

## TD2021IRMS version 2.0

### Summary of Major Modifications

The *Technical Document* on Detection of Synthetic Forms of *Prohibited Substances* by GC/C/IRMS, TD2021IRMS version 2.0, has been corrected as follows:

#### Article 1.0

It has been clarified that the general technical recommendations and method validation requirements described in this TD shall also be applied to the analysis of 19-NA and 19-NE, notwithstanding that these two substances are subjects of a separate TD (TD NA).

#### Article 1.1.3

It has been clarified that the same endogenous reference compounds (ERCs) and target compounds (TCs) that led to the “A” *Sample AAF* shall be measured in the “B” *Sample*.

#### Article 2.1.1

It has been clarified as a comment that *Sample* volume adjustment may allow the analysis of TCs and ERCs present in the *Sample* at a concentration lower than (<) the respective LOQ as long as the signal is within the validated linearity range of the IRMS instrument for that particular Analyte.

#### Article 2.1.2

The reference to the monitoring of the precision of the 45/44 and 46/44 ratios as a check for the stability of CO<sub>2</sub> pulses has been removed, since this is not applicable to users of Thermo instruments.

Correction of a minor mistake that has been detected in the description of the equation 3 for the mass balance adjustment of measured  $\delta^{13}\text{C}$  values of acetylated compounds. The term “C” in Eq. 3 had been misquoted as representing the concentration; in fact, “C” stands for “carbon” in  $\delta^{13}\text{C}$  and there is no concentration value represented in Eq. 3.

#### Article 2.3.2 - iv

It has been clarified as a comment that the criterion for the ERC-A and ERC-Etio pairs may be applied only if no other criterion based on the measurement of T, 5 $\alpha$ -Adiol and/or 5 $\beta$ -Adiol can be used.

#### Article 2.4.2

It has been clarified that in those cases when pregnanediol (PD) is not measurable in the *Sample*, or when in the Laboratory's opinion a negative or inconclusive result has been caused by a  $\delta^{13}\text{C}$  (ERC<sub>1</sub>) value that is not consistent with an endogenous origin, the

Laboratory shall explain in the comments section of the Test Report in *ADAMS* why PD could not be used as the ERC<sub>1</sub>.

Once more, it is stressed that the two (2) ERCs that allowed concluding the *AAF* for the “A” *Sample* shall also be measured during the “B” Sample CP.