

## Urgent Field Safety Notice

10818055, Rev. A

March 2014

ADVIA Centaur®  
 ADVIA Centaur® XP  
 ADVIA Centaur® CP

### Information Regarding ADVIA Centaur Folate Assay

Our records indicate that your facility has received the following product:

**Table 1. ADVIA Centaur Systems Affected Products**

Assay	Siemens Material Number (SMN)	Kit Lots Ending in	Expiration Date
ADVIA Centaur Folate	10310308 (100 tests) 10325366 (500 tests) 10331250 (500 tests) 10340209 (2500 tests)	219	25 March, 2014
		222	12 May, 2014
		223	12 May, 2014
		224	04 August, 2014
		225	04 August, 2014
		226	12 September, 2014
		227	06 October, 2014

### Reason for Correction

Siemens Healthcare Diagnostics has confirmed two different sets of customer complaints regarding the ADVIA Centaur® Systems Folate assay.

#### ADVIA Centaur Folate Lot 223

Siemens confirmed one set of customer complaints with the ADVIA Centaur Systems Folate assay, kit lots ending in 223. Customers observed that the Bio-Rad Liquichek™ and Bio-Rad Lyphochek™ Immunoassay Plus Folate serum controls may go out of range low with the ADVIA Centaur Systems Folate assay, kit lots ending in 223. Serum patient results were consistent with previous reagent lots.

Whole blood controls with the ADVIA Centaur Systems Folate assay, kit lots ending in 223, are within range and whole blood patient results are consistent with previous reagent lots.

#### ADVIA Centaur Folate Lot 219 Whole Blood Bias

In addition, Siemens has confirmed a second set of customer complaints regarding a shift in whole blood patient results when moving from ADVIA Centaur Systems Folate assay, kit lots ending in 219 to kit lots ending in 222, 224, 225, 226 and/or 227. Siemens performed a whole blood normal range study that shows kit lots ending in 219 demonstrate a normal range distribution pattern different from other reagent lots as shown in Figure 1. However, this normal

range study showed that kit lots ending in 222, 224, 225, 226 and 227 demonstrate a consistent pattern.

Serum controls and patients show normal lot to lot variation with all in date lots of ADVIA Centaur Folate assay reagents.

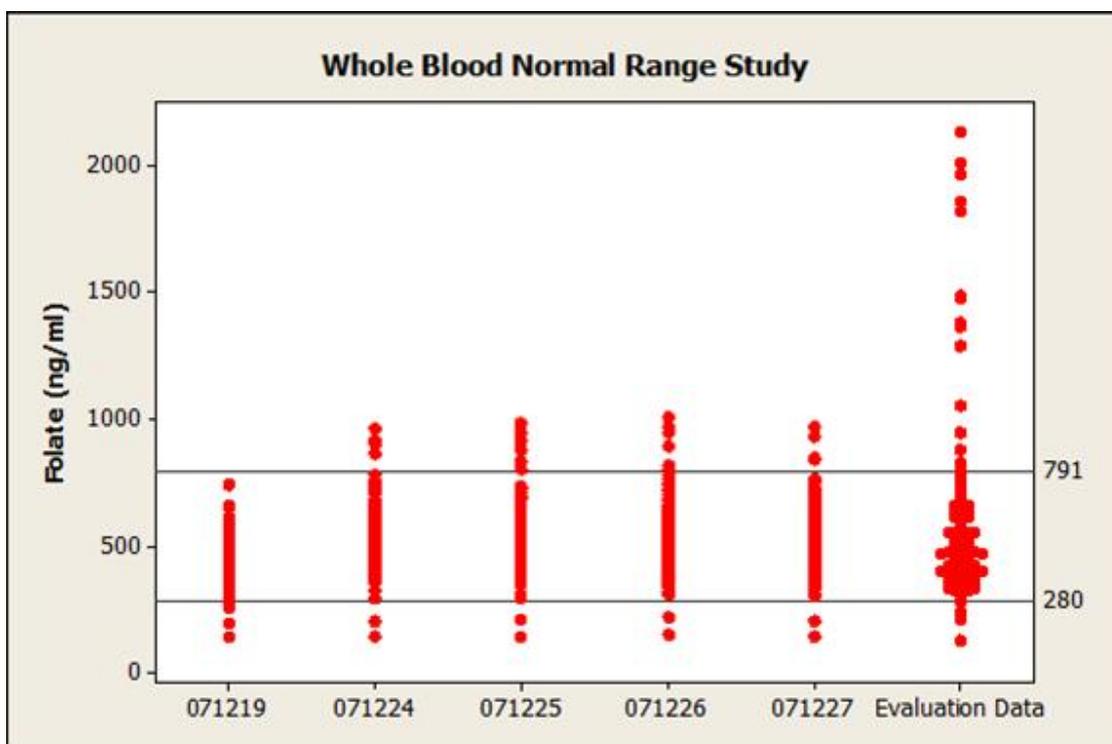
The observed shifts in whole blood samples are as follows:

- When comparing whole blood patient sample results from lot 219, customers may observe a mean bias of -21% to all current lots. Refer to Table 2 for the demonstrated range of bias.
- Whole blood patient samples for lots 224, 225, 226 and 227 show a bias of -1.2% to +1.4% from the mean. Refer to Table 2 for the demonstrated range of bias.
- Current lots show consistent agreement for lots ending in 224 through 227. Refer to Figure 1 for normal range data.
- Whole blood controls show a bias between 219 versus lots 224, 225, 226 and 227. When the mean dose for each level of control for 219 is compared to the mean dose for 224, 225, 226 and 227, the range of bias observed is from -13% to -21%. This bias is similar to the bias shown with the whole blood patient samples. Refer to Table 3 for current whole blood control ranges for kit lots 222 and above.

**Table 2. Summary of Bias From 071219 to 224, 225, 226 and 227 Using 150 Whole Blood Patient Samples**

<b>Lot Number</b>	<b>Mean Dose (ng/mL)</b>	<b>% Bias Compared to 224-227 Average</b>
219	423	-21%
224	530	-1.2%
225	538	0.3%
226	544	1.4%
227	535	-0.3%
224-227 Average	537	-

**Figure 1. Whole Blood Normal Range Study**



**NOTE:** The ADVIA Centaur Folate evaluation data in Figure 1 is a representation of the normal range study published in the Instructions For Use (IFU).

**Table 3. Whole Blood Control Bias From 071219 to 224, 225, 226 and 227**

Bio-Rad Lyphocek Immunoassay Plus Whole Blood Controls, Lot 73210					
Sample	Current Target	Current Specification	Mean Dose (ng/mL) (071219)	Mean Dose (ng/mL) (017224-071227)	% Bias Compared to 224-227 Average
73211	0.985	0.394-1.58	0.85	1.02	20.6%
73212	4.39	2.46-6.32	4.44	5.01	12.9%
73213	7.37	4.42-10.3	6.48	7.64	17.8%

### Risk to Health

#### ADVIA Centaur Folate Lot 223

If serum controls are out of range with ADVIA Centaur Systems Folate assay, kit lots ending in 223, patient results cannot be reported. A delay in reporting Folate results does not constitute a risk to health. If results have been reported with controls in acceptable ranges, then these results are considered clinically acceptable and no further review is required.

### **ADVIA Centaur Folate Lot 219 Whole Blood Bias**

Siemens has determined that the ADVIA Centaur Systems Folate assay, kit lots ending in 222, 224, 225, 226 and/or 227, meets the published claims in the IFU for normal whole blood cell Folate. The whole blood cell Folate results for lots ending in 219 may show a negative bias of up to 41.2% relative to new lots 224, 225, 226, and 227. This negative bias may result in a patient receiving Folate supplementation; however, this supplementation does not constitute a risk to health.

### **Actions to be Taken by the Customer**

- Discontinue use of the ADVIA Centaur Systems Folate assay, kit lots ending in 219 and 223.
- Continue to use ADVIA Centaur Systems Folate assay for serum and whole blood determinations with kit lots ending in 222, 224, 225, 226, 227 and above as they provide accurate Folate results.
- Updated control value assignment ranges will be posted on Bio-Rad's QCnet.com website. Please refer to the website for updated package inserts.
- Please review this letter with your Medical Director.
- Review your inventory of these products to determine your laboratory's replacement needs and to provide information to Siemens for reporting to the authorities.
- If you have received any complaints of illness or adverse events associated with the products listed in Table 1, immediately contact your local Siemens Customer Care Center or your local Siemens 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 has caused. If you have any questions, please contact your Siemens Customer Care Center or your local Siemens technical support representative.

Siemens Healthcare Diagnostics  
333 Coney Street  
East Walpole, MA 02032  
[www.siemens.com/diagnostics](http://www.siemens.com/diagnostics)

ADVIA Centaur is a trademark of Siemens Healthcare Diagnostics.

Liquichek and Lyphocheck are trademarks of Bio-Rad Laboratories.