



### NAME AND INTENDED USE

The OraQuick® HCV Rapid Antibody Test is a single-use immunoassay for the qualitative detection of antibodies to hepatitis C virus (anti-HCV) in venipuncture whole blood specimens (EDTA, sodium heparin, lithium heparin, and sodium citrate) from individuals 15 years or older. The OraQuick® HCV Rapid Antibody Test results, in conjunction with other laboratory results and clinical information, may be used to provide presumptive evidence of infection with HCV (state of infection or associated disease not determined) in persons with signs or symptoms of hepatitis and in persons at risk for hepatitis C infection.

**Warning: This assay has not been FDA approved for use in patient populations without signs, symptoms, or not at risk for hepatitis C infection.**

**Not for use in screening whole blood, plasma, or tissue donors. Performance characteristics have not been established for testing a pediatric population less than 15 years of age or for pregnant women.**

### SUMMARY AND EXPLANATION OF THE TEST

Hepatitis C virus is a single-stranded ribonucleic acid RNA virus responsible for most, if not all non-A, non-B hepatitis.<sup>2</sup> HCV is primarily transmitted by contact with contaminated blood, blood products, or through other close personal contact.<sup>3</sup> The presence of antibodies to HCV indicates that the individual may be currently infected and capable of transmitting the virus.

The OraQuick® HCV Rapid Antibody Test utilizes an indirect lateral flow immunoassay method to detect antibodies to both structural and non-structural HCV proteins. The device utilizes synthetic peptides and recombinant antigens from the core, NS3, and NS4 regions of the HCV genome, that are immobilized as a single test line on the assay strip. Antibodies reacting with these peptides and antigens are visualized by colloidal gold labeled with protein A generating a visible line in the test zone for a reactive sample.

### PRINCIPLES OF THE TEST

The OraQuick® HCV Rapid Antibody Test is a manually performed, visually read immunoassay for the qualitative detection of HCV antibodies in human venipuncture whole blood. The OraQuick® HCV Rapid Antibody Test is comprised of both a single-use test device and vial containing a pre-measured amount of a buffered developer solution. The test consists of a sealed pouch with two separate compartments for each component. The OraQuick® HCV Rapid Antibody Test utilizes a proprietary lateral flow immunoassay procedure.

The assay test strip, which can be viewed through the test device result window, contains synthetic peptides and recombinant proteins from the core, NS3, and NS4 regions of the HCV genome (test) and a goat anti-human IgG (procedural control) immobilized onto a nitrocellulose membrane at the Test (T) and the Control (C) Zone, respectively.

A venipuncture whole blood specimen is collected using a specimen loop and transferred into the developer solution vial, followed by the insertion of the device. The developer solution facilitates the capillary flow of the specimen into the device and onto the assay strip. As the specimen flows through the device, antibodies from the specimen are bound to the protein A gold colorimetric reagent present on the assay strip. If the specimen contains anti-HCV antibodies, the resulting labeled complexes contain HCV antibody and bind to immobilized HCV antigens at the HCV Test Zone (T Zone) resulting in a reddish-purple line. If the specimen does not contain anti-HCV antibodies, the labeled complexes do not bind at the HCV Test Zone and no line is observed in the T Zone. The intensity of the line color is not directly proportional to the amount of HCV antibody present in the specimen. The remaining labeled complexes are transported to the Control Zone (C Zone) binding to a goat anti-human antibody fragment. The presence of IgG antibodies in the sample (regardless of their specificity) results in a reddish-purple line at the C Zone. This procedural control serves to demonstrate that a specimen was added to the vial and that the fluid has migrated adequately through the device. A reddish-purple line will appear at the C Zone during the performance of all valid tests; whether or not the sample is positive or negative for HCV antibodies (refer to the *Test Result and Interpretation of Test Result* section in this package insert).

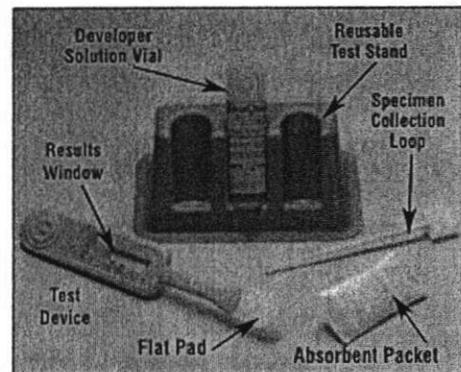
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The test results are interpreted after 20 minutes, but not more than 40 minutes following the introduction of the device into the developer solution vial. No precision pipetting, pre-dilutions, or specialized instrumentation are required to perform the OraQuick® HCV Rapid Antibody Test.

**MATERIALS PROVIDED**

OraQuick® HCV Rapid Antibody Test Kits are available in the following packaging configurations:

| Components of Kit<br>Catalog Number   | 25 Count Kit<br>1001-0181 | 100 Count Kit<br>1001-0180 |
|---|---------------------------|----------------------------|
| Divided Pouch,<br>Each containing:<br>Test Device (1)<br>Absorbent Packet (1)<br>Developer Solution Vial (1)<br>(each vial contains 0.750 mL of a buffered saline solution with an antimicrobial agent) | 25                        | 100                        |
| Reusable Test Stands  | 5                         | 10                         |
| Specimen Collection Loops   | 25                        | 100                        |
| Package Insert  | 1                         | 1                          |



**MATERIALS REQUIRED AND AVAILABLE AS AN ACCESSORY TO THE KIT**

**OraQuick® HCV Rapid Antibody Test Kit Controls 1001-0182**

Package contains:

- HCV Positive Control (1 vial, purple cap, 0.2 mL),
- HCV Negative Control (1 vial, white cap, 0.2 mL), and
- Package Insert.

**MATERIALS REQUIRED BUT NOT PROVIDED**

- Timer or watch capable of timing 20 to 40 minutes
- Biohazard waste container
- Materials required for venipuncture whole blood specimen collection

**RESTRICTIONS**

- Sale of the OraQuick® HCV Rapid Antibody Test is restricted to healthcare professionals:
  - that have an adequate quality assurance program, including planned systematic activities to provide adequate confidence that requirements for quality will be met; and
  - where there is assurance that operators will receive and use the instructional materials.
- This assay has not been FDA approved for use in patient populations without signs, symptoms, or not at risk for hepatitis C infection.
- Not for use in screening whole blood, plasma, or tissue donors. Performance characteristics have not been established for testing a pediatric population less than 15 years of age or for pregnant women.

**WARNINGS**

For *in vitro* Diagnostic Use

- This package insert must be read completely before using the product.
- Follow the instructions carefully when performing the OraQuick® HCV Rapid Antibody Test, failure to do so may cause an inaccurate test result.
- Before proceeding with testing, all operators MUST read and become familiar with Universal Precautions for Prevention of Transmission of Human Immunodeficiency Virus, Hepatitis A Virus, Hepatitis B Virus, and other Bloodborne Pathogens in Health-Care Settings.<sup>1</sup>
- This kit has been approved for use with venipuncture whole blood specimens only. Use of this test kit with specimen types other than those specifically approved for this device may cause inaccurate test results.
- This test is not intended to be used to monitor individuals who are undergoing treatment.
- This test should be performed at temperatures in the range of 15°-37°C (59°-99°F). If stored refrigerated, ensure that the Divided Pouch is brought to operating temperature (15°-37°C, 59°-99°F) before performing testing.
- Do not use if the test kit is exposed to temperatures outside of the recommended storage temperature (2°-30°C, 36°-86°F), or is tested outside of the operating temperature (15°-37°C, 59°-99°F).

## PRECAUTIONS

### Safety Precautions

- Handle specimens and materials in contact with specimens as if capable of transmitting infectious agents.
- Wear disposable gloves while handling and testing blood specimens. Change gloves and wash hands thoroughly after performing each test. Dispose of used gloves in a biohazard waste container.
- For additional information on biosafety, refer to "Universal Precautions for Prevention of Transmission of Human Immunodeficiency Virus, Hepatitis A Virus, Hepatitis B Virus, and other Bloodborne Pathogens in Health-Care Settings"<sup>1</sup> and "Updated U.S. Public Health Service Guidelines for the Management of Occupational Exposures to HBV, HCV, and HIV and Recommendations for Postexposure Prophylaxis".<sup>4</sup>

### Device Handling Precautions

- Use all Specimen Collection Loops, Test Devices, and Developer Solution Vials only once and dispose of properly (see *Safety Precautions*). **Do not reuse any of these test components.**
- Do not use the test beyond the expiration date printed on the Divided Pouch. Always check expiration date prior to testing.
- Inspect the Divided Pouch. If the Divided Pouch has been damaged, discard the Divided Pouch and its contents and select a new Divided Pouch for testing.
- Do not interchange Test Devices and Developer Solution Vials from kits with different lot numbers.
- Avoid microbial contamination and exercise care in handling the kit components.
- To ensure accurate results, the Test Device must be inserted into the Developer Solution Vial within 60 minutes after introducing the specimen into the Developer Solution.
- Adequate lighting is required to read a test result.

## STORAGE INSTRUCTIONS

Store unused OraQuick<sup>®</sup> HCV Rapid Antibody Tests unopened at 2°-30°C (36°-86°F). Do not open the Divided Pouch until you are ready to perform a test. If stored refrigerated, ensure that the Divided Pouch is brought to operating temperature (15°-37°C, 59°-99°F) before opening.

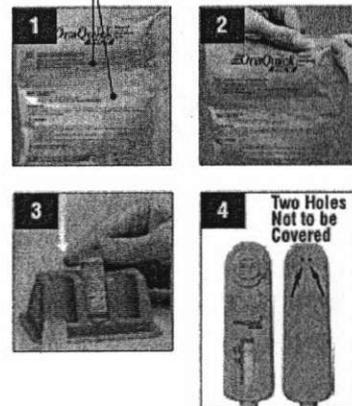
## DIRECTIONS FOR USE

### GENERAL TEST PREPARATION

1. Follow *Safety Precautions* section in this package insert.
2. Gather the materials you will need.
3. Allow the OraQuick<sup>®</sup> HCV Rapid Antibody Tests to come to operating temperature (15°-37°C; 59°-99°F) before use.
4. Refer to the *External Quality Control* section in this package insert to determine when the Kit Controls should be run.
5. Set an OraQuick<sup>®</sup> Reusable Test Stand at your workspace, using only the stand provided.
6. Open the two chambers of the OraQuick<sup>®</sup> Divided Pouch by tearing at the top notches located on each side of the Pouch (see *picture 1 and 2*). To prevent contamination, leave the Test Device in the Pouch until you are ready to use it.
7. Remove the Developer Solution Vial from the Pouch. Hold the Developer Vial firmly in your hand. Remove the cap from the Developer Vial by gently rocking the cap back and forth while pulling it off. Set the cap aside. Slide the vial into the top of the slots in the Reusable Test Stand. (see *picture 3*).

**NOTE: DO NOT cover the two holes in the back of the Device with labels or other materials, as it may cause an Invalid test result (see *picture 4*).**

Two Chambers



## SPECIMEN COLLECTION AND TESTING

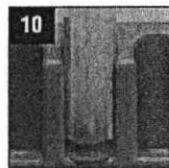
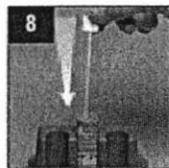
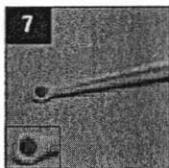
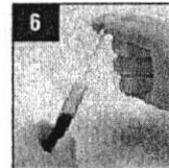
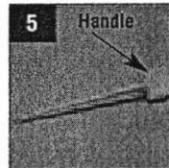
### PROCEDURE

The OraQuick<sup>®</sup> HCV Rapid Antibody Test can be used for testing venipuncture whole blood specimens. Refer to the specific testing procedure below.

### VENIPUNCTURE WHOLE BLOOD PROCEDURE

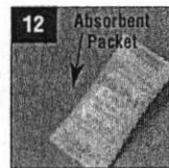
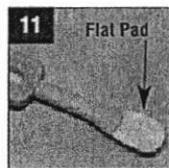
#### STEP 1: COLLECT

1. Using standard venous phlebotomy procedures collect a whole blood specimen using a tube containing any of the following anticoagulants: EDTA, sodium heparin, lithium heparin, or sodium citrate. **Other anticoagulants have not been tested and may cause an inaccurate result.** If the specimens are not tested at the time of collection, the whole blood may be stored at 2°-8°C (36°-46°F) for up to 7 days or at 15°-30°C (59°-86°F) for up to 3 days.
2. Prior to testing, mix the blood tube gently by inversion several times to ensure a homogeneous specimen. Obtain an unused Specimen Collection Loop by the handle (see picture 5). Insert the rounded end of the Loop into the tube of blood (see picture 6), and verify that the Loop is completely filled with blood (see picture 7). **NOTE:** If the Loop is dropped or comes in contact with any other surface, discard it in a biohazard waste container. Use a new Loop for the collection of the blood specimen.



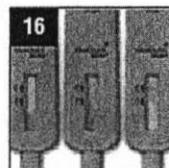
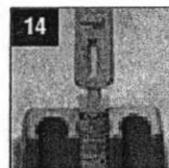
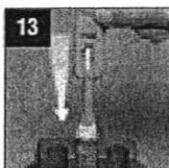
#### STEP 2: MIX

1. Immediately insert the blood-filled end of the Loop all the way into the Developer Vial (see picture 8). Use the Loop to stir the blood sample in the Developer Solution (see picture 9). Remove the used Loop from the Solution and discard in a biohazard waste container.
2. Verify that the Solution is pink in color, indicating that the blood was thoroughly mixed into the Solution (see picture 10). If the Solution is not pink, discard all test materials in a biohazard waste container. Start the test over using a new Pouch and a new blood sample. **NOTE:** To ensure accurate results, the Test Device must be inserted into the Developer Solution Vial within 60 minutes after introducing the specimen into the Developer Solution.



#### STEP 3: TEST

1. Remove the Device from the Pouch. **DO NOT** touch the Flat Pad (see picture 11). Verify that an Absorbent Packet is included with the Device (see picture 12). If no Absorbent Packet is present, or if the Absorbent Packet is damaged, discard the Device and obtain a new Pouch for testing.
2. Insert the Flat Pad of the Device all the way into the Developer Vial containing the blood sample (see picture 13), and verify that the Flat Pad touches the bottom of the Developer Vial. The Result Window on the Device should be facing towards you (see picture 14).
3. Start timing the test (see picture 15). **DO NOT** remove the Device from the Developer Vial while the test is running. A pink color will migrate up the Result Window, and will gradually disappear as the test develops (see picture 16). Read the result in a fully lighted area after 20 minutes, but no more than 40 minutes.
4. Refer to the *Test Result and Interpretation of Test Result* section in this package insert.



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## GENERAL TEST CLEAN-UP

1. Dispose of the used test materials and gloves in a biohazard waste container.
2. When using gloves, change your gloves between each test to prevent contamination.
3. Use a freshly prepared 10% solution of bleach to clean up any spills.<sup>5</sup>

## QUALITY CONTROL

### *Built-in Control Features*

The OraQuick<sup>®</sup> HCV Rapid Antibody Test has a built-in procedural control that demonstrates assay validity. A reddish-purple line in the Control Zone (C Zone) of the Result Window indicates that a specimen was added and that the fluid migrated appropriately through the Device. The Control line will appear on all valid tests, whether or not the sample is reactive or non-reactive for anti-HCV (Refer to *Test Result and Interpretation of Test Result* section in this package insert).

### *External Quality Control*

OraQuick<sup>®</sup> HCV Rapid Antibody Test Kit Controls are available separately for use only with the OraQuick<sup>®</sup> HCV Rapid Antibody Test. The Kit Controls are specifically formulated and manufactured to ensure proper performance of the test. The HCV Positive Control will produce a reactive reddish-purple line at the Test Zone (T Zone). The HCV Negative Control will generate a non-reactive test result (no reddish-purple line at the T Zone). Refer to *Test Result and Interpretation of Test Result* section in this package insert. Use of kit control reagents manufactured by any other source will not meet the requirements for an adequate quality assurance program for the OraQuick<sup>®</sup> HCV Rapid Antibody Test.

Additional controls may be tested according to guidelines or requirements of local, state, and /or federal regulations or accrediting organizations.

### **Run the Kit Controls under the following circumstances:**

- Each new operator prior to performing testing on patient specimens,
- When opening a new test kit lot,
- Whenever a new shipment of test kits is received,
- If the temperature of the test kit storage area falls outside of 2°-30°C (36°-86°F),
- If the temperature of the testing area falls outside of 15°-37°C (59°-99°F), and
- At periodic intervals as dictated by the user facility.

Refer to the OraQuick<sup>®</sup> HCV Rapid Antibody Test Kit Controls package insert for instructions on the use of these reagents. It is the responsibility of each laboratory using the OraQuick<sup>®</sup> HCV Rapid Antibody Test to establish an adequate quality assurance program to ensure the performance of the device under its specific locations and conditions of use. Contact OraSure Technologies' Customer Service if the Kit Control reagents do not produce the expected results.

## TEST RESULT AND INTERPRETATION OF TEST RESULT

Refer to the Result Window on the Test Device.

### **NON-REACTIVE**

A test is **Non-Reactive** if:

a line appears in the C Zone and NO line appears in the T Zone (see picture 17).

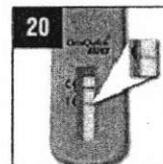
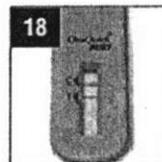
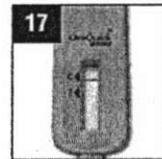
A **Non-Reactive** test result means that HCV antibodies were not detected in the specimen. Patient is presumed not to be infected with HCV.

### **REACTIVE**

A test is **Reactive** if:

a line appears in the C Zone and a line appears in the T Zone. Lines may vary in intensity. The test is reactive regardless of how faint these lines appear (see pictures 18, 19 and 20).

A **Reactive** test result means that HCV antibodies have been detected in the specimen. Patient is presumed to be infected with HCV.



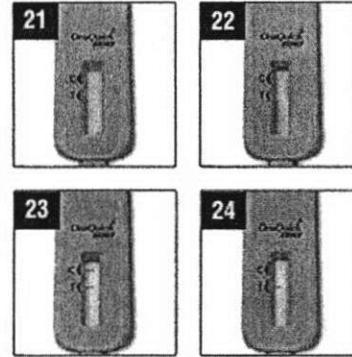
Individuals with a reactive result in the OraQuick® HCV Rapid Antibody Test should undergo appropriate clinical follow-up, according to CDC recommendations for supplemental testing.<sup>6</sup>

### INVALID

A test is **Invalid** if:

- **NO** line appears in the C Zone (see picture 21), or
- a pink background obscures the results during the 20- to 40 minute read times (see picture 22), or
- any partial line on one side of the C or T Zones (see pictures 23 and 24).

An **Invalid** test result means that there was a problem running the test, either related to the specimen or to the Device. An **Invalid result cannot be interpreted**. Repeat the test with a new Pouch and a new specimen. Contact OraSure Technologies' Customer Service if you are unable to get a valid test result upon repeat testing.



### LIMITATIONS OF THE TEST

1. The OraQuick® HCV Rapid Antibody Test must be used in accordance with the instructions in this package insert to obtain an accurate result.
2. Reading test results earlier than 20 minutes or later than 40 minutes may yield inaccurate test results.
3. This test is approved for use with venipuncture whole blood specimens only. Use of other types of specimens, or venipuncture whole blood specimens collected using a tube containing anticoagulants other than EDTA, lithium heparin, sodium heparin, or sodium citrate may yield inaccurate results.
4. Clinical data has not been collected to demonstrate the performance of the OraQuick® HCV Rapid Antibody Test in individuals under 15 years of age.
5. A reactive result using the OraQuick® HCV Rapid Antibody Test suggests the presence of HCV antibodies in the specimen, and the intensity of the test line does not necessarily correlate with the HCV antibody titer in the specimen. The OraQuick® HCV Rapid Antibody Test is intended as an aid in the diagnosis of HCV infection.
6. A non-reactive result does not exclude the possibility of exposure to HCV or infection with HCV. An antibody response to recent exposure may take several months to reach detectable levels.
7. A person who has HCV antibodies is presumed to be infected with the virus. Additional testing and medical evaluation is required to determine the state or associated disease.

## EXPECTED RESULTS FOR THE INTENDED USE POPULATION

Of the 1207 specimens tested in two OraQuick® HCV Rapid Antibody Test clinical studies, 88.2% (1064/1207) were from subjects at risk for hepatitis C infection but were asymptomatic and reported no current signs or symptoms of hepatitis, and 11.8% (142/1207) were from subjects with current signs or symptoms of hepatitis. One (1/1207) pregnant subject was enrolled without signs or symptoms of hepatitis or risk factors for hepatitis C infection. The 1207 individuals were enrolled from the following collection locations:

- 50.3% from Miami, FL
- 24.7% from Ft. Lauderdale, FL
- 13.6% from Fall River, MA
- 10.7% from Allentown, PA
- 0.7% from College Park, MD, San Francisco, CA, Dallas TX, and Philadelphia, PA.

The OraQuick® HCV Rapid Antibody Test was reactive in 36.7% (443/1207) of subjects tested. There were no invalid OraQuick® HCV Rapid HCV Antibody Test results reported for the 1207 specimens tested (0/1207, 95% CI: 0.0%, 0.3%). Of the 1207 individuals tested, 33.1% (400/1207) were also self-reported HIV positive. The table below summarizes the distribution of OraQuick® HCV Rapid Antibody Test reactive, non-reactive, and invalid results for the 1207 specimens tested.

| Age Range  | Gender | OraQuick® HCV Rapid Antibody Test Results in High Risk Individuals |                       |                  |              |                   |                       |                  |              |
|------------|--------|--|-----------------------|------------------|--------------|-------------------|-----------------------|------------------|--------------|
|            |        | No signs or symptoms   |                       |                  |              | Signs or symptoms |                       |                  |              |
|            |        | Reactive<br>n (%)  | Non-Reactive<br>n (%) | Invalid<br>n (%) | Total<br>(n) | Reactive<br>n (%) | Non-Reactive<br>n (%) | Invalid<br>n (%) | Total<br>(n) |
| 0-9        | F      | 0 (0.0)  | 0 (0.0)               | 0 (0.0)          | 0            | 0 (0.0)           | 0 (0.0)               | 0 (0.0)          | 0            |
|            | M      | 0 (0.0)  | 0 (0.0)               | 0 (0.0)          | 0            | 0 (0.0)           | 0 (0.0)               | 0 (0.0)          | 0            |
| 10-19      | F      | 0 (0.0)  | 11 (1.0)              | 0 (0.0)          | 11           | 1 (0.7)           | 1 (0.7)               | 0 (0.0)          | 2            |
|            | M      | 0 (0.0)  | 14 (1.3)              | 0 (0.0)          | 14           | 0 (0.0)           | 1 (0.7)               | 0 (0.0)          | 1            |
| 20-29      | F      | 11 (1.0)   | 29 (2.7)              | 0 (0.0)          | 40           | 1 (0.7)           | 9 (6.3)               | 0 (0.0)          | 10           |
|            | M      | 7 (0.7)  | 50 (4.7)              | 0 (0.0)          | 57           | 1 (0.7)           | 3 (2.1)               | 0 (0.0)          | 4            |
| 30-39      | F      | 15 (1.4)   | 44 (4.1)              | 0 (0.0)          | 59           | 4 (2.8)           | 18 (12.7)             | 0 (0.0)          | 22           |
|            | M      | 23 (2.2)   | 56 (5.3)              | 0 (0.0)          | 79           | 4 (2.8)           | 4 (2.8)               | 0 (0.0)          | 8            |
| 40-49      | F      | 53 (5.0)   | 97 (9.1)              | 0 (0.0)          | 150          | 7 (4.9)           | 8 (5.6)               | 0 (0.0)          | 15           |
|            | M      | 82 (7.7)   | 191 (18.0)            | 0 (0.0)          | 273          | 6 (4.2)           | 8 (5.6)               | 0 (0.0)          | 14           |
| 50-59      | F      | 41 (3.9)   | 38 (3.6)              | 0 (0.0)          | 79           | 3 (2.1)           | 5 (3.5)               | 0 (0.0)          | 8            |
|            | M      | 137 (12.9)   | 117 (11.0)            | 0 (0.0)          | 254          | 23 (16.2)         | 9 (6.3)               | 0 (0.0)          | 32           |
| 60-69      | F      | 6 (0.6)  | 5 (0.5)               | 0 (0.0)          | 11           | 1 (0.7)           | 9 (6.3)               | 0 (0.0)          | 10           |
|            | M      | 13 (1.2)   | 19 (1.8)              | 0 (0.0)          | 32           | 2 (1.4)           | 5 (3.5)               | 0 (0.0)          | 7            |
| 70-79      | F      | 0 (0.0)  | 0 (0.0)               | 0 (0.0)          | 0            | 0 (0.0)           | 4 (2.8)               | 0 (0.0)          | 4            |
|            | M      | 1 (0.1)  | 4 (0.4)               | 0 (0.0)          | 5            | 1 (0.7)           | 2 (1.4)               | 0 (0.0)          | 3            |
| 80-89      | F      | 0 (0.0)  | 0 (0.0)               | 0 (0.0)          | 0            | 0 (0.0)           | 1 (0.7)               | 0 (0.0)          | 1            |
|            | M      | 0 (0.0)  | 0 (0.0)               | 0 (0.0)          | 0            | 0 (0.0)           | 1 (0.7)               | 0 (0.0)          | 1            |
| 90-100     | F      | 0 (0.0)  | 0 (0.0)               | 0 (0.0)          | 0            | 0 (0.0)           | 0 (0.0)               | 0 (0.0)          | 0            |
|            | M      | 0 (0.0)  | 0 (0.0)               | 0 (0.0)          | 0            | 0 (0.0)           | 0 (0.0)               | 0 (0.0)          | 0            |
| Total (N)* |        | 389 (36.6)   | 675 (63.4)            | 0 (0.0)          | 1064         | 54 (38.0)         | 88 (62.0)             | 00 (0.0)         | 142          |

\* Does not include one pregnant woman enrolled without signs or symptoms of hepatitis or at risk for hepatitis C infection.

## PERFORMANCE CHARACTERISTICS

### Clinical Performance

Two multi-center prospective studies were conducted to evaluate the clinical performance of the OraQuick® HCV Rapid Antibody Test in subjects with signs or symptoms of hepatitis and subjects at risk for hepatitis C infection. These risk factors included past or present intravenous drug use, having received a blood transfusion or organ transplant prior to 1992, evidence of high-risk sexual behavior, being born to an HCV positive mother, having been on long-term hemodialysis, history of incarceration, and positive for HIV. Clinical performance was evaluated in venipuncture whole blood specimens from subjects prospectively enrolled at 8 geographically dispersed centers within the United States.

The population tested was African American (43.0%), Caucasian (37.7%), Hispanic/Latino (17.1%), as well as a small proportion of other ethnic groups (2.2%). The mean age was 45 years (age range: 15 to 84 years). Of the 1207 subject specimens tested, 436 were HCV infected, 762 were negative, and 9 specimens had the status of "Unable to Determine". HCV status was determined for each subject by EIA, with supplemental RIBA® and PCR

assays as required. The table below summarizes the distribution of OraQuick® HCV Rapid Antibody Test reactive, non-reactive, and invalid results in subject with HCV infected status per the reference laboratory testing algorithm.

| OraQuick® HCV Rapid Antibody Test Results | Subject HCV Infected Status |          |                                     |
|---|-----------------------------|----------|-------------------------------------|
|   | Positive                    | Negative | Unable to Determine Infected Status |
| Positive                                  | 435                         | 0        | 8                                   |
| Negative                                  | 1                           | 762      | 1                                   |
| Invalid                                   | 0                           | 0        | 0                                   |

#### Positive and Negative Agreement Calculations

Percent positive and percent negative agreement between the OraQuick® HCV Rapid Antibody Test and HCV status were calculated overall for the population (n=1207), as well as for subjects with signs or symptoms of hepatitis and subjects at risk for hepatitis C infection.

$$\text{Percent Positive Agreement} = \frac{\text{Number of OraQuick® HCV Rapid Antibody Test Reactive Results}}{\text{Total number of HCV Infected status}} \times 100$$

$$\text{Percent Negative Agreement} = \frac{\text{Number of OraQuick® HCV Rapid Antibody Test Non-Reactive Results}}{\text{Total number of HCV Not Infected status}} \times 100$$

For the purposes of calculating percent agreement, OraQuick® HCV Rapid Antibody Test reactive results for samples whose HCV status was "Unable to Determine" following EIA with supplemental RIBA® and PCR testing were considered "HCV Not Infected", and OraQuick® HCV Rapid Antibody Test non-reactive results for samples whose HCV status was "Unable to Determine" following EIA with supplemental RIBA® and PCR testing were considered "HCV Infected".

#### Positive and Negative Agreement

The percent positive and negative agreement between the OraQuick® HCV Rapid Antibody Test and the subject HCV Infected Status was calculated for the population (n=1207). Percent of invalid OraQuick® HCV Rapid Antibody Test results was 0% (0/1207) with 95% CI: 0.0% to 0.3%. Percent positive and negative agreement was also calculated for individuals with signs or symptoms of hepatitis (n=142); and for individuals at risk for hepatitis C infection (n=1064). Percent positive and negative agreement according to risk factors for HCV infection was also calculated. The risks for HCV were ranked on a clinical evaluation of the likelihood of acquiring hepatitis C, with the most common given higher rankings.<sup>7</sup> Each subject was assigned only one risk (the highest). Results with the 95% confidence intervals are summarized in the following table.

| Study Subjects                    | Total | Positive Percent Agreement | 95% Exact Confidence Interval | Negative Percent Agreement | 95% Exact Confidence Interval |
|-----------------------------------|-------|----------------------------|-------------------------------|----------------------------|-------------------------------|
| Overall                           | 1207  | 99.5%*<br>(435 / 437)      | 98.4, 99.9                    | 99.0%*<br>(762 / 770)      | 98.0, 99.6                    |
| Overall with signs or symptoms    | 142   | 100.0%<br>(54 / 54)        | 93.4, 100.0                   | 100.0%<br>(88 / 88)        | 95.9, 100.0                   |
| Overall without signs or symptoms | 1064  | 99.5%*<br>(381 / 383)      | 98.1, 99.9                    | 98.8%*<br>(673 / 681)      | 97.7, 99.5                    |
| IVDU                              | 456   | 99.3%<br>(291 / 293)       | 97.6, 99.9                    | 98.2%<br>(160 / 163)       | 94.7, 99.6                    |
| Dialysis                          | 6     | 100%<br>(1 / 1)            | 2.5, 100.0                    | 100.0%<br>(5 / 5)          | 47.8, 100.0                   |
| Transfusion/Transplant            | 63    | 100.0%<br>(16 / 16)        | 79.4, 100.0                   | 100.0%<br>(47 / 47)        | 92.5, 100.0                   |
| High Risk sex                     | 461   | 100.0%<br>(58 / 58)        | 93.8, 100.0                   | 98.8<br>(398 / 403)        | 97.1, 99.6                    |
| HCV positive mother               | 2     | 100.0%<br>(1 / 1)          | 2.5, 100.0                    | 100.0%<br>(1 / 1)          | 2.5, 100.0                    |
| Prior history of incarceration    | 56    | 100%<br>(11 / 11)          | 71.5, 100.0                   | 100.0%<br>(45 / 45)        | 92.1, 100.0                   |
| HIV positive <sup>§</sup>         | 17    | 100.0<br>(2 / 2)           | 15.8, 100.0                   | 100.0%<br>15 / 15          | 78.2, 100.0                   |
| None specified                    | 3     | 100.0%<br>(1 / 1)          | 2.5, 100.0                    | 100.0%<br>(2 / 2)          | 15.8, 100.0                   |

\*Includes subjects with "unable to determine" status.

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<sup>6</sup>Does not include 377 additional HIV positive subjects enrolled but included in higher ranked risk categories, and 6 HIV positive subjects enrolled with signs or symptoms of hepatitis.

**RESULTS OF SUPPLEMENTAL TESTING OF SPECIMENS REACTIVE IN THE ORAQUICK<sup>®</sup> HCV RAPID ANTIBODY TEST**  
 The table below shows the results obtained when subjects reactive in the OraQuick<sup>®</sup> HCV Rapid Antibody Test were tested by recombinant immunoblot assay (RIBA<sup>®</sup>).

| Number of OraQuick Reactive Results | RIBA <sup>®</sup> Results |               |          |
|-------------------------------------|---------------------------|---------------|----------|
|                                     | Positive                  | Indeterminate | Negative |
| 443                                 | 418                       | 25*           | 0        |

\*Seventeen (17) of the RIBA<sup>®</sup> indeterminate results were positive for HCV RNA when tested by PCR.

Of the subjects reactive in the OraQuick<sup>®</sup> HCV Rapid Antibody Test 94.4% (418/443) were positive by RIBA<sup>®</sup>. Seventeen (17) of the RIBA<sup>®</sup> indeterminate results were positive for HCV RNA when tested by PCR; therefore, 98.2% (435/443) were positive by either RIBA<sup>®</sup> or PCR.

**REACTIVITY WITH HCV SEROCONVERSION PANELS**

Eighteen panels containing sequential plasma specimens from individuals undergoing seroconversion as a result of HCV infection were evaluated with the OraQuick<sup>®</sup> HCV Rapid Antibody Test and compared with an FDA approved anti-HCV EIA test. The OraQuick<sup>®</sup> HCV Rapid Antibody Test and the reference anti-HCV assay results are summarized in the following table. The sensitivity of the OraQuick<sup>®</sup> HCV Rapid Antibody Test to detect seroconversion was similar to that of the comparator EIA. The OraQuick<sup>®</sup> HCV Rapid Antibody Test detected anti-HCV antibodies earlier than EIA in 9 of the 18 seroconversion panels (50%) and by an overall average of 3.6 days (95% CI = 1.2 to 5.9).

| Seroconversion Panel | Days to Evidence of HCV Infection             |                |                           |                | Difference (OraQuick <sup>®</sup> - EIA) |
|----------------------|---|----------------|---------------------------|----------------|--|
|                      | OraQuick <sup>®</sup> HCV Rapid Antibody Test |                | FDA approved anti-HCV EIA |                |  |
|                      | Last Non-Reactive                             | First Reactive | Last Non-Reactive         | First Reactive |  |
| HCV 6213             | 35  | 37             | 30                        | 37             | 0  |
| HCV 6214             | 18  | 23             | 23                        | 25             | -2                                       |
| HCV 6227             | 46  | 74             | 46                        | 74             | 0  |
| HCV 9041             | 31  | 62             | 31                        | 62             | 0  |
| HCV 9046             | 0   | 69             | 0                         | 69             | 0  |
| HCV 9047             | 21  | 28             | 21                        | 28             | 0  |
| PHV 901              | 65  | 97             | 65                        | 97             | 0  |
| PHV 905              | 7   | 11             | 11                        | 18             | -7                                       |
| PHV 907              | 7   | 13             | 13                        | 18             | -5                                       |
| PHV 910 (M)          | 0   | 4              | 0                         | 4              | 0  |
| PHV 911 (M)          | 0   | 11             | 0                         | 11             | 0  |
| PHV 914              | 9   | 12             | 19                        | 24             | -12                                      |
| PHV 916 (M)          | 7   | 9              | 9                         | 23             | -14                                      |
| PHV 917 (M)          | 22  | 85             | 22                        | 85             | 0  |
| PHV 920              | 7   | 13             | 7                         | 16             | -3                                       |
| PHV 921              | 0   | 4              | 4                         | 14             | -10                                      |
| RP 006               | 388   | 461            | 461                       | 469            | -8                                       |
| RP 038               | 47  | 52             | 52                        | 55             | -3                                       |
| <b>Average</b>       |   | <b>59.2</b>    |                           | <b>62.7</b>    | <b>- 3.6 (-5.9 to -1.2)</b>              |

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**REACTIVITY WITH HCV SPECIMENS FROM VARIOUS GENOTYPES AND SUBTYPES**

The ability of the OraQuick® HCV Rapid Antibody Test to detect infection derived from various genotypes and subtypes was assessed using two commercially available Worldwide HCV Performance panels. Thirty-two HCV-positive plasma specimens derived from multiple geographies, representing four genotypes (1, 2, 3, 4) were tested. All specimens were reactive with the OraQuick® HCV Rapid Antibody Test. Three HCV-negative samples were included in the panel and all were non-reactive with the OraQuick® HCV Rapid Antibody Test.

**INTERFERING SUBSTANCES**

The OraQuick® HCV Rapid Antibody Test was evaluated with the following interfering substances present in whole blood, samples in order to assess their potential effect on the assay performance as per CLSI guidelines EP7-A2. Testing was completed on ten HCV-negative whole blood, samples and ten HCV-positive spiked matched whole blood samples. All matched samples were spiked according to one of the following conditions as per the table below:

| Interfering Substances | Concentration |
|------------------------|---------------|
| Bilirubin              | 10 mg/dL      |
| Hemoglobin             | 500 mg/dL     |
| Lipid (Triolein)       | 3500 mg/dL    |
| Protein                | 12 g/dL       |

None of these interfering substances had any impact on the OraQuick® HCV Rapid Antibody Test assay performance at the concentrations evaluated.

**MEDICAL CONDITIONS UNRELATED TO HCV INFECTION**

The performance of the OraQuick® HCV Rapid Antibody Test was evaluated with commercially available HCV negative plasma and serum specimens derived from medical conditions unrelated to HCV infection. Results are summarized in the table below.

| Medical Condition                      |            | Non-Reactive (%) | Reactive (%) |
|--|------------|------------------|--------------|
| <b>Autoimmune Diseases</b>             |            |                  |              |
| Myasthenia Gravis                      | 4          | 4 (100)          | 0 (0)        |
| Rheumatoid Arthritis                   | 10         | 10 (100)         | 0 (0)        |
| Systemic Lupus Erythematosus (SLE)     | 10         | 10 (100)         | 0 (0)        |
| <b>Other Medical Conditions</b>        |            |                  |              |
| Influenza Vaccination                  | 10         | 10 (100)         | 0 (0)        |
| Hepatitis A Virus (HAV)                | 59         | 59 (100)         | 0 (0)        |
| Hepatitis B Virus (HBV)                | 58         | 58 (100)         | 0 (0)        |
| Hepatitis D Virus (HDV)                | 2          | 2 (100)          | 0 (0)        |
| Hepatitis E Virus (HEV)                | 8          | 8 (100)          | 0 (0)        |
| Epstein-Barr Virus (EBV)               | 10         | 10 (100)         | 0 (0)        |
| Cytomegalovirus (CMV)                  | 10         | 10 (100)         | 0 (0)        |
| Herpes Simplex Virus (HSV)             | 10         | 10 (100)         | 0 (0)        |
| Parvovirus B19                         | 10         | 10 (100)         | 0 (0)        |
| Rubella                                | 10         | 10 (100)         | 0 (0)        |
| Syphilis                               | 10         | 10 (100)         | 0 (0)        |
| Toxoplasmosis                          | 10         | 10 (100)         | 0 (0)        |
| Human Immunodeficiency Virus (HIV-1/2) | 154        | 154 (100)        | 0 (0)        |
| Heterophilic antibodies                | 10         | 10 (100)         | 0 (0)        |
| Multiparous Female                     | 10         | 10 (100)         | 0 (0)        |
| <b>Total Samples Tested</b>            | <b>405</b> | <b>405</b>       | <b>0</b>     |

None of the medical conditions tested produced false positive results in the OraQuick® HCV Rapid Antibody Test. Performance characteristics in scleroderma, Sjögren's Syndrome and Human T-Cell Lymphotropic Virus (HTLV I/II) have not been established.

**SAMPLE STABILITY**

The OraQuick® HCV Rapid Antibody Test was evaluated with whole blood stored at various storage conditions over numerous days. Results are summarized in the table below.

| Specimen Type | Days at Storage Condition |                     |
|---------------|---------------------------|---------------------|
|               | 2°-8°C (36°-46°F)         | 15°-30°C (59°-86°F) |
| Whole Blood   | 7                         | 3                   |

Storing whole blood for up to 7 days refrigerated or 3 days incubated at 15°-30°C (59°-86°F) did not impact the performance of the OraQuick® HCV Rapid Antibody Test.

**SPECIMEN TYPES**

The OraQuick® HCV Rapid Antibody Test was evaluated with whole blood samples collected in various types of anticoagulants including Ethylenediaminetetracetic Acid (EDTA), sodium heparin, lithium heparin, and sodium citrate. Testing was performed with twenty anti-HCV negative whole blood samples and twenty anti-HCV-spiked positive whole blood samples. All samples produced acceptable assay performance. The recommended anticoagulant types for use with the OraQuick® HCV Rapid Antibody Test in whole blood are vacutainers containing EDTA, sodium heparin, lithium heparin, sodium citrate.

**LIMIT OF DETECTION**

The limit of detection (LoD), defined as the EIA signal to cutoff ratio which yielded reactive results 95% of the time in the OraQuick® Rapid HCV Antibody Test device, was calculated for each of three (3) lots separately and for three (3) lots combined. The LoD was calculated to be 0.75 s/co using an FDA approved EIA.

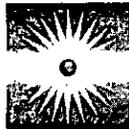
**REPRODUCIBILITY**

The reproducibility of the OraQuick® HCV Rapid Antibody Test was tested at 3 sites using 3 lots of test devices twice a day for 5 days with 9 operators (3 per site). Three whole blood panel member types (negative, limit of detection (LoD), and low positive) were tested in 5 unique test kit types. Each test kit consisted of eight (8) blinded panel members that had various combinations of the 3 panel members in a randomized sequence. Panel members were blinded per operator, run, and device lot to ensure that the results of the panel member types were unpredictable to the operator. LoD specimen was determined to be a 0.75 s/co by an FDA approved EIA. Overall concordance across operators, sites, and device lots was 98.9% for the negative specimen, 98.7% for the specimen at the limit of detection and 99.7% for the low positive specimen.

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