

## Urine Drug Testing Methods<sup>3-5</sup>

Type of Test	Logistics	Pearls
<b>Initial Screening Test:</b> Immunoassay	<ul style="list-style-type: none"> <li>• Inexpensive</li> <li>• Fast</li> <li>• Widely available</li> </ul>	<ul style="list-style-type: none"> <li>• High sensitivity, low specificity (higher potential for false positives)</li> <li>• Opiate screen not sensitive for semisynthetic (e.g. oxycodone) or synthetic opioids (e.g. fentanyl)</li> </ul>
<b>Confirmatory Test:</b> Gas chromatography-mass spectrometry (GCMS) <sup>+</sup> or Liquid chromatography-mass spectrometry (LCMS)	<ul style="list-style-type: none"> <li>• Expensive</li> <li>• Time consuming</li> </ul>	<ul style="list-style-type: none"> <li>• High sensitivity, high specificity</li> <li>• Expensive</li> <li>• Detects medication even if concentration is low</li> </ul>

<sup>+</sup> GCMS is considered the criterion standard for confirmatory testing; Immunoassay tests have high predictive values for marijuana and cocaine, but lower predictive values for opiates and amphetamines

### Urine Drug Testing Specimen Validity<sup>3-4</sup>

- Urine samples that are adulterated, substituted, or diluted may avoid detection of drug use<sup>4</sup>
- Urine collected in the early morning is most concentrated and most reliable
- Excessive water intake and diuretic use can lead to diluted urine samples (Creatinine < 20)<sup>3-4</sup>
- THC assays are sensitive to adulterants (e.g. Visine eyedrops)

### Normal Characteristics of a Urine Sample<sup>3-5</sup>

Temperature within 4 minutes of voiding: 90-100<sup>o</sup>F

pH: 4.5-8.0

Creatinine: > 20 mg/dL

Specific gravity: > 1.003

Nitrates: < 500 mcg/dL

Volume: ≥ 30 mL

## Urine Drug Testing (UDT) Federal Work Place Cut Off Values<sup>3-9</sup>

		Initial Drug Test Level (immunoassay) (ng/mL)	Confirmatory Drug Test Level (GC-MS) (ng/mL)	Confirmatory Test Analyte <sup>3,7</sup>	Detection Period After Last Dose (Days)*	
Extended UDT	Regular UDT	Marijuana Metabolites	50	15	THCA	2-8 single use 20-30 chronic use <sup>+</sup>
		Cocaine Metabolites	300	150	BEG	1-3
		Opioid Metabolites	2000 <sup>§</sup>	2000 <sup>§</sup>	Codeine, Morphine, 6-MAM	2-3 days opiates 3-5 minutes heroin 12-24 hours 6-MAM
		Oxycodone	N/A	N/A		2-4
		Amphetamines	1000	500	Amphetamine, Methamphetamine MDMA, MDA, MDEA	1-3
	Methamphetamine	Incomplete data	500		3-4	
	Benzodiazepines	300	200		3 short-acting 30 long-acting	
	Barbiturates	300	200		1 short-acting 21 long-acting	
	Methadone	300	200	EDDP	3-6	
	Alcohol	N/A	N/A	EtG, EtS	12 hours	

THCA = delta-9-tetrahydrocannabinol-9-carboxylic acid; BEG = benzyolycgonine; 6-MAM = 6-monoacetylmorpine; EDDP = 2-ethylidene-1,5-dimethyl-3,3-diphenylpyrrolidine; EtG = ethyl glucuronide; EtS = ethyl sulfate; \* Detection time for most drugs in urine is 1-3 days; <sup>+</sup> Long-term use of lipid-soluble drugs (THC, diazepam, ketamine) can be detected for a longer period of time;

§ Testing levels for opiates were raised from 300 ng/mL to 2000 ng/mL to reduce detection from foods containing poppy seeds

Agent	Summary of Agents Potentially Contributing to False Positives <sup>3-8</sup>				
Marijuana metabolites	<ul style="list-style-type: none"> <li>dronabinol</li> <li>efavirenz</li> </ul>	<ul style="list-style-type: none"> <li>NSAIDs*</li> <li>proton pump inhibitors</li> </ul>	<ul style="list-style-type: none"> <li>hemp foods: tea, oil<sup>+</sup></li> </ul>		
Cocaine metabolites	<ul style="list-style-type: none"> <li>coca leaf teas</li> </ul>	<ul style="list-style-type: none"> <li>topical anesthetics containing cocaine</li> </ul>			
Opioid metabolites	<ul style="list-style-type: none"> <li>dextromethorphan</li> <li>flouroquinolones</li> </ul>	<ul style="list-style-type: none"> <li>levofloxacin</li> <li>ofloxacin</li> </ul>	<ul style="list-style-type: none"> <li>poppy seeds</li> <li>poppy oil</li> </ul>	<ul style="list-style-type: none"> <li>rifampin</li> <li>quinine</li> </ul>	
Amphetamines/ Methamphetamine (high rate of false positives)	<ul style="list-style-type: none"> <li>amantadine</li> <li>benzphetamine</li> <li>brompheniramine</li> <li>bupropion</li> <li>chlorpromazine</li> <li>desipramine</li> </ul>	<ul style="list-style-type: none"> <li>dextroamphetamine</li> <li>doxepin</li> <li>ephedrine</li> <li>fluoxetine</li> <li>isometheptene</li> <li>isoxsuprine</li> </ul>	<ul style="list-style-type: none"> <li>labetalol</li> <li>l-methamphetamine (OTC nasal inhaler)</li> <li>methylphenidate</li> <li>MDMA</li> <li>phentermine</li> </ul>	<ul style="list-style-type: none"> <li>phenylephrine</li> <li>phenyl-propanolamine</li> <li>promethazine</li> <li>pseudoephedrine</li> </ul>	<ul style="list-style-type: none"> <li>ranitidine</li> <li>selegiline</li> <li>thioridazine</li> <li>trazodone</li> <li>trimethobenzamide</li> <li>trimipramine</li> </ul>
Benzodiazepines	<ul style="list-style-type: none"> <li>oxaprozin</li> </ul>	<ul style="list-style-type: none"> <li>sertraline</li> </ul>			
Barbiturates	<ul style="list-style-type: none"> <li>ibuprofen</li> </ul>	<ul style="list-style-type: none"> <li>naproxen</li> </ul>			
Methadone	<ul style="list-style-type: none"> <li>chlorpromazine</li> <li>clomipramine</li> <li>diphenhydramine</li> </ul>	<ul style="list-style-type: none"> <li>doxylamine</li> <li>ibuprofen</li> <li>quetiapine</li> </ul>	<ul style="list-style-type: none"> <li>thioridazine</li> <li>verapamil</li> </ul>		
Alcohol	<ul style="list-style-type: none"> <li>mouthwash</li> </ul>	<ul style="list-style-type: none"> <li>short-chain alcohols</li> </ul>	<ul style="list-style-type: none"> <li>OTC cough products (isopropyl alcohol)</li> </ul>		

\* NSAIDs resulting in false-positive for marijuana mainly consist of ibuprofen and naproxen and modern tests **do not** result in false positives ; <sup>+</sup> THC concentrations in hemp products are low enough to prevent positive immunoassay results

## Interpreting Urine Drug Testing<sup>2,3-5</sup>

Drug or Class	Expected Results	Considerations
Alcohol	Alcohol	<ul style="list-style-type: none"> <li>• Testing for ETOH metabolites, ethyl glucuronide or ethyl sulfate, can identify alcohol up to 80 hours after consumption</li> </ul>
Amphetamines	<b>Immunoassay</b> –amphetamines, methamphetamines or MDMA <b>Confirmatory</b> –amphetamines, methamphetamines or MDMA	<ul style="list-style-type: none"> <li>• Immunoassay tests are highly cross-reactive; therefore confirmatory testing is required and can identify which amphetamine is present</li> </ul>
Benzodiazepines	<b>Immunoassay</b> –unconjugated oxazepam or its metabolites <b>Confirmatory</b> –alprazolam, diazepam, clonazepam, lorazepam, etc.	<ul style="list-style-type: none"> <li>• Immunoassays for benzodiazepines have a 28% overall false negative rate</li> <li>• Confirmatory testing is needed when use is expected or suspected (alprazolam, clonazepam and lorazepam often not detected by immunoassay)</li> </ul>
Barbiturates	<b>Immunoassay</b> –barbiturates	<ul style="list-style-type: none"> <li>• N/A</li> </ul>
Cocaine metabolites	<b>Immunoassay</b> –cocaine or benzoylecgonine (BEG)	<ul style="list-style-type: none"> <li>• Cocaine’s primary metabolite, BEG, has low cross-reactivity with other substances and is highly predictive of cocaine use</li> <li>• A positive result should be interpreted as recent exposure to cocaine</li> </ul>

Interpreting Urine Drug Testing <sup>2,3-5</sup>		
Drug or Class	Expected Results	Considerations
<b>Opioids or "opiates"- Natural (from opium)</b>		
Codeine (Tylenol # 2,3,4)	<b>Opiates Immunoassay</b> –positive <b>Confirmatory</b> –codeine, possibly morphine & hydrocodone	<ul style="list-style-type: none"> <li>Immunoassays for "opiates" are responsive to morphine and codeine but do not distinguish which</li> <li>Codeine is metabolized to morphine and small quantities of hydrocodone</li> </ul>
Morphine (Avinza, Embeda, MS Contin, Kadian)	<b>Opiates Immunoassay</b> –positive <b>Confirmatory</b> –morphine, possibly hydromorphone	<ul style="list-style-type: none"> <li>Immunoassays for "opiates" are responsive to morphine and codeine but do not distinguish which</li> <li>Morphine (&lt;10%) may be metabolized to hydromorphone</li> </ul>
Heroin	<b>Opiates Immunoassay</b> –positive <b>Confirmatory</b> –heroin (6-MAM), morphine, possibly codeine	<ul style="list-style-type: none"> <li>6-MAM is pathognomic for heroin use, detection 12–24 hrs</li> <li>Heroin is metabolized to morphine</li> </ul>
<b>Opioid Metabolic Pathways</b> <pre> graph LR     Codeine --&gt; Morphine     Morphine --&gt; 6MAM[6-MAM]     Heroin --&gt; 6MAM     6MAM --&gt; Morphine     Codeine --&gt; Hydrocodone     Hydrocodone --&gt; Hydromorphone     Oxycodone --&gt; Oxymorphone     Morphine --&gt; Hydromorphone     </pre> <p>The diagram illustrates the metabolic pathways of opioids. It shows Codeine, Morphine, 6-MAM, Heroin, Hydrocodone, Hydromorphone, Oxycodone, and Oxymorphone in red and green rounded rectangles. Arrows indicate the direction of conversion: Codeine to Morphine, Morphine to 6-MAM, Heroin to 6-MAM, 6-MAM to Morphine, Codeine to Hydrocodone, Hydrocodone to Hydromorphone, Oxycodone to Oxymorphone, and Morphine to Hydromorphone. Downward arrows from Morphine and 6-MAM indicate conversion to Hydromorphone with percentages &lt;15% and &lt;10% respectively.</p>		

continued

Interpreting Urine Drug Testing <sup>2,3-5</sup>		
Drug or Class	Expected Results	Considerations
<b>Opioids-Semisynthetic (derived from opium)</b>		
Hydrocodone (Lorcet, Lortab, Norco, Vicodin)	<b>Opiates Immunoassay</b> –positive <b>Confirmatory</b> –hydrocodone, possibly hydromorphone	<ul style="list-style-type: none"> <li>• “Opiates” immunoassay may detect semisynthetic opioids               <ul style="list-style-type: none"> <li>◦ hydrocodone &gt; hydromorphone &gt; oxycodone</li> </ul> </li> <li>• Negative result does not exclude use and confirmatory testing (GC/MS) is required</li> <li>• Hydrocodone is metabolized in small amounts to hydromorphone, both may be found in urine</li> <li>• Oxycodone is metabolized to oxymorphone, both may be found in urine</li> <li>• Hydromorphone and oxymorphone use does not result in positive screens for hydrocodone and oxycodone, respectively</li> </ul>
Hydromorphone (Dilaudid, Exalgo)	<b>Opiates Immunoassay</b> –may be positive <b>Confirmatory</b> –hydromorphone	
Oxycodone (Roxicet, OxyCotonin)	<b>Opiates Immunoassay</b> –may be positive <b>Oxycodone Immunoassay</b> –positive <b>Confirmatory</b> –oxycodone possibly oxymorphone	
Oxymorphone (Opana)	<b>Oxycodone Immunoassay</b> –positive <b>Confirmatory</b> –oxymorphone	
<b>Opioids-Synthetic (man-made)</b>		
Fentanyl	<b>GC/MS</b> –fentanyl and norfentanyl	<ul style="list-style-type: none"> <li>• Current “opiates” immunoassays do not detect synthetic opioids</li> <li>• Confirmatory testing (GC/MS) is needed</li> </ul>
Meperidine (Demerol)	<b>GC/MS</b> –normeperidine, possibly meperidine	
Methadone (Methadose)	<b>Methadone Immunoassay</b> –positive <b>GC/MS</b> –methadone, EDDP	
Propoxyphene (Darvon, Darvocet)	<b>Propoxyphene Immunoassay</b> –positive <b>GC/MS</b> –propoxyphene & norpropoxyphene	

Confirmatory testing: Chromatography (gas chromatography-mass spectrometry (GC/MS) or liquid chromatography-mass spectrometry (LC/MS)) Note: Each facility may have its own order sets and lab policies and procedures. Contact your lab for additional details.