

English

**ACTALYKE®**

## Clotting Time Test System

**REF** Cat. No. C-ACT, K-ACT, G-ACT



### INTENDED USE

Actalyke Activated Clotting Time Test Tubes are intended for use in the performance of the activated clotting time test, a whole blood coagulation assay commonly used to monitor heparin anticoagulation during various medical and surgical procedures. Actalyke ACT test tubes can be used in conjunction with the Actalyke, the Actalyke MINI, and the Actalyke XL instruments. Actalyke ACT test tubes can also be used in conjunction with Hemochron® instruments.

### SUMMARY

Actalyke ACT test tubes employ clot-promoting reagents (celite, kaolin or glass beads) to standardize and quicken fibrin formation induced by contact activation of the coagulation protein Factor XII. ACT test results reflect the ability of a blood sample to clot in this fashion, and they are prolonged in the presence of heparin. The more prolonged the ACT result is from baseline or normal values, the greater the degree of anticoagulation. As a result, the ACT test is widely used to assess an individual patient's response to a given heparin dose. Especially when moderate to high levels of heparin are administered and during extra-corporeal circulation, the ACT is a practical, accurate method for controlling anticoagulation therapy.

### PRINCIPLE

The ACT test was first described by Paul Hattersley, M.D., in 1966. Although the test was originally a manual method, automated ACT instruments were later introduced. These improved the accuracy, reproducibility and convenience of ACT test results. Automated test systems like Actalyke involve pre-loaded, disposable test tubes to which a blood sample is added; the test tube is then inserted into the instrument where the tube is rotated and warmed to 37°C (± 0.5°C) until a fibrin clot is mechanically detected. Upon clot detection the test terminates, a beeper is sounded and the instrument displays the ACT result in seconds.

Each Actalyke test tube has a barcode label affixed to it, which may be read by the Actalyke instrument to determine the test tube type (i.e. celite, kaolin or glass bead activated). [Barcode reader not available on Actalyke MINI models]. The lot number and expiration date of each tube are also identified on the barcode label.

### REAGENTS

**WARNING: FOR IN VITRO DIAGNOSTIC USE. CAUTION:** Some data suggests that the micro crystalline silica in C-ACT is a possible carcinogen. Avoid contact.

Each box contains 50 Actalyke test tubes:

| Cat. No. | Use   | Activator           | Description   |
|----------|---|---------------------|---|
| C-ACT    | Bypass and vascular surgery and PTCA        | Celite (14 mg)      | Glass tube w/ black flip-top flip-top and barcode label. 2.0 cc blood sample. |
| K-ACT    | Bypass surgery in the presence of aprotinin | Kaolin (14 mg)      | Glass tube w/ yellow flip-in and barcode label. 2.0 cc blood sample.          |
| G-ACT    | Renal Dialysis, ECMO                        | Glass (50 mg) beads | Plastic tube w/ clear flip-top and barcode label. 0.4 cc blood sample.        |

*Note: Used Actalyke test tubes should be disposed of in accordance with each institution's policy regarding biohazardous material.*

**Recommended Use:** Extracorporeal circulation, angioplasty, thoracic and vascular surgery (in the presence or absence of antifibrinolytics).

**Storage and Stability:** Actalyke ACT test tubes can be stored at room temperature (15-30°C). They must not be used past the expiration date that appears on the barcode label affixed to each test tube and its corresponding box.

### INSTRUMENTS

The Actalyke test tubes should be used with the Actalyke, Actalyke MINI, Actalyke XL or Hemochron® instruments. Refer to the appropriate Operator's Manual's for detailed instructions.

## SPECIMEN COLLECTION AND HANDLING

### C-ACT and K-ACT:

The specimen collection and test procedure for both the C-ACT and K-ACT test tubes are exactly the same. Use a 5.0 cc syringe to obtain the blood specimen as follows:

**Extracorporeal line:** Using a two-syringe technique, flush the extracorporeal blood access line by withdrawing 5.0 cc of blood into a syringe and discarding it. Then use a second syringe to obtain a 3.0 cc sample for testing.

**In-dwelling line:** Discontinue fluids drip, if necessary. Using a two-syringe technique, withdraw 5.0 cc of blood into a syringe and discard it. Then use a second syringe to obtain a 3.0 cc sample for testing.

**Venipuncture:** Using a two-syringe technique, withdraw 2.0 cc of blood into a syringe and discard it. Then use a second syringe to obtain a 3.0 cc sample for testing.

### G-ACT:

Use a 1.0 cc tuberculin syringe to obtain the blood specimen as follows:

**Extracorporeal line:** Do not take a blood sample from a line through which heparin has been administered. Using a two-syringe technique, flush the blood access line by withdrawing 3.0 cc of blood into a syringe and discarding it. Then use a 1.0 cc syringe to obtain a 0.4 cc sample for testing.

## PROCEDURE

### Materials Provided:

|                            | Cat. No. |
|----------------------------|----------|
| Celite Test Tubes (50/box) | C-ACT    |
| Kaolin Test Tubes (50/box) | K-ACT    |
| Glass Bead Tubes (50/box)  | G-ACT    |

### Materials Required:

|                                     |        |
|-------------------------------------|--------|
| Actalyke XL Analyzer                | 5770   |
| Actalyke XL Barcode Reader & Cable  | 5758   |
| Actalyke MINI Analyzer              | 5752   |
| Actalyke MINI Analyzer with printer | 5750   |
| Actalyke Clotting Time Analyzer     |        |
| Single Well                         | A1     |
| Single Well with printer            | A1P    |
| Dual Well                           | A2     |
| Dual Well with printer              | A2P    |
| Actalyke Thermometer                | 5757   |
| Actalyke QC Kit (Level I & II)      | AQC-LP |
| Actalyke QC Kit (Level I & III)     | AQC-HP |
| Electronic Clotting Tube            | XL-ECT |
| Syringes for sample collection      |        |

## STEP-BY-STEP METHOD

**Note: Refer to the appropriate Operator's Manual for detailed instrument instructions. Refer to the SAMPLE COLLECTION section for detailed collection instructions.**

1. Open the flip top of the selected Actalyke test tube.
2. Dispense the following amount of blood into the appropriate tube or fill to line indicated on the tube.

| Tube  | Blood Volume |
|-------|--------------|
| C-ACT | 2.0 cc       |
| K-ACT | 2.0 cc       |
| G-ACT | 0.4 cc       |

3. Perform the next steps quickly and in order depending on the type of instrument settings used.

◇ Press the Start button at the same time that the blood is added to the tube.

◇ Close the flip-top. Mix the blood in the tube according to the appropriate instructions.

| Tube         | Mixing   |
|--------------|--|
| C-ACT/ K-ACT | Shake the tube vigorously from end-to-end ten times to disperse the activator. Tap the bottom of the tube lightly on the table top to ensure that the magnet moves freely. |
| G-ACT        | While holding the tube in a vertical position, tap the bottom sideways five times to ensure proper mixing.   |

**NOTE:** This mixing procedure eliminates protein loss to the tube walls and assists in standardizing the mixing procedure from operator to operator.

- ◇ Insert the tube into the instrument test well.
  - ◇ Rotate the tube 4-5 times; the green detector light will illuminate or "Tube In" will display.
4. Upon clot detection, the buzzer will sound and the ACT test result will be displayed in seconds. (Results can be printed if using Actalyke Models A1P, A2P, Actalyke XL, or Actalyke MINI with printer.)

### Quality Control

Routine quality control testing and tracking should be part of a comprehensive quality assurance program. Actalyke Quality Control products are available to make routine QC convenient and affordable. In the US, according to the CLIA '88 regulations, the frequency of quality control testing is mandated. Since the ACT is categorized as moderately complex, ACT users must (1) perform two levels of quality control during each shift in which the test system is used clinically, and (2) subscribe to a Proficiency Testing program. Additionally, CLIA (42 CFR 493) requires that biological controls be used weekly to verify system function. To help users comply with these regulations, Actalyke Quality Control Material is available (or other commercial coagulation products can also be used), and ACT Proficiency Testing Programs are available through the American Proficiency Institute. Contact Helena's Technical Services Department for details.

Each box of Actalyke Activated Clotting Time tubes contains 50 unitized reaction tubes from a single manufactured lot and should be validated once, initially upon arrival. This can be accomplished using the appropriate Actalyke QC Kit. Acceptable performance ranges for the various Actalyke coagulation assays are included in each kit. After a box from each lot number of Actalyke Activated Clotting Time tubes has been verified with the QC kit, that lot number of tubes should be marked as "VERIFIED", along with the date of verification and the initials of the operator. This box is now "IN CONTROL" and will not require any further investigation until a new box of tubes is received, unless the tubes are not stored to manufacturers recommendations or unless a shift in clinical results is experienced. Since Actalyke test tubes contain non-biological, inert reagents and are not susceptible to environmental or temperature changes, Actalyke ACT test tubes have exceptional room temperature stability and longevity.

Quality control results that fall outside the established expected values should be repeated. If the problem persists, the source should be investigated and corrected prior to continued use of the test system.

### Operating Cautions and Limitations

1. Do not use Actalyke ACT test tubes that are past the expiration date marked on the tube barcode label and the corresponding test tube box.
2. All guidelines pertaining to the handling of fresh whole human blood should be adhered to when handling Actalyke test tubes and instruments.
3. Specimen contamination and inappropriate technique can affect ACT results.
4. The ACT results may be affected by hemodilution, hypothermia, pharmacologic compounds, and various coagulopathies. Test results should be interpreted with respect to the patient's condition and the clinical circumstances. Those results which do not agree with expected values should be repeated and further evaluated by other diagnostic methods, if indicated.
5. When using the ACT test as a monitor for heparin anticoagulation effect, a baseline (non-heparinized) sample may be run so that a differential ratio can be used as a comparison.

### REFERENCE RANGES

Actalyke ACT test tubes were run on Actalyke instruments (Model A2P, MINI and XL) and Hemochron Instrument (Model 8000). Quality control tests were performed on each instrument prior to testing of Actalyke tubes for this study. Each tube type was run with normal volunteers on each instrument. The results are as follows:

| Test  | Instrument    | N  | Mean | 2SD | Reference Range |
|-------|---------------|----|------|-----|-----------------|
| C-ACT | Actalyke      | 30 | 123  | 26  | 97-149 sec.     |
|       | Actalyke MINI | 30 | 123  | 29  | 94-152 sec.     |
|       | Actalyke XL   | 66 | 126  | 21  | 105-148 sec.    |
|       | Hemochron     | 15 | 126  | 15  | 111-141 sec.    |

|       |               |    |     |    |              |
|-------|---------------|----|-----|----|--------------|
| K-ACT | Actalyke      | 35 | 123 | 28 | 93-150 sec.  |
|       | Actalyke MINI | 35 | 125 | 25 | 100-150 sec. |
|       | Actalyke XL   | 64 | 132 | 21 | 102-153 sec. |
| G-ACT | Hemochron     | 15 | 129 | 23 | 106-152 sec. |
|       | Actalyke      | 44 | 185 | 38 | 147-223 sec. |
|       | Actalyke MINI | 44 | 181 | 34 | 147-215 sec. |
|       | Actalyke XL   | 63 | 189 | 42 | 147-233 sec. |
|       | Hemochron     | 15 | 167 | 20 | 147-187 sec. |

## PERFORMANCE CHARACTERISTICS

### Precision Studies

All precision studies were done according to NCCLS EP-5 guidelines.

#### Actalyke MINI

The precision of the Actalyke Activated Clotting Time Test System was evaluated by performing multiple replicates on twenty separate days with heparinized and non-heparinized Actalyke Whole Blood Control samples. C-ACT and K-ACT tests were run with levels I and III; G-ACT with levels I and II. The coefficient of variation for each test type was less than 10%.

| Level | C-ACT |       | K-ACT |       | G-ACT |       |
|-------|-------|-------|-------|-------|-------|-------|
|       | I     | III   | I     | III   | I     | II    |
| mean  | 127.0 | 388.2 | 125.1 | 389.9 | 146.8 | 286.3 |
| sd    | 4.60  | 18.44 | 4.97  | 16.40 | 6.76  | 16.37 |
| % cv  | 3.6   | 4.8   | 4.0   | 4.2   | 4.6   | 5.7   |

#### Actalyke XL

The precision of the Actalyke Activated Clotting Time Test System was evaluated by performing multiple replicates on twenty separate days with heparinized and non-heparinized Actalyke QC samples. C-ACT and K-ACT tests were run with levels I and III; G-ACT with levels I and II. The coefficient of variation for each test type was less than 10%.

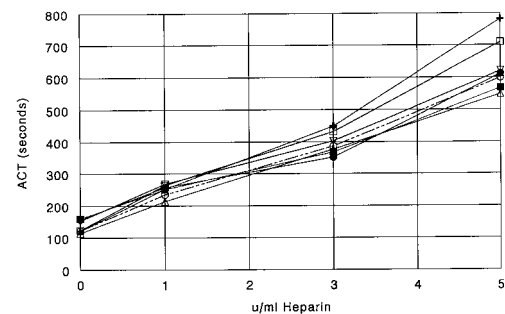
| Level | C-ACT |       | K-ACT |       | G-ACT |       |
|-------|-------|-------|-------|-------|-------|-------|
|       | I     | III   | I     | III   | I     | II    |
| mean  | 173.5 | 277.8 | 172.5 | 287.7 | 176.7 | 245.6 |
| sd    | 11.3  | 17.1  | 11.1  | 14.7  | 9.2   | 17.2  |
| % cv  | 6.5   | 6.2   | 6.4   | 5.1   | 5.2   | 7.0   |

### Accuracy Data

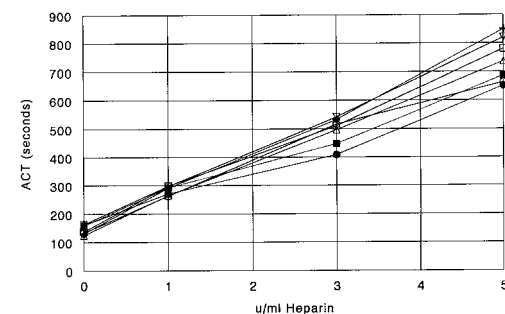
The accuracy of the Actalyke Activated Clotting Time Test System was evaluated by performing multiple Activated Clotting Time tests using varied combinations of instruments and tubes. Citrated pooled blood was heparinized to create the below concentrations yielding the following results:

### Heparin Sensitivity

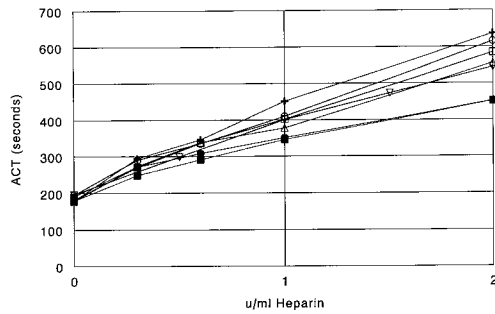
#### Celite Response to Heparin



#### Kaolin Response to Heparin



### Glass Beads Response to Heparin



#### Key to Graphs



#### Instrument/Tube

- A-A\* = Actalyke /Actalyke
- M-A\* = MINI /Actalyke
- XL-A\* = Actalyke XL /Actalyke
- H-H = Hemochron /Hemochron
- H-A = Hemochron /Actalyke
- A-H = Actalyke /Hemochron
- A-A = Actalyke /Actalyke

\*run with different set of donors

#### Correlation Data

Patient samples from all clinical sites were tested using each tube type on the MINI, the XL and the Actalyke. The data was as follows.

| Instrument | Tube | n   | Y=                | r=    |
|------------|------|-----|-------------------|-------|
| C-ACT      | MINI | 84  | $Y=0.948X + 15$   | 0.979 |
|            | XL   | 104 | $Y=0.943X + 11.7$ | 0.960 |
| G-ACT      | MINI | 97  | $Y=0.967X + 2$    | 0.979 |
|            | XL   | 92  | $Y=0.998X - 3.0$  | 0.985 |
| K-ACT      | MINI | 90  | $Y=0.984X + 10$   | 0.966 |
|            | XL   | 97  | $Y=0.978X + 6.9$  | 0.988 |

#### Linearity

Linearity studies were done according to NCCLS EP-6 guidelines with the following results.

| Heparin u/mL blood | C-ACT | K-ACT | G-ACT |
|--------------------|-------|-------|-------|
|                    | 5.0   | 5.0   | 2.0   |

#### BIBLIOGRAPHY

##### Literatur/Bibliografia/Bibliographie/Bibliografia

- Miale JB: Laboratory Medicine - Hematology, Fourth Edition. The Mosby Company, St. Louis, Missouri, p 1267, 1972.
- Hill JD, Dontigny L, deLaval M, Mielke CH: A simple method of heparin management during prolonged extracorporeal circulation. Ann Thorac Surg 17:129-134, 1974.
- Bull BS, Korpman RA, Huse WM, Briggs BD: Heparin therapy during extracorporeal circulation: I. Problems inherent in existing protocols. J Thorac Cardiovasc Surg 69: 674-684, 1975.
- Esposito RA, Culliford AT, Colvin SB, Thomas SJ, Lackner H, Spencer FC: The role of the activated clotting time in heparin administration and neutralization for cardiopulmonary bypass. J Thorac Cardiovasc Surg 85:174-185, 1983.
- Gambino R: Monitoring heparin therapy. Lab Report for Physicians 4: 17-20,1982.
- Hattersley P: Activated coagulation time of whole blood. JAMA 136:436,1966.
- Plonsey R, Collin RE. Magnetic field in material bodies. In: Principles and applications of electromagnetic fields. New York: McGraw-Hill Book Co. 1961 226-57.
- Lee RI, White PD: A clinical study of the coagulation time of blood. Am J Med Sci 145:495-503, 1913.
- Schriever HG, Epstein SE, Mintz MD : Statistical correlation and heparin sensitivity of activated partial thromboplastin time, whole blood

coagulation time, and an automated coagulation time. Am J Clin Path 60:323-329. 1973.

- Doty DB, Knott HW, Hoyt JL, Koepke JA: Heparin dose for accurate anticoagulation in cardiac surgery. J Cardiovasc Surg 20:597-604, 1979.
- National Institutes of Health Consensus Development Conference Statement: Prevention of venous thrombosis and pulmonary embolism. 1986.
- Ogilby JD, Kopelman HA, Klein LW, Agarwal JB: Adequate heparinization during PCTA: Assessment using activated clotting time. JACC 11: 237A, 1988.
- Hattersley PG: Heparin anticoagulation. In: Koepke JA (editor): Laboratory Hematology. Churchill Livingstone, 789-818, 1984.
- Lindsay RM. Practical use of anticoagulants. In: Drukker W, Parsons FW, Maher JF (editors): Replacement of Renal Function by Dialysis. Martinus Nijhof Publishers, 201-22, 1983.

### ACTALYKE CLOTTING TIME SYSTEM

|                                    | Cat. No. |
|------------------------------------|----------|
| C-ACT (Celite) Test Tubes (50/box) | C-ACT    |
| K-ACT (Kaolin) Test Tubes (50/box) | K-ACT    |
| G-ACT (Glass) Bead Tubes (50/box)  | G-ACT    |
| MAX-ACT Test Tubes (50/box)        | MAX-ACT  |

#### Other Supplies and Equipment

The following items, needed for performance of the Actalyke System must be ordered individually.

|   |        |
|---|--------|
| Actalyke XL   | 5770   |
| Actalyke XL Barcode Reader & Cable                  | 5758   |
| Actalyke MINI Analyzer                              | 5752   |
| Actalyke MINI Analyzer with printer                 | 5750   |
| Actalyke Clotting Time Analyzer                     |        |
| Single Well   | A1     |
| Single Well with printer                            | A1P    |
| Dual Well   | A2     |
| Dual Well with printer                              | A2P    |
| Actalyke Thermometer                                | 5757   |
| Electronic Clotting Tube                            | XL-ECT |
| Actalyke QC Kit (Level I & II) (for use with G-ACT) | AQC-LP |
| Actalyke QC Kit (Level I & III)                     | AQC-HP |

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