{21}H_{23}NO_5$



DiacetyImorphine or Diamorphine

Heroin



dangerous street drug

Heroin Diacetylmorphine





dangerous street drug

medicine

Am. J. Ph.] 7 [December, 1901 BAYER Pharmaceutical Products HEROIN-HYDROCHLORIDE

is pre-eminently adapted for the manufacture of cough elixirs, cough balsams, cough drops, cough lozenges, and cough medicines of any kind. Price in 1 oz. packages, \$4.85 per ounce; less in larger quantities. The efficient dose being very small (1-48 to 1-24 gr.), it is

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* marketing name from the German "heroisch" for "heroic, strong"

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