#### A Lawyer's Guide to Drug Testing: What the Various Tests Do & Don't Tell

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ACDLA 10/9/2021

#### Tests Tell Us Nothing Test Data Do Not Speak For Themselves We Draw Conclusions From Test Data

23°

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# PV=nRT

n = 6.02214076×10<sup>23</sup> mol<sup>-1</sup> R = 8.21×10<sup>-2</sup> L atm/mol K

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- Purpose
- Venue
- Execution

- Purpose
  - Health
  - Performance
  - Ingestion/Exposure
- Venue
- Execution

- Purpose
- Venue
  - Personal
  - Diagnostic
  - Administrative
  - Evidential/Forensic
- Execution

- Purpose
- Venue
- Execution
  - Analysis
  - Analytes
    - Substances detected with analysis
  - Specimens
  - Interpretation

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- Presumptive Testing
  - Bulk specimen analysis
  - Hand-held, spot tests
  - "Strip" immunoassays
  - Typically limited to urine testing
    - Blood
      - Evidential/Forensic





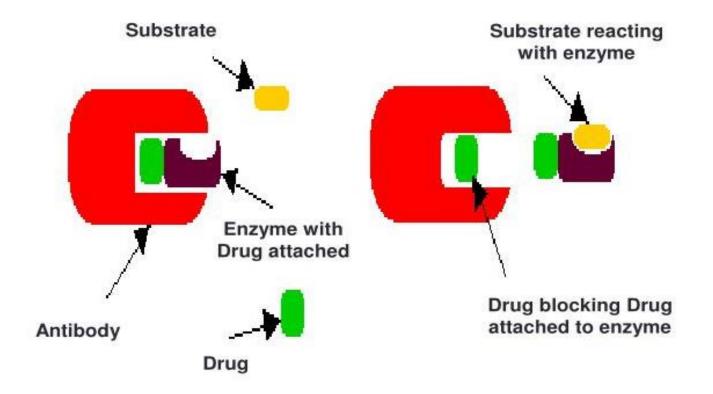
- Presumptive Testing
  - Instrumental immunoassays
    - Large-scale testing





- Presumptive Testing
  - Immunoassay
    - EMIT, EIA, ELISA
    - Antibody directed to a target analyte structure
    - Antibody fixed to a solid sorbent
    - Substances with structures similar to the target analyte bind with the antibody-sorbent
      - Substances other than the target analyte may bind
    - Unbound materials and waste are washed away

- Presumptive Testing
  - Immunoassay example (homogeneous)



- Presumptive Testing
  - Immunoassay
    - Sum total of sorbent-bound substances are "quantified" relative to a single target analyte concentration
      - Substances other than the target analyte may bind
    - If response meets assigned cutoff POS
    - If response fails to meet assigned cutoff NEG

- Presumptive Testing
  - Immunoassay
    - Pros
      - Fast
      - Cheap
      - No expertise required (non-instrumental methods)
        - » Easy to interpret NO/YES
    - Cons
      - Class-specific only
      - Substances other than the target analyte may bind
        - » Antigen-antibody recognition is not definitive
      - Interferents

#### THC - Substances or Conditions which can cause false positives

Dronabinol (Marinol) Ibuprofen (Advil, Nuprin, Motrin, Excedrin IB etc) Ketoprofen (Orudis KT) Kidney infection (Kidney disease, diabetes) Liver Disease Naproxen (Aleve) Promethazine (Phenergan, Promethegan) Riboflavin (B2, Hempseed Oil)

#### Amphetamines - Substances or Conditions which can cause false positives

Ephedrine, pseudoephedrine, propylephedrine, phenylephrine, or desoxyephedrine (Nyquil, Contact, Sudafed, Allerest, Tavist-D, Dimetapp, etc) Phenegan-D, Robitussin Cold and Flu, Vicks Nyquil Over-the-counter diet aids with phenylpropanolamine (Dexatrim, Accutrim) Over-the-counter nasal sprays (Vicks inhaler, Afrin) Asthma medications (Marax, Bronkaid tablets, Primatine Tablets) Prescription medications (Amfepramone, Cathne, Etafediabe, Morazone, phendimetrazine, phenmetrazine, benzphetamine, fenfluramine, dexfenfluramine, dexdenfluramine, Redux, mephentermine, Mesocarb, methoxyphenamine, phentermine, amineptine, Pholedrine, hydroymethamphetamine, Dexedrine, amifepramone, clobenzorex, fenproyorex, mefenorex, fenelylline, Didrex, dextroamphetamine, methphenidate, Ritalin, pemoline, Cylert, selegiline, Deprenyl, Eldepryl, Famprofazone) Kidney infection, kidney disease, Liver disease, diabetes

#### **Opiates - Substances or Conditions which can cause false positives**

Poppy Seeds Tylenol with codeine Most prescription pain medications Cough suppressants with Dextromethorphan (DXM) Nyquil Kidney infection, Kidney Disease Diabetes, Liver Disease

#### **Cocaine - Substances or Conditions which can cause false positives**

Kidney infection (kidney disease) Liver infection (liver disease) Diabetes Amoxicillin, tonic water

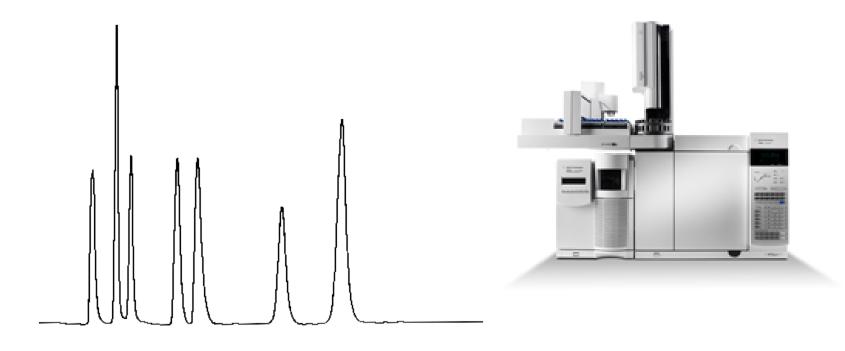
- Presumptive Testing
  - Immunoassay
    - Response may be due to <u>multiple</u> bound substances, not a single substance
      - Could include the target analyte, could include interferents
    - <u>Quantification is solely intended to meet a cutoff</u> <u>concentration for positive/negative, not to quantify a</u> <u>specific target</u>

- Definitive Testing
  - Analyte resolution
    - Sample preparation
      - LLE, SPE
    - Chromatography
      - GC, LC
  - Analyte identification
    - Molecular identification
      - MS, MS/MS
    - GC/MS, LC/MS/MS

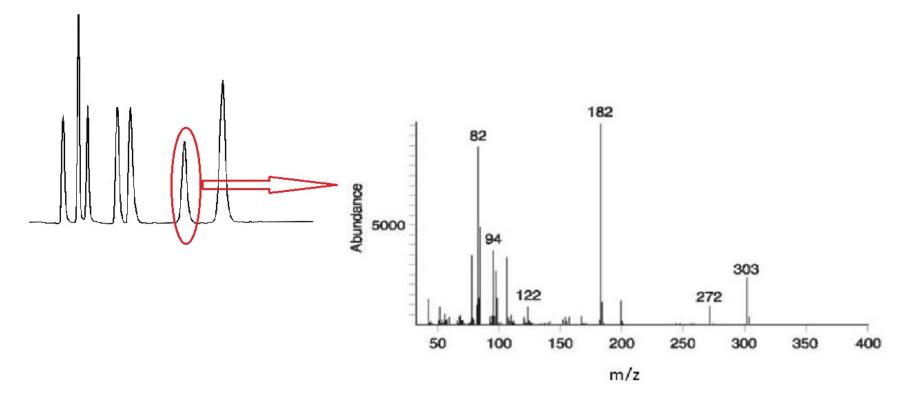
- Analyte Resolution
  - Sample preparation
    - LLE, SPE
    - Specimens extracted
      - Organic solubility
      - Ionic transformation (pH)
    - Unwanted materials discarded



- Analyte Resolution
  - Chromatography
    - GC/MS, LC/MS
    - Separation of substance mixtures



- Analyte Identification
  - Chromatography/Mass spectrometry
    - Resolution then fragmentation then detection



- Analyte Identification
  - Mass spectrometry



- Definitive Testing
  - Pros
    - Definitive
      - Identify individual substances
    - Can be calibrated
      - Quantification
    - Controllable

- Definitive Testing
  - Cons
    - Not fast
    - Expensive \$\$\$\$
    - Labor-intensive
    - Laboratory facilities required
    - Expertise required

- Purpose
  - Health
  - Performance
  - Ingestion/Exposure
- Venue
- Execution

#### Purpose

- Health
  - Explanation of symptoms
    - Cause of death
    - Legal considerations
  - Guide to treatment
    - Compliance with treatment program
    - Avoid adverse drug interactions
    - Successful patient outcome renders test results moot
      - False negative invokes alternate efforts
      - False positive invokes alternate efforts

#### Purpose

- Performance
  - Explanation of actions
  - Results used in adjudication
    - False negative is harmless to the defendant
      - Homicide victim?
    - False positive is devastating to the defendant

#### Purpose

- Ingestion/Exposure
  - Results used in adjudication
    - Solely to indicate exposure
      - Compliance
    - False negative is harmless to the defendant
    - False positive is devastating to the defendant

- Purpose
- Venue
  - Personal
  - Diagnostic
  - Administrative
  - Evidential/Forensic
- Execution

- Personal
  - Immunoassays

**ALWAYS TEST CLEAN** 

**URINE - BLOOD - HAIR** 

**GUARANTEED** 

I just have to know





A drug test is commonly a technical examination of urine, blood, semen, sweat, or oral fluid samples to determine the presence or absence of specified drugs or their metabolized traces

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- Diagnostic
  - Immunoassays
    - Guide to treatment
    - Class-specific test is sufficient
    - Avoid adverse drug interactions
    - Successful patient outcome renders test results moot
      - False negative invokes alternate efforts
      - False positive invokes alternate efforts

- Diagnostic
  - Clinical laboratories
    - CLIA 88
    - Urine



- Drugs of abuse (DAU) by immunoassay kit
- Serum
  - Derivative of blood
  - Ethanol
  - Therapeutic drug monitoring (TDM) by clinical methods
    - » Antibiotics, anticonvulsants, anticoagulants



- Diagnostic
  - Urine drug testing
    - Immunoassays class-specific only
      - No confirmation of positive results
      - No definitive analyte identification
    - Sufficient to guide treatment
  - Serum drug testing (derived from clotted blood)
    - Clinical method
      - No confirmation of positive results
      - No molecular identification
    - Sufficient to guide treatment

- Diagnostic
  - Serum/plasma ethanol testing
    - Serum derived from clotted blood
    - Plasma derived from unclotted blood
    - Enzyme specificity is demonstrated
      - Serum/plasma-to-blood ratio =  $1.18 \pm 0.04$
    - Sufficient to guide treatment
    - Results may be reliable
      - Sufficient QA
    - Often used in adjudication
      - Must be rendered evidential

- Administrative
  - Non-regulated testing
    - Presumptive
    - Immunoassays
      - Hand-held devices, spot tests
      - No confirmation of positive results
      - Not evidential, sufficient for probable cause
  - Regulated testing
    - Presumptive + confirmation of positives
      - Immunoassay + GC/MS
    - Evidential

- Regulated Testing
  - Federal Drug-Free Workplace Program
    - Executive Order 12564, Sept. 15, 1986
      - Each Executive agency shall establish a program to test for the use of illegal drugs by Federal employees in sensitive positions and the Secretary shall promulgate scientific and technical guidelines for drug testing programs
    - Public Law 100-71 (July 11, 1987)
    - Intended to incorporate presumptive and confirmation testing otherwise recognized for legally acceptable toxicological testing
      - Frye, Fed. Rule 702

- Regulated Testing
  - Health and Human Services (HHS)
    - Substance Abuse and Mental Health Services Administration (SAMHSA)
      - Center for Substance Abuse Prevention
        - » Division of Workplace Programs (DWP)
          - Oversight of the program
          - Oversight of the National Laboratory Certification Program (NLCP)
        - » DOT, 19 CFR Part 40

https://www.transportation.gov/odapc/part40

- Regulated Testing
  - Urine presumptive testing
  - Hair
    - Alternate specimens under investigation
  - Confirmation of positive results
    - Definitive molecular identification, quantification
  - Evidential

- Regulated Testing
  - Conceived to deter illicit drug use amongst safetysensitive federal employees
  - Intended <u>solely</u> to identify intentional illicit drug exposure
  - Results are NEG/POS
    - Medical Review Officer (MRO)
    - <u>Positive</u> laboratory finding is reported as <u>negative</u> if the subject has legitimate access

- Regulated Testing
  - Limited scope popular in 1980's California
    - Marijuana metabolites
      - Δ<sup>9</sup>-THC-COOH, <u>NOT</u> Δ<sup>9</sup>-THC
    - Cocaine metabolites
      - Benzoylecgonine, <u>NOT</u> cocaine
    - Opiates
      - Morphine, 6-MAM, codeine, hydrocodone, oxycodone
    - Phencyclidine
    - Amphetamines
      - Methamphetamine, amphetamine, MDMA, MDA

#### Analyte urine cutoff concentrations (ng/mL)

Δ<sup>9</sup>-THC-9-COOH benzoylecgonine morphine/codeine 6-monoacetylmorohine (6-MAM) hydrocodone/hydromorphone oxycodone/oxymorphone phencyclidine meth/amphetamine MDMA/MDA

50/15 300/150 2000/2000 10/10300/100 100/100 25/25 250/250 500/250

- Administrative
  - Court-ordered compliance
    - Non-regulated
      - Non-evidential
    - Regulated
      - Evidential

- Evidential/Forensic
  - Criminal investigations
    - Explanation of actions
      - Suspect
      - Victim
  - Death investigations
    - Cause
    - Manner

- Evidential/Forensic
  - [fo-ren' sik] *adj* belonging to courts of law
    - L *forensis*-forum, market place
  - Belonging to, used in, or suitable to courts of judicature, or to public discussion and debate

Merriam Webster's Dictionary

- Evidential/Forensic
  - Legal requirements
    - Relevance
    - Collection and identification
    - Preservation of evidential value
    - Competent examination
      - Analysis
      - Interpretation

- Evidential/Forensic
  - Scientific requirements
    - Quantity
    - Maintenance
    - Analysis
      - Technology/methodology
      - Reliability/precision/accuracy
        - » Standardization

- Evidential/Forensic
  - Standardization
    - Calibration
      - Standardization of process
      - Establish parameters for quantitative evaluation
    - Control
      - Verification of accuracy of process
      - Positive
      - Negative
      - Alternate substances

- Evidential/Forensic
  - Initial presumptive testing followed by confirmation of positive results
    - Definitive molecular identification
    - Quantification ±
  - Specimens limited only by method validation
  - Analytes limited only by method validation
  - Methodology limited only by validation
  - Quantification limited only by method validation

- Evidential/Forensic
  - Laboratory results not intended to be used without interpretation
  - Interpretation requires knowledge of pharmacology and toxicology

- Evidential/Forensic
  - Admissibility rather than regulation
  - Pre-dates regulated testing standards
    - Regulated testing standards established to facilitate introduction into to legal proceedings
    - Frye v. US, 1923
      - General acceptance standard
    - Federal Rules of Evidence 702, 1973
    - Daubert v. Merrell Dow, 1993
      - Reliability AND Relevance standards

- Evidential/Forensic
  - Analyst certification
    - Forensic Specialties Accreditation Board
      - www.thefsab.org
  - Laboratory accreditation
    - ANAB (ANSI National Certification Board)
      - <u>https://anab.ansi.org/laboratory-accreditation</u>
  - Method standardization
    - AAFS/ASB
      - <a href="https://www.asbstandardsboard.org/">https://www.asbstandardsboard.org/</a>

- Purpose
- Venue
- Execution
  - Analysis
  - Analytes
    - Substances detected with analysis
  - Specimens
  - Interpretation

- Analysis
  - Presumptive screening
    - Immunoassay
    - Advanced MS/TOF methods
  - Confirmation
    - GC/MS, LC/MS/MS

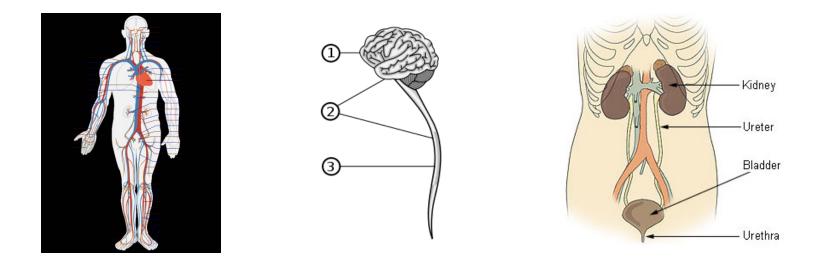


- Analytes
  - Selection based upon purpose and venue
  - Drugs of abuse (DAU), misuse, non-use
  - Parent drugs
    - Pharmacodynamics
  - Metabolites
    - Pharmacodynamics

- Specimens
  - Selection based on purpose and venue

Blood	Gastric contents
Serum	Hair
Plasma	Nails
Urine	Breath
Vitreous humor	Oral fluid
Bile	Sweat
Tissues	"Other"

- Specimens
  - Relevance
    - Does the specimen reflect a target organ?
    - Does it matter?



- Specimens
  - Relevance
    - CNS, cardiovascular, pulmonary are is the target organs for toxicity-based pharmacodynamic effects
    - CNS is the target organ for performance-based
      pharmacodynamic effects
    - Blood is the most documented specimen to reflect CNS

- Specimens
  - Blood
    - Circulates throughout the body
    - Reflects presence of analytes in organs based upon pharmacokinetic parameters (volume of distribution V<sub>d</sub>)
    - Drugs in blood vs. performance
    - Presence of active analytes in blood may be used to establish impairment
  - Serum/Plasma
    - Derived from blood
      - Largely limited to ethanol

- Specimens
  - Urine
    - Excreted from the circulating body water
    - Reflects time-averaged presence of analytes in blood filtered in the kidneys
    - Analytes are detectable for longer periods in urine than in blood
    - Less invasive to collect than blood

- Specimens
  - Urine
    - Presence of analytes in urine may be used to establish consumption of the analyte or a substance giving rise to the analyte
    - Presence of analytes in urine do NOT necessarily reflect a concurrent presence in blood or organs
    - Presence of analytes in urine CANNOT be used to establish impairment
    - Concentrations are essentially uninterpretable due to variable solute composition

- Specimens
  - Hair
    - Limited to chronic exposure
      - May provide timeline based upon position of drugs in hair (proximal vs. distal)
      - Growth ~1 cm/month (head)
  - Nails
    - Similar to hair
    - Slower growth



- Specimens
  - Oral fluid
    - Limited to drugs of abuse
    - May reflect blood
      - Under investigation
  - Sweat
    - Limited to drugs of abuse
    - May reflect blood
      - Under investigation

- Specimens
  - Vitreous humor
    - Limited to ethanol
    - May substitute for putrefied blood
  - Breath
    - Limited to ethanol

- Purpose
  - Health assessment
    - Presumptive test result is sufficient
    - Evidential test result is necessary for COD/MOD
  - Performance assessment
    - Presumptive test result is PC
    - Evidential test result is necessary
  - Ingestion/exposure assessment
    - Presumptive test result is PC
    - Evidential test result is necessary

- Venue
  - Personal
    - Presumptive test result is sufficient
  - Diagnostic
    - Presumptive test result is sufficient
  - Administrative
    - Presumptive test should NOT be used as evidential
    - Evidential test should be necessary
  - Evidential/Forensic
    - Name says it all

- Execution
  - Analysis
    - Presumptive + Confirmation = Evidence
  - Analytes
    - Class-specific for diagnostic use only
    - Individual substances necessary for evidential use
  - Specimens
    - Blood exposure, performance
    - All others exposure only (maybe)

Specimen	Exposure	Effect
Urine	Yes	No
Hair	Yes	No
Oral fluid	Yes	TBD
Sweat	Yes	TBD
Blood	Yes	Yes
Vitreous Humor	Yes	Maybe
		(ethanol)
Tissues	Yes	Maybe

- Significant Considerations
  - Pharmacological activity
    - Metabolites may/may not retain activity of parent
  - Concentration
    - Blood, vitreous humor (ethanol)
    - Generalities
      - Tolerance
  - Time relative to incident
    - Residence
      - Absorption, Distribution, Metabolism, Elimination
      - Retrograde extrapolation

- Drug test findings are generally an unreliable sole basis for establishing specific degrees of individual impairment.
- Drug test findings are most useful for explaining observed signs/symptoms.
  - Performance
  - Cause of death
- *Per Se* limits represent cutoff for likelihood of misadventure, not degree of impairment.

Testing	Reliable?	Active substance?	Relevant specimen?
Diagnostic			
Drugs	No	No	Urine
Ethanol	Yes*	Yes	Serum
Administrative			
Non-regulated	No	No	Urine
Regulated	Yes	Yes/No	Urine+
Personal	No	No	Urine
Evidential	Yes	Yes	Blood+

Testing	Health?	Performance?	Exposure?
Diagnostic			
Drugs	Yes	No	No
Ethanol	Yes	Yes*	Yes
Administrative			
Non-regulated	Yes	No	No
Regulated	Yes	No	Yes
Personal	caveat emptor	No	No
Evidential	Yes	Yes*	Yes