

PRODUCT ALERT NOTICE

ACTION REQUIRED

Alert Ref No: CI Bilirubin Total Gen. 3: Introduction of a new reagent formulation according to the REACH regulation requiring new Cat.No., updates in instrument settings and new Carry-Over evasion steps QN-RDS-CoreLab-2025-009

Date: 06/06/2025

Product Affected: BILT3 (05795648190) cobas c 111 4 x 100 tests
 BILT3 (05795397190) cobas c 311/501/502, Integra 250 tests
 BILT3 (05795419190) cobas c 701/702 600 tests
 BILT3 (08056960190) cobas c 303/503/703 1050 tests

System Affected: COBAS Integra 400 plus
 cobas c 111
 cobas c 311
 cobas c 501
 cobas c 701 cobas c 702 cobas c 502
 cobas c 303
 cobas c 503 cobas c 703

Summary of Issue:

Significant Updates:

- Commencement of the OPE-free BILT3 formal launch scheduled for June 2025.
- Adherence to REACH regulatory standards.
- Implementation of new BILT3 material identification codes.
- Introduction of revised ACN codes, Test IDs, and Short names.
- Demonstration of patient results equivalence between the existing and OPE-free BILT3 assays.
- Utilisation of identical calibrator (C.f.a.s) and QC materials for both assay types.
- Compatibility allowing for concurrent use of current and OPE-free BILT3 reagents on the analyser.

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Transition to BILT3 OPE-free reagent

Due to Global bottlenecks we are holding extremely limited stock of BILT3 (containing OPE), which necessitates a shorter than usual transition period to the OPE-free reagent. **A swift conversion is highly recommended due to the bottleneck.** We understand this is a short timeline, so we're committed to providing all the support you need.

Date	Action
As soon as is practically possible	Customer verification tasks to be completed
31 st August 2025	Current assay no longer available.

2 x Free of Charge (FOC) BILT3 reagent packs will proactively be sent to you for this task with the PO of BILT3 REACH. If you don't receive these reagent packs by 13th June 2025 please reach out to your sales rep.

Reason for Notice

We are pleased to formally launch the new OPE-free Bilirubin Total Gen.3 assay (BILT3). In accordance with the REACH regulation, the R1 reagent in the current BILT3 assay has been reformulated and this assay is now available for you to order.

Following on from our previous customer notification **CI REACH (NPE/OPE) launch of BILT3 and general updates MN-RDS-CoreLab-2025-026** published at the end of March 2025. Please review the information in the tables below that provides you with analyser and application changes for all affected applications:

Table 1 – Provides an overview of the changes depending on the analyser and application:

	Immediate Action Required
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	Information Only

PRODUCT ALERT NOTICE

ACTION REQUIRED

Current Cat.No. & reagent Short Name	Analyzer	Current Test Short name	current ACN / Test ID	New Cat.No. & System ID	New Test Short Name / ACN / Test ID	Measuring Range	Post-Dilution	Prozone Check/ Kinetic unstable check	OBS	Blank update
05795397190 BILT3	cobas c 311	BILT3	712	x	x	x	x	unchanged	unchanged	unchanged
		SBILT3	711							
	cobas c 501	BILT3	712							
		SBILT3	711							
	cobas c 502	BILT3	8712							
		SBILT3	8711							
08056960190 BILT3	cobas c 303	BILT3	20310	x	x	x	x	x	x	x
								x		
								unchanged		
05795419190 BILT3	cobas c 701/702	BILT3	8712	x	x	unchanged	unchanged	x	unchanged	unchanged
		SBILT3	8711					x		

On **cobas® c 111**, no changes in Cat.No., ACN or settings are required for the REACH conforming reagent.

On COBAS INTEGRA 400 plus using cobas c pack small, the new Cat.No., new Test ID and new cassette definition / cassette size applies, but all application settings remain identical (measuring range, calibration interval and onboard stability).

Table 2 – Provides application Short names / Application Code numbers and Test IDs for current and new applications:

	Immediate Action Required
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	Information Only

PRODUCT ALERT NOTICE

ACTION REQUIRED

Analyzer	Current Test Short name	current Cat.No.	current ACN / Test ID	new Cat.No.	new Test Short Name	new ACN / Test ID		
cobas c 311	BILT3	05795397190	712	10421410190	BIT3	299		
	SBILT3		711		SBIT3	300		
cobas c 501	BILT3		712		BIT3	299		
	SBILT3		711		SBIT3	300		
cobas c 502	BILT3		8712		BIT3	8299		
	SBILT3		8711		SBIT3	8300		
COBAS INTEGRA 400 plus	BILT3		0-712		BIT3	0-411		
cobas c 303 cobas c 503 cobas c 703	BILT3		08056960190		20310	10421436190	BILT-3	20312
cobas c 701 cobas c 702	BILT3		05795419190		8712	10421428190	BIT3	8299
	SBILT3				8711		SBIT3	8300
cobas c 111	BILT3	05795648190	712	unchanged	unchanged	unchanged		

Most important changes in the updated BILT3 OPE-free assay

Following on from the significant points in the Summary section above, please note the following:

- On **cobas c 311/501/502/503/703** analysers: Reduction of measuring range and change of post dilution factor:
 - Lowering the upper measurement range from 650 µmol/L to 500 µmol/L.
 - Changing the post-dilution factor from 1:2 to 1:2.37.

Please note that reducing the measuring range is counterbalanced by increasing the post dilution factor accordingly thus ensuring measurement of samples up to 1300 µmol/L being unchanged.

- Only on **cobas c 701/702/303/503**: adjustment of the Prozone check/ kinetic unstable check.
- On **cobas c 311/501/502** and **cobas c 701/702**, a Prozone check has been set up for BILT3 to mark gammopathic samples. The adjustments made to this check harmonises the prozone / kinetic unstable check between all **cobas c** analysers.

	Immediate Action Required
✓	Action Required
	Information Only

PRODUCT ALERT NOTICE ACTION REQUIRED

cobas® c 111 and COBAS INTEGRA 400 plus are not affected.

Method comparison, performance data and test characteristics

There is a good method comparability between the current BILT3 and the new BILT3 OPE-free assay. Method comparison graphs can be found in previously communicated customer notice **CI REACH (NPE/OPE) launch of BILT3 and general updates MN-RDS-CoreLab-2025-026**

Performance Data and test characteristics of the new BILT3 OPE-free assay can also be found in respective Method Sheets (IFU). You can download these from the cobas® link or from e-LabDoc on the navify® Portal .

Implementation of 26 weeks Onboard Stability instead of 6 weeks for cobas c pack green

For the REACH conform reagent the onboard stability (OBS) in **cobas c** pack green has been increased from 6 to 26 weeks.

The OBS on all other analysers remain unchanged.

Implementation of a blank update every 6 weeks

Based on the implementation of an increased onboard stability, a blank update becomes necessary for **cobas® c 303/ 503 and c 703** every 6 weeks.

Action Required

Below you will find the technical instructions that support the application and technical updates and considerations:

Method Sheets and Supplemental Value Sheet information

BILT3 OPE-free has a new method sheet (IFU) and calibrator and control supplemental value sheet (SVS) version numbers which can be found in the table below. These may be downloaded from your cobas link or from eLabDoc on the navify® Portal.

	Immediate Action Required
▼	Action Required
	Information Only

PRODUCT ALERT NOTICE

ACTION REQUIRED

BILT3 OPE-free reagents	Method Sheets (IFU)	Supplemental Value Sheets (SVS)
cobas® c 311/501/502	Version 1.0	Version 11
COBAS INTEGRA 400 plus	Version 1.0	Version 1
cobas® c 701/702	Version 1.0	Version 11
cobas® c 503/303/703	Version 1.0	Version 14
cobas® c 111**	Version 10	N/A

** for cobas® c 111, only a new version of the Method Sheet is necessary as an update to the chapter for Precautions and warnings was needed.

A detailed overview of individual calibrator and control lot numbers and expiry dates is given in the attached *Supplemental Values Sheets (SVS)*.

Lot numbers with shorter shelf life as mentioned in the table above, which will not get updated e-library packages, are indicated in the SVS by the symbol "-" instead of the e-barcode version in column "ebc Ver."

Application information cobas® c 111

For cobas® c 111 no change in ACN nor calibrator or control values is necessary.

Application information COBAS INTEGRA 400 plus

For COBAS INTEGRA 400 plus, the calibrator set-points and control target values for the **new Test ID** must be entered **manually** for all lots based on the SVS attached to this document. Test Application Software (TAS 36.11) must be installed to support the running of the BILT3-OPE free assay on the COBAS INTEGRA 400 plus. This includes the new application (BIT3) and a new cassette definition (BILT3) for the new BILT3 reagent formulation:

	Immediate Action Required
✓	Action Required
	Information Only

PRODUCT ALERT NOTICE

ACTION REQUIRED

Current reagent formulation				NEW reagent formulation acc to REACH regulation			
Cat.No.	Test Short Name	Test ID	Cassette Definition System ID	Cat.No.	Test Short Name	Test ID	Cassette Definition System ID
05795397190	BILT3	0-712	BILT3 07 7483 9	10421410190	BIT3	0-411	BILT3 07 7678 5

For the cobas c pack kit itself, the Cassette Short Name “BILT3” remains unchanged due to regulatory reasons.

The cassette Long Name remains also unchanged.

Only the System ID will change.

In consequence, there are two Cassette Definitions “BILT3” included in the TAS file

- the current one used for the current test BILT3 Test ID 0-712
- the new one used for the new test BIT3 Test ID 0-411

For each test, the corresponding Cassette Definition will be loaded automatically. In the Reagent Status Overview screen cobas c packs with both Cat.Nos / System IDs will be displayed as “BILT3”.

Both tests can be installed in parallel on the analyser (e.g. during the verification period).

Please reach out to us if you require TAS file and we will send this out to you.

Application information cobas® c 311/501/502/701/702/303/503 and 703 systems:

Kindly download the new BILT3-OPE-free application to your systems referring to the information in Table 1 for applicable ACN.

For calibrator/control lots that are transmitted via an updated e-library package, please note that the setpoints and target values for the newly added ACN in calibrator/control e-BC versions must be installed first and then the new

	Immediate Action Required
▼	Action Required
	Information Only

PRODUCT ALERT NOTICE

ACTION REQUIRED

REACH-conform -application can be installed.

Please see **QN-RDS-CoreLab-2025-009 Attachment** for updated e-barcode numbers and if you have old stock of Cfas and QC please use the attachment information for lot number information with updated e-barcode compatibility.

Updated e-Library packages for zero calibrator for all systems

The new ACNs for BILT3 reagent formulation are included in following e-barcode versions for the zero calibrator:

Analyser / Module	e-barcode version
cobas® c 311	04-16
cobas® c 501	04-26
cobas® c 701	02-25
cobas® c 702	02-25
cobas® c 502	02-26
cobas® c 503	01-13
cobas® c 303	01-07
cobas® c 703	01-04

Patient results

Sample results are not affected.

Extra Wash Cycles

Kindly ensure you have followed the instructions in customer notification **CI Carry-over evasion SN-RDS-CoreLab-2025-029** for the appropriate extra wash cycles required to be in place before using this assay. Please ensure the extra wash cycles are activated and in effect before running the new application.

Attachments

QN-RDS-CoreLab-2025-009 Attachment – method sheets and SVS

	Immediate Action Required
▼	Action Required
	Information Only