

Interpretation of Drug Testing Results in Medication Assisted Treatment

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Disclaimer

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- ❑ Points of views or opinions in this document are those of the author and do not necessarily represent the official position or policies of the U.S. Department of Justice.

What Does This Result Mean?

Does the use of MAT drugs in an effort to promote recovery complicate the interpretation of drug testing results?

Two-Step Testing Approach

- screening test – designed to separate negative samples from samples that are “presumptively” positive
 - on-site screening devices
 - lab/court-based screening instrumentation
- confirmation test – lab-based follow-up procedure designed to validate positive test results
 - GC/MS
 - LC/MS/MS
- why can't you adjudicate based on the screening test results?
- FALSE POSITIVES

Drug tests & cross reactivity:

- ❑ screening tests can and do react to “non-target” compounds
 - ❑ amphetamines
 - ❑ benzodiazepines
- ❑ obtain list of interfering compounds from lab or on-site test vendor
- ❑ study results have demonstrated accuracy rates for initial screening tests as low as 70%
- ❑ confirm positive results

Typical Cutoff Levels

screening & confirmation

| | | |
|-------------------------|----------------|-----------|
| □ amphetamines * | 500 ng/mL | 250 ng/mL |
| □ benzodiazepines | 300 ng/mL | variable |
| □ cannabinoids * | 20 & 50 ng/mL | 15 ng/mL |
| □ cocaine (crack)* | 150 ng/mL | 100 ng/mL |
| □ opiates (heroin) * | 300/2000 ng/mL | variable |
| □ phencyclidine (PCP) * | 25 ng/mL | 25 ng/mL |
| □ alcohol | 20 mg/dL | 10 mg/dL |

□ * SAMHSA (formerly NIDA) drugs

What is a “cutoff” level ?

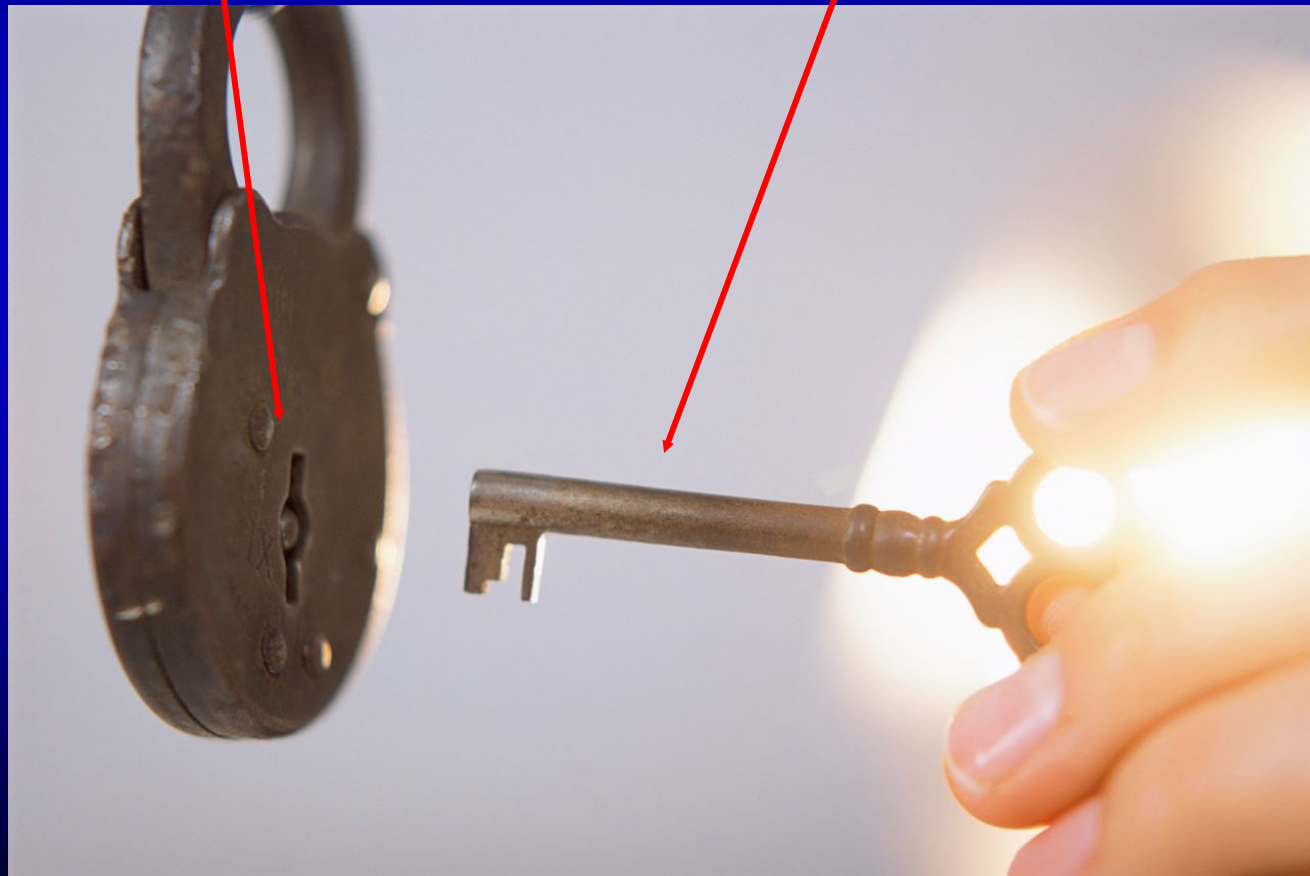
- ❑ cutoffs are not designed to frustrate CJ professionals
- ❑ a drug concentration, *administratively* established for a drug test that allows the test to distinguish between negative and positive sample - “threshold”
- ❑ cutoffs provide important safeguards:
 - ❑ scientific purposes (detection accuracy)
 - ❑ legal protections (evidentiary admissibility)
- ❑ measured in ng/mL = ppb

How Do Screening Drug Tests Work?

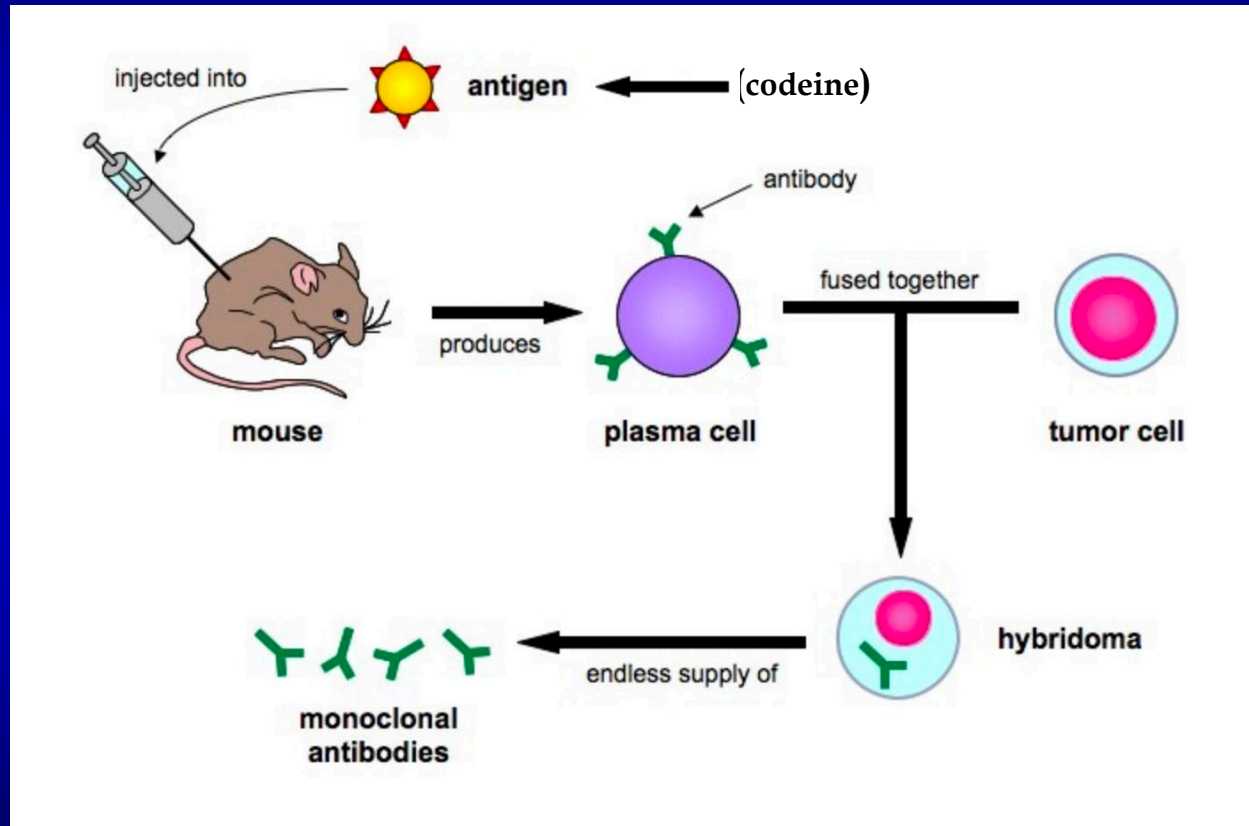
Drug tests & cross reactivity:

Drug Test

Drug



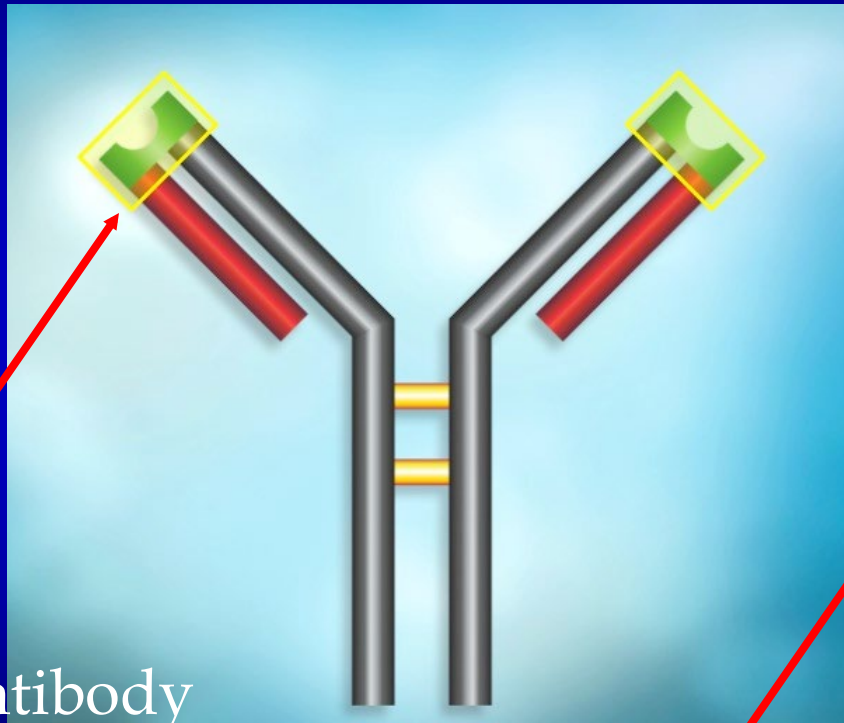
Let's Build an Opiate Screening Drug Test:



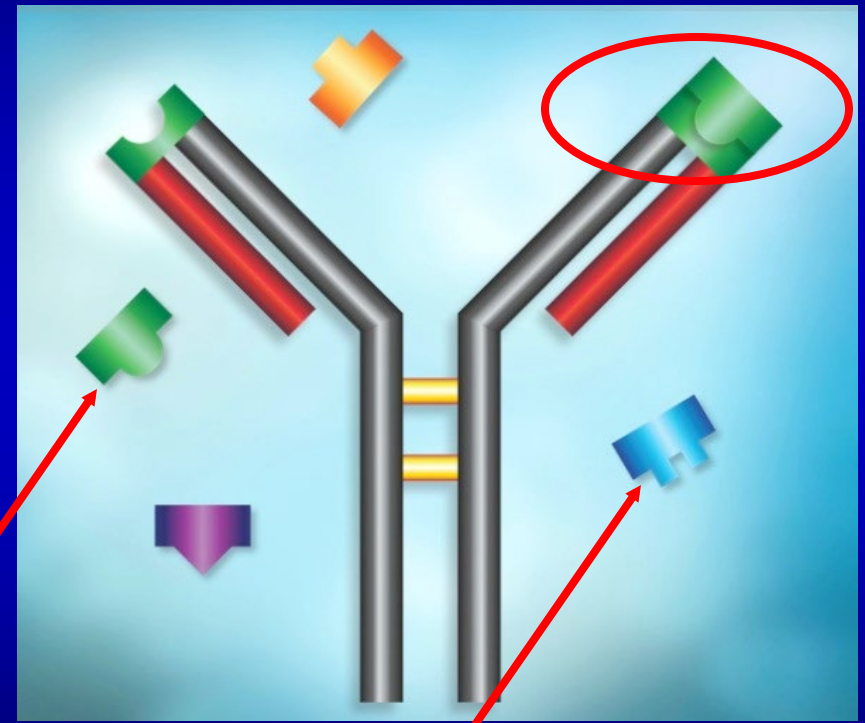
Creating antibodies using codeine

Drug tests & cross reactivity:

Immunoassay screening tests



opiates antibody



opiates (codeine) fit = positive test

non-opiates don't fit = negative test

Drug tests & cross reactivity:

| | |
|---------------|------|
| morphine | 100% |
| codeine | 200% |
| heroin | 80% |
| hydrocodone | 75% |
| hydromorphone | 45% |
| oxycodone | 20% |

Drug tests & cross reactivity:



(300 ng/mL opiate cutoff test)



150 ng/mL codeine



1500 ng/mL oxycodone



If oxycodone is a major substance of abuse in your jurisdiction, you should consider a separate drug test for oxycodone as part of your initial screening analysis.

Result Interpretation for MAT Drugs

MAT Drugs

- Medications for Alcohol Dependence
 - Naltrexone: (ReVia[®], Vivitrol[®], Depade[®])
 - Disulfiram: (Antabuse[®])
 - Acamprosate: (Campral[®])
- Medications for Opioid Dependence
 - Methadone:
 - Buprenorphine: (Suboxone[®] and Subutex[®])
 - Naltrexone: (ReVia[®], Vivitrol[®], Depade[®])

What is Naltrexone?

- ❑ belongs to a class of drugs known as opiate antagonists
- ❑ block the brain's neurotransmitters
- ❑ displaces opiates from their binding site
- ❑ diminishes physical effects of opiates
- ❑ will naltrexone test positive on an opiate drug test?

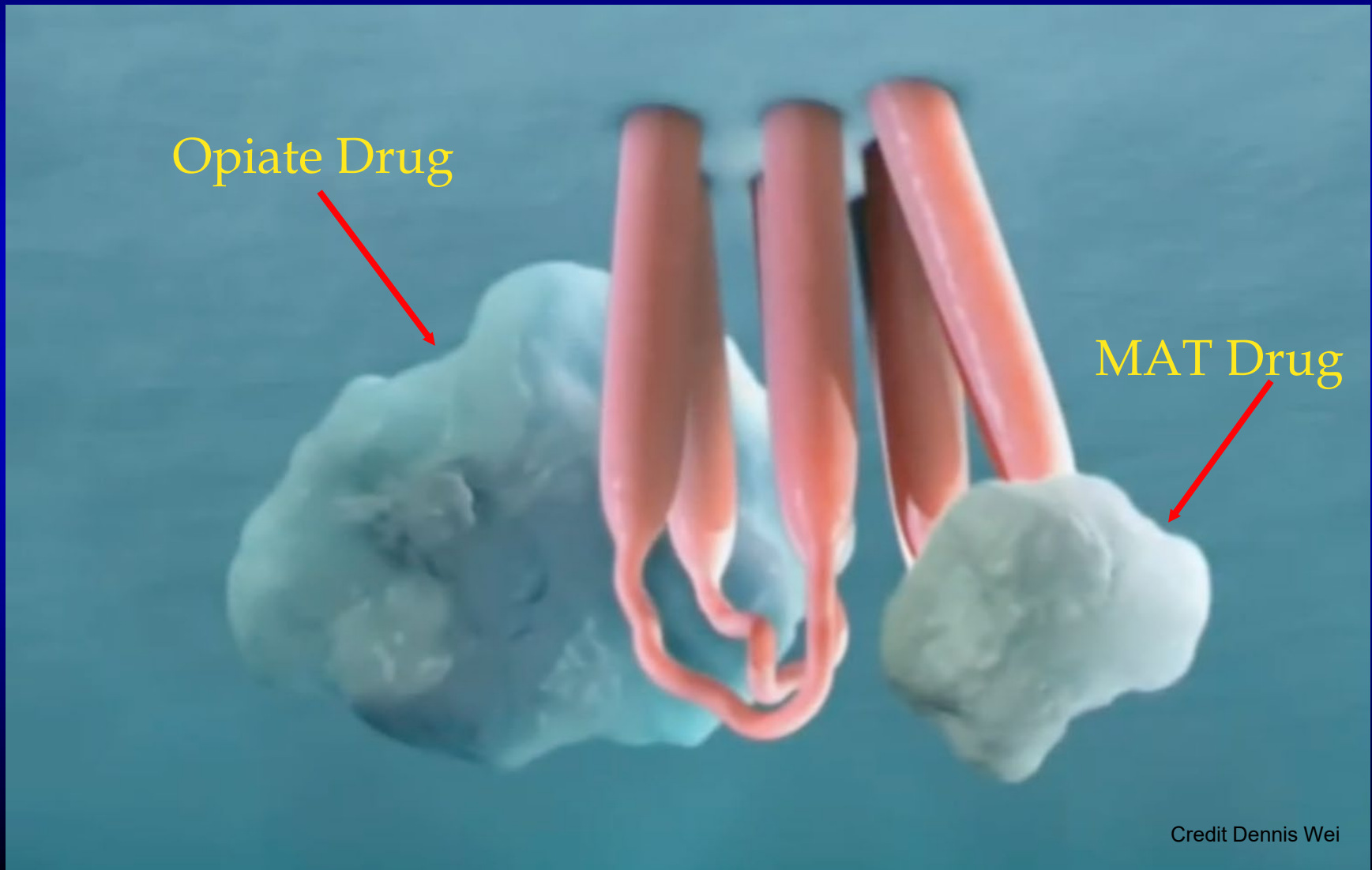
Neuron Transmission



Neural Surface Membrane

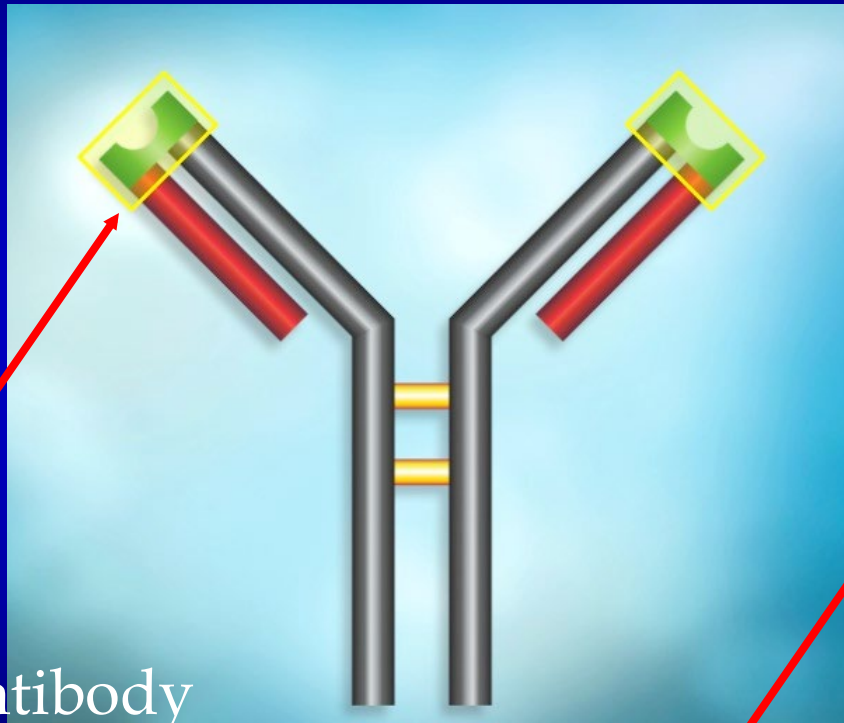


Ligand (MAT drug) Binds to Receptor

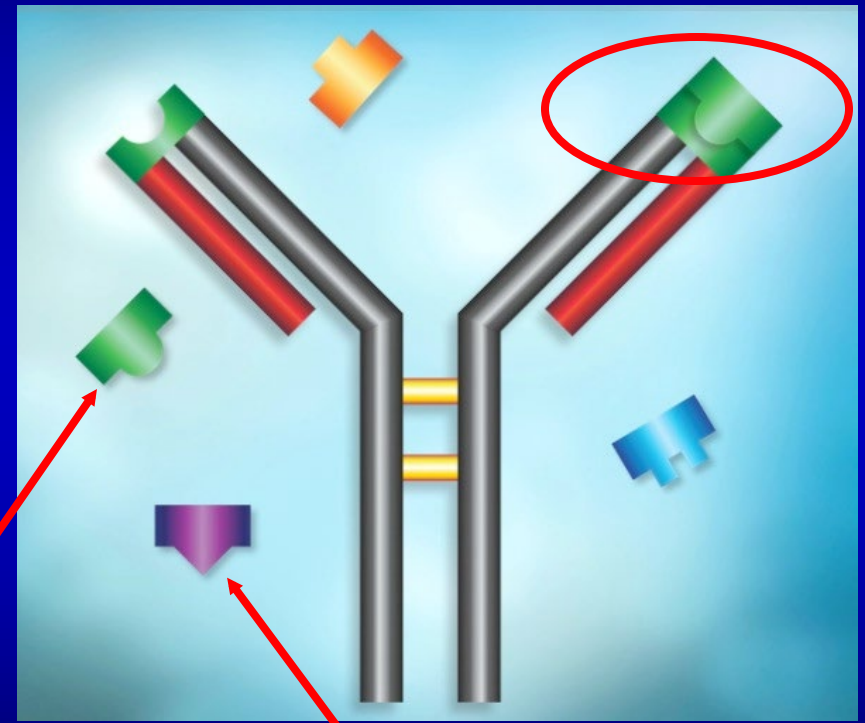


Drug tests & cross reactivity:

Immunoassay screening tests

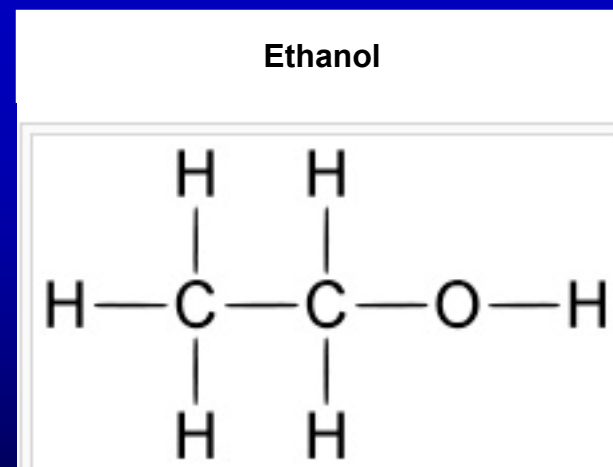
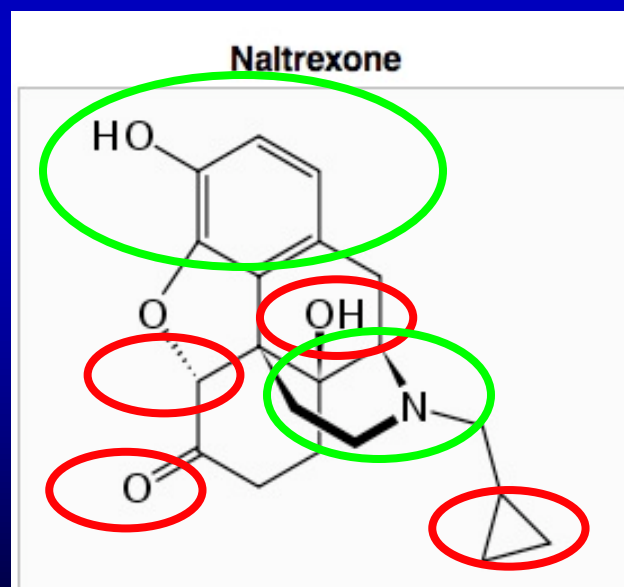
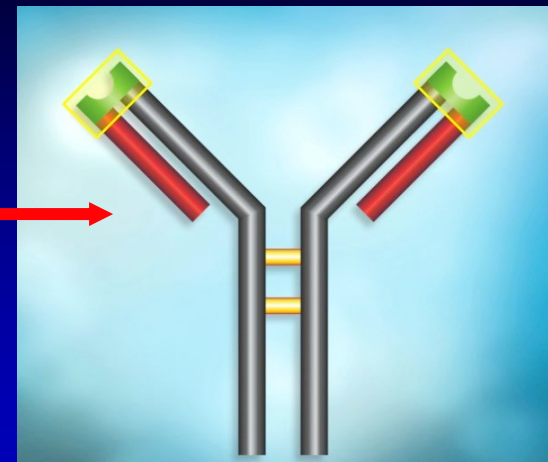
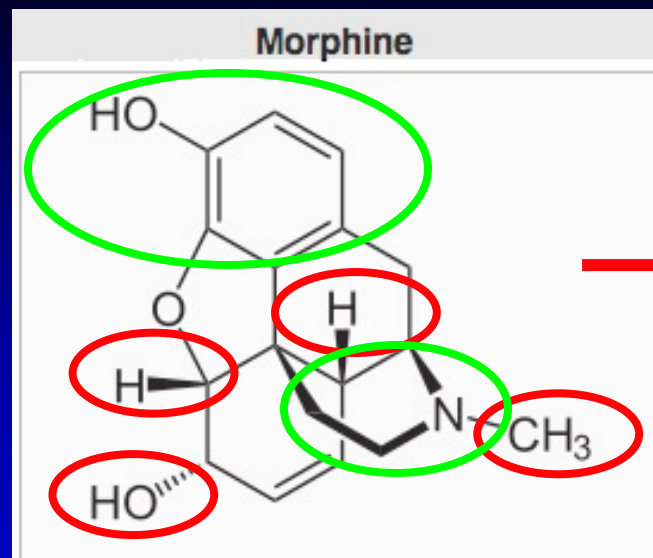


opiates antibody



opiates fit = positive test

does naltrexone "fit" and produce a "false positive" result



EMIT® II PLUS – OPIATE

Negative

The compounds listed below were negative for the Opiate 300 and 2000 cutoffs at the concentrations shown except where noted. Concentrations listed are in µg/mL.

| | | | |
|---|------|---|--------------|
| Acetaminophen | 1000 | EMDP | 100 |
| Acetylsalicylic Acid | 1000 | Enalapril Maleate | 1000 |
| Albuterol | 1000 | Ephedrine | 1000 |
| Alendronate | 1000 | Escitalopram | 1000 |
| Alprazolam | 1000 | Esomeprazole | 1000 |
| 5-Aminosalicylic Acid | 1000 | Eszopiclone | 1000 |
| Amitriptyline @ 300 | 500 | Ezetimibe | 1000 |
| Amitriptyline @ 2000 | 1000 | Fentanyl | 1000 |
| Amlodipine | 1000 | Fexofenadine | 1000 |
| Amoxicillin | 1000 | Fluoxetine | 900 |
| d-Amphetamine | 1000 | Fluticasone Propionate | 1000 |
| Atomoxetine | 1000 | Furosemide | 1000 |
| Atorvastatin | 1000 | Gabapentin | 1000 |
| Azithromycin | 1000 | Glutethimide | 500 |
| AZT (Zidovudine) | 2000 | Glyburide | 1000 |
| Benazepril | 1000 | Goldenseal | tea solution |
| Benzoylcegonine | 1000 | Griseofulvin | 1000 |
| Buprenorphine | 1000 | Hydrochlorothiazide | 1000 |
| Bupropion | 1000 | Ibuprofen | 1000 |
| Caffeine | 1000 | d,l-Isoproterenol | 1000 |
| Carisoprodol | 1000 | Isoxsuprine | 1000 |
| Celecoxib | 1000 | Ketamine | 100 |
| Cephalexin | 1000 | Ketoprofen | 1000 |
| Cetirizine | 1000 | Ketorolac Tromethamine | 1000 |
| Chlorpheniramine | 1000 | LAAM (l-α-Acetylmethadol) | 25 |
| Chlorpromazine | 125 | dinor LAAM (α-Acetyl-N, N-dinormethadol) | 25 |
| Cimetidine | 1000 | Lamotrigine | 1000 |
| Ciprofloxacin | 1000 | Lansoprazole | 1000 |
| Citalopram | 1000 | Lidocaine | 1000 |
| Clomipramine | 2.5 | Lisinopril | 1000 |
| Clonazepam | 1000 | Loperamide | 1000 |
| Clonidine | 1000 | Lormetazepam | 1 |
| Clopidogrel Hydrogen Sulfate | 1000 | LSD (Lysergic acid diethylamide) | 0.01 |
| Clotrimazole | 1000 | MDA (Methylenedioxymphetamine) | 5 |
| Cotinine | 100 | MDMA (Methylenedioxymethamphetamine) | 200 |
| Cyclobenzaprine | 63 | Meloxicam | 1000 |
| Desipramine | 800 | Meprobamate | 1000 |
| Dextromethorphan | 63 | Metaproterenol | 1000 |
| Dezocine | 1000 | Metformin | 1000 |
| Diazepam | 1000 | Methadone | 100 |
| Diclofenac | 1000 | d-Methamphetamine | 35 |
| Dihydroergotamine | 1000 | Methaqualone | 1500 |
| Diltiazem | 1000 | Metoprolol Tartrate | 1000 |
| Diphenhydramine | 1000 | Metoprolol | 1000 |
| Dothiepin | 100 | Myoglobin | 287 |
| Doxepin | 10 | Naltrexone | 1000 |
| Doxycycline | 1000 | NAPA (N-Acetylprocainamide) | 400 |
| Doxylamine | 500 | Naproxen | 1000 |
| Droperidol | 1000 | Nefazodone | 1000 |
| EDDP 2-Ethylidene-1, 5-dimethyl-3, 3-diphenylpyrrolidine | 1000 | Nortriptyline | 250 |
| | | Nylidrin | 1000 |

Siemens EMIT Assay Cross-Reactivity Data

| | |
|-----------------------------|------|
| Myoglobin | 287 |
| Naltrexone | 1000 |
| NAPA (N-Acetylprocainamide) | 400 |
| Naproxen | 1000 |

= 1,000,000 ng/mL

CEDIA® Opiate Cross-Reactivity Table
For catalog #s 100089, 100098 & 1661248

POSITIVE COMPOUNDS

The following compounds tested POSITIVE on the CEDIA® DAU Opiate assay at the 300 ng/mL cutoff.

| Positive Compounds | Trade Name | Concentration Tested (ng/mL) |
|------------------------|---------------------------------------|------------------------------|
| 6-Monoacetylmorphine | | 370 |
| Clomipramine HCl | Anafranil | 500,000 |
| Codeine | | 240 |
| Cyclazocine | | 500,000 |
| Cyamemazine | | 31,125 |
| Diacetylmorphine | Heroin | 570 |
| Dihydrocodeine | DHC Plus, Synalgos-DC | 600 |
| Hydrocodone | Lortab, Vicodin | 625 |
| Hydromorphone | Dilaudid | 530 |
| Levorphanol tartrate | Levo-Dromoran | 100,000 |
| Morphine | | 300 |
| Morphine SO4 | MS Contin, MSIR, Oramorph SR, Roxanol | 100,000 |
| Morphine-3-glucuronide | | 370 |
| Morphine-6-glucuronide | | 640 |
| Nalorphine HCl | | 100,000 |
| Naloxone | Narcan | 6,000 |
| Naltrexone HCl | Depade, ReVia | 50,000 |
| Ofloxacin | Floxin | 100,000 |
| Oxycodone | OxyContin | 320,000 |
| Pholcodine | | 500 |
| Rifampin | Rifadin | 65,000 |
| Thebaine | | 1,250 |

Abstract: A clinical evaluation of the naltrexone, a biodegradable sustained-release dosage was carried out in 4 healthy normal males.

Subjects were given an intravenous dose of 10 mg naltrexone and approximately 1 week later a 63-mg dose of naltrexone by subcutaneous administration.

Urine levels for naltrexone were 79-215 ng/mL.

| | | |
|----------------|---------------|---------|
| Naloxone | Narcan | 6,000 |
| Naltrexone HCl | Depade, ReVia | 50,000 |
| Ofloxacin | Floxin | 100,000 |

MAT Drugs

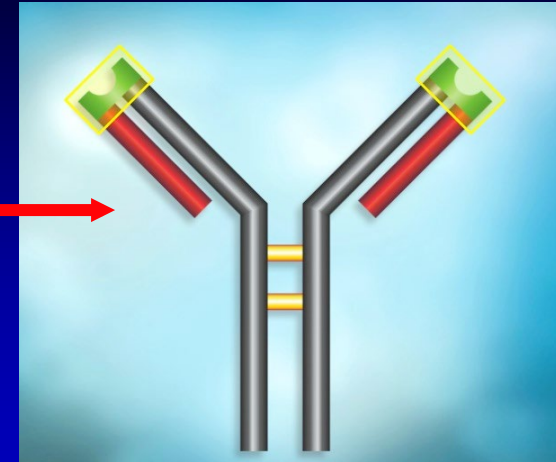
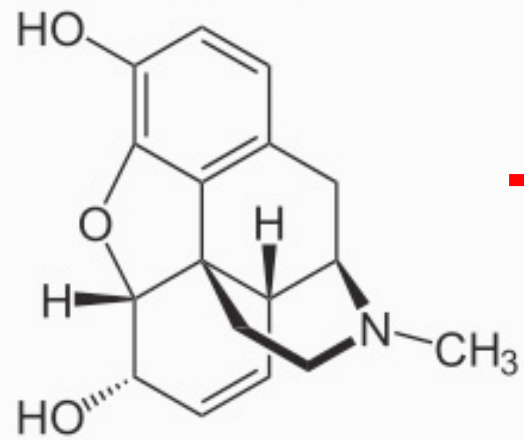
□ Medications for Alcohol Dependence

- Naltrexone: False Positive with Opiate Assay - NO!
- Disulfiram: (Antabuse®)
- Acamprosate: (Campral®)

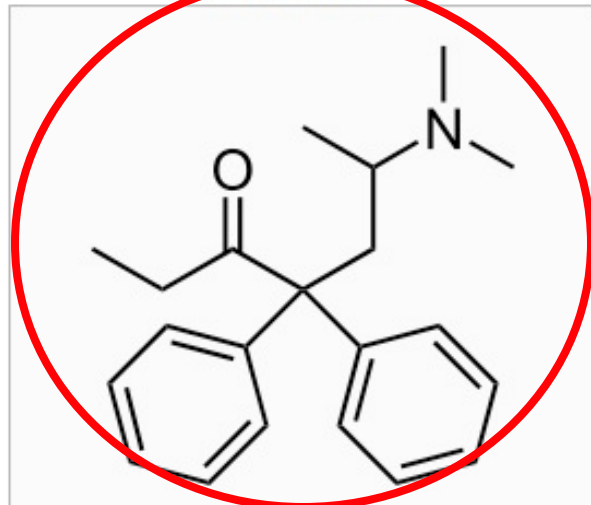
□ Medications for Opioid Dependence

- Methadone:
- Buprenorphine: (Suboxone® and Subutex®)
- Naltrexone: False Positive with Opiate Assay - NO!

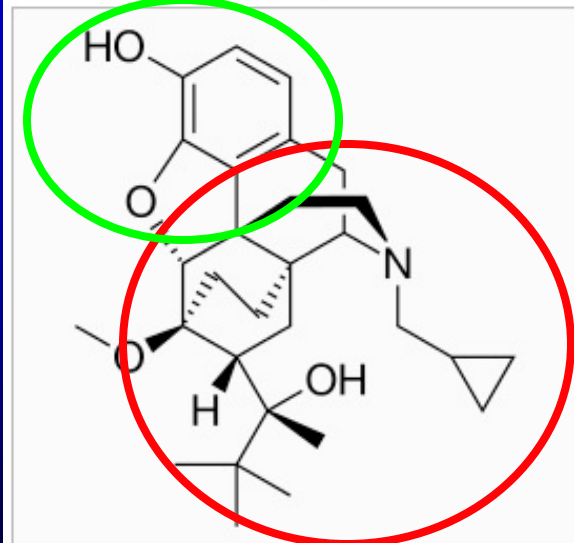
Morphine



Methadone



Buprenorphine



Siemens Negative Reactivity Data

| | |
|------------------|------|
| Azithromycin | 1000 |
| AZT (Zidovudine) | 2000 |
| Benazepril | 1000 |
| Benzoyllecgonine | 1000 |
| Buprenorphine | 1000 |
| Bupropion | 1000 |
| Caffeine | 1000 |

= 1,000,000 ng/mL

Thermo-Fisher Negative Reactivity Data

| Negative Compounds | Trade Name | Concentration Tested (ng/mL) |
|------------------------|--|------------------------------|
| Bromocriptine mesylate | Ergoset, Parlodel | 500,000 |
| Brompheniramine | Dimetane, Dimetapp, Nasahist, ND-Stat, Oraminic II | 500,000 |
| Bupivacaine | Marcaine, Sensorcaine | 500,000 |
| Buprenorphine | Buprenex | 100,000 |
| Bupropion | Wellbutrin, Zyban | 100,000 |

Siemens Negative Reactivity Data

| | |
|-------------------|------|
| Metaproterenol | 1000 |
| Metformin | 1000 |
| Methadone | 100 |
| d-Methamphetamine | 35 |
| Methaqualone | 1500 |

= 100,000 ng/mL

Thermo-Fisher Negative Reactivity Data

| | | |
|---------------------------------|---|---------|
| Metaproterenol hemisulfate salt | Alupent, Metaprel | 500,000 |
| Metaraminol bitartrate | Aramine | 500,000 |
| Methadone HCl | Dolophine | 500,000 |
| Methamphetamine | Desoxyn | 500,000 |
| Methaqualone HCl | Normi-Nox, Pallidan, Somnomed, Quaalude | 100,000 |

MAT Drugs

□ Medications for Alcohol Dependence

- Naltrexone: False Positive with Opiate Assay - NO!
- Disulfiram: (Antabuse®)
- Acamprosate: (Campral®)

□ Medications for Opioid Dependence

- Methadone: NO! with Opiate Assay
- Buprenorphine: NO! with Opiate Assay
- Naltrexone: False Positive with Opiate Assay - NO!

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| <input type="checkbox"/> Latest Update | <input type="checkbox"/> Pharmacologic Actions | |

Publication Year - -

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MAT Drugs

n Medications for Alcohol Dependence

- u Naltrexone: False Positive with Opiate Assay - NO!
- u Disulfiram: NO! with drug tests reviewed
- u Acamprosate: NO! with drug tests reviewed

n Medications for Opioid Dependence

- u Methadone: NO! with Opiate Assay
- u Buprenorphine: NO! with Opiate Assay
- u Naltrexone: False Positive with Opiate Assay - NO!



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Home › Q & A › Questions › Why does naltrexone post a...



chet1

Why does naltrexone post a positive opiate result on a UA test?

Posted: 4 Nov 2010 by chet1

Topics: [naltrexone](#), [opiate](#)

does naltrexone cause a positive opiate or benzo result on a UA test?

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Responses (1)



Marvell

5 Nov 2010

Because naltrexone is actually a special narcotic drug that blocks the effects of other narcotic [medicines](#) and alcohol. That's why it comes up in a urinary analysis as an opiate.

<http://www.drugs.com/mtm/naltrexone.html>

Votes: +0

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symptomfind.com/UrinaryProblems

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Opiates - Results Interpretation

- all opiates are narcotic analgesics
 - relieve pain & controlled substances
- not all narcotic analgesics are opiates
 - meperidine (Demerol[®])
 - propoxyphene (Darvon[®])
 - methadone
 - pentazocine (Talwin[®])
 - fentanyl (Sublimaze[®])
 - buprenorphine: (Suboxone[®])
 - naltrexone: (ReVia[®], Vivitrol[®], Depade[®])

Result Interpretation for Therapeutic/OTC Drugs

Very Difficult Task

- not all drug tests are created equal
 - laboratory-based tests (numerous products)
 - on-site, instant, POC tests (dozens of products)
 - each test has unique selectivity (i.e. ability to distinguish between similar compounds)
- hundreds of therapeutic drugs
- hundreds of OTC medications

Court's Obligation

- limit use of therapeutic drugs
 - court must be notified
- prohibit the use of OTC medications without prior approval
- prohibit the use of dietary supplements, energy drinks, homeopathic substances, herbal products, sports nutrition powders, anything not regulated by FDA (anything from GNC)

Drug Metabolism Guide

Opiate Metabolites

Parent Drug: Codeine

Metabolites: Norcodeine, Morphine,
(hydrocodone potential minor metabolite in
high codeine doses)

Parent Drug: Morphine

Metabolites: Normorphine

Parent Drug: Heroin

Metabolites: 6-monoacetyl morphine (6-AM),
Normorphine, Morphine

Parent Drug: Oxycodone

Metabolites: Oxymorphone, Noroxycodone,
Noroxymorphone

Opiate Metabolites

Parent Drug: Hydrocodone

Metabolites: Hydromorphone, Norhydrocodone

Parent: Hydromorphone (may only as parent drug)

Metabolites: undetectable conjugated metabolites

Benzo Metabolites

Parent: Alprazolam

Metabolites: alpha-hydroxyalprazolam

Parent: Lorazepam

Metabolites: Detected as parent drug;
undetectable metabolites

Parent: Clonazepam

Metabolites: 7-aminoclonazepam

Parent: Diazepam

Metabolites: Temazepam, Nordiazepam,
Oxazepam

Benzo Metabolites

Parent: Temazepam

Metabolites: Oxazepam

Parent: Chlordiazepoxide

Metabolites: Norchlordiazepoxide,
Nordiazepam, Oxazepam

Parent: Triazolam

Metabolites: only as parent drug;
undetectable metabolites

Parent: Clorazepate

Metabolites: Nordiazepam, Oxazepam

Therapeutic/OTC Drugs

Drug/Class

Potential F/P Results

- | | |
|--|---------------------|
| □ antihistamines/decongestants | amphetamines |
| □ Adderall | amphetamines |
| □ confirm by GC/MS - ensure no methamphetamine | |
| □ chlordiazepoxide | benzodiazepine |
| □ confirm by GC/MS - look for other benzos not metabolites of chlordiazepoxide | |
| □ dextromethorphan | phencyclidine (PCP) |
| □ <i>l</i> -methamphetamine (OTC nasal inhaler) Vick' s | amphetamines |
| □ diet pills (eg, clobenzorex, fenproporex) | amphetamines |
| □ quinolone antibiotics (eg, levofloxacin) | opiates |
| □ antidepressants (Stertraline) | benzodiazepine |

How to Drive a Toxicologist Crazy

My client claims he is testing positive for THC because he takes ibuprofen (Advil).

Urine Drug Screening: Practical Guide for Clinicians

KAREN E. MOELLER, PHARM D, BCPP; KELLY C. LEE, PHARM D, BCPP; AND JULIE C. KISSACK, PHARM D, BCPP

Drug testing, commonly used in health care, workplace, and criminal settings, has become widespread during the past decade. Urine drug screens have been the most common method for analysis because of ease of sampling. The simplicity of use and access to rapid results have increased demand for and use of immunoassays; however, these assays are not perfect. False-positive results of immunoassays can lead to serious medical or social consequences if results are not confirmed by secondary analysis, such as gas chromatography-mass spectrometry. The Department of Health and Human Services' guidelines for the workplace require testing for the following 5 substances: amphetamines, cannabinoids, cocaine, opiates, and phencyclidine. This article discusses potential false-positive results and false-negative results that occur with immunoassays of these substances and with alcohol, benzodiazepines, and tricyclic antidepressants. Other pitfalls, such as adulteration, substitution, and dilution of urine samples, are discussed. Pragmatic concepts summarized in this article should minimize the potential risks of misinterpreting urine drug screens.

Mayo Clin Proc. 2008;83(1):66-76

Our goal is to provide clinically relevant information that can be used to interpret urine drug screens (UDSs) for commonly abused drugs (ie, alcohol, amphetamines, benzodiazepines, opioids, marijuana, cocaine, phencyclidine [PCP], and tricyclic antidepressants [TCAs]). Proper evaluation of urine specimens, including detection times, are discussed, as well as false-positive results and potential false-negative results. Interpretation of tests for performance-enhancing drugs is beyond the scope of this article and is not discussed.

METHODS OF DRUG TESTING

Urine, blood, hair, saliva, sweat, and nails (toenails and fingernails) are some biological specimens used to perform laboratory drug testing, and they provide different levels of

TABLE 3. Summary of Agents Contributing to Positive Results by Immunoassay^a

| Substance tested via immunoassay | Potential agents causing false-positive result | Substance tested via immunoassay | Potential agents causing false-positive result |
|-------------------------------------|---|-------------------------------------|---|
| Alcohol ²⁰ | Short-chain alcohols (eg, isopropyl alcohol) | Cannabinoids ^{1,8,43-48} | Dronabinol |
| Amphetamines ²¹⁻⁴⁰ | Amantadine | | Efavirenz |
| | Benzphetamine | | Hemp-containing foods |
| | Bupropion | | NSAIDs |
| | Chlorpromazine | | Proton pump inhibitors |
| | | | Tolmetin |

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Investigation of Interference by Nonsteroidal Anti-Inflammatory Drugs in Urine Tests for Abused Drugs

Douglas E. Rollins,¹ Thomas A. Jennison,² and Graham Jones³

Anecdotal and uncontrolled studies have suggested that nonsteroidal anti-inflammatory drugs produce false-positive results in immunoassay urine tests for some drugs of abuse. This study was performed in 60 volunteers who took ibuprofen as either a single 400-mg dose, or 200 mg three times a day, or 400 mg three times a day, and in 42 patients taking ibuprofen, naproxyn, or fenoprofen in therapeutic regimens for more than 30 days. Of the 510 urines collected from 102 individuals during these dosage regimens, two gave false-positive tests for cannabinoid by enzyme-mediated immunoassay (EMIA), one after 1200 mg of ibuprofen in three divided doses for one day and one in a patient taking naproxyn on a chronic basis; none was falsely positive for

falsely positive report.

Conversely, adulterants (e.g., acids or bases or substances with a high ionic strength) added to a urine specimen may give falsely negative immunoassay results (1). Moreover, the excretion of drugs, drug metabolites, or food substances in the urine could also interfere with immunoassays and cause a false-positive or false-negative result for a urine drug assay. Ibuprofen and other commonly used nonsteroidal anti-inflammatory drugs (NSAIDs) reportedly cause false-positive test results with the EMIA (EMIT™; Syva Co., Palo Alto, CA) for cannabinoids (2-4), false-negative mass-spectrometric confirmation for cannabinoids (5), and false-positive results for barbiturates and benzodiazepines by the FPIA (TDx™; Abbott Laboratories, Diagnostic Division, Abbott Park, IL) (6,7).

oid tests (14).

Brunk (5) describes a false-negative GC/MS cannabinoid confirmation caused by high concentrations of urinary ibuprofen that competed with the analyte for the cannabinoid-derivatizing reagent. This is an unlikely explanation for the data presented in this study. In his GC/MS tests, Brunk extracted 10 mL of urine, which would contain large amounts of potentially competing substances. He used tetramethyl ammonium hydroxide as a derivatizing reagent, and did not use a deuterated internal standard. For cannabinoid confirmation in this study we used only 1 mL of urine, hexafluoroisopropanol and pentafluoropropionic anhydride as the derivatizing reagents, and deuterated COOH-THC as an internal standard. If competition between ibuprofen and COOH-THC for the derivatizing reagents had occurred, no peak for the internal standard would have been observed—a situation that did not occur for any specimen in this study.

In conclusion, these data demonstrate that ibuprofen taken as either a single dose or in acute multiple doses or ibuprofen, naproxyn, or fenoprofen taken as chronic doses is unlikely to result in a positive immunoassay test for urine cannabinoids, benzodiazepines, or barbiturates. All positive immunoassay results should be considered as presumptively positive. A second chemical test such as GC/MS, performed properly, will markedly reduce the possibility of falsely accusing of substance abuse someone who was taking NSAIDs.

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1. Article used by the Mayo paper claiming ibuprofen could cause a false positive cannabinoid test is 25 years old.
2. Even though the Rollins paper is 25 years old, it concludes “unlikely”.
3. Assay used to conduct the 25-year old paper has not been available commercially for two decades
4. Confirmation testing resolves potential “false positive” concerns.
5. Doesn't prohibit Mayo from publishing a misleading paper.

Commonly prescribed medications and potential false-positive urine drug screens

NANCY C. BRAHM, LYNN L. YEAGER, MARK D. FOX, KEVIN C. FARMER, AND TONY A. PALMER

The potential for false-positive urine drug screen (UDS) results for substances of abuse presents a therapeutic selection dilemma for the treating health care professional. While this problem is reported with specific medications, the extent of the problem in a clinic serving indigent patients and the medically underserved has not been evaluated. In particular, the use of medications with the potential for false-positive UDS results may present a significant liability for individuals required to undergo random or periodic UDSs as a component of a recovery or court-ordered monitoring program^{1,2} or as a condition of employment.^{1,3,4} In addition, false-positive UDS results may affect the clinician-patient relationship by raising issues of trust.⁵ This article identifies commonly used medications associated with reports of false-positive UDSs.

Literature review

A comprehensive literature review

Purpose. The implications of potential false-positive urine drug screen (UDS) results for patients receiving commonly prescribed medications were evaluated.

Summary. A comprehensive literature review was conducted to identify false-positive UDSs associated with all clinic formulary medications, as well as common nonprescription medications. The references of each report describing a medication whose use was associated with false-positive UDS results were also reviewed. If a class effect was suspected, additional agents in the category were searched. A total of 25 reports of false-positive UDS results were identified. Categories of medications included antihistamines, antidepressants, antibiotics, analgesics, antipsychotics, and nonprescription agents. Reports of false-positive results were found for the following formulary and nonprescription medications: brompheniramine, bupropion, chlorpromazine, clomipramine, dextromethorphan, diphenhydramine, doxylamine, ibuprofen, naproxen, promethazine, quetiapine, quinolones (ofloxacin and gatifloxacin), ranitidine, sertraline, thioridazine, trazodone, venlafaxine,

verapamil, and a nonprescription nasal inhaler. False-positive results for amphetamine and methamphetamine were the most commonly reported. False-positive results for methadone, opioids, phencyclidine, barbiturates, cannabinoids, and benzodiazepines were also reported in patients taking commonly used medications. The most commonly used tests to screen urine for drugs of abuse are immunoassays, even though false-positive results for drugs of abuse have been reported with a number of these rapid-screening products. Results from such tests should be confirmed using additional analytical methods, including gas chromatography-mass spectrometry.

Conclusion. A number of routinely prescribed medications have been associated with triggering false-positive UDS results. Verification of the test results with a different screening test or additional analytical tests should be performed to avoid adverse consequences for the patients.

Index terms: Drug abuse; Drugs, over the counter; Drugs; False positive reactions; Tests, laboratory; Urine levels
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Table 1.

Reports of False-Positive Results of Urine Drug Screens for Selected Formulary Agents⁶⁻³⁰

| Medication | False-Positive Result | | | | | | |
|-------------------------------------|--------------------------------|---------------|-----------|---------|-----------------|--------------|--------------|
| | Amphetamine or Methamphetamine | Phencyclidine | Methadone | Opiates | Benzodiazepines | Cannabinoids | Barbiturates |
| Antihistamines/decongestants | | | | | | | |
| Brompheniramine | X | | | | | | |
| Diphenhydramine | | | X | | | | |
| Doxylamine | | | X | | | | |
| Phenylpropanolamine | X | | | | | | |
| Nonprescription nasal inhaler | X | | | | | | |
| Antidepressants | | | | | | | |
| Bupropion | X | | | | | | |
| Clomipramine | | | X | | | | |
| Sertraline | | | | | X | | |
| Trazodone | X | | | | | | |
| Venlafaxine | | X | | | | | |
| Antibiotics | | | | | | | |
| Quinolones (selected agents) | | | | X | | | |
| Analgesics | | | | | | | |
| Ibuprofen | | X | | | | X | X |
| Naproxen | | | | | | X | X |
| Antipsychotics | | | | | | | |
| Chlorpromazine | X | | X | | | | |
| Promethazine | X | | | | | | |
| Quetiapine | | | X | | | | |
| Thioridazine | | | X | | | | |
| Other agents | | | | | | | |
| Dextromethorphan | | X | | | | | |
| Ranitidine | X | | | | | | |
| Verapamil | | | X | | | | |

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Confirmation: Best Practice

- gas or liquid chromatography-mass spectrometry - GC/MS or LC/MS/MS
 - drug molecules separated by physical characteristics
 - identified based on chemical “finger-print”
 - considered “gold standard”
- refer to NADCP Adult Drug Court Best Practice Standards - Volume II

CONCLUSIONS

- ❑ Using standard instrument-based screening immunoassay drugs tests (in-lab or in-court), MAT drugs do not cross-react to produce “false positive” results
- ❑ When using on-site testing devices the cross-reactivity toward MAT drugs is largely unstudied. Contact product vendor.
- ❑ Confirmation testing (GC/MS or LC/MS) resolves nearly all cross-reactivity “false-positive” issues

No Substitute for Knowledge/Expertise

- ❑ unethical to adjudicate based upon misinformation - violation of due process
- ❑ develop a relationship with your laboratory
- ❑ develop a relationship with your on-site device vendor
- ❑ don't be afraid to “call the company”
- ❑ seek expert advice

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