

Public declaration concerning the manufacture and use of in-house devices by health institutions

Name of health institution:

The Center for Predictive and Preventive Genetics (CGPP), Institute for Molecular and Cell Biology (IBMC).

Address:

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CGPP-IBMC declares that the devices listed in the accompanying table are exclusively manufactured and used within our Center and comply with the applicable General Safety and Performance Requirements (GSPR) of the In Vitro Diagnostic Medical Devices Regulation (EU 2017/746). Where applicable General Safety and Performance Requirements are not fully met, a reasoned justification is provided.

Since 2014, CGPP-IBMC has been accredited to ISO 15189:2014 by the Portuguese Accreditation Institute (IPAC) for various tests using methodologies such as Sanger sequencing, MLPA, fragment analysis, and whole-exome sequencing (WES). This accreditation encompasses all related equipment, reagents, and software.

For the in-house devices components marked with (*), there is no IVD version (revision 24-05-2024).

Date and location:

24-05-2024, Porto

Name, function and signature of responsible person(s):



Jorge Oliveira, Laboratory Director

Table of in-house devices:

Device identification	Device type (IVD/MD)	Risk class of the device	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
A. Genetic tests based on conventional methodology (Sanger sequencing; MLPA; fragments analysis)	IVD	Class C	Diagnostic, prenatal and presymptomatic or carrier testing	Y	Not applicable
B. Genetic tests based on whole-exome sequencing - WES (analysis of targeted multigene panels)	IVD	Class C	Diagnostic, prenatal and presymptomatic or carrier testing	Y	Not applicable
C. Genetic tests based on whole-exome sequencing – WES (total WES analysis - trio; singleton)	IVD	Class C	Diagnostic, prenatal and presymptomatic or carrier testing	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
A. Genetic tests based on conventional methodology (Sanger sequencing; MLPA; fragments analysis)				
Hi-Di™ Formamide for 3500 Dx/ 3500xL Dx Genetic Analyzers – Applied Biosystems	IVD	denaturing DNA samples prior to capillary electrophoresis	Y	Not applicable
POP-7™ Polymer for 3500 Dx/ 3500xL Dx Genetic Analyzers – Applied Biosystems	IVD	resolving DNA fragments (Sanger sequencing products and fragment analysis)	Y	Not applicable
Conditioning Reagent for 3500 Dx/ 3500xL Dx Genetic Analyzers – Applied Biosystems	IVD	prepare and maintain the capillaries of the 3500 Dx Genetic Analyzer	Y	Not applicable
Anode Buffer Container for 3500 Dx/ 3500xL Dx Genetic Analyzers – Applied Biosystems	IVD	provides the necessary buffer environment for the separation of nucleic acids during the electrophoretic process	Y	Not applicable
Cathode Buffer Container for 3500 Dx/ 3500xL Dx Genetic Analyzers – Applied Biosystems	IVD	ensures that the capillaries are supplied with the appropriate buffer solution at the cathode end, maintaining the necessary conditions for effective DNA separation and analysis	Y	Not applicable
3500 Dx Genetic Analyzer 8-Capillary Array, 50 cm – Applied Biosystems	IVD	conducts simultaneous, parallel capillary electrophoresis runs, enabling the high-throughput analysis of DNA samples	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
QIAsymphony DSP DNA Kits - QIAGEN	IVD	automated extraction of high-quality DNA from blood samples	Y	Not applicable
AmpliTaq Gold DNA Polymerase with Gold Buffer (*) – Applied Biosystems	MD	polymerase chain reaction (PCR) amplification requiring high specificity and robust amplification	Y	Not applicable
GeneScan 500 LIZ dye Size Standard (*) – Applied Biosystems	MD	reference standard used for DNA fragment analysis	Y	Not applicable
BigDye Terminator v1.1 Cycle Sequencing Kit (*) – Applied Biosystems	MD	Sanger sequencing of DNA products, by incorporating fluorescently-labeled dideoxynucleotides (ddNTPs) during DNA synthesis	Y	Not applicable
PowerUp SYBR Green Master Mix for qPCR (*) – Applied Biosystems	MD	DNA intercalation dye used to quantify the levels of target regions during the amplification process in quantitative polymerase chain reaction (qPCR) assays	Y	Not applicable
RNAlater Stabilization Solution (*) - Invitrogen	MD	preserves RNA integrity in biological samples by rapidly inactivating RNases and stabilizing RNA molecules	Y	Not applicable
Glycogen, molecular biology grade (20mg/m) (*) - ThermoFisher Scientific	MD	carrier for the precipitation of DNA and RNA; enhances the efficiency of precipitation	Y	Not applicable
GC-RICH PCR System, dNTPack (*) – Sigma-Aldrich	MD	PCR amplification of GC-rich DNA templates	Y	Not applicable
Dimethyl sulfoxide for molecular biology (*) – Sigma-Aldrich	MD	PCR enhancer to improve the efficiency and specificity of PCR reactions	Y	Not applicable
dNTP Mix, 10mM each (*) - ThermoFisher Scientific	MD	serves as the building blocks for DNA synthesis	Y	Not applicable
NanoDrop CF-1 Calibration Fluid, 0.5mL ampule (*) - ThermoFisher Scientific	MD	verifies the NanoDrop spectrophotometer calibration status	Y	Not applicable
TaKaRa LA Taq DNA polymerase, hot-start version (*) - TaKaRa	MD	for long-range PCR amplification (DNA fragments)	Y	Not applicable
Phusion high-fidelity DNA polymerase (*) - ThermoFisher Scientific	MD	amplification of DNA fragments with high fidelity and accuracy.	Y	Not applicable
Exo/SAP Go PCR Purification Kit (*) - Grisp	MD	Purification of PCR products by removing excess primers, nucleotides, enzymes, and salts from the PCR reaction mixture	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
AmplideX PCR C9orf72 (*) - Asuragen	MD	genetic testing for amyotrophic lateral sclerosis (ALS) and frontotemporal dementia (FTD) caused by expansions in the C9orf72 gene	Y	Not applicable
ROX 1000 size ladder (*) - ThermoFisher Scientific	MD	reference standard used in DNA fragment analysis	Y	Not applicable
High Sensitivity D5000 ScreenTape Assays & Reagents (*) - Agilent	MD	automated sizing, quantification, and quality control of DNA samples	Y	Not applicable
QIAamp DNA Mini Kit (50) (*) - QIAGEN	MD	DNA extraction kit	Y	Not applicable
DyeEx 96 Kit (*) - QIAGEN	MD	purify dye-labeled DNA fragments generated during cycle sequencing reactions	Y	Not applicable
RNase-Free Water (*) - QIAGEN	MD	used in reactions where the presence of RNases can compromise the results (e.g., PCR)	Y	Not applicable
HotStar TAQ Master mix Kit, 1000U (*) - QIAGEN	MD	PCR amplification of DNA fragments	Y	Not applicable
QIAGEN Multiplex PCR Kit (*) - QIAGEN	MD	amplification of multiple DNA targets simultaneously in a single reaction	Y	Not applicable
Matrix Standard BT5 for Multi-Capillary Instruments (*) - QIAGEN	MD	calibration standard for the 3500 Dx genetic analyzer	Y	Not applicable
QIAxcel DNA Screening Kit (*) - QIAGEN	MD	sizing and quantification of DNA fragments	Y	Not applicable
N2 cylinder (*) - QIAGEN	MD	provides the pressure necessary to drive the sample loading process on the QIAxcel system	Y	Not applicable
Primer 25 nmoles, purification DSL (*) - ThermoFisher Scientific	MD	specific DNA amplification	Y	Not applicable
Ligase Buffer A, 360 µl - MRC Holland	IVD	contains components necessary for the DNA ligase enzyme to catalyze the ligation of the probe pairs	Y	Not applicable
Ligase Buffer B, 360 µl - MRC Holland	IVD	provides the necessary conditions for efficient DNA ligation	Y	Not applicable
SALSA PCR Primer FAM, 240µl - MRC Holland	IVD	allows the detection and quantification of the amplified products	Y	Not applicable
SALSA Polymerase, 65 µl - MRC Holland	IVD	amplification step of MLPA assays		
SALSA MLPA buffer - MRC Holland	IVD	necessary components, such as ligase and buffer salts, to facilitate the ligation of specific probe pairs to the target DNA sequences	Y	Not applicable
SALSA Ligase-65, 115 µl - MRC Holland	IVD	ligation of probe oligos that hybridized to immediately adjacent target sequences.	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P010 POLG (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the genes <i>POLG</i> , <i>POLG2</i> , <i>TWINK</i> and <i>SLC25A4</i>	Y	Not applicable
SALSA MLPA Probemix P013 ATRX (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ATRX</i> gene	Y	Not applicable
SALSA MLPA Probemix P015 MECP2 – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>MECP2</i> , <i>CDKL5</i> , <i>ARX</i> and <i>NTNG1</i> genes	Y	Not applicable
SALSA MLPA Probemix P018 SHOX – MRC Holland	IVD	assay for the detection of deletions or duplications in the human short stature homeobox (<i>SHOX</i>)	Y	Not applicable
SALSA MLPA Probemix P021 SMA – MRC Holland	IVD	assay for the detection of deletions or duplication in the <i>SMN1</i> , <i>SMN2</i> and exon 5 of <i>NAIP</i> genes	Y	Not applicable
SALSA MLPA Probemix P022 PLP1 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>PLP1</i> gene and the Xq22 region	Y	Not applicable
SALSA MLPA Probemix P026 Sotos (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>NSD1</i> and <i>NFIX</i> genes	Y	Not applicable
SALSA MLPA Probemix P031 FANCA mix 1 (*) – MRC Holland	MD	assays for the detection of deletions or duplications in the <i>FANCA</i> gene	Y	Not applicable
SALSA MLPA Probemix P032 FANCA mix 2 (*) – MRC Holland	MD	assays for the detection of deletions or duplications in the <i>FANCA</i> gene	Y	Not applicable
SALSA MLPA Probemix P033 CMT1 – MRC Holland	IVD	assay for the detection of deletions or duplications in the human <i>PMP22</i> and <i>KIF1b</i> genes	Y	Not applicable
SALSA MLPA Probemix P034 DMD-1 – MRC Holland	IVD	assays for the detection of deletions or duplications in the <i>DMD</i> gene	Y	Not applicable
SALSA MLPA Probemix P035 DMD-2 – MRC Holland	IVD	assays for the detection of deletions or duplications in the <i>DMD</i> gene	Y	Not applicable
SALSA MLPA Probemix P041 ATM-1 – MRC Holland	IVD	assays for the detection of deletions or duplications in the <i>ATM</i> gene	Y	Not applicable
SALSA MLPA Probemix P042 ATM-2 – MRC Holland	IVD	assays for the detection of deletions or duplications in the <i>ATM</i> gene	Y	Not applicable
SALSA MLPA Probemix P044 NF2 – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>NF2</i> gene	Y	Not applicable
SALSA MLPA Probemix P046 TSC2 – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>TSC2</i> gene	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P047 RB1 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>RB1</i> gene and of methylation status of the <i>RB1</i> gene promoter and imprinted locus in a DNA sample.	Y	Not applicable
SALSA MLPA Probemix P048 LMNA/MYOT/ZMPSTE24 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>LMNA</i> , <i>MYOT</i> , <i>CAV3</i> and <i>ZMPSTE24</i> genes	Y	Not applicable
SALSA MLPA Probemix P050 CAH – MRC Holland	IVD	assay2 for the detection of large deletions and large gene conversions in the <i>CYP21A2</i> gene and its surrounding region located on chromosome 6p21.3	Y	Not applicable
SALSA MLPA Probemix P051 Parkinson mix 1 – MRC Holland	IVD	assay for the detection of deletions or duplication in <i>SNCA</i> , <i>PARK2</i> , <i>UCHL1</i> , <i>PINK1</i> , <i>PARK7</i> , <i>ATP13A2</i> , <i>LRRK2</i> , <i>GCH1</i> genes, and the presence of two disease-causing single nucleotide variants: p.A30P in the <i>SNCA</i> gene and p.G2019S in the <i>LRRK2</i> gene	Y	Not applicable
SALSA MLPA Probemix P054 <i>FOXL2-TWIST1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>TWIST1</i> , <i>FOXL2</i> , <i>FOXC1</i> , <i>FOXC2</i> , <i>ATR</i> , <i>PITX2</i> , <i>PISRT1</i> and <i>GPR143</i> genes	Y	Not applicable
SALSA MLPA Probemix P058 <i>IGHMBP2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>IGHMBP2</i> gene	Y	Not applicable
SALSA MLPA Probemix P059 Dystonia (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>TOR1A</i> , <i>THAP1</i> , <i>ATP1A3</i> , and <i>PRKRA</i> gene	Y	Not applicable
SALSA MLPA Probemix P061 Lissencephaly (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>PAFAH1B1</i> , <i>DCX</i> , <i>POMT1</i> , <i>POMGNT1</i> and <i>FLNA</i> genes	Y	Not applicable
SALSA MLPA Probemix P062 <i>LDLR</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>LDLR</i>	Y	Not applicable
SALSA MLPA Probemix P064 Microdeletion Syndromes-1B – MRC Holland	IVD	assay for the detection of a distinct subset of recurrent microdeletions and microduplications	Y	Not applicable
SALSA MLPA Probemix P065 Marfan Syndrome-1 – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>FBN1</i> gene	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P066 Marfan Syndrome-2 – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>FBN1</i> gene	Y	Not applicable
SALSA MLPA Probemix P071 <i>LMNB1-PLP1-NOTCH3</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>LMNB1</i> , <i>PLP1</i> and <i>NOTCH3</i> genes	Y	Not applicable
SALSA MLPA Probemix P073 <i>IKBKG</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the chromosome Xq28 region, including <i>IKBKG</i>	Y	Not applicable
SALSA MLPA Probemix P075 <i>TCF4-FOXP1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>TCF4</i> and <i>FOXP1</i> genes	Y	Not applicable
SALSA MLPA Probemix P080 Craniofacial (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>FGFR1</i> , <i>FGFR2</i> , <i>FGFR3</i> , <i>TWIST1</i> , <i>MSX2</i> , <i>ALX4</i> , <i>ALX3</i> , <i>ALX1</i> , <i>RUNX2</i> and <i>EFNB1</i> genes	Y	Not applicable
SALSA MLPA Probemix P081 <i>NF1</i> mix 1 – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>NF1</i> gene	Y	Not applicable
SALSA MLPA Probemix P082 <i>NF1</i> mix 2 – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>NF1</i> gene	Y	Not applicable
SALSA MLPA Probemix P091 <i>CFTR</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>CFTR</i> gene and the wild type allele of the <i>CFTR</i> p.Phe508del and p.Ile507del variants	Y	Not applicable
SALSA MLPA Probemix P093 <i>HHT/HPAH</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>ENG</i> , <i>ACVRL1</i> and <i>BMPR2</i> genes	Y	Not applicable
SALSA MLPA Probemix P098 Wilson disease (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ATP7B</i> gene	Y	Not applicable
SALSA MLPA Probemix P099 <i>GCH1-TH-SGCE-PRRT2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>GCH1</i> , <i>TH</i> , <i>SGCE</i> and <i>PRRT2</i> genes	Y	Not applicable
SALSA MLPA Probemix P104 Menkes <i>ATP7A</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ATP7A</i> gene	Y	Not applicable
SALSA MLPA Probemix P109 <i>ABCB4</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ABCB4</i> gene	Y	Not applicable
SALSA MLPA Probemix P114 Long-QT – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>KCNQ1</i> , <i>KCNH2</i> , <i>KCNE1</i> , <i>KCNE2</i> and <i>KCNJ2</i> genes	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P116 SGC (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>SGCA</i> , <i>SGCB</i> , <i>SGCD</i> , <i>SGCG</i> and <i>FKRP</i> genes	Y	Not applicable
SALSA MLPA Probemix P120 <i>PANK2/PLA2G6</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>PANK2</i> and <i>PLA2G6</i> genes	Y	Not applicable
SALSA MLPA Probemix P124 <i>TSC1</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>TSC1</i> gene	Y	Not applicable
SALSA MLPA Probemix P128 <i>CYP450</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>GSTM1</i> , <i>CYP1B1</i> , <i>CYP3A4</i> , <i>CYP3A5</i> , <i>CYP2C19</i> , <i>CYP2C9</i> , <i>CYP2E1</i> , <i>GSTP1</i> , <i>CYP1A2</i> , <i>CYP1A1</i> , <i>CYP2A6</i> , <i>CYP2B6</i> , <i>GSTT1</i> and <i>CYP2D6</i> genes	Y	Not applicable
SALSA MLPA Probemix P137 <i>SCN1A</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in <i>SCN1A</i> gene	Y	Not applicable
SALSA MLPA Probemix P138 <i>SLC2A1-STXBP1</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>SLC2A1</i> and <i>STXBP1</i> genes	Y	Not applicable
SALSA MLPA Probemix P141 <i>NIPBL-1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>NIPBL</i> gene	Y	Not applicable
SALSA MLPA Probemix P142 <i>NIPBL-2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>NIPBL</i> gene	Y	Not applicable
SALSA MLPA Probemix P143 <i>MFN2-MPZ</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>MFN2</i> and <i>MPZ</i> genes	Y	Not applicable
SALSA MLPA Probemix P148 <i>TGFBR1-TGFBR2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>TGFBR1</i> and <i>TGFBR2</i> genes	Y	Not applicable
SALSA MLPA Probemix P151 <i>ABCA4</i> mix-1 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ABCA4</i> gene	Y	Not applicable
SALSA MLPA Probemix P152 <i>ABCA4</i> mix-2 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ABCA4</i> gene	Y	Not applicable
SALSA MLPA Probemix P154 <i>GPC3-GPC4</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>GPC3</i> and <i>GPC4</i> genes	Y	Not applicable
SALSA MLPA Probemix P158 <i>JPS</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>BMPR1A</i> and <i>SMAD4</i> genes	Y	Not applicable
SALSA MLPA Probemix P159 <i>GLA</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>GLA</i> gene	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P160 <i>STS</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>STS</i> gene	Y	Not applicable
SALSA MLPA Probemix P163 <i>GJB-WFS1-POU3F4</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>GJB2, GJB3, GJB6, WFS1, and POU3F4</i> genes, microdeletions upstream of <i>POU3F4</i> , and the presence of six specific disease-causing variants in the <i>GJB2</i> gene	Y	Not applicable
SALSA MLPA Probemix P165 HSP mix-1 – MRC Holland	IVD	assay for the detection of deletions or duplications in <i>ATL1 and SPAST</i> genes	Y	Not applicable
SALSA MLPA Probemix P166 <i>KCNQ2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>KCNQ2</i> gene	Y	Not applicable
SALSA MLPA Probemix P169 Hirschsprung-1 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>RET, ZEB2, EDN3 and GDNF</i> genes	Y	Not applicable
SALSA MLPA Probemix P170 <i>APP</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>APP</i> gene	Y	Not applicable
SALSA MLPA Probemix P176 <i>CAPN3</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CAPN3</i> gene	Y	Not applicable
SALSA MLPA Probemix P177 <i>CASR</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CASR</i> gene	Y	Not applicable
SALSA MLPA Probemix P180 Limb malformations-2 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>SALL1, SALL4, and TBX5</i> genes	Y	Not applicable
SALSA MLPA Probemix P181 Centromere mix 1 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in genes close to the centromeres of all chromosomes, with the exception of the Y-chromosome	Y	Not applicable
SALSA MLPA Probemix P184 <i>JAG1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>JAG1</i> gene	Y	Not applicable
SALSA MLPA Probemix P185 Intersex (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>NR0B1 (DAX1), SOX9, SRY, WNT4 and NR5A1</i> genes	Y	Not applicable
SALSA MLPA Probemix P189 <i>CDKL5/ARX/FOXP1</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the human <i>CDKL5</i> and <i>FOXP1</i> genes	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P193 <i>NPC1-NPC2-SMPD1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>NPC1</i> , <i>NPC2</i> and <i>SMPD1</i> genes	Y	Not applicable
SALSA MLPA Probemix P197 <i>KCNQ3</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>KCNQ3</i> , <i>CHRNA4</i> , <i>EPM2A</i> , <i>NHLRC1</i> , and <i>CHRNA2</i> genes	Y	Not applicable
SALSA MLPA Probemix P198 <i>FH</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the fumarate hydratase (<i>FH</i>) gene	Y	Not applicable
SALSA MLPA Probemix P201 <i>CHARGE</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CHD7</i> gene	Y	Not applicable
SALSA MLPA Probemix P213 HSP mix-2 – MRC Holland	IVD	assay for the detection of deletions or duplication in <i>REEP1</i> and <i>SPG7</i> genes	Y	Not applicable
SALSA MLPA Probemix P215 <i>EXT</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>EXT1</i> and <i>EXT2</i> genes	Y	Not applicable
SALSA MLPA Probemix P219 <i>PAX6</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the 11p13-14 region, which includes the <i>PAX6</i> and <i>WT1</i> genes, and the <i>SOX2</i> gene on 3q26	Y	Not applicable
SALSA MLPA Probemix P225 <i>PTEN</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in <i>PTEN</i> gene	Y	Not applicable
SALSA MLPA Probemix P226 <i>SDH</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in <i>SDHB</i> , <i>SDHC</i> , <i>SDHD</i> , <i>SDHAF1</i> , and <i>SDHAF2</i> genes	Y	Not applicable
SALSA MLPA Probemix P241 <i>MODY</i> Mix 1 – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>HNF4A</i> , <i>GCK</i> , <i>HNF1A</i> , and <i>HNF1B</i> genes	Y	Not applicable
SALSA MLPA Probemix P254 <i>PSEN1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>PSEN1</i> gene	Y	Not applicable
SALSA MLPA Probemix P258 <i>SMARCB1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>SMARCB1</i> gene	Y	Not applicable
SALSA MLPA Probemix P266 <i>CLCNKB</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CLCNKB</i> and <i>CLCNKA</i> genes	Y	Not applicable
SALSA MLPA Probemix P271 <i>COL1A1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>COL1A1</i> gene	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P272 <i>COL1A2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>COL1A2</i> gene	Y	Not applicable
SALSA MLPA Probemix P275 <i>MAPT-GRN</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>MAPT</i> , <i>GRN</i> , and <i>CRHR1</i> genes	Y	Not applicable
SALSA MLPA Probemix P279 <i>CACNA1A</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the genes <i>CACNA1A</i>	Y	Not applicable
SALSA MLPA Probemix P285 <i>LRP5</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>LRP5</i> , <i>DKK1</i> , <i>NDP</i> and <i>FZD4</i> genes	Y	Not applicable
SALSA MLPA Probemix P295 <i>SPRED1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>SPRED1</i> gene	Y	Not applicable
SALSA MLPA Probemix P306 <i>SPG11</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>SPG11</i> gene	Y	Not applicable
SALSA MLPA Probemix P307 <i>SEPT9</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>SEPT9</i> gene	Y	Not applicable
SALSA MLPA Probemix P310 <i>TCOF1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>TCOF1</i> gene	Y	Not applicable
SALSA MLPA Probemix P313 <i>CREBBP</i> (*) – MRC Holland	MD	assay for the detection of copy number variations in the <i>CREBBP</i> and <i>EP300</i> genes	Y	Not applicable
SALSA MLPA Probemix P316 Recessive Ataxias (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>APTX</i> , <i>SETX</i> , and <i>FXN</i> genes	Y	Not applicable
SALSA MLPA Probemix P319 Thyroid (*) – MRC Holland	MD	assay for the detection of deletions or duplications in <i>TPO</i> , <i>PAX8</i> , <i>FOXE1</i> , <i>NKX2-1</i> and <i>TSHR</i> genes	Y	Not applicable
SALSA MLPA Probemix P325 <i>OCA2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>OCA2</i> and <i>TYR</i> genes	Y	Not applicable
SALSA MLPA Probemix P333 <i>EP300</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>EP300</i> gene	Y	Not applicable
SALSA MLPA Probemix P336 <i>UBE3A</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>UBE3A</i> gene	Y	Not applicable
SALSA MLPA Probemix P337 <i>TSC2</i> Confirmation – MRC Holland	IVD	assay for the confirmation of deletions or duplications in the <i>TSC2</i> gene as initially observed using the SALSA MLPA P046 <i>TSC2</i> probemix	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P340 <i>EHMT1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>EHMT1</i> gene	Y	Not applicable
SALSA MLPA Probemix P347 Hemochromatosis (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>HFE</i> , <i>TFR2</i> , <i>HFE2</i> (<i>HJV</i>), <i>HAMP</i> , and <i>SLC40A1</i> genes	Y	Not applicable
SALSA MLPA Probemix P348 <i>ATP1A2-CACNA1A-PRRT2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ATP1A2</i> , <i>CACNA1A</i> and <i>PRRT2</i> genes	Y	Not applicable
SALSA MLPA Probemix P350 <i>CLCN1-KCNJ2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CLCN1</i> and <i>KCNJ2</i> genes	Y	Not applicable
SALSA MLPA Probemix P352 <i>PKD1-PKD2</i> – MRC Holland	IVD	assays for the detection of deletions or duplications in the <i>PKD1</i> and <i>PKD2</i> genes	Y	Not applicable
SALSA MLPA Probemix P353 <i>CMT4</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>GDAP1</i> , <i>MTMR2</i> , <i>SBF2</i> , <i>SH3TC2</i> , <i>EGR2</i> , and <i>PRX</i> genes	Y	Not applicable
SALSA MLPA Probemix P355 Microcephaly (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>MCPH1</i> , <i>MCPH2</i> (<i>WDR62</i>), <i>MCPH3</i> (<i>CDK5RAP2</i>), <i>MCPH5</i> (<i>ASPM</i>), <i>MCPH6</i> (<i>CENPJ</i>), and <i>MCPH7</i> (<i>STIL</i>) genes	Y	Not applicable
SALSA MLPA Probemix P369 Smith-Magenis (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the 17p11.2 chromosomal region, including <i>RAI1</i>	Y	Not applicable
SALSA MLPA Probemix P378 <i>MUTYH</i> – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>MUTYH</i> gene, as well as the presence of the two most common disease-causing single nucleotide variants among people of European descent, c.536A>G (p.Tyr179Cys) and c.1187G>A (p.Gly396Asp)	Y	Not applicable
SALSA MLPA Probemix P386 <i>DOCK8-STAT3</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>DOCK8</i> and <i>STAT3</i> genes	Y	Not applicable
SALSA MLPA Probemix P387 <i>NPHP1</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>NPHP1</i> gene	Y	Not applicable
SALSA MLPA Probemix P389 <i>MLL2</i> (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>MLL2</i> gene	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P397 SCN4A-CACNA1S (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CACNA1S</i> and <i>SCN4A</i> genes	Y	Not applicable
SALSA MLPA Probemix P398 CASK (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CASK</i> gene	Y	Not applicable
SALSA MLPA Probemix P405 CMT1 – MRC Holland	IVD	assay for the detection of deletions or duplications in the human <i>PMP22</i> , <i>MPZ</i> and <i>GJB1</i> genes	Y	Not applicable
SALSA MLPA Probemix P410 GRIN2A GRIN2B (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>GRIN2A</i> and <i>GRIN2B</i> genes	Y	Not applicable
SALSA MLPA Probemix P411 Porphyria mix 1 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ALAD</i> , <i>HMBS</i> and <i>PPOX</i> genes	Y	Not applicable
SALSA MLPA Probemix P419 CDKN2A/2B-CDK4 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CDKN2A</i> , <i>CDKN2B</i> and <i>CDK4</i> genes	Y	Not applicable
SALSA MLPA Probemix P436 ANO5 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ANO5</i> gene	Y	Not applicable
SALSA MLPA Probemix P439 COL4A3 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>COL4A3</i> gene	Y	Not applicable
SALSA MLPA Probemix P441 SACS (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>SACS</i> gene	Y	Not applicable
SALSA MLPA Probemix P445 KDM6A (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>KDM6A</i> gene	Y	Not applicable
SALSA MLPA Probemix P454 CGD (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CYBA</i> , <i>CYBB</i> , <i>NCF2</i> and <i>NCF4</i> genes	Y	Not applicable
SALSA MLPA Probemix P457 DHCR7 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>DHCR7</i> gene	Y	Not applicable
SALSA MLPA Probemix P460 SMA (Silent) Carrier – MRC Holland	IVD	assay for the detection of 1. copy number changes of exons 7 and 8 of the <i>SMN1</i> gene for carrier testing and 2. the presence of the g.27134T>G and g.27706-27707delAT variants in <i>SMN1</i>	Y	Not applicable
SALSA MLPA Probemix P461 STRC-CATSPER2-OTOA – MRC Holland	IVD	assay for the detection of deletions or duplications in the <i>STRC</i> , <i>CATSPER2</i> and <i>OTOA</i> genes	Y	Not applicable

Device's component identification	Device type (IVD/MD)	Intended purpose	Applicable GSPR fully met? (Y/N)	Information about, and justification for, applicable GSPR that are not fully met
SALSA MLPA Probemix P463 MRKH (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>TBX6</i> , <i>LHX1</i> , <i>HNFB1B</i> , and <i>TBX1</i> genes	Y	Not applicable
SALSA MLPA Probemix P465 ACADM (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ACADM</i> gene	Y	Not applicable
SALSA MLPA Probemix P466 CDC73 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>CDC73</i> gene	Y	Not applicable
SALSA MLPA Probemix P471 EOFAD (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>PSEN1</i> , <i>PSEN2</i> , and <i>APP</i> genes	Y	Not applicable
SALSA MLPA Probemix P490 ADA2 (*) – MRC Holland	MD	assay for the detection of deletions or duplications in the <i>ADA2</i> gene	Y	Not applicable
QIASymphony SP - QIAGEN	IVD compatible: at CGPP we use all the available IVD reagents	automated extraction and purification of DNA from blood samples	Y	Not applicable
QIAxcel (*) - QIAGEN	MD	analyzing DNA fragments generated by PCR; verify (rapidly and accurately) the size and quantity of PCR products	Y	Not applicable
3500 Dx Genetic Analyzer - Applied Biosystems	IVD compatible: at CGPP we use all the available IVD reagents	capillary electrophoresis and automated analysis of Sanger sequencing reactions and other fluorescently labelled PCR products	Y	Not applicable
QIAgility (*) – QIAGEN	MD	DNA working dilutions	Y	Not applicable
Biomtra TProfessional Thermal cycler (*) - Biomtra	MD	PCR, Sanger sequencing, sequences purification	Y	Not applicable
Veriti 96-well Thermal cycler (*) - Applied Biosystems	MD	PCR, Sanger sequencing, sequences purification	Y	Not applicable
Biomtra Tadvanced Thermal cycler (*) – Biomtra	MD	PCR, Sanger sequencing, sequences purification	Y	Not applicable
Biomtra Tone Thermal cycler (*) – Biomtra	MD	PCR, Sanger sequencing, sequences purification	Y	Not applicable
VeritiPro 96-well Thermal Cycler (*) - Applied Biosystems	MD	PCR, Sanger sequencing, sequences purification	Y	Not applicable
TapeStation 2200 (*) - Agilent	MD	assess the quality and integrity of DNA samples	Y	Not applicable
Nanodrop 2000/2000C (*) – ThermoFisher Scientific	MD	DNA quantification	Y	Not applicable
QuantStudio 5 (*) - Applied Biosystems	MD	relative quantification of DNA molecules	Y	Not applicable
SeqScape (*) - Applied Biosystems	MD	variant identification by Sanger sequencing	Y	Not applicable
GeneMapper (*) - Applied Biosystems	MD	DNA sizing and quality allele calls – fragment analysis	Y	Not applicable

GeneMarker (*) - SoftGenetics	MD	analysis of copy number variants – MLPA	Y	Not applicable
B. Genetic tests based on whole-exome sequencing – WES (analysis of targeted multigene panels)				
C. Genetic tests based on whole-exome sequencing – WES (total WES analysis - trio; singleton)				
Emedgene (*) - Illumina	MD	variant annotation, prioritization and filtering	Y	Not applicable
GATK (*) - Broad Institute	MD	genome Analysis ToolKit (variant caller - SNVs/InDels)	Y	Not applicable
Ensembl VEP (*) - Ensembl	MD	variant effect predictor (annotations)	Y	Not applicable
GEMINI (*) - Quinlan Lab, Department of Public Health Sciences and Center for Public Health Genomics, University of Virginia, Charlottesville, Virginia, United States of America	MD	variant annotations	Y	Not applicable
VarSeq (CNV caller) – Golden Helix	IVD	CNV caller	Y	Not applicable
bcbio-nextgen (*) – open source	MD	pipeline runner and orchestrator	Y	Not applicable
A. Genetic tests based on conventional methodology (Sanger sequencing; MLPA; fragments analysis)				
B. Genetic tests based on whole-exome sequencing – WES (analysis of targeted multigene panels)				
C. Genetic tests based on whole-exome sequencing – WES (total WES analysis - trio; singleton)				
HGMD Online, Clinical Use - QIAGEN	IVD	variants literature review	Y	Not applicable
Alamut Visual Plus (*) – SOPHiA GENETICS	MD	variant annotation and interpretation	Y	Not applicable

Adapted from the document: “Guidance on the health institution exemption under Article 5(5) of Regulation (EU) 2017/745 and Regulation (EU) 2017/746”