



ichroma™ CRP

INTENDED USE

ichroma™ CRP is a fluorescence Immunoassay (FIA) for the quantitative determination of CRP in human whole blood/serum/plasma. It is useful as an aid in management and monitoring of autoimmune diseases and infectious processes, such as rheumatoid arthritis.

For in vitro diagnostic use only.

INTRODUCTION

C-reactive protein (CRP) is a protein found in blood; the level of which rises in response to inflammation. CRP is first acute-phase protein to be described and is an exquisitely sensitive systemic marker of inflammation and tissue damage. The serum CRP level may rise from a normal level of < 5 mg/L to 500 mg/L during the body's general, nonspecific response to infections and other acute inflammatory conditions. Measurement of CRP concentration has been widely used as a clinical tool for monitoring the status of inflammation, effectiveness of treatment of various infections and autoimmune diseases such as rheumatoid arthritis. And high sensitive CRP (hsCRP) has been suggested that along with serum cholesterol, may be marker of diagnosis in cardiovascular diseases (CVD). And hsCRP is emerging as the strongest and most independent predictive risk factor for CVD.

PRINCIPLE

The test uses a sandwich immunodetection method.

The detector antibodies in buffer bind to antigens in the sample, forming antigen-antibody complexes, and migrate onto nitrocellulose matrix to be captured by the other immobilized-antibodies on a test strip.

More antigens in the sample will form more antigen-antibody complexes which lead to stronger fluorescence signal by detector antibodies, which is processed by the instrument for ichroma™ tests to show CRP concentration in the sample.

COMPONENTS

ichroma™ CRP consists of 'cartridges' and 'detection buffers'.

- The cartridge contains the membrane called a test strip which has anti human CRP at the test line, human C-reactive protein at the antigen line and streptavidin at the control line. All cartridges are individually sealed in an aluminum foil pouch containing a desiccant, and they are further packaged in a cartridge box.
- The detection buffer contains anti-human CRP fluorescence conjugate, anti-human CRP, BSA-biotin-fluorescence conjugate and bovine serum albumin (BSA) as a stabilizer and sodium azide as a preservative in phosphate buffered saline (PBS). It is pre-dispensed in a tube. The detection buffer tubes are packaged in a detection buffer box and further packed in a styrofoam box with ice-pack for the shipment.

WARNINGS AND PRECAUTIONS

- *For in vitro* diagnostic use only.
 - Follow instructions and procedures described in this 'Instructions for use'.
 - Use only fresh samples and avoid direct sunlight.
 - Lot numbers of all the test components (cartridge, detection buffer and ID chip) must match each other.
 - Do not interchange the test components between different lots or use the test components after the expiration date, either of which might yield incorrect test result(s).
 - Do not reuse cartridges or detection buffer tubes. A cartridge should be used for testing one sample only. A detection buffer tube should be used for processing of one sample only.
 - The cartridge should remain sealed in its original pouch until just before use. Do not use the cartridge, if pouch is damaged or has already been opened.
 - Frozen sample should be thawed only once. For shipping, samples must be packed in accordance with local regulations. Sample with severe hemolysis and/or hyperlipidemia must not be used.
 - If test components and/or sample are stored in refrigerator, then allow cartridge, detection buffer and sample to be at room temperature for approximately 30 minutes before use.
 - The instrument for ichroma™ tests may generate slight vibration during use.
 - Used cartridges, detection buffers and sample collectors should be handled carefully and discarded by an appropriate method in accordance with relevant local regulations.
 - The detection buffer contains sodium azide (NaN₃), and they may cause certain health issues like convulsions, low blood pressure and heart rate, loss of consciousness, lung injury and respiratory failure. Avoid contact with skin, eyes, and clothing. In case of contact, rinse immediately with running water.
 - Biotin interference was not observed in **ichroma™ CRP** when biotin concentration in the sample was up to 3,500 ng/mL. In reference, if biotin is taken at a level of 300 mg/day, the biotin concentration in the body is about 1,200 ng/mL. However, if a patient has been taking biotin at dosage of more than 300 mg a day, it is recommended to collect blood again 24 hours after discontinuation of biotin intake.
 - **ichroma™ CRP** will provide accurate and reliable results subject to the below conditions.
 - **ichroma™ CRP** should be used only in conjunction with the instrument for ichroma™ tests.
 - Have to use recommended anticoagulant.
- | Recommended anticoagulant |
|--|
| K ₂ EDTA, K ₃ EDTA,
Sodium heparin, Lithium heparin, Sodium citrate |
- **The sample collector should be used when the following conditions are met.**
 - The sample collector provided with the kit is recommended to obtain correct test result.
 - Whole blood should be immediately tested after collection.
 - Excess the sample around the sample collector should be wiped off.

- In order to avoid cross-contamination, please do not re-use the sample collector for multiple samples.

LIMITATION OF THE TEST SYSTEM

- The test may yield false positive result(s) due to the cross-reactions and/or non-specific adhesion of certain sample components to the capture/detector antibodies.
- The test may yield false negative result(s) due to the non-responsiveness of the antigen to the antibodies which is the most common if the epitope is masked by some unknown components, so therefore not being able to be detected or captured by the antibodies. The instability or degradation of the antigen with time and/or temperature may also cause false negative result as it makes antigen unrecognizable by the antibodies.
- Other factors may interfere with the test and cause erroneous results, such as technical/procedural errors, degradation of the test components/reagents or presence of interfering substances in the test samples.
- Any clinical diagnosis based on the test result must be supported by a comprehensive judgment of the concerned physician in conjunction with clinical symptoms and other relevant test results.

STORAGE AND STABILITY

Storage condition			
Component	Storage Temperature	Shelf life	Note
Cartridge	2 - 30 °C	20 months	Disposable
Detection buffer	2 - 8 °C	20 months	Disposable

- After the cartridge pouch is opened, the test should be performed immediately.

MATERIALS SUPPLIED

REF I-CHROMA CRP-25

Components of **ichroma™ CRP**

- Cartridge box:
 - Cartridge 25
 - Sample collector 25
 - ID chip 1
 - Instructions for use 1
- Detection buffer box:
 - Detection buffer 25

MATERIALS REQUIRED BUT SUPPLIED ON DEMAND

Following items can be purchased separately from **ichroma™ CRP**.

Please contact our sales division for more information.

- Instrument for **ichroma™** tests
 - **ichroma™ Reader** **REF** FR203
 - **ichroma™ II** **REF** FPRR021
 - **ichroma™ III** **REF** FPRR037
 - **ichroma™ M3** **REF** FPRR035
 - **ichroma™-50** **REF** FPRR022
 - **ichroma™-50 Plus** **REF** FPRR036
- Printer **REF** FPRR007
- **Boditech CRP Control** **REF** CFPO-100

SAMPLE COLLECTION AND PROCESSING

The sample type for **ichroma™ CRP** is human whole blood/serum/plasma.

- It is recommended to test the sample within 24 hours after collection.
- The sample (serum, plasma) should be separated from the clot by centrifugation within 3 hours after the collection of whole blood.
- The samples (whole blood, serum, plasma) may be stored for a week at 2-8 °C prior to being tested. If testing will be delayed more than a week, samples (serum, plasma) should be frozen at below -20 °C.
- The samples (serum, plasma) stored frozen below -20 °C for 3 months showed no performance difference.
- However, the whole blood sample should not be kept in a freezer in any case.
- As a repeated freeze-thaw cycle may affect the test result, do not refreeze previously frozen samples.
- Whole blood sample may be used to collect according to below:
 - ① Wear disposable gloves and protective equipment for safety.
 - ② Open the zipper bag which has sample collectors.
 - ③ Take out the sample collector and check for damage or contamination.
 - ④ Touch the surface of blood with the sample collector.
 - ⑤ Fill it with blood completely.
 (Make sure that no air bubbles are present in the sample collector. Do not get blood on the surface of the sample collector. If the blood gets on the surface of the sample collector, remove it gently with gauze.)

TEST SETUP

- Check the contents of **ichroma™ CRP**: Sealed cartridges, detection buffers, sample collectors, an ID chip and an instructions for use.
- Ensure that the lot number of the cartridges matches that of the detection buffers as well as an ID chip.
- If the sealed cartridge and the detection buffer have been stored in a refrigerator, place them on a clean and flat surface at room temperature for at least 30 minutes before testing.
- Turn on the instrument for **ichroma™** tests.
- Insert the ID chip into the 'ID chip port'.

※ Please refer to the instrument for **ichroma™** tests operation manual for complete information and operating instructions.

TEST PROCEDURE

► **ichroma™ Reader, ichroma™ II, ichroma™ M3**

Multi test mode

- 1) Make a puncture on the top of a detection buffer by inserting an empty sample collector.
- 2) Take 10 µL of sample (whole blood/serum/plasma/control) using a sample collector.
- 3) Assemble the sample collector and the detection buffer into one.
- 4) Shake 10 times or more until the sample out of the

sample collector by inversion. The mixture of the buffer and the sample has to be used within 30 seconds.

- 5) Remove the cap off the top of assembled tube. Discard two drops of reagent onto the paper towel before applying to a cartridge.
- 6) Apply only two drops of the mixture into the sample well of the cartridge.
- 7) Leave the cartridge at room temperature for 3 minutes before inserting the cartridge into the holder.
△ Scan the sample-loaded cartridge immediately when the incubation time is over. If not, it will cause inaccurate test result.
- 8) To scan the sample-loaded cartridge, insert it into the cartridge holder of the instrument for ichroma™ tests. Ensure proper orientation of the cartridge before pushing it all the way inside the cartridge holder. An arrow is marked on the cartridge especially for this purpose.
- 9) Press the 'Select' or tap the 'Start' button on the instrument for ichroma™ tests to start the scanning process.
 (ichroma™ M3 is tested automatically after inserting.)
- 10) The instrument for ichroma™ tests will start scanning the sample-loaded cartridge immediately.
- 11) Read the test result on the display screen of the instrument for ichroma™ tests.

Single test mode

- 1) The test procedure is same with "Multi test mode 1) – 6)".
- 2) Insert the sample-loaded cartridge into the holder of the instrument for ichroma™ tests. Ensure proper orientation of the cartridge before pushing it all the way inside the cartridge holder. An arrow is marked on the cartridge especially for this purpose.
- 3) Press the 'Select' or tap the 'Start' button on the instrument for ichroma™ tests.
 (ichroma™ M3 is tested automatically after inserting.)
- 4) The cartridge goes inside the instrument for ichroma™ tests and will automatically start scanning the sample-loaded cartridge after 3 minutes.
- 5) Read the test result on the display screen of the instrument for ichroma™ tests.

▶ ichroma™ III

- 1) The test procedure is same with the 'Single test mode'.

▶ ichroma™-50, ichroma™-50 Plus

- 1) Insert the tip array in the tip station.
- 2) Insert the detection buffer tubes in the reagent station and cover the reagent station to hold the detection buffer tubes in place.
- 3) Insert the cartridge magazine with the cartridges into the magazine station.
- 4) Insert the sample tube into the blood collection tube rack and load the blood collection tube rack into the sampling station (loading part).
- 5) Tap the button located in the upper side of the No. of test cartridge region to select the ID chip that you want to use.
- 6) When the selected cartridge slot is activated, set the number of the detection buffer tube by tapping.
- 7) Tap the number of pipette tips by tapping.

- 8) Tap the 'Start' button on the left upper of the main screen to start test.

INTERPRETATION OF TEST RESULT

- The instrument for ichroma™ tests calculates the test result automatically and displays CRP concentration of the test sample in terms of mg/L.
- Cut-off value: 10 mg/L
- Working range: 2.5-300 mg/L
- Effect of Hematocrit
 The CRP whole blood of the instrument for ichroma™ tests is calibrated to read the CRP serum concentration of a blood sample with a hematocrit of 40%. If the actual hematocrit value deviates from 40%, the result should be corrected by multiplying with the respective factor in the table: deviates from 40%, the result should be corrected by multiplying with the respective factor in the table:

Hct (%)	Factor	Hct (%)	Factor
20-29	0.8	56-58	1.4
30-36	0.9	59-61	1.5
37-42	1.0	62-63	1.6
43-47	1.1	64-65	1.7
48-51	1.2	66-67	1.8
52-55	1.3	68-69	1.9

Reference range, HCT:

- Woman: 35-44%
- Men: 39-48%

QUALITY CONTROL

- Quality control tests are a part of the good testing practice to confirm the expected results and validity of the assay and should be performed at regular intervals.
- Quality control tests should also be performed whenever there is any question concerning the validity of the test results.
- Control materials are provided on demand with **ichroma™ CRP**. For more information regarding obtaining the control materials, contact Boditech Med Inc.'s Sales Division for assistance.
 (Please refer to the instructions for use of control material.)

PERFORMANCE CHARACTERISTICS

- **Analytical sensitivity**
 - Limit of Blank (LoB) 0.22 mg/L
 - Limit of Detection (LoD) 0.37 mg/L
 - Limit of Quantitation (LOQ) 0.50 mg/L
- **Analytical Specificity**
 - Interference
 Interferents such as below the ones in the table were added to the test sample(s) the same as the below concentrations. **ichroma™ CRP** test results did not show any significant interference with these materials.

Interferent	Conc.
Ascorbic acid	350 µmol/L
Bilirubin (conjugated)	475 µmol/L
Albumin	60 g/L
Glucose	1,000 mg/dL

Triglyceride mixture (Triglycerides, total)	1,500 mg/dL
Hemoglobin	10 g/L
Biotin	3,500 ng/ml
K2-EDTA	5.45mg/mL
K3-EDTA	5.45mg/mL
Na-Heparin	10.41mg/mL
Li-Heparin	10.44mg/mL
Sodium Citrate	32 mg/mL

- Cross-reactivity

Biomolecules such as below the ones in the table were added to the test sample(s) at concentrations much higher than their normal physiological levels in the blood. **ichroma™ CRP** test results did not show any significant cross-reactivity with these biomolecules.

Cross reactive substance	Conc.
IL-6	50 µg/mL
PCT	50 µg/mL
Serum amyloid P	500 µg/mL
Serum amyloid A	500 µg/mL
ferritin	500 µg/mL

■ Precision

- Single-site study

Three lots of **ichroma™ CRP** was tested for 20 days at one site by one person. Each standard material was tested 2 runs per day. For each test, each standard material was tested in duplicate. (20 x 2 x 2)

Standard material	Repeatability				Within-laboratory precision			
	N	Mean [mg/L]	SD	CV (%)	N	Mean [mg/L]	SD	CV (%)
1	40	5.10	0.29	5.8	80	5.07	0.30	5.9
2	40	10.04	0.53	5.2	80	10.07	0.53	5.3
3	40	151.08	9.46	6.3	80	152.64	11.05	7.2
Standard material	Lot to Lot precision							
	N	Mean [mg/L]	SD	CV (%)				
1	240	5.05	0.32	6.4				
2	240	10.06	0.52	5.2				
3	240	153.75	8.55	5.6				

- Multi-site study

One lot of **ichroma™ CRP** was tested for 5 days at three sites (each site by one tester person, one analyzer). Each standard material was tested in 5 replicates per day. (3 x 5 x 5 x 1)

Standard material	Reproducibility			
	N	Mean [mg/L]	SD	CV (%)
1	75	5.03	0.18	3.56
2	75	9.75	0.37	3.80
3	75	151.21	5.32	3.52

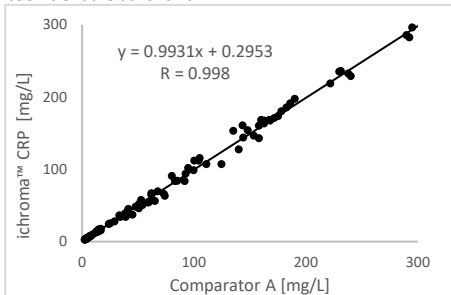
■ Accuracy

The accuracy was confirmed by testing 3 different lots, three times at each concentration of the international standard (ERM-DA474 IFCC HUMAN SERUM).

	LOT 1		LOT 2		LOT 3	
	Result [mg/L]	Bias	Result [mg/L]	Bias	Result [mg/L]	Bias
Run 1	42.16	2.3%	40.88	-0.8%	41.68	1.2%
Run 2	41.09	-0.3%	42.33	2.7%	41.35	0.4%
Run 3	41.53	0.8%	41.15	-0.1%	40.09	-2.7%

■ Comparability













CRP concentrations of 100 clinical samples were quantified independently with **ichroma™ CRP (ichroma™ II)** and **comparator A** as per prescribed test procedures. Test results were compared, and their comparability was investigated with linear regression and correlation coefficient (R). The regression equation and correlation coefficient are as follows.



REFERENCES


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Note: Please refer to the table below to identify various symbols.

	Sufficient for <n> tests
	Read instruction for use
	Use by Date
	Batch code
	Catalog number
	Caution
	Manufacturer
	Authorized representative of the European Community
	In vitro diagnostic medical device
	Temperature limit
	Do not reuse
	This product fulfills the requirements of the Directive 98/79/EC on in vitro diagnostic medical devices

For technical assistance, please contact:
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