

## **Description of Additional Supplementary Files**

File Name: Supplementary Data 1

Description: Summary of microarray-based karyotyping data of all iPSC lines used.

File Name: Supplementary Data 2

Description: List of peptide hits and statistical analysis of proteomics data from wild-type and mutant iPSCs untreated, treated with etoposide or subjected to starvation.

File Name: Supplementary Data 3

Description: Genes significantly up-/downregulated in 3' end RNA-seq data from wildtype, patient-derived or monoallelic mutant iPSCs after starvation or etoposide treatment.

File Name: Supplementary Data 4

Description: Sequences of oligonucleotides used as gRNAs, repair templates or PCR primers.

File Name: Supplementary Data 5

Description: Genes significantly up/downregulated in ribominus RNA-seq data from patient-derived or monoallelic mutant CMs or NCCs compared to wild-type counterparts.

File Name: Supplementary Data 6

Description: Lists of significant CHD6 ChIP-seq peaks and their genomic coordinates (hg19) from all cell types and genetic backgrounds.