# Urgent Field Safety Notice

*SBN-RDS-MolecularLab-2021-011*

RDS/**cobas**® EGFR Mutation Test v2
Version 2
May 2024

# cobas® EGFR Mutation Test v2: Potential for False Mutation Detected results for exon20 Insertion

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|---|---|
| **Product Name** | **cobas**® EGFR Mutation Test v2 |
| **BASIC UDI-DI/GMMI / Part No** | GMMI: 07248563190 |
| **Device Identifier (UDI)** | Device Identifier: 00875197005448 |
| **Production Identifier (Lot No./Serial No.)** | Not Applicable (not kit lot specific) |
| **SW Version** | Not Applicable |
| **Type of Action** | Field Safety Corrective Action (FSCA) |

Dear Valued Customer,

## Description of Situation

Roche is pleased to announce the availability of the updated ASAP (SW c4800 EGFR Tissue P1 AP v1.0.1.2311) for use with the **cobas**® EGFR Mutation Test v2 in CE-mark accepting countries. The updated ASAP utilizes an additional Ex20Ins parameter to reduce the risk of reporting false positive Ex20Ins Mutation Detected results with the **cobas**® EGFR Mutation Test v2. Installation of the updated ASAP is mandatory and must be completed by 29-Nov-2024.

Additionally, the **cobas**® EGFR Mutation Test v2, lot K27769, was the 1st kit lot manufactured utilizing an additional Quality Control function test for the enzyme raw material. The additional functional test for the enzyme raw material was implemented to screen enzyme batches prior to utilizing them in **cobas**® EGFR Mutation Test v2 test kit manufacturing.

As previously communicated, Roche received complaints from customers reporting the generation of false Mutation Detected results for the EGFR exon 20 insertion (EGFR Ex20Ins) mutation when using the **cobas**® EGFR Mutation Test v2 (GMMI: 07248563190).

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## Actions taken by Roche Diagnostics (if applicable)

Despite extensive root cause investigative testing, a definitive root cause for the early Ex20Ins Ct values and increased variability in the non-specific amplification of Ex20Ins mutation could not be determined. Ct results for false positive Ex20Ins Mutation Detected results were earlier and more variable for all batches of EGFR MMX3 v2 that used a specific lot of enzyme (raw material). It is plausible that the earlier Ex20Ins Ct values, coupled with the increased variability in Ct results, led to an increase in false positive Ex20Ins Mutation Detected results and the complaints in the field.

The investigation identified additional contributing factors that may increase the frequency of the false positive Ex20Ins mutation results, including the complexity of the enzyme manufacturing process as well as off-label practices such as the use of non-validated DNA quantitation methods (e.g., fluorometer).

Roche will schedule service visits to install the updated ASAP version, which reduces the risk of reporting false positive Ex20Ins Mutation Detected results with the **cobas**® EGFR Mutation Test v2. Installation of the updated ASAP is mandatory and must be completed by 29-Nov-2024.

## Actions to be taken by the customer/user

Customers must follow the **cobas**® DNA Sample Preparation Kit (M/N 05985536190) IFU for sample input.

Until the updated ASAP has been installed at customer sites:

- If an Ex20Ins Mutation Detected result is generated with the **cobas**® EGFR Mutation Test v2, customers must confirm the result with another method (e.g., sequencing or other PCR-based tests).
- Clinical laboratories should consider the availability and approval status of amivantamab in their country as well as eligibility for immunotherapy as part of SOC in the presence of any EGFR mutation when determining the date range of test result reports (TRR) of the **cobas**® EGFR Mutation Test v2 that must be reviewed retrospectively, and should follow local guidelines and procedures.
- Clinical laboratories located in the United States may consider reviewing results generated since May 2021 (Amivantamab was approved by the US FDA for NSCLC patients with EGFR Ex20Ins on May 21, 2021). TRRs with Ex20Ins mutation detected may be considered for confirmatory testing using sequencing or other PCR-based tests, upon the discretion of a CAP/CLIA laboratory director.

Once the updated ASAP is installed at customer sites, the confirmation testing for Ex20Ins Mutation Detected results is no longer necessary.

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## Communication of this Field Safety Notice (if appropriate)

*<If the recipient needs to forward the FSN to additional organizations/individuals then one or more of the following statements may be included:*

This notice must be passed on to all those who need to be aware within your organization or to any organization/individual where the potentially affected devices have been distributed/supplied. (If appropriate).

Please transfer this notice to other organizations/individuals on which this action has an impact. (If appropriate).

Please maintain awareness of this notice and resulting action for an appropriate period to ensure the effectiveness of the corrective action. (If appropriate).>

**The following statement is mandatory in FSNs for EEA countries but is not required for the rest of the World:**

*Include if applicable:* The undersigned confirms that this notice has been notified to the appropriate Regulatory Agency.

We apologize for any inconvenience this may cause and hope for your understanding and your support.

<closing salutations>,

**Contact Details**

*To be completed locally:*

Name
Title
Company Name
Address
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Email name@roche.com

Roche Molecular Systems, Inc.- SRN: US-MF-000018066 (legal manufacturer)
Roche Diagnostics GmbH- SRN: DE-AR-000006262 (EU authorized representative)