

OUTLINE

- History and Evolution of Opioid Agonist Therapy(OAT)
- Evidence of Effectiveness of Maintenance
- · Safety Issues
- Methadone Medical Maintenance (OBOT)
- Stigma Issues
- Barriers to Long Term Maintenance
- Conclusions
- Discussion

Treatment of Opioid Addiction

- Medication Assisted: Therapy, Treatment, Recovery
- Opioid Full/Partial Agonist Therapy (OAT): Methadone, Buprenorphine, (LAAM)
- Opioid Antagonist Therapy: Naltrexone Tablets and Depot I.M. Injection
- Medication Plus Psychosocial--Optimal
- Drug Free Recovery-"Abstinence Based"
- Mutual Help, CBT, DBT, MI, etc.

PC MAT TRAINING SS PROVIDERS' CLINICAL SUPPORT SYSTEM

MEDICATION ASSISTED ADDICTION TREATMENT

"All Treatments Work For Some People/Patients" "No One Treatment Works for All People/Patients"

> Alan I. Leshner, Ph.D Former Director NIDA

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MEDICATION ASSISTED ADDICTION TREATMENT

For Emphasis and Clarity,

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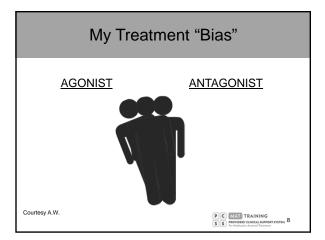
P C MAT TRAINING S S PROVIDERS' CLINICAL SUPPORT SYSTE For Medication Assisted Transment

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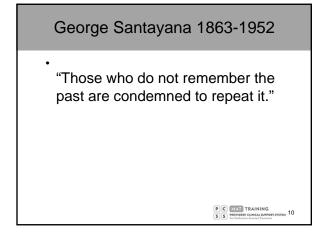
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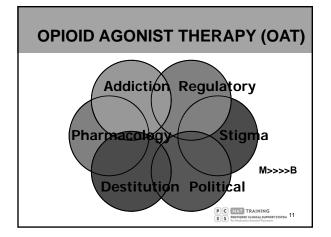




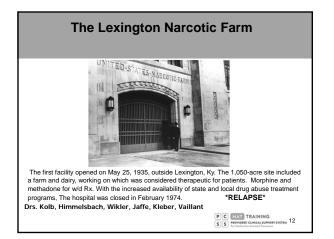
Webinars; PCSS-MAT, PCSS-O

- Drs. Bisaga and Sullivan: Naltrexone, PCSS-MAT 7/21/14, 1/13/15
- Dr. David Fiellin: Buprenorphine, PCSS-O 12/15/14
- Dr. Judith Martin: Methadone, PCSS-O 1/21/15
- Dr. Kevin Sevarino: Neuroadaptations to Opioids, PCSS-MAT 10/9/14
- Dr. Daniel Alford: Managing Acute and Chronic Pain in Patients Maintained on OAT, PCSS-MAT 8/12/14

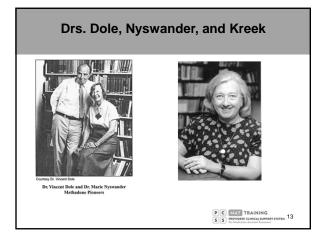


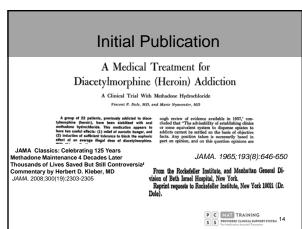


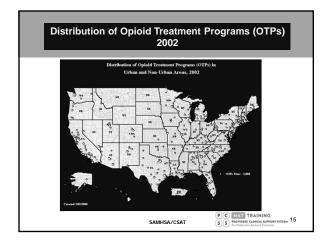


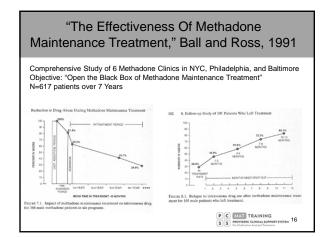




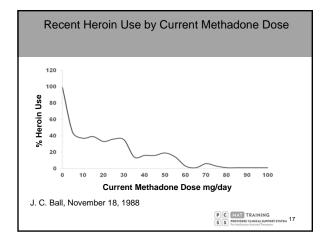




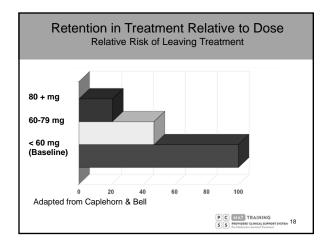












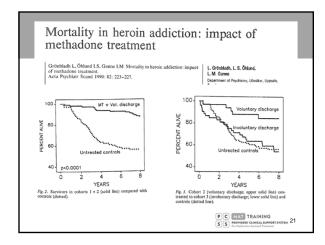






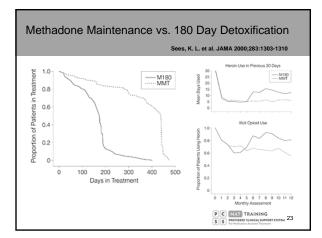
DATA 2000: Buprenorphine

- Major Paradigm Shift: OBOT vs MMTP
- Mechanism of Action: Similar to methadone
- Partial Agonist: Safety Implications
- 12 years of use in USA
- Now, more patients treated with Bupe than methadone
- Some of the same issues developing:
- 1. Diversion, Misuse, Abuse
- 2. Dosage
- 3. Duration
- 4. Other Drug Use Disorders
- 5. Access6. Insurance Coverage, Prior Authorizations

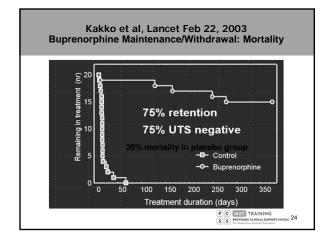




Psychosociall	Maintenance vs 180-Day y Enriched Detoxification
for Treatment	t of Opioid Dependence
A Randomized Co	
Karen I., Sees, DO	
Kevin L. Delucchi, PhD	 Context Despite evidence that methadone maintenance treatment (MMT) is effective for opioid dependence, it remains a controversial therapy because of its indefinit
Carmen Masson, PhD	provision of a dependence-producing medication.
Amy Rosen, PsyD	Objective To compare outcomes of patients with opioid dependence treated wit MMT vs an alternative treatment, psychosocially enriched 180-day methadone
H. Westley Clark, MD	
Helen Robillard, RN, MSN, MA	Design Randomized controlled trial conducted from May 1995 to April 1999.
Peter Banys, MD	Setting Research clinic in an established drug treatment service.
Sharon M. Hall, PhD	Patients Of 858 volunteers screened, 179 adults with diagnosed opioid depen
Sharon M. Hall, PhD	Patients Of 858 volunteers screened, 179 adults with diagnosed opioid dep



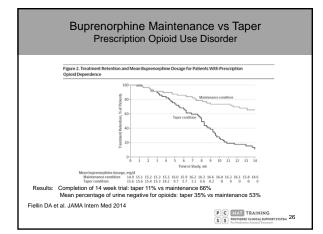




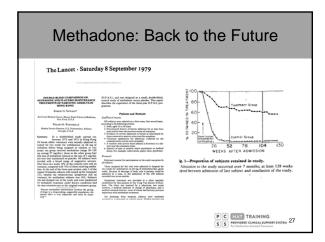


F	POATS	Study			
	ORIGINAL	ARTICLE			
ONLINE FIEST Adjunctive Co Buprenorphine Opioid Depend A 2-Phase Randomized Rept D. Weis, MD, Pandy R. Hilay S. Consery, MD, FMD, W Mar N. Conservid, MD, FMD, W Mar N. Conservid, MD, FMD, Reb Jafry Selwa, JDD, Rader Co.	2-Naloxone T dence I Controlled Trial arpe Potter, PhD; David A. F Iliam Dichuson, DO; John G Delovah L. Haller, PhD, Ab UsinBhad, MD; Eliware F. N	reatment for idlin, MD; Marilyn Byrne, M andin, PhD; Margaret L Grij ert L Hasson, MSW; Zhen Ha (Cance-Katz, MD; Scott E P	Prescriptie	on	
Table 2. Successful Opioid Use Outco			ime Points		
Time Point	Observed, No./Tota	al No. (%) (95% CI) SMM+0DC	GEE Model-Based OR (95% CII ^A	P Value	
End of phase 1 Phase 2, end of treatment Phase 2, 8-wk positireatment follow-up	24024 (7.4) [48-10.8] 84/18((46.7) [392-54.2] 13/18((7.2) [3.9-12.0]	19329 (5.8) [8.5-8.9] 93/189 (51.7) [44.1-59.2] 18/189 (10.0) [6.0-15.3]	1.3 (0.7-2.4) ^b 0.8 (0.5-1.2) ^c 0.7 (0.3-1.3) ^c	.36 .27 .22	
Abbreviations: GEE, generalized estimating e ³ The reference category is SMM + ODC. ⁹ Adjusted for chronic pain at baseline and lift ⁶ Adjusted for chronic pain at baseline, lifetin	fetime history of heroin use.		I, standard medical manage	ment.	
			SS PC	MIDERS' CLINICAL SUPPO	RT SYSTEM







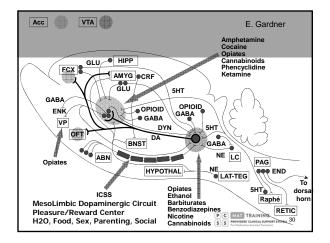


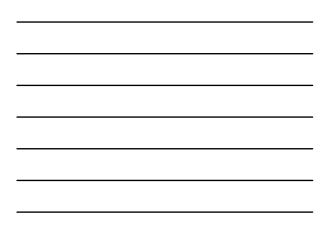


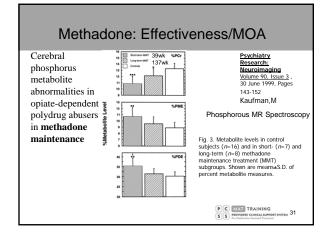
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	Bu	prer	orphine:	Recu	rrent	Relap	ose
30	vo male. B	upreno	rphine was effe	ctive. Sian	ificant ps	vchosocia	l problems.
			job, and many				
			or unwilling to a				
ove	er maintena	ance pa	radigm. Advise	d to return	for treatn	nent. Lost	to F/U.
				End	Start		
	ICHAEL ICHAEL 2	12/22/79	Suboxone 8mg qd Suboxone 8mg ad Film	7/23/10	6/19/09	XXX drop out 1 XXX drop out 2	oxycodone, oxycontin Relapse oxycodone IR ER
	ICHAEL 3 ICHAEL 4	12/22/79	Suboxone 12 mg Film Suboxone 12mg Film	2/21/12	12/21/11	XXX drop out 3 XXX drop out 4	Relapse oxycodone IR ER Relapse oxycodone IR ER
	ICHAEL 4	10201919	Suboxone 12mg Film	5/13/2012	4/13/2012	XXX drop out 4	Relapse oxycodone IR ER
					_		
					1		LINICAL SUPPORT SYSTEM 28



Buprenorphine: Dosage Issue
Addiction 33
Treatment retention among patients randomized to buprenorphine/aloxone compared to methadone in a multi-site trial
The lag from, halves 1, taxas 1, chank 1, takas
Review
Buprenorphine maintenance and <i>mu</i> -opioid receptor availability in the treatment of opioid use disorder: Implications for clinical use and policy
Mark K. Greenwald ^{a,} *, Sandra D. Comer ^b , David A. Fiellin ^c
Drug and Alcohol Dependence, 144, 2014
B C MART TRAINING S Prometer consciounders were service and the service servi









Methadone: Effectiveness/MOA

From these data, we conclude that polydrug abusers in MMT have 31P-MRS results consistent with abnormal brain metabolism and phospholipid balance. The nearly normal metabolite profile in long-term MMT subjects suggests that prolonged MMT may be associated with improved

neurochemistry.

Psychiatry Research: Neuroimaging Volume 90, Issue 3, 30 June 1999, Pages 143-152

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Methadone: Effectiveness/MOA

Article

Acute Effect of Methadone Maintenance Dose on Brain fMRI Response to Heroin-Related Cues

Daniel D. Langleben, M.D.	Objective: Environmental drug-related cues have been implicated as a cause of	Results: The heightened responses to heroin-related stimuli in the insula.
Kosha Ruparel, M.S.E.	illicit heroin use during methadone main- tenance treatment of heroin depen-	amygdala, and hippocampal complex, but not the orbitofrontal and ventral an-
Igor Elman, M.D.	dence. The authors sought to identify the functional neuroanatomy of the brain re-	terior cingulate cortices, were acutely re- duced after administration of the daily
Samantha Busch-Winokur, B.A.	sponse to visual heroin-related stimuli in methadone maintenance patients.	methadone dose.
Ramapriyan Pratiwadi, B.S.E.	Method: Event-related functional mag- netic resonance imaging was used to	Conclusions: The medial prefrontal cor- tex and the extended limbic system in
James Loughead, Ph.D.	compare brain responses to heroin-re- lated stimuli and matched neutral stimuli	methadone maintenance patients with a history of heroin dependence remains re-
Charles P. O'Brien, M.D., Ph.D.	in 25 patients in methadone mainte- nance treatment. Patients were studied	sponsive to salient drug cues, which sug- gests a continued vulnerability to relapse.
Anna R. Childress, Ph.D.	before and after administration of their regular daily methadone dose.	Vulnerability may be highest at the end of the 24-hour interdose interval.
		(Am J Psychiatry 2008; 165:390–394)

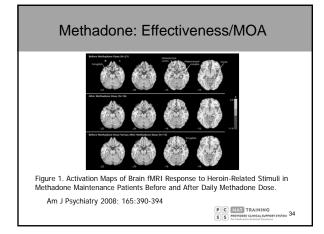
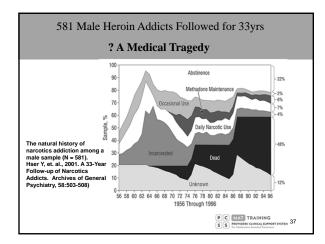




TABLE 3. Stress Response Hormones	HPA AXIS	;
	Adrenocorticotropic hormone	Cortisol
Short-acting opiates Opiate withdrawal	ł	ļ
Methadone Buprenorphine	**	
Naltrexone (oral) Naltrexone (extended release)	ţ	ţ
† = stimulate; ↓ = suppress, ↔ = no charge. Gavin Bart MD, FACP, FASAM (2012) Maintenance Medication f Journal of Addictive Diseases, 31	or Opiate Addiction: The Foundation	of Recovery,

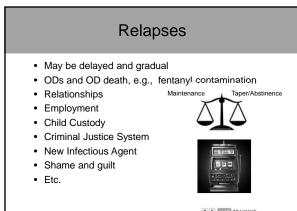
Infection New York City:	alence of HIV-1 (AIDS Virus) on in Intravenous Drug Users 1983 - 1984 Study: Protective Effect of done Maintenance Treatment
50 - 60%	Untreated, street heroin addicts: Positive for HIV-1 antibody
9%	Methadone maintained since<1978 (beginning of AIDS epidemic): less than 10% positive for HIV-1 antibody
Kreek , 1984; Des Jarlai	is et al., 1984; 1989
	PC MAT TRAINING 5 Promotion Councel Approved Particle 36 Kreek, 1984; Des Jariais et al., 1984; 1989





Hser Y, et. al., 2001. A 33-Year Follow-up of Narcotics Addicts. Archives of General Psychiatry, 58:503-508

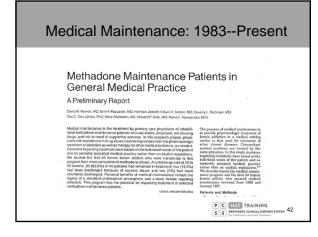
- California cohort of heroin addicted males-CJS
- After 15 years of abstinence, 25% relapsed to heroin
- Participation rates in methadone maintenance were <10% in any given year



Duration:Safety	y
Drug and Ainshol Dependence, 33 (1969) 225 - 245 Eliterner Sciettille Publishers terland LM.	225
The medical status of methadone maintenanc treatment for 11–18 years	ce patients in N=111
David M. Novick ^{a,b} , Reverly L. Richman ^c , Jeffrey M. Fried Friedman [*] , Christine Fried [*] , Janifer P. Wilson [*] , Anita Townle Kreek [*]	man ^a , Jacqueline E.
*The Eucleptiter University, New York, NY 19821 and "Department of Medicine and 'Media. Program, Bath Isreal Medical Center, New York, NY 19021 (55, (Account) Aug. 1, 1983).	
As compared to active IV heroin users the methadone patients g sexual dysfunction, Chronic liver disease was common, and anto "No clusters of unusual medical complications were observations we	edated methadone treatment.
	P C MAT TRAINING S S Paroneers concerns support system 40

OAT Duration: Safety

- Avoid OD: Induction Methadone Deaths→Pain Rx
- Drug/Drug Interactions: M>B
- Constipation
- Sweating
- Secondary Hypogonadism; ?M>B
- QTc Prolongation: M
- Other: Nausea, arousal, sedation, etc.
- No Organ Damage: Compare to Alcohol, Cocaine and Tobacco
- "Rots Teeth and Bones:" An enduring myth



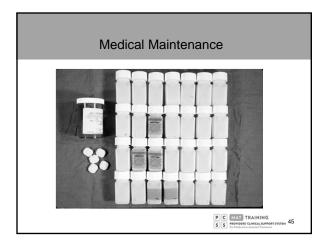
Medical Maintenance Admission Criteria

- At least 4 years in MMTP
- Negative urines for last 3 years
- Working/School etc.
- Adequate income for fees
- Recommendation from clinic
- Not in military reserves
- Stable and safe storage environment

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Medical Maintenance Procedures

- Patient given 28 day supply of methadone, by MD, in disket/tablet form, every 4 weeks.
- Medication prepared by hospital pharmacy in usual Rx type bottle and label
- Routine urine toxicology
- Patient returns before "run out" date
- Primary care provided







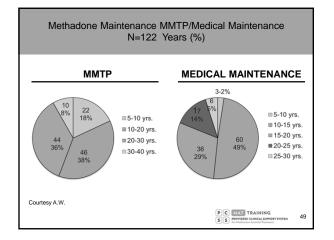




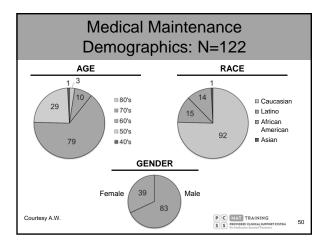




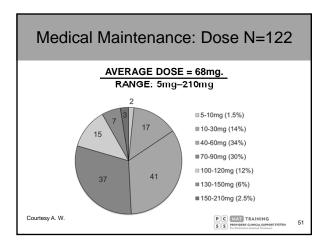




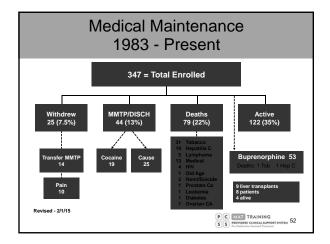








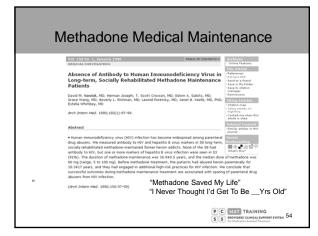




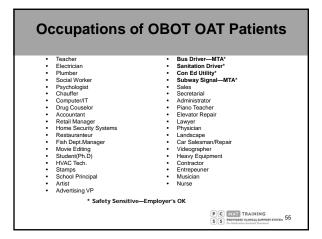


	eaths (22%)	
# of Patients	Cause	
31	Tobacco	
19	Hepatitis C	
5	Lymphoma	
13	Medical	
4	HIV	
1	Old Age	
2	Homicide/Suicide	
1	Prostate Cancer	
1	Leukemia	
1	Diabetes	
1	Ovarian Cancer	
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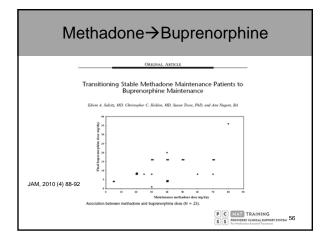








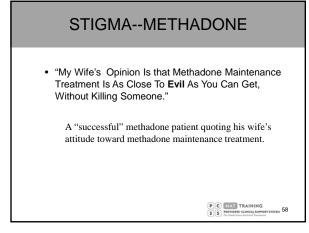


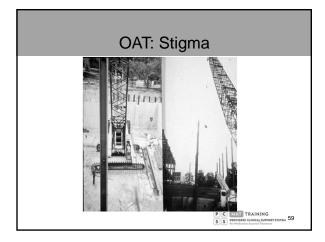


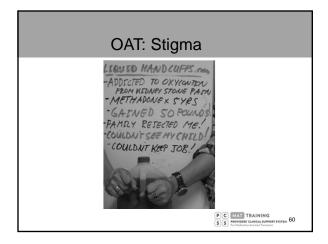


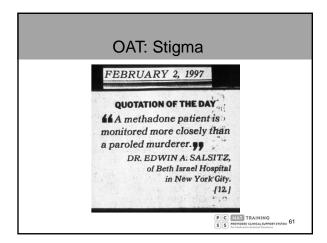


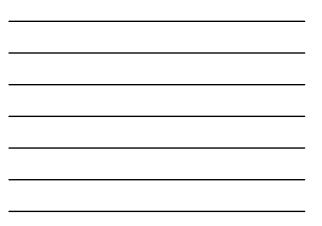


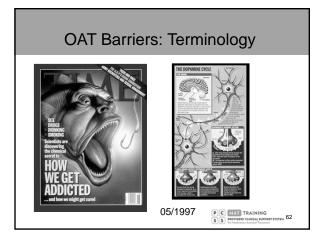




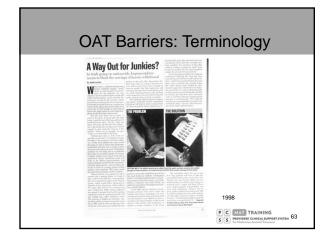


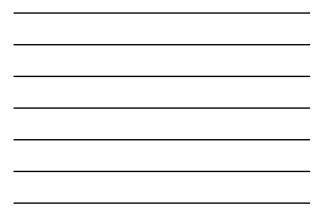




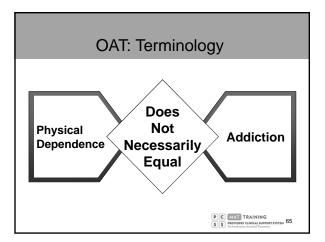




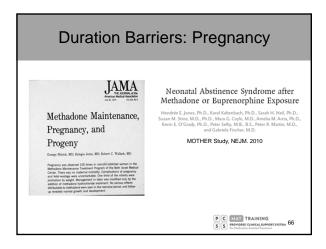


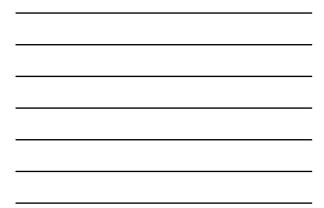


"Substitution Treatment" "OST" Standard terminology in Europe and Australia ?? Accurate ?? Helpful ?? Harmful "Aren't you just substituting one drug or addiction for another??" Why not just call it "Treatment for Opioid Use Disorder?"





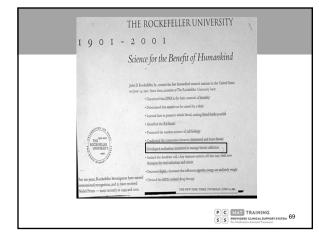


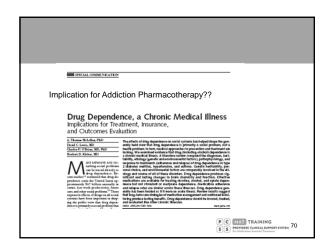


Duration Barriers: Policy

Mayor Steps Up His Criticism Of Methadone	e Maine governor proposes to eliminate Medicaid funding for OTPs
One day after detailing his plan to wean 2,000 heroin addicts off methadone at city hospitals, Mayor Rudolph W. Giuliani stepped up his attack on methadone treatment providers yesterday, accusing hem of enalaving former drug users instead of pushing them toward hostinence.	Gog Rhall Lebrages who has been try- impt to limit tourneers with media. Under his personal, Leir done, and bapersonphine in Maine file, several years, this month pro- posed to interact and the several several several ing, for epicetary land tearneers pregnant ing for epicetary land tearneers pregnant of 1071973 and methadnee, and to tam- if er praters to offici-bard barges form the folder government.
TURNAROUND RUDY PUTS \$5M IN Methadone clinics	
By Susan Rubinovitz October 6, 1999 4:00em	
By Soun Rhilowitz Ooker 6. 1999 400wn Mayor Giuliani has backed off further from his vow to end methadone treatment for herein addicts - funding a 55 million expansion of the city's clinics.	
Mayor Giuliani has backed off further from his vow to end methadone treatment for	

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"Ass Backwards" Paradigm

- The patients who have responded well to OAT, are the patients who are urged to "get off" their medication. They are often not rewarded with the Federal and State regulations for which they are entitled.
- The patients doing well, feel the most stigmatized.
- Protracted Abstinence Syndrome Both Physiologic(RR, T) and Psychological s/sx
- No other chronic medical disease is viewed this way by providers—asthma, hypertension, diabetes, depression
- No acceptance by insurers of long term maintenance, no longer requiring weekly UDTs or documented counseling

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What If There Were a Methadone or Buprenorphine for:

- Methamphetamine and Cocaine Addiction?
- Alcohol Addiction?
- Tobacco Addiction?
- Benzodiazepine Addiction?
- Food Addiction?
- Pathological Gambling?

PC MAT TRAINING SS PROVIDERS CLINICAL SUPPORT SYSTEM For Medication Assisted Transmet

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Final Comments: OAT Duration

- The scientific evidence base, and 50 years of clinical experience overwhelmingly support maintenance in the OAT treatment paradigm.
- The goal of OAT maintenance is not to see how fast a patient can "get off" medication.
- The goal is normalization and stabilization of the brain, establishing durable and safe hedonic tone, and functioning at maximal potential at home and at work.
- Like most chronic medical therapies, the medication only works, when it is taken.
- "If It Ain't Broke, Why Fix It?

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MEDICATION ASSISTED ADDICTION TREATMENT

"All Treatments Work For **Some** People/Patients" "No One Treatment Works for All People/Patients"

If your treatment is working, keep doing the treatment If your treatment is not working, change your treatment!!

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Why Is This So Important?



Actor Philip Seymour Hoffman, who was found dead February 2, 2014 on the bathroom floor of his New York apartment with a syringe in his left arm, died of acute mixed drug intoxication, **including heroin**, cocaine, benzodiazepines and amphetamine, the New York medical examiner's office said Friday

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Vincent Dole, Albert Lasker Award JAMA, 1988 *27 Years Ago

"It is postulated that the high rate of relapse of addicts after detoxification from heroin use is due to persistent derangement of the endogenous ligand-narcotic receptor system and that methadone in an adequate daily dose compensates for this defect. Some patients with long histories of heroin use and subsequent rehabilitation on a maintenance program do well when the treatment is terminated. The majority, unfortunately, experience a return of symptoms after maintenance is stopped. The treatment, therefore, is corrective but not curative for severely addicted persons. A major challenge for future research is to identify the specific defect in receptor function and to repair it. Meanwhile, specific defect in receptor function and to repair it. Meanwhile, methadone maintenance provides a safe and effective way to normalize the function of otherwise intractable narcotic addicted patients."

> P C MAT TRAINING TSYSTEM 76

References

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