

April 2020

Important Note

therascreen® EGFR RGQ PCR Kit (24), V2, REF 874111

therascreen® EGFR Plasma RGQ PCR Kit (24), REF 870311

Dear, valued *therascreen* EGFR RGQ PCR Kit customer.

QIAGEN has identified a risk for a false mutation positive result in rare cases resulting from a fluorescence artefact that requires an update of the Instructions for Use (Handbook).

Description of the Issue

In said rare cases, the controls pass correctly, while a fluorescence artefact in the mutation assay causes an incorrect valid mutation positive result. The software and the protocol for manual sample assessment for the interpretation of all mutations claimed in the assay do not currently distinguish such fluorescence artefact from a real amplification obtained with valid mutation positive samples.

Inadequate reagent mixing at each mixing step during the assay setup may contribute to the occurrence of fluorescence artefacts.

The calling of false positive mutation samples due to the fluorescence artefact can be avoided by introducing a lower cut-off value for the delta C_T (ΔC_T).

Actions to be taken by the customer:

- For use of the *therascreen* EGFR RGQ PCR Kit (24), V2, REF 874111 and the *therascreen* EGFR Plasma RGQ PCR Kit (24), REF 870311, check the delta C_T values (ΔC_T) (in the sample result table for automated interpretation of results). For mutation positive results with a ΔC_T below -10.00 , these should be considered as invalid and retested.
- To implement the lower cut-off for the delta C_T (ΔC_T), the following new instruction replaces the Table 17 (p.94) of the *therascreen* EGFR RGQ PCR Kit (24), V2, REF 874111 handbook:

Assay	C_T range	Cutoff (ΔC_T)
T790M	0.00 to 40.00	$-10.00 \geq \text{to} \leq 7.40$
Deletions	0.00 to 40.00	$-10.00 \geq \text{to} \leq 8.00$
L858R	0.00 to 40.00	$-10.00 \geq \text{to} \leq 8.90$
L861Q	0.00 to 40.00	$-10.00 \geq \text{to} \leq 8.90$
G719X	0.00 to 40.00	$-10.00 \geq \text{to} \leq 8.90$
S768I	0.00 to 40.00	$-10.00 \geq \text{to} \leq 8.90$
Insertions	0.00 to 40.00	$-10.00 \geq \text{to} \leq 8.00$

The following new instruction replaces Table 8 (p.36) of *therascreen* EGFR Plasma RGQ PCR Kit (24), REF 870311 handbook:

Mutation assay	ΔC_T cutoffs
T790M	$-10.00 \geq \text{to} \leq 7.40$
Deletions	$-10.00 \geq \text{to} \leq 8.00$
L858R	$-10.00 \geq \text{to} \leq 8.90$

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- To obtain a valid qPCR result for the controls and samples, strict attention must be paid to thorough reagent mixing at each mixing step during the assay setup in accordance to the instructions for use.
 - Forward this information to all individuals and departments within your organization who are using *therascreen* EGFR RGQ PCR Kit (24), V2, REF 874111 and/or *therascreen* EGFR Plasma RGQ PCR Kit (24), REF 870311. If you are not the end user, please forward this notice to the product end user.
 - Review this notice with your laboratory/medical director.

Actions taken by QIAGEN

With this Important Note, QIAGEN is revising the instructions for use in order to reduce any risk resulting from fluorescence artefacts leading to invalid runs or false mutation positive results and to emphasize the importance of thorough mixing.

QIAGEN is updating the automatic calling of results in the Rotor-Gene Software to match the revised instructions for use. You will be informed as soon as such updated software is available. Until then, we advise you to manually check the result tables for entries in the delta C_T (ΔC_T) column as outlined above.

Translations of this Important Note

QIAGEN provides the printed Important Note in the English language. Translations of the Important Note are electronically available in a portable document format (PDF). You may access the translated documents in the Product Resources tab of the corresponding product page in www.qiagen.com.

If you have any questions or concerns, please contact your local QIAGEN Technical Service Department. For contact information, visit the following webpages:

QIAGEN Subsidiaries:

<https://www.qiagen.com/about-us/contact/global-contacts/subsidiaries/>

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With kind regards,

QIAGEN

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