MAYO CLINIC LABORATORIES

Test Definition: 6MAMM

6-Monoacetylmorphine (6-MAM), Confirmation, Meconium

Overview

Useful For
Detection of in utero heroin exposure up to 5 months before birth

Special Instructions
- Clinical Toxicology CPT Code Client Guidance

Method Name
Liquid Chromatography Tandem Mass Spectrometry (LC-MS/MS)

NY State Available
Yes

Specimen

Specimen Type
Meconium

Ordering Guidance
For chain-of-custody testing, order MAMMX / 6-Monoacetylmorphine (6-MAM) Confirmation, Chain of Custody, Meconium.

Specimen Required
Supplies: Stool container, Small (Random), 4 oz (T288)
Container/Tube: Stool container
Specimen Volume: 1 g (approximately 1 teaspoon)
Collection Instructions:
1. Collect entire random meconium (newborn's first bowel movements) specimen.
2. Send specimen frozen.
Additional Information: When refrigerated, a significant percentage of 6-monoacetylmorphine (MAM) will convert to morphine in less than 24 hours.

Forms
If not ordering electronically, complete, print, and send a Therapeutics Test Request (T831) with the specimen.

Specimen Minimum Volume
0.3 g (approximately 1/4 teaspoon)

Reject Due To

Grossly bloody	Reject; Pink OK
Stool; diapers	Reject

Specimen Stability Information

Specimen Type	Temperature	Time	Special Container
Meconium	Frozen	14 days	

Clinical & Interpretive

Clinical Information

Heroin (diacetylmorphine) is a semisynthetic opiate that is closely related to morphine. It is no longer used clinically in the United States, though it is used elsewhere for rapid relief of pain.(1) Like morphine and other opiates, its relaxing and euphoric qualities make heroin a popular drug of abuse. Heroin is commonly injected intravenously, although it can be administered by other means such as snorting, smoking, or inhaling vapors.

Heroin shares the core structure of morphine with the addition of 2 acetyl groups, which are thought to enhance its permeation into the central nervous system.(2,3) Heroin is metabolized by sequential removal of these acetyl groups; loss of the first acetyl converts heroin into 6-monoacetylmorphine (6-MAM).(2,3) Heroin is rarely found in meconium since only 0.1% of a dose is excreted unchanged. 6-MAM is a unique metabolite of heroin, and its presence is a definitive indication of heroin use. Like heroin, 6-MAM has a very short half-life; however, its detection time in meconium, the first fecal material passed by the neonate, is uncharacterized. 6-MAM is further metabolized into morphine, the dominant metabolite of heroin, and morphine will typically be found in a specimen containing 6-MAM.

Opiates, including heroin, have been shown to readily cross the placenta and distribute widely into many fetal tissues.(4) Opiate use by the mother during pregnancy increases the risk of prematurity and being small for gestational age. Furthermore, infants who have been exposed to heroin exhibit an early onset of withdrawal symptoms compared with infants who have been exposed to methadone. Infants exposed to heroin demonstrate a variety of symptoms including irritability, hypertonia, wakefulness, diarrhea, yawning, sneezing, increased hiccups, excessive sucking, and seizures. Long-term intrauterine drug exposure may lead to abnormal neurocognitive and behavioral development as well as an increased risk of sudden infant death syndrome.(5)

The disposition of drug in meconium is not well understood. The proposed mechanism is that the fetus excretes the drug into bile and amniotic fluid. The drug accumulates in meconium either by direct deposit from bile or through swallowing of amniotic fluid.(6) The first evidence of meconium in the fetal intestine appears at approximately the 10th to 12th week of gestation, and it slowly moves into the colon by the 16th week of gestation.(7) Therefore, the presence of drugs in meconium has been proposed to be indicative of in utero drug exposure during the final 4 to 5 months of pregnancy, a longer historical measure than is possible by urinalysis.(6)

Reference Values

Negative
Positive results are reported with a quantitative liquid chromatography tandem mass spectrometry result.
Cutoff concentration: 5 ng/g

Interpretation

The presence of 6-monoacetylmorphine (6-MAM) in meconium is definitive for heroin use by the mother. However, the absence of 6-MAM does not rule-out heroin use because of its short half-life and stability.

Cautions

The short half-life and stability of 6-monoacetylmorphine (6-MAM) may prevent its detection in heroin users.

6-MAM is metabolized to morphine, but the presence of morphine alone is not sufficient evidence to prove heroin use. 6-MAM is the only definitive metabolite of heroin.

Clinical Reference

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2. Levine B, ed. Principles of Forensic Toxicology. 4th ed. AACC Press; 2013
3. Brunton LL, Hilal-Dandan R, Knollmann BC, eds. Goodman and Gilman's: The Pharmacological Basis of Therapeutics. 13th ed. McGraw-Hill; 2018
4. Szeto HH. Kinetics of drug transfer to the fetus. Clin Obstet Gynecol. 1993;36:246-254
5. Kwong TC, Ryan RM. Detection of intrauterine illicit drug exposure by newborn drug testing. Clin Chem. 1997;43(1):235-242
6. Ostrea EM Jr, Brady MJ, Parks PM, et al. Drug screening of meconium in infants of drug-dependent mothers: an alternative to urine testing. J Pediatr. 1989;115(3):474-477
7. Ahanya SN, Lakshmanan J, Morgan BL, Ross MG. Meconium passage in utero mechanisms, consequences, and management. Obstet Gynecol Surv. 2005;60(1):45-56; quiz 73-74
8. Langman LJ, Bechtel LK, Holstege CP. Clinical toxicology. In: Rifai N, Chiu RWK, Young I, Burnham CAD, Wittwer CT, eds. Tietz Textbook of Laboratory Medicine. 7th ed. Elsevier; 2023:chap 43
9. Baselt RC. Disposition of Toxic Drugs and Chemical in Man. 12th ed. Biomedical Publications; 2020

Performance

Method Description

Meconium is mixed with internal standard and extracted with methanol. The methanolic extract is further processed by solid-phase extraction. The extract is analyzed by liquid chromatography tandem mass spectroscopy.(Unpublished Mayo method)

PDF Report

No

Day(s) Performed

Monday through Sunday

Report Available

2 days

Specimen Retention Time
2 weeks

Performing Laboratory Location
Mayo Clinic Laboratories - Rochester Superior Drive

Fees & Codes

Fees
- Authorized users can sign in to Test Prices for detailed fee information.
- Clients without access to Test Prices can contact Customer Service 24 hours a day, seven days a week.
- Prospective clients should contact their account representative. For assistance, contact Customer Service.

Test Classification
This test was developed and its performance characteristics determined by Mayo Clinic in a manner consistent with CLIA requirements. It has not been cleared or approved by the US Food and Drug Administration.

CPT Code Information
G0480
80356 (if appropriate for select payers)
Clinical Toxicology CPT Code Client Guidance

LOINC® Information

Test ID	Test Order Name	Order LOINC® Value
6MAMM	6-MAM Confirmation, M	29345-6

Result ID	Test Result Name	Result LOINC® Value
31874	6-Monoacetylmorphine	29345-6
31875	Interpretation	69050-3