

ACHC20-04.B.OUS.CHC
May 2020

ADVIA® Chemistry 1800
ADVIA Chemistry 2400
ADVIA Chemistry XPT

Reassignment of the ADVIA® Chemistry CardioPhase High Sensitivity C-Reactive Protein (hsCRP) Calibrator Lots 484707 and 516407

Our records indicate that your facility may have received the following products:

Table 1. ADVIA Chemistry Systems Affected Product(s)

| Calibrator | Reference Number | Siemens Material Number (SMN) | Lot Number | Expiration Date (YYYY-MM-DD) | Manufacturing /1 st Distribution Date (YYYY-MM-DD) |
|--|------------------|-------------------------------|------------------|------------------------------|---|
| CardioPhase High Sensitivity C-Reactive Protein (hsCRP) Calibrator | 05006455 | 10335897 | 484707 516407 | 2020-11-28 2021-08-28 | 2019-05-28 / 2019-11-19 2020-02-28 / 2020-04-17 |

Reason for Communication

Siemens Healthcare Diagnostics Inc. issued Urgent Field Safety Notice (UFSN) ACHC20-04.A.OUS.CHC on November 19, 2019 to inform customers of a positive bias for patient samples and quality control (QC) material when using the ADVIA Chemistry hsCRP Calibrator lot 453610 (expired).

As a follow up to ACHC20-04, we have completed our investigation and this communication provides additional information. The commutability from European Reference Material ERM470 to ERM DA474/IFCC is not equivalent for high sensitivity CRP assays. Further evaluation of the alignment of the ADVIA Chemistry hsCRP calibrators to the European Reference Material ERM DA474/IFCC has demonstrated a bias for patient and QC material that is greater than previously communicated,

Siemens has confirmed that the current assigned values for ADVIA Chemistry hsCRP Calibrator lots 484707 and 516407 demonstrate a positive bias of approximately 15% for hsCRP patient samples and QC material when compared to the European Reference Material ERM DA474/IFCC for CRP measurement. Depending on the QC concentrations and ranges used by the laboratory, QC may not detect this issue.

The values for the current ADVIA Chemistry hsCRP Calibrator lots 484707 and 516407 have been reassigned to align with reference material ERM DA474/IFCC (see Tables 2 and 3 below). Going

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forward, future commercial calibrator lots will be traceable to this new reference material and will demonstrate comparable performance to the reassigned values.

Table 2. Current and Re-assigned values for hsCRP calibrator lot 484707

| Calibrator Level | Current Calibrator Values, mg/L | Reassigned Calibrator Values, mg/L |
|------------------|---------------------------------|------------------------------------|
| CAL 1 | 0.00 | 0.00 |
| CAL 2 | 0.47 | 0.40 |
| CAL 3 | 0.95 | 0.81 |
| CAL 4 | 1.44 | 1.22 |
| CAL 5 | 5.23 | 4.45 |
| CAL 6 | 10.68 | 9.08 |

Table 3. Current and Re-assigned values for hsCRP calibrator lot 516407

| Calibrator Level | Current Calibrator Values, mg/L | Reassigned Calibrator Values, mg/L |
|------------------|---------------------------------|------------------------------------|
| CAL 1 | 0.00 | 0.00 |
| CAL 2 | 0.52 | 0.44 |
| CAL 3 | 1.03 | 0.88 |
| CAL 4 | 1.56 | 1.33 |
| CAL 5 | 5.36 | 4.56 |
| CAL 6 | 10.70 | 9.10 |

Patient and QC results are expected to shift approximately -15% when using the reassigned calibrator values for lots 484707 and 516407. Based on the negative shift in recovery, it may be necessary to adjust your laboratory's QC ranges. Refer to Table 4 for representative QC recovery data. Future commercial calibrator lots will be traceable to this new reference material and will demonstrate comparable performance to the reassigned values.

Table 4. Quality Control Recovery

| Quality Control Material | Mean (mg/L) | Range (mg/L) |
|--|-------------|---------------|
| Bio-Rad Cardiac Markers Plus Control Lot 23691 Level 1 | 0.614 | 0.407 - 0.821 |
| Bio-Rad Cardiac Markers Plus Control Lot 23692 Level 2 | 1.865 | 1.510 - 2.220 |
| Bio-Rad Cardiac Markers Plus 67601 Level 1 | 0.666 | 0.459 – 0.873 |
| Bio-Rad Cardiac Markers Plus 67602 Level 2 | 1.917 | 1.554 – 2.287 |
| Bio-Rad Cardiac Markers Plus 67611 Level 1 | 0.666 | 0.437 – 0.895 |
| Bio-Rad Cardiac Markers Plus 67612 Level 2 | 1.872 | 1,49 5– 2.257 |
| Bio-Rad Cardiac Markers Plus (Tube) 67611T Level 1 | 0.681 | 0.451 – 0.910 |
| Bio-Rad Cardiac Markers Plus (Tube) 67612T Level 2 | 1.909 | 1.532 – 2.287 |
| Bio-Rad Cardiac Markers Plus Control Lot 67621 Level 1 | 0.585 | 0.363 – 0.807 |
| Bio-Rad Cardiac Markers Plus Control Lot 67622 Level 2 | 1.835 | 1.458 – 2.205 |
| Bio-Rad Cardiac Markers Plus (Tube) 67621T Level 1 | 0.592 | 0.437 – 0.747 |

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| | | |
|--|-------|---------------|
| Bio-Rad Cardiac Markers Plus (Tube) 67622T Level 2 | 1.865 | 1.576 – 2.146 |
| Bio-Rad Cardiac Markers Plus Control Lot 99561 Level 1 | N/A* | N/A* |
| Bio-Rad Cardiac Markers Plus Control Lot 99562 Level 2 | 1.968 | 1.576 – 2.361 |

*Bio-Rad does not publish a target value/range for this QC level.

Please see **Additional Information** section for patient sample correlation before and after calibrator value reassignment.

The information provided in this letter supersedes the information in the current ADVIA Chemistry hsCRP assay Instructions for Use (IFU) and the Declaration of Traceability and Uncertainty until these documents are updated, as well as the previous UMDc ACHC20-04.A.OUS.CHc dated November 2019.

Risk to Health

The risk to health due to this issue is negligible. Increased hsCRP results would be considered with other information, such as clinical history and symptomology, as well as other laboratory results. Siemens is not recommending a review of previously generated results.

Actions to be Taken by the Customer

- Please review this letter with your Medical Director.
- Calibrate using the reassigned calibrator values for hsCRP Calibrator Lots 484707 and 516407 provided in Table 2 and 3 above.
- QC targets and ranges should be reviewed and adjusted accordingly.
- Complete and return the Field Correction Effectiveness Check Form attached to this letter within 30 days.
- If you have received any complaints of illness or adverse events associated with the products listed in Table 1, immediately contact your local Siemens Healthineers Customer Care Center or your local Siemens Healthineers technical support representative.

Please retain this letter with your laboratory records and forward this letter to those who may have received this product.

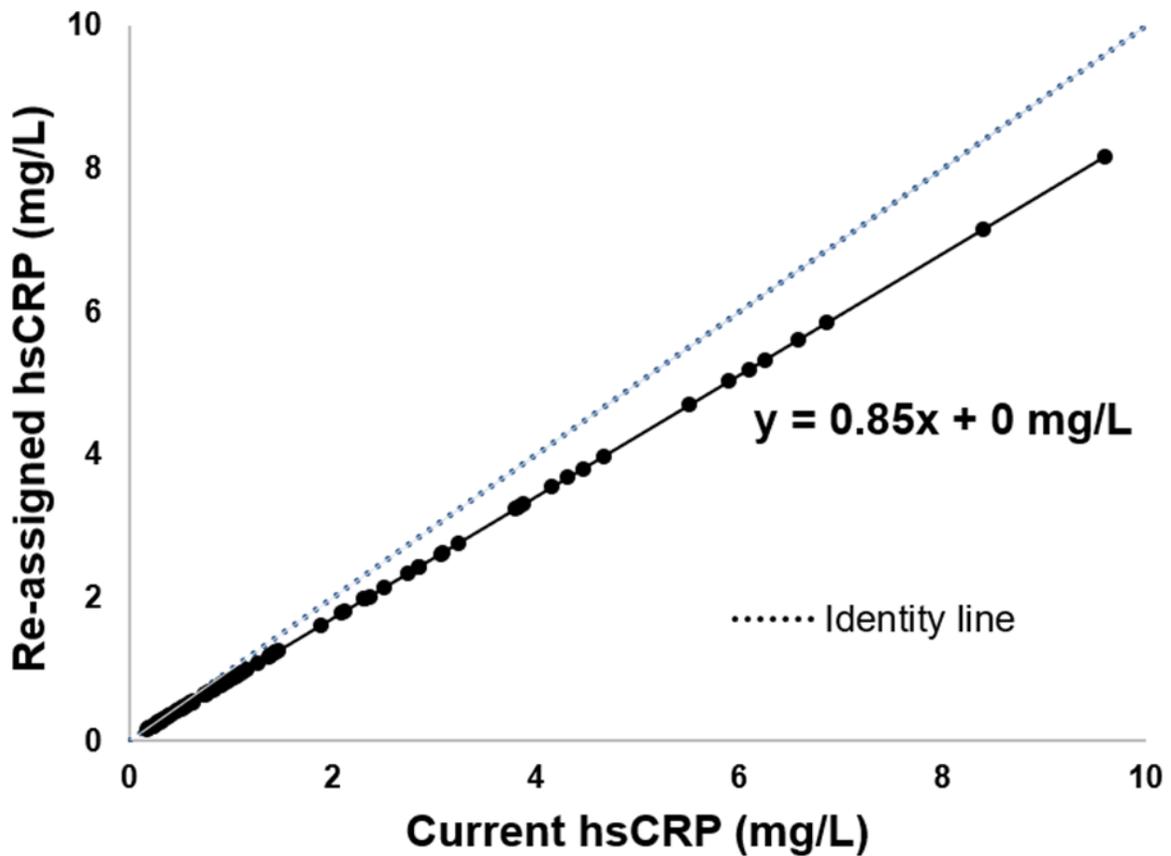
We apologize for the inconvenience this situation may cause. If you have any questions, please contact your Siemens Healthineers Customer Care Center or your local Siemens Healthineers technical support representative.

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Additional Information

An hsCRP correlation study comparing patient sample recovery using current and re-assigned calibrator values can be seen in Figure 1 below.

Figure 1. Patient sample correlation comparing results using current and re-assigned calibrator values.



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FIELD CORRECTION EFFECTIVENESS CHECK

ADVIA Chemistry Systems Reassignment of the ADVIA® Chemistry CardioPhase High Sensitivity C-Reactive Protein (hsCRP) Calibrator Lost 484707 and 516407.

This response form is to confirm receipt of the enclosed Siemens Healthcare Diagnostics Urgent Field Safety Notice, ACHC20-04.B.OUS.CHC dated May 2020 regarding ADVIA Chemistry Systems Reassignment of the ADVIA® Chemistry CardioPhase High Sensitivity C-Reactive Protein (hsCRP) Calibrator Lots 484707 and 516407. Please read each question and indicate the appropriate answer.

Return this completed form to Siemens Healthcare Diagnostics as per the instructions provided at the bottom of this page.

1. I have read and understood the Urgent Field Safety Notice instructions provided in this letter. Yes No

Name of person completing questionnaire: _____

Title: _____

Institution: _____ Instrument Serial Number: _____

Street: _____

City: _____ State: _____

Phone: _____ Country: _____

Please fax this completed form to the Customer Care Center at (###) ###-####. If you have any questions, contact your local Siemens technical support representative.