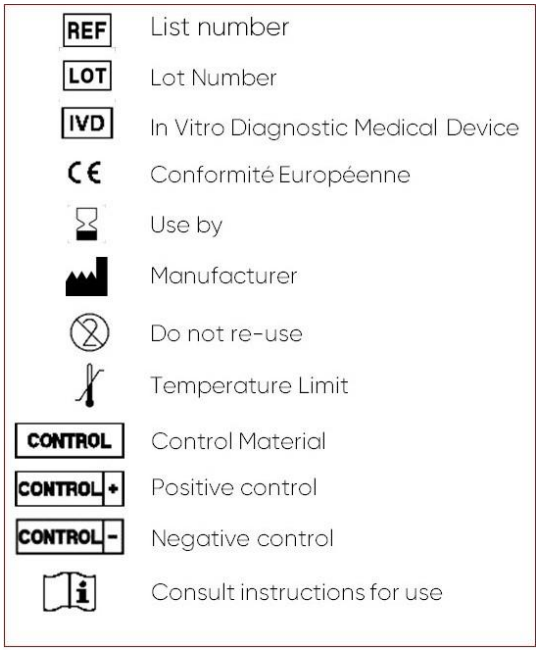
NAME

Idylla™ MSI FFPE Reference Set

Key to Symbols Used



Preface and important Notes

Thank you for purchasing your Idylla™ MSI FFPE Reference Set from SensID GmbH. Your product Idylla™ MSI FFPE Reference Set is a quality control material based on DNA and serves the purpose of a control for the Idylla™ MSI Assay from Biocartis NV using the Biocartis Idylla™ System.

WARNING! Improper handling and use can cause danger and damage. Therefore, we ask you to read these instructions for use and follow them carefully. Always keep them nearby. To avoid personal injury and damage to property, please also observe the safety instructions.

If you have any questions about the contents of these instructions for use or about the use of the product, please contact us.

Intended Purpose

Idylla™ MSI FFPE Reference Set is a qualitative control for the Idylla™ MSI Assay (Biocartis NV). Relevant literature shows, approximately, 15-20 % of colorectal cancers (CRC) display microsatellite instability (MSI) which is a unique molecular alteration and hyper mutable phenotype, resulting from a defective DNA mismatch repair (MMR)[1].

Intended Use

Idylla™ MSI FFPE Reference Set is intended as commutable, qualitative control material for microsatellite instability (MSI) markers in FFPE format based on human cell lines. It consists of MSI FFPE MSI-H control with all MSI markers at a frequency of 1x LOD for the Idylla™ MSI Test (refer to Idylla™ MSI Test package insert) and MSI FFPE MSS control with 0% frequency for all MSI markers. This Reference set enables monitoring of the performance, procedures and workflow of laboratory tests using FFPE in combination with the automatic workflow of the Idylla™ platform (Biocartis NV).
The product is intended to be used only with the Idylla™ MSI Test by professionals.

Content

Product	REF
Idylla™ MSI 0% FFPE Reference Standard (SID-000116)	SID-000114
Idylla™ MSI 16.67 % FFPE Reference Standard (SID-000117)	

One 10 µm FFPE curl per vial.

Storage

The Idylla™ MSI FFPE Reference Set should be stored at +2°C to +8°C upon arrival in qualified equipment. The Idylla™ MSI FFPE Reference Set is stable until the expiration date when stored under these conditions.

Instruction for Use

Information: Each curl within its vial has to go through the extraction process automatically performed by Idylla™ MSI Test. Refer to the manufacturer's instructions for the Idylla™ MSI Test. Test procedures for Idylla™ MSI Test provided by manufacturers must be followed closely as deviations from procedures recommended for Idylla™ MSI Test by manufacturers may produce unreliable results.

Preparation: Allow the product vials contained in the Idylla™ MSI FFPE Reference Set to come to room temperature (~15-25 °C) before use. Avoid contamination of the product when opening and closing the vial (see Warnings and Precautions).

1. Open the tube by twisting and lifting the lid.
2. Please remove carefully the curl without destruction by using a suitable forceps (e.g. anatomic forceps or other with flat tip) and transfer the curl in the appropriate position according to Idylla™ MSI Test instructions.
3. For further analysis, follow the Idylla™ MSI Test instructions.
4. Assuming a qualified device and validated process, the following is expected:

	Idylla™ MSI 0% FFPE Reference Standard (SID-000116)	Idylla™ MSI 16.67 % FFPE Reference (SID-000117)
MSI Status	MSS	MSI-H
MSI Score Range	< 0.15	≥ 0.15

Product limitations

Please note that samples of Idylla™ MSI FFPE Reference Set are not calibrators and should not be used for assay calibration.
Attention should be paid to expiration dates and storage conditions printed on the box and labels of all components.
Do not use Idylla™ MSI FFPE Reference Set Material beyond the expiry date. Do not use incorrectly stored components.
Check primary packaging for integrity before first opening.
Do not use products from damaged primary packaging.
The product is intended for use in the Idylla™ MSI workflow that should be performed in laboratory or clinical environment by professional and trained staff. It is not intended for self-testing or for use in the home or at the bedside.

Metrological traceability

In accordance with the quality management system of SensID GmbH, each lot of Idylla™ MSI FFPE Reference Set is tested against predetermined specifications to ensure consistent product quality.
In quality control, DNA is extracted from the FFPE and the amount of DNA for each batch is determined metrologically traceable to internationally certified reference material (ERM_AD442K). The fraction of MSI is determined by ddPCR (of selected marker) as qualified reference method[2].

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DNA quality in terms of integrity (DIN) is determined by automated electrophoresis. For assessment of the DIN no certified, metrologically traceable reference material is available. The method for determining DIN using automated electrophoresis is suitable as a quality control of DNA integrity.

Warnings and Precautions

The application of the in vitro diagnostic must be carried out by medical or scientific personnel. A safety data sheet for the product is provided by the manufacturer.
Caution: Use the required protective clothing and observe hygiene when using the set. The product contains immortalized and formalin-fixed paraffin embedded cells from cell lines categorized into biosafety level 1. Use universal precautions for handling reference materials and human specimens as recommended by Health Care Infection Control Practices Advisory Committee (HICPAC) in the guidelines from 2007 (Siegel et a., 2007). The safety precautions include the following rules:

- Mechanical pipetting only (no mouth pipetting allowed).
- Safe sharps handling.
- Avoidance of splashes or aerosols (by working under a clean bench/safety hood).
- Daily decontamination of all work surfaces when work is complete.
- Hand washing & disinfection.
- Prohibition of food, drink and smoking materials in lab setting.
- Personal protective equipment, such as; eye protection, gloves and a lab coat or gown.
- Consult package insert instructions for use.
- Do not use beyond expiration date.
- The Control + and Control - vials are intended for single use only; remaining reagents after first use should be discarded.
- For more safety information, please consult the appropriate material safety data sheets (MSDS).

Customer Service and Technical Assistance

Report all serious incidents (damage, injury, infection, etc.) that have occurred in connection with the product to the manufacturer and their competent national authority where you are established. If it is a serious incident, then the manufacturer informs the national authorities (e.g. EMA, BfArM, FDA) and initiates product recalls.

Reporting incidents to the manufacturer:
SensID GmbH – Schillingallee 68 – D-18057 Germany
Web: www.sens-id.com
E-Mail: tech@sens-id.com
Tel: +49 0381 377 18201

In Germany, the competent authority is the BfArM (Bundesinstitut für Arzneimittel und Medizinprodukte).
For incident reporting, the BfArM provides PDF forms on their website. https://www.bfarm.de/EN/Home/

Technical Service Assistance

Our Technical Service Assistance is staffed by experienced scientists with extensive practical and theoretical expertise with our products. If you have any questions or experience any difficulties regarding the Idylla™ MSI FFPE Reference Set or SensID GmbH products in general, please do not hesitate to contact us. For technical assistance and more information, please see our website www.sens-id.com or call one of the SensID GmbH Technical Service Assistance Tel. +49038137718201.

Abbreviations

Abbreviation	Description
BfArM	Bundesinstitut für Arzneimittel und Medizinprodukte (German Federal Institute for Drugs and Medical Devices)
ddPCR	Droplet digital polymerase chain reaction
DIN	DNA integrity number
DNA	Desoxyribonucleic acid
EMA	European Medicines Agency
FDA	Food and Drug Administration (U.S.)
FFPE	Formalin-fixed paraffin-embedded tissue
GmbH	Gesellschaft mit beschränkter Haftung (limited company)
Idylla™ platform	Trademark by Biocartis NV
Idylla™ MSI Test	
MSDS	Material safety data sheet
MSI	Microsatellite instability
MSS	Microsatellite stability
SID-...	SensID Catalog number
%	Percent
°C	Degree Celsius

Bibliography

1) Nojadeh JN, Behrouz Sharif S, Sakhinia E. Microsatellite instability in colorectal cancer. EXCLI J. 2018;17: 159-168. Published 2018 Jan 22. doi:10.17179/excli2017-948 MSI

2) Whale, Alexandra S., et al. "Assessment of digital PCR as a primary reference measurement procedure to support advances in precision medicine." Clinical chemistry 64.9 (2018): 1296-1307.