Oratect®
Oral Fluid Drug Screen Device

Catalog number: AOT-06

The Alere Oratect® Oral Fluid Drug Screen Device is a one-step lateral flow immunoassay device for the qualitative detection of Methamphetamine (ME), Marijuana (TH), Cocaine (CO), Amphetamine (AM), Opiate (OP) and Phencyclidine (PC) in human oral fluid. The Alere Oratect® Test detects these drugs at the cut-off concentration listed below and their metabolites. The test is a prescription assay. This product is for in vitro diagnostic use and it can be used at the Point-of-Care site.

The Alere Oratect® Test device detects these drugs at the following cut-off concentrations:

<table>
<thead>
<tr>
<th>Drug</th>
<th>Cut-off</th>
</tr>
</thead>
<tbody>
<tr>
<td>Methamphetamine (ME)</td>
<td>50 ng/mL</td>
</tr>
<tr>
<td>Marijuana (TH)</td>
<td>40 ng/mL</td>
</tr>
<tr>
<td>Cocaine (CO)</td>
<td>20 ng/mL</td>
</tr>
<tr>
<td>Amphetamine (AM)</td>
<td>50 ng/mL</td>
</tr>
<tr>
<td>Opiate (OP)</td>
<td>40 ng/mL</td>
</tr>
<tr>
<td>Phencyclidine (PC)</td>
<td>10 ng/mL</td>
</tr>
</tbody>
</table>

The Alere Oratect® Oral Fluid Drug Screen Device provides only preliminary drug test results. For a quantitative result or for a confirmation of a presumptive positive result obtained by the Alere Oratect® Oral Fluid Drug Screen Device, a more specific alternative method must be used. GC/MS or LC/MS/MS is the preferred confirmatory method. The samples for confirmatory testing should be collected with the oral fluid confirmation tube provided.

Summary and Explanation

Illegal drug consumption contributes to many accidents, injuries and medical conditions. Screening individuals for drugs of abuse is an important method in identifying those who may cause harm to themselves and to others.

The Alere Oratect® Oral Fluid Drug Screen Device is developed to detect active drugs-of-abuse present in saliva. Studies on methamphetamine, cannabinoids, cocaine, amphetamine, opiates, and phencyclidine show that all of these drugs are detectable in oral fluids. The Alere Oratect® Oral Fluid Drug Screen Device is designed to integrate oral fluid collection and lateral flow immunoassay screen testing for drugs-of-abuse in one single device.

Test Principle

The Alere Oratect® Oral Fluid Drug Screen Device is based on a competitive immunoassay procedure in which drug derivatives immobilized on the membrane compete with the drug(s) which may be present in oral fluid for limited antibody binding sites on the colored colloidal gold antibody conjugate. During testing, oral fluid is collected at the collection pad and migrates across the membrane. If no drug is present in the oral fluid, the colored colloidal gold antibody conjugate will bind to the drug derivatives on the membrane to form visible bands at specific test regions. Therefore, the presence of a purple-red band at a specific test region indicates a negative result. If any drug(s) is (are) present in the oral fluid, it competes with the immobilized drug conjugate for limited antibody binding sites of the colored colloidal gold conjugate. When a sufficient amount of drug is present, the drug will saturate the antibodies, and the colored colloidal gold conjugate cannot bind to the drug derivative on the membrane. Therefore, the absence of a purple-red band at the test region indicates a presumptive positive result for that particular test.

The presence of a blue line in each window indicates that the device is unused. The movement of the blue lines indicates that a sufficient amount of oral fluid has been collected. A control band at the control region (C) indicates the test has performed properly. This control band should always appear regardless of the presence of drug or metabolite.

Reagents

The Alere Oratect® Oral Fluid Drug Screen Device contains one or two membrane strips and a collection pad. Each strip consists of a membrane, a colloidal gold conjugate pad, a sample pad and an absorbent pad.

The number of drugs per strip may vary depending on the selected product catalogue number.

Membrane:

- ME/TH/CO test strip: Methamphetamine, THC and Cocaine-protein conjugates are coated onto specific region on the membrane known as the “Test Region”.
- AM/OP/PC test strip: Amphetamine, Opiate, Phencyclidine protein conjugates are coated onto the test region of the membrane.

Colloidal Gold Conjugate Pad:


Collection Pad:

- The collection pad consists of an absorbent material.

Materials Provided

Each Alere Oratect® Oral Fluid Drug Screen Device kit contains:

- 25 test devices. Each device consists of a plastic holder and a detachable cap. The devices are packaged individually in a foil pouch with a desiccant.
- 1 Package Insert
- 1 Oral Fluid Collection Tube (50 mL polypropylene tube) for confirmation shipping. The Alere Oratect® Oral Fluid Collection Tube provided in this kit should only be used for the confirmation sample.

Warnings and Precautions

- For in vitro diagnostic use only
- The test device should remain in its original sealed pouch until ready for use.
- Discard the test device if package is ripped or torn.
- Do not use the test device beyond the expiration date indicated on the kit.
- Handle all oral specimens as potentially infectious. Proper handling and disposal methods should be established.

Product Storage

The Alere Oratect® Oral Fluid Drug Screen Device pouch should be stored at room temperature 15°C-30°C (59°F-86°F). Do not open pouch until ready to perform the assay.

Specimen Collection and Handling

IMPORTANT: At least 10 minutes prior to administering the test, instruct the donor not to eat, drink, smoke or chew tobacco products.

Presumptive positive samples are collected in the 50 ml collection tube supplied and mailed immediately to confirm the test.

Confirmation laboratory may keep samples for up to 2 weeks when stored at 2 – 8 °C or up to 24 months when stored below -15°C.

Test Procedure

1. Remove the test device from the sealed pouch.
2. Carefully remove the cranberry color cap by holding the sides and pull gently. This will expose the collection pad.
3. Ensure that the blue line is present in each test window.
4. The oral fluid collection process must be observed. Instruct the donor to hold the top portion of the device (above the test windows).
5. When placing device into the mouth, keep head level.
   a. Open mouth and rub the collection pad inside mouth against one cheek gently in a circular motion several (approximately 15-20) times. (Fig. b)
   b. Still keeping head level, gently rub the collection pad against the opposite cheek in a circular motion (approximately 15-20) several times. (Fig. b)
CO test regions

Opiate and Cocaine

In the control band. Any band that can be seen visually, no matter how faint, the color of the test band may be slightly darker or lighter than the example Fig e.

For each of the test windows, purple-red colored bands should be specific drug abbreviation (e.g. AM, OP, CO) in the test region. See

Negative Results

Interpreting Test Results

6. Place the collection pad underneath the tongue for approximately 30 seconds to collect saliva. Instruct the donor to hold the device in place with their hand.

7. The movement of the blue lines indicates the collection of a sufficient amount of saliva has occurred. If blue lines are still stationary after placing the collection pad underneath the tongue for 30 seconds, repeat the procedure in steps 5 and 6 until the blue lines move.

8. Remove the device from mouth as soon as the blue lines start moving at both test windows.

Note: The flow of the blue lines should appear in the test windows within 5 minutes. If no flow is observed after 5 minutes in the mouth, discard the device, review procedures 4-7 above with the donor and repeat the test using a new device.

9. Re-cap the device, lay it on a flat surface and read results in approximately 5 minutes after removing device from mouth. Do not read results after 15 minutes.

Invalid Results

When no band appears in the control (C) region, the test is invalid regardless of the results in the test region. If the test is invalid, check testing procedures. Repeat the test using a new device. In Fig. g below, the test is invalid because there are no bands in the control regions.

Instructions for collecting a confirmation oral fluid sample.

1. If a user of this product obtains a presumptive positive test result, the user should obtain confirmation testing using a more specific test method such as Gas Chromatography/Mass Spectrometry (GC/MS) or Liquid Chromatography/Mass Spectrometry (LC/MS/MS). The test sample for this confirmation testing should be collected immediately after obtaining the presumptive positive test result(s).

2. Use only the Alere Oracet® Oral Fluid Collection Tube (50ml polypropylene tube) provided in this kit. Note: If additional sample collection tubes are needed, please contact Alere Toxicology customer service at 1-800-340-4029. Remove cap from collection tube and carefully spit into tube several times until half of the bottom cone (~ 2.5 mLs) is collected.

3. Tightly re-cap sample collection tube.

4. Complete the tube label affixed to sample collection tube with the requested information.

5. Avoid high temperatures and sunlight pending shipment.

6. Mail the sample immediately to a license test laboratory for GC/MS or LC/MS/MS confirmation testing. The sample should be shipped by an overnight courier service using a small shipping box or padded envelope.

Quality Control

Internal control: The Alere Oracet® Oral Fluid Drug Screen Device provides a built-in control bands in each window at the control regions (C) to indicate that the test has performed properly. These control bands should always appear regardless of the presence of drugs. The flow of the blue lines indicates that a sufficient amount of oral fluid has been collected. The presence of the purple-red bands in the control regions verifies that proper flow was obtained. If the control bands do not appear, the test device should be discarded.

External control: It is recommended that negative and positive saliva controls be used to initially test each new lot of product to ensure proper kit performance. The use of the Alere Oracet® Oral Fluid Drug Screen Device Controls under catalogue number OC001 may be used
as external quality control material. The pipette test procedure provided with the Alere Oratect Oral Fluid Drug Screen Device Controls package insert should be followed. Each Laboratory should establish and run its own QC program as it is familiar with its own environment. When external controls do not produce the expected results, repeat with a new unopened bottle of controls.

Quality control testing at regular intervals is a good laboratory practice and laboratories should comply with all federal, state, and local laws, guidelines and regulations. Always check with the appropriate licensing or accrediting bodies to ensure that the quality program employed meets the established standards.

Limitations of Procedure
- The assay is designed for human oral fluid use only.
- Positive results only indicate the presumptive presence of drugs and do not indicate or measure intoxication.
- Technical or procedural errors as well as substances in certain foods and certain medications may interfere with the test and cause false results.
- Do not use the device past expiration date
- Read instructions before testing
- Subjects with dry mouth symptoms have difficulty with this test

Performance Characteristics Comparison and Accuracy
The accuracy of the Alere Oratect Oral Fluid Drug Screen Device was evaluated by testing Oratect product with clinical saliva samples which were subsequently analyzed by GC/MS or LC/MS/MS method. A minimum of forty negative samples and forty positive samples were tested. Of the forty positive samples, at least 4 samples were near negative (between 50% to 100%) and 4 samples were near positive (between 100% to 150%). The results are summarized below:

<table>
<thead>
<tr>
<th>Drug Name</th>
<th>GC/MS negative</th>
<th>GC/MS positive</th>
<th>Near cutoff negative ≥50% to &lt;100%</th>
<th>Near cutoff positive ≥100% to &lt;150%</th>
<th>GC/MS positive ≥150%</th>
<th>% Agreement with GC/MS</th>
</tr>
</thead>
<tbody>
<tr>
<td>ME</td>
<td>+</td>
<td>-</td>
<td>0</td>
<td>0</td>
<td>5</td>
<td>58</td>
</tr>
<tr>
<td>TH</td>
<td>+</td>
<td>-</td>
<td>0</td>
<td>0</td>
<td>10</td>
<td>36</td>
</tr>
<tr>
<td>CO</td>
<td>+</td>
<td>-</td>
<td>0</td>
<td>0</td>
<td>3</td>
<td>38</td>
</tr>
<tr>
<td>AM</td>
<td>+</td>
<td>-</td>
<td>0</td>
<td>0</td>
<td>7</td>
<td>12</td>
</tr>
<tr>
<td>OP</td>
<td>+</td>
<td>-</td>
<td>0</td>
<td>0</td>
<td>4</td>
<td>55</td>
</tr>
<tr>
<td>PC</td>
<td>+</td>
<td>-</td>
<td>0</td>
<td>0</td>
<td>1</td>
<td>2</td>
</tr>
</tbody>
</table>

Precision
For each specific drug test, pooled oral fluid solution was spiked with a drug standard at various concentrations (0%, 25%, 50%, 75%, 100%, 125%, 150%, 175%, and 200%). Three lots were tested by at least 3 operators at 3 different sites to validate the test performance. The results for each drug of the Alere Oratect Oral Fluid Drug Screen Device Tests are summarized below:

<table>
<thead>
<tr>
<th>Cut-off level</th>
<th>ME</th>
<th>TH</th>
<th>CO</th>
<th>AM</th>
<th>OP</th>
<th>PC</th>
</tr>
</thead>
<tbody>
<tr>
<td>0%</td>
<td>360</td>
<td>0</td>
<td>360</td>
<td>0</td>
<td>360</td>
<td>0</td>
</tr>
<tr>
<td>25%</td>
<td>45</td>
<td>0</td>
<td>45</td>
<td>0</td>
<td>45</td>
<td>0</td>
</tr>
<tr>
<td>50%</td>
<td>45</td>
<td>0</td>
<td>45</td>
<td>0</td>
<td>45</td>
<td>0</td>
</tr>
<tr>
<td>75%</td>
<td>43</td>
<td>2</td>
<td>43</td>
<td>3</td>
<td>43</td>
<td>4</td>
</tr>
<tr>
<td>100%</td>
<td>26</td>
<td>19</td>
<td>25</td>
<td>20</td>
<td>27</td>
<td>18</td>
</tr>
<tr>
<td>125%</td>
<td>3</td>
<td>42</td>
<td>2</td>
<td>43</td>
<td>4</td>
<td>41</td>
</tr>
<tr>
<td>150%</td>
<td>0</td>
<td>45</td>
<td>0</td>
<td>45</td>
<td>0</td>
<td>45</td>
</tr>
<tr>
<td>175%</td>
<td>0</td>
<td>45</td>
<td>0</td>
<td>45</td>
<td>0</td>
<td>45</td>
</tr>
<tr>
<td>200%</td>
<td>0</td>
<td>45</td>
<td>0</td>
<td>45</td>
<td>0</td>
<td>45</td>
</tr>
</tbody>
</table>

Specificity
The specificity study for each drug test was evaluated by adding structurally related compounds to pooled oral fluid sample. The results are expressed as the amount of the compound, in ng/ml, that produced a positive result.

Interference
The Alere Oratect Oral Fluid Drug Screen Test performance at ± 50% cut-off levels is not affected by any oral fluid samples with pH range of 4.0 to 8.5. The following compounds were spiked into ± 50% oral fluid controls and found not to cross-react with the Alere Oratect Oral Fluid Drug Screen Device when tested at concentration of 10 µg/ml (10,000ng/ml).
Acetaminophen      Hemoglobin
α-Amylase           Human IgA
Albumin from human serum Human IgG
l-Ascorbic Acid     Human IgM
Aspartame           Ibuprofen
Benzillic acid      Ketamine
Benzocaine          Lidocaine
Benzoic acid        Meperidine
Bilirubin           Naltrexone hydrochloride
Butether            Naloxone
Caffeine            d-Naproxen
d-Chlorpheniramine   Papaverine
Cholesterol         Pentazocine
Dextromethorphan    Promazine
Diphenhydramine     Promethazine
Doxylamine          Ranitidine
1R, 2S l- Ephedrine (except ME assay) Riboflavin
1S, 2R d-Ephedrine  Salicylic acid
l-Epinephrine       Serotonin
Erythromycin        Tetracycline
Ethanol             Thiamine
Glutethimide        d,l-Tryptophan

Food/Beverage/Hygiene Products Interference
Foods, drinks and hygiene products were spiked at 1% concentration in ±25% and ±50% oral fluid controls to evaluate the interference with Oratect® test results. For interference of cigarette, oral fluid samples were collected from 6 subjects within 15 minutes after consuming a cigarette and then spiked with drug standards (MET, THC, COC, AMP, OPI, and PCP). The following substances were found not to interfere with Alere Oratect® Oral Fluid Drug Screen Test performance.

<table>
<thead>
<tr>
<th>Mouth wash</th>
<th>Orange juice</th>
<th>Alcohol</th>
</tr>
</thead>
<tbody>
<tr>
<td>MSG</td>
<td>Sugar</td>
<td>Cranberry juice</td>
</tr>
<tr>
<td>Salt</td>
<td>Food color: Red</td>
<td>Carbonated Cola</td>
</tr>
<tr>
<td>Toothpaste</td>
<td>Food color: Green</td>
<td>Baking Soda</td>
</tr>
<tr>
<td>Gum</td>
<td>Food color: Blue</td>
<td>Cigarette</td>
</tr>
<tr>
<td>Cough Syrup</td>
<td>Tea</td>
<td></td>
</tr>
</tbody>
</table>

Bibliography of Suggested Reading