EDDP Rapid Test Cassette (Whole Blood/Serum/Plasma) Package Insert

A rapid test for the qualitative detection of EDDP in human whole blood or serum or plasma.
For medical and other professional in vitro diagnostic use only.

【INTENDED USE】

The EDDP Rapid Test Cassette (whole blood/serum/plasma) is a lateral flow chromatographic immunoassay for the detection of EDDP in whole blood or serum or plasma at a cut-off concentration of 50ng/ml. This test will detect other related compounds, please refer to the analytical Specificity table in this package insert.

This assay provides only a qualitative, preliminary analytical test result. A more specific alternate chemical method must be used in order to obtain a confirmed analytical result. Gas chromatography/mass spectrometry (GC/MS) is the preferred confirmatory method. Clinical consideration and professional judgment should be applied to any drug of abuse test result, particularly when preliminary positive results are used.

【SUMMARY】

Methadone is an unusual drug in that its primary urinary metabolites (EDDP and EMDP) are cyclic in structure, making them very difficult to detect using immunoassays targeted to the native compound.[1]Exacerbating this problem. there is a subsection of the population classified as "extensive metabolizers" of methadone. In these individuals, a whole blood/serum/plasma specimen may not contain enough parent methadone to yield a positive drug screen even if the individual is in compliance with their methadone maintenance. EDDP represents a better whole blood/serum/plasma marker for methadone maintenance than unmetabolized methadone.

【PRINCIPLE】

The EDDP Rapid Test Cassette (whole blood/serum/plasma) is an immunoassay based on the principle of competitive binding. Drugs that may be present in the whole blood/serum/plasma specimen compete against the drug conjugate for binding sites on the antibody.

During testing, a whole blood/serum/plasma specimen migrates upward by capillary action. EDDP, if present in the whole blood/serum/plasma specimen below the cut-off level, will not saturate the binding sites of the antibody in the test. The antibody coated particles will then be captured by immobilized EDDP-protein conjugate and a visible colored line will show up in the test line region. The colored line will not form in the test line region if the EDDP level exceeds the cut-off level because it will saturate all the binding sites of anti-EDDP antibodies.

A drug-positive whole blood/serum/plasma specimen will not generate a colored line in the test line region because of drug competition, while a drug-negative whole blood/serum/plasma specimen or a specimen containing a drug concentration less than the cut-off will generate a line in the test line region. To serve as a procedural control, a colored line will always appear at the control line region indicating that proper volume of specimen has been added and membrane wicking has occurred.

【REAGENTS】

The test contains mouse monoclonal anti-EDDP antibody coupled particles and EDDP-protein conjugate. A goat antibody is employed in the control line system.

【PRECAUTIONS】

- For professional in vitro diagnostic use only. Do not use after the expiration date.
- Do not eat, drink or smoke in the area where the specimens or kits are handled.
- Do not use test if pouch is damaged
- Handle all specimens as if they contain infectious agents. Observe established precautions against microbiological hazards throughout testing and follow the standard procedures for proper disposal of specimens.
- Wear protective clothing such as laboratory coats, disposable gloves and eye protection when specimens are being tested.
- The used test should be discarded according to local regulations.
- Humidity and temperature can adversely affect results.

【STORAGE AND STABILITY】

Store as packaged in the sealed pouch at room temperature or refrigerated (2-30°C). The test is stable through the expiration date printed on the sealed pouch. The test must remain in the sealed pouch until use. **DO NOT FREEZE**. Do not use beyond the expiration date.

【SPECIMEN COLLECTION AND PREPARATION】

- The EDDP Rapid Test Cassette can be performed using whole blood (from venipuncture or fingerstick)/serum/plasma.
- To collect **Fingerstick Whole Blood specimens**:
  - Wash the patient's hand with soap and warm water or clean with an alcohol swab. Allow to dry.
  - Massage the hand without touching the puncture site by rubbing down the hand towards the fingertip of the middle or ring finger.
  - Puncture the skin with a sterile lancet. Wipe away the first sign of blood.
  - Gently rub the hand from wrist to palm to finger to form a rounded drop of blood over the puncture site.
  - Add the Fingerstick Whole Blood specimen to the test by using **a capillary tube**:
    - Touch the end of the capillary tube to the blood until filled to approximately 40 μl. Avoid air bubbles.
    - Place the bulb onto the top end of the capillary tube, then squeeze the bulb to dispense the whole blood to the specimen area of the test cassette.
- Testing should be performed immediately after the specimens have been collected. Do not leave the specimens at room temperature for prolonged periods. For long term storage, specimens should be kept below -20°C. Whole blood/serum/plasma collected by venipuncture should be stored at 2-8°C if the test is to be run within 2 days of collection. Do not freeze whole blood or serum or plasma specimens. Whole blood/serum/plasma collected by fingerstick should be tested immediately.
- Bring specimens to room temperature prior to testing. Frozen specimens must be completely thawed and mixed well prior to testing. Specimens should not be frozen and thawed repeatedly.
- If specimens are to be shipped, they should be packed in compliance with local regulations covering the transportation of etiologic agents.

【MATERIALS】

Materials Provided

• Test cassettes • Droppers • Buffer • Package insert

Materials Required But Not Provided

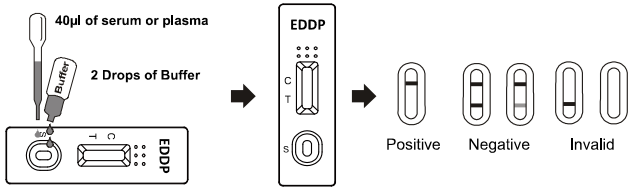
• Specimen collection containers • Centrifuge
• lancets (for fingerstick whole blood only) • Timer
•Heparinized capillary tubes and dispensing bulb (for fingerstick whole blood only)

【DIRECTIONS FOR USE】

Allow the test, specimen, buffer and/or controls to reach room temperature (15-30°C) prior to testing.

For serum or plasma specimen:

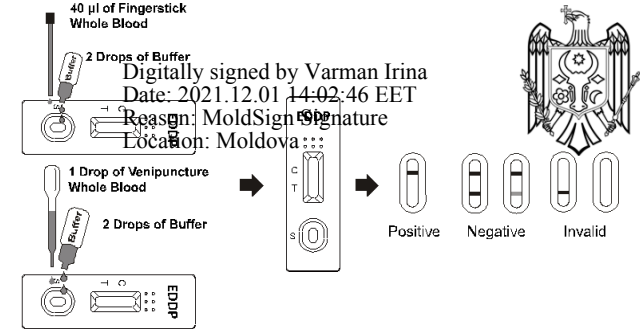
1. Bring the pouch to room temperature (15-30℃) before opening it. Remove the cassette from the sealed pouch and use it within one hour.
2. Place the cassette on a clean and level surface. Hold the dropper vertically and transfer **1 full drop of serum or plasma** (approximately 40ul), then add **2 drops of buffer** (approximately 80 μl) to the specimen well of the cassette, and then start the timer. Avoid trapping air bubbles in the specimen well. See illustration below.
3. Wait for the colored line(s) to appear. **Read the result at 5 minutes**. Do not interpret the result after 10 minutes.



For whole blood specimen:

1. Bring the pouch to room temperature before opening it. Remove the test cassette from the sealed pouch and use it as soon as possible.
2. Place the cassette on a clean and level surface.
   For **Venipuncture Whole Blood** specimen:
   - Hold the dropper vertically and transfer **1 drop of whole blood** (approximately 40μl) to the specimen well, then **add 2 drops of buffer** (approximately 80 μl), and start the timer. See illustration below.

   For **Fingerstick Whole Blood** specimen:
   - To use a capillary tube: Fill the capillary tube and **transfer approximately 40μl of fingerstick whole blood specimen** to the specimen area of test cassette, then add**2 drops of buffer**(approximately 80 μl) and start the timer. See illustration below.
3. Wait for the colored line(s) to appear. **Read results at 5 minutes.** Do not interpret the result after 10 minutes.



Digitally signed by Varman Irina
Date: 2021.12.01 14:02:46 EET
Reason: MoldSign Signature
Location: Moldova

【INTERPRETATION OF RESULTS】

(Please refer to the illustration above)

NEGATIVE:* Two lines appear. One colored line should be in the control line region (C), and another apparent colored line should be in the test line region (T). This negative result indicates that the EDDP concentration is below the detectable cut-off level.

***NOTE:** The shade of color in the test line region (T) may vary, but it should be considered negative whenever there is even a faint colored line.

POSITIVE: One colored line appears in the control line region (C). No line appears in the test line region (T). This positive result indicates that the EDDP concentration exceeds the detectable cut-off level.

INVALID: Control line fails to appear. Insufficient specimen volume or incorrect procedural techniques are the most likely reasons for control line failure. Review the procedure and repeat the test with a new test. If the problem persists, discontinue using the test kit immediately and contact your local distributor.

【QUALITY CONTROL】

A procedural control is included in the test. A colored line appearing in the control region (C) is the internal procedural control. It confirms sufficient specimen volume and correct procedural technique. Control standards are not supplied with this kit; however, it is recommended that positive and negative controls be tested as a good laboratory practice to confirm the test procedure and to verify proper test performance.

【LIMITATIONS】

1. The EDDP Rapid Test Cassette (whole blood/serum/plasma) provides only a qualitative, preliminary analytical result. A secondary analytical method must be used to obtain a confirmed result. Gas chromatography/ mass spectrometry (GC/MS) is the preferred confirmatory method.[2]
2. It is possible that technical or procedural errors, as well as other interfering substances in the Whole blood or serum or plasma specimen may cause erroneous results.
3. A positive result indicates presence of the drug or its metabolites but does not indicate level of intoxication, administration route or concentration in Whole blood or serum or plasma.
4. A negative result may not necessarily indicate drug-free Whole blood/serum/plasma. Negative results can be obtained when drug is present but below the cut-off level of the test.
5. Test does not distinguish between drugs of abuse and certain medications.

【EXPECTED VALUES】

This negative result indicates that the EDDP concentration is below the detectable level of 50ng/ml. Positive result means the concentration of EDDP is above the level of 50ng/ml. The EDDP Rapid Test Cassette has a sensitivity of 50ng/ml

【PERFORMANCE CHARACTERISTICS】

Accuracy

A side-by-side comparison was conducted using The EDDP Rapid Test Cassette and GC/MSat the cut-off of 50ng/ml. Testing was performed on 90 clinical specimens previously collected from subjects present for Drug Screen Testing. The following results were tabulated:

Clinic Result of Whole Blood

Method		GC/MS		Total Results
EDDP Rapid Test Cassette	Results	Positive	Negative	
	Positive	18	2	20
	Negative	2	68	70
Total Results		20	70	90
% Agreement		90%	97.1%	95.6%

Clinic Result of Serum or Plasma

Method		GC/MS		Total Results
EDDP Rapid Test Cassette	Results	Positive	Negative	
	Positive	18	2	20
	Negative	2	68	70
Total Results		20	70	90
% Agreement		90%	97.1%	95.6%

Analytical Sensitivity

A drug-free whole blood/serum/plasma pool was spiked with EDDP at the following concentrations of ±50%cutoff and 3xcutoff. The data are summarized below:

For whole blood:

EDDP Concentration (ng/ml)	Percent of Cut-off	n	Visual Result	
			Negative	Positive
0	0	30	30	0
25	-50%	30	30	0
50	Cut-off	30	15	15
75	+50%	30	0	30
150	3X	30	0	30

For serum or plasma:

EDDP Concentration (ng/ml)	Percent of Cut-off	n	Visual Result	
			Negative	Positive
0	0	30	30	0
25	-50%	30	30	0
50	Cut-off	30	15	15
75	+50%	30	0	30
150	3X	30	0	30

Analytical Specificity

The following table lists compounds that are positively detected in whole blood/serum/plasma by The EDDP Rapid Test Cassette (whole blood/serum/plasma) at 5 minutes.

Compound	Concentration (ng/ml)
2-Ethylidene-1,5-dimethyl-3,3-diphenylpyrrolidine (EDDP)	50

Precision

A study was conducted at three hospitals by untrained operators using three different lots of product to demonstrate the within run, between run and between operator precision. An identical panel of coded specimens, containing no EDDP and 50% EDDP above and below the 50ng/ml cut-off was provided to each site. The following results were tabulated:

EDDP Concentration (ng/ml)	n per Site	Site A		Site B		Site C	
		-	+	-	+	-	+
0	10	10	0	10	0	10	0
25	10	8	2	9	1	9	1
75	10	1	9	1	9	2	8

Cross-Reactivity

A study was conducted to determine the cross-reactivity of the test with compounds in either drug-free whole blood/serum/plasma or determine positive whole blood/serum/plasma. The following compounds show no cross-reactivity when tested with The EDDP Rapid Test Cassette (whole blood/serum/plasma) at a concentration of 100 μg/ml.

Non Cross-Reacting Compounds

4-Acetaminophenol	4-Dimethylaminoantipyrine	Loperamide	Prednisolone
Acetone	Diphenhydramine	Maprotiline	Prednisone
Acetophenetidin	5,5-Diphenylhydantoin	Meperidine	Procaine
N-Acetylprocainamide	Disopyramide	Meprobamate	Promazine
Acetylsalicylic acid	Doxylamine	d-Methamphetamine	Promethazine
Albumin	Ecgonine	l-Methamphetamine	l-Propoxyphene
Amitriptyline	Ecgonine methylester	Methaqualone	d,l-Propranolol
Amobarbital	EMDP	Methadone	d-Pseudoephedrine
Amoxapine	Ephedrine	Methoxyphenamine	Quinacrine
Amoxicillin	l-Ephedrine	(+)-3,4-Methylendioxy-methamphetamine	Quinidine
Ampicillin	l-Epinephrine		Quinine
Ascorbic acid	(±)-Epinephrine	Methylphenidate	Ranitidine
Aminopyrine	Erythromycin	Mephentermine	Riboflavin
Apomorphine	β-Estradiol	Metoprolol	Salicylic acid
Aspartame	Estrone-3-sulfate	Morphine-3-β-D-glucuronide	Secobarbital
Atropine	Ethanol (Ethyl alcohol)	Morphine sulfate	Serotonin
Benzilic acid	Ethyl-p-aminobenzoate	Methyprylon	(5-Hydroxytryptamine)
Benzoic acid	Etodolac	Nalidixic acid	Sodium chloride
Benzphetamine	Famprofazone	Nalorphine	Sulfamethazine
Bilirubin	Fenfluramine	Naloxone	Sulindac
Brompheniramine	Fenoprofen	Naltrexone	Sustiva (Efavirenz)
Buspirone	Fentanyl	α-Naphthaleneacetic acid	Temazepam
Caffeine	Fluoxetine	Naproxen	Tetracycline
Cannabidiol	Furosemide	Niacinamide	Tetrahydrocortexolone
Cannabinol	Gentisic acid	Nifedipine	Tetrahydrocortisone, 3-acetate
Cimetidine	d-Glucose	Nimesulide	
Chloral hydrate	Guaiacol glyceryl ether	Norcodeine	Tetrahydrozoline
Chloramphenicol	Hemoglobin	Normorphine	Thebaine
Chlordiazepoxide	Hydralazine	Norethindrone	Theophylline
Chloroquine	Hydrochlorothiazide	d-Norpropoxyphene	Thiamine
Chlorothiazide	Hydrocodone	Noscapine	Thioridazine
(+)-Chlorpheniramine	Hydrocortisone	d,l-Octopamine	l-Thyroxine
(±)-Chlorpheniramine	o-Hydroxyhippuric acid	Orphenadrine	Tolbutamide
Chlorpromazine	p-Hydroxymethamphetamine	Oxalic acid	cis-Tramadol
Chlorprothixene	Hydromorphone	Oxazepam	trans-2-Phenylcyclopropylamine
Cholesterol	3-Hydroxytyramine	Oxolinic acid	
Clomipramine	(Dopamine)	Oxycodone	Trazodone

Clonidine	Hydroxyzine	Oxymetazoline	Trimethobenzamide
Codeine	Ibuprofen	Oxymorphone	Triamterene

Interfering Substances

The EDDP Rapid Test Cassette (whole blood/serum/plasma) has been tested for possible interference from visibly hemolyzed and lipemic specimens. In addition, no interference was observed in specimens containing up to 100 mg/dl hemoglobin;up to 100 mg/dl bilirubin; and up to 200 mg/dl human serum albumin.

【BIBLIOGRAPHY】

1. Hardman JG, limbird lE. Goodman and Gilman's: The Pharmacological Basis for Therapeutics. 10th Edition. McGraw Hill Medical Publishing, 2001; 208-209.
2. Baselt RC. Disposition of Toxic Drugs and Chemicals in Man.2nd Ed. Biomedical Publ., Davis, CA. 1982; 488

Number: 145318101
Effective date: 2017-06-22