Cell Line Information Pack (CLIP)

Cell line name	BIONi010-C-6

Purpose

The purpose of this Cell Line Information Pack (CLIP) is to communicate cell line specific information to potential users of the cell line, and to confirm that a User has received it upon the purchase of an EBiSC cell line.

Information

The CLIP may provide a variety of types of information related to an individual cell line. Of particular importance are Third Party Obligations (TPOs), which are ethical or legal obligations of a Depositor related to the use of the cell line. TPOs may impose ethical or legal limitations on the ability of a User to use the cell line, or require steps to be taken before it can be used. TPOs are likely to be:

- Obligations under license to an intellectual property rights (patent) holder, or
- Restrictions on use imposed by the donor of the primary tissue from which the cell line was made.

Third Party Obligations: donor consent provisions

None.

Third Party Obligations: IP or license provisions

iPS-AJ: This EBiSC Cell line was generated under the technology disclosed in patents related to iPS cells which are owned by Kyoto University and are licensable from iPS Academia Japan., Inc.("iPS AJ"). Commercial user (for-profit entity) acknowledges that, prior to receipt and use of this EBiSC Cell line, such commercial user needs to have an appropriate patent license from iPS AJ even for its research use. Academic user (academic or not-for-profit entity) acknowledges that such academic user does not need a patent license from iPS AJ for its research use, provided, however, that when such academic user uses this EBiSC Cell line for other than its independent research use, such academic user acknowledges that the academic user might need to obtain an appropriate patent license from iPS AJ. For inquiries to iPS AJ, please contact at license@ips-ac.co.jp.

Lonza: The following text was provided by the original provider of the primary material (Catalogue code CC-2511) from which this iPSC line was generated. Details on the primary material can be found here: "The customer is allowed to use cells purchased from Lonza, and any cells or information derived from those original cells, for any type of research (i.e. no type of research is excluded, so genetic testing is ok). They may share the original cells and any cells or information derived from those original cells as long as whomever they share them free of cost (i.e. they may not sell the cells) and whomever they share the cells/information with is also only using them for research purposes."

CRISPR: The lines have been generated using CRISPR-Cas9 technology under a license with ERS Genomics. Commercial user (for-profit entity) acknowledges that, prior to receipt and use of such EBiSC Cell line, such commercial user needs to have an appropriate patent license from ERS Genomics even for its research use. For inquiries to ERS Genomics, please contact: legalnotices@ersgenomics.com.

Other information

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Any publications or public dissemination of results using EBiSC iPSCs should be accompanied by the following acknowledgement: "The EBiSC Bank acknowledges Bioneer A/S as the source of the human induced pluripotent cell line BIONi010-C-6 which was generated with support from the EBiSC project. The EBiSC has received support from the Innovative Medicines Initiative (IMI) Joint Undertaking (JU) under grant agreement n°115582 and from the IMI-2 JU under grant agreement No 821362, resources of which are composed of financial contribution from the European Union's Seventh Framework Programme (FP7/2007-2013), European Union's Horizon 2020 research and innovation programme and EFPIA."

This APOE2/E2 (BIONi010-C-6) line) line has only a single functional APOE allele due to the presence of on-target unintended insertions caused by CRISPR gene editing in one copy of the alleles. BIONi010-C-6 is hemizygous due to a 3.4 kb insertion from the selection plasmid (used during gene editing) in the coding part of one of the two APOE alleles. The remaining (non-disrupted) allele is correctly transcribed and does drive expression of the expected APOE 2 isoform. This is characterised in Schmid B, et al. Corrigendum to "Generation of a set of isogenic, gene-edited iPSC lines homozygous for all main APOE variants and an APOE knock-out line". Stem Cell Res. 2020 Oct; 48:102005. doi: 10.1016/j.scr.2020.102005. Jan;34:101349. PMID: 32971461.

SIGN AND RETURN THIS DOCUMENT WITH YOUR COMPLETED ACCESS AND USE AGREEMENT

User acknowledgement

Please sign below to indicate that you have read and acknowledge the information contained in this CLIP.

Name _____ **Position** _____

Signature _____ **Date** _____