IMPORTANT PRODUCT NOTICE

Date: xx Mar. 2009

Attention: **Laboratory Director**
PCR Laboratory

Subject: Factor II (Prothrombin) G20210A Macro (M/N 04586930 011) used with Factor II (Prothrombin) G20210A Kit for use with the LightCycler® 2.0 Instrument (M/N 03610195 001)

Issue

Roche Molecular Diagnostics has confirmed that the Factor II (Prothrombin) G20210A Macro (M/N 04586930 011) intended to be used with FACTOR II (PROTHROMBIN) G20210A KIT **for use with the LightCycler® 2.0 Instrument** may erroneously identify a mutation at position 20209 within the Factor II gene as wild-type genotype instead of unknown genotype as stated in the package insert. Although unconfirmed, similar erroneous results may occur with mutations at positions 20207, 20218 and 20221.

Impact on Results

The C20209T mutation is an uncommon sequence variant that occurs in < 1% of individuals of African descent and is very rare in Caucasians. The frequency of this mutation in other ethnic groups has not been determined. Currently, the functional significance of the C20209T mutation is uncertain. While some case reports have suggested an increased thrombotic risk in these patients [1], recent commentary has concluded that "the clinical utility of testing for and reporting this variant remains unresolved." [2]. Therefore the clinical significance in the evaluation of patients with suspected thrombophilia is uncertain.

In those rare occasions where the C20209T mutation is present, erroneous wild-type genotypic results may be reported.

Detection of the G20210A mutation is not impacted.

Corrective Actions

- Should you suspect the presence of the C20209T mutation or are uncertain of the accuracy of the wild-type genotypic result the melting curve should be examined to determine if the specimen is homozygous wild-type, heterozygous, or homozygous mutant genotype.

1) Dunn, St et al., *Blood Coagul Fibrinolysis*, 2006
2) Quigley, DI et al., *J Assoc Genet Technol*, 2007