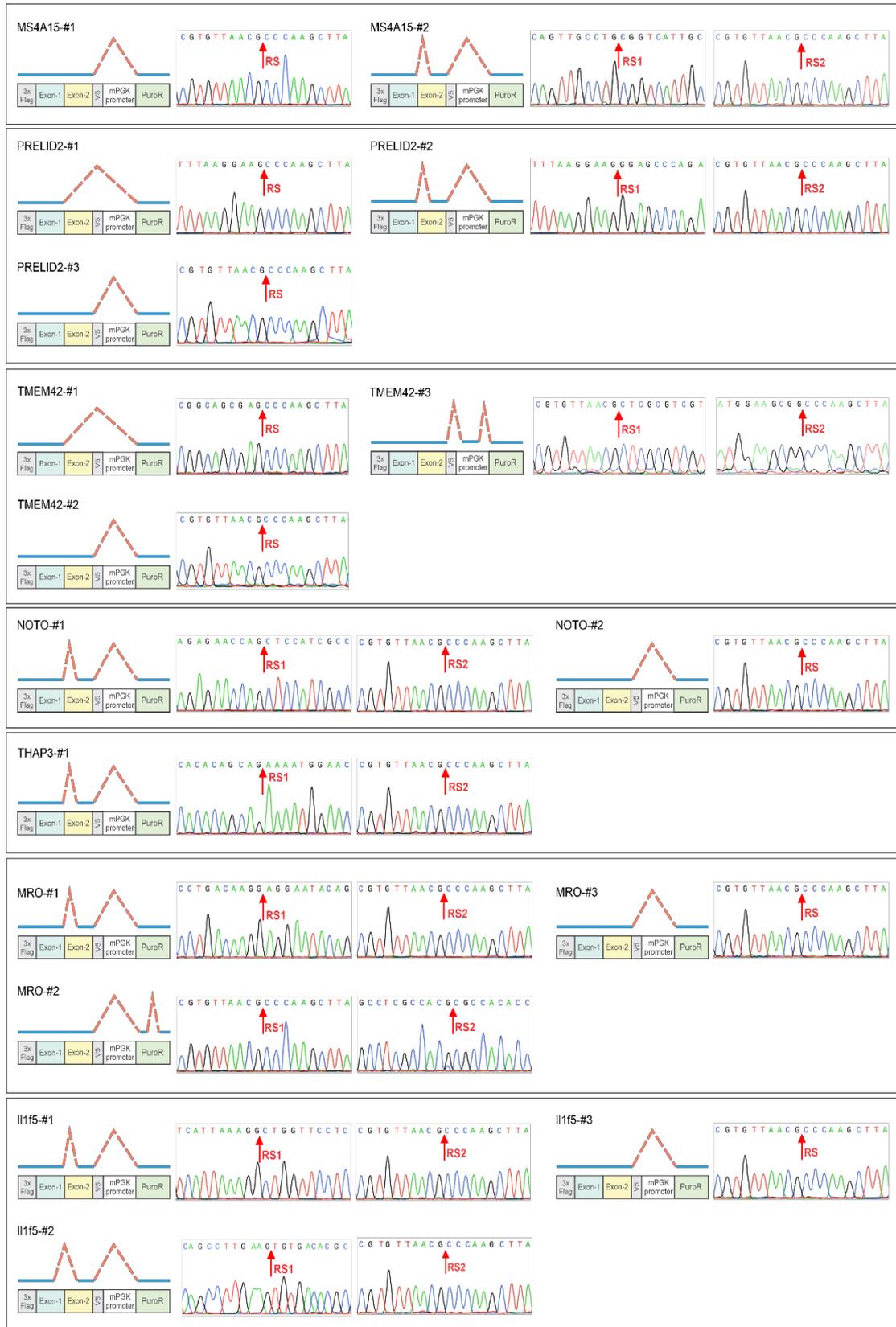


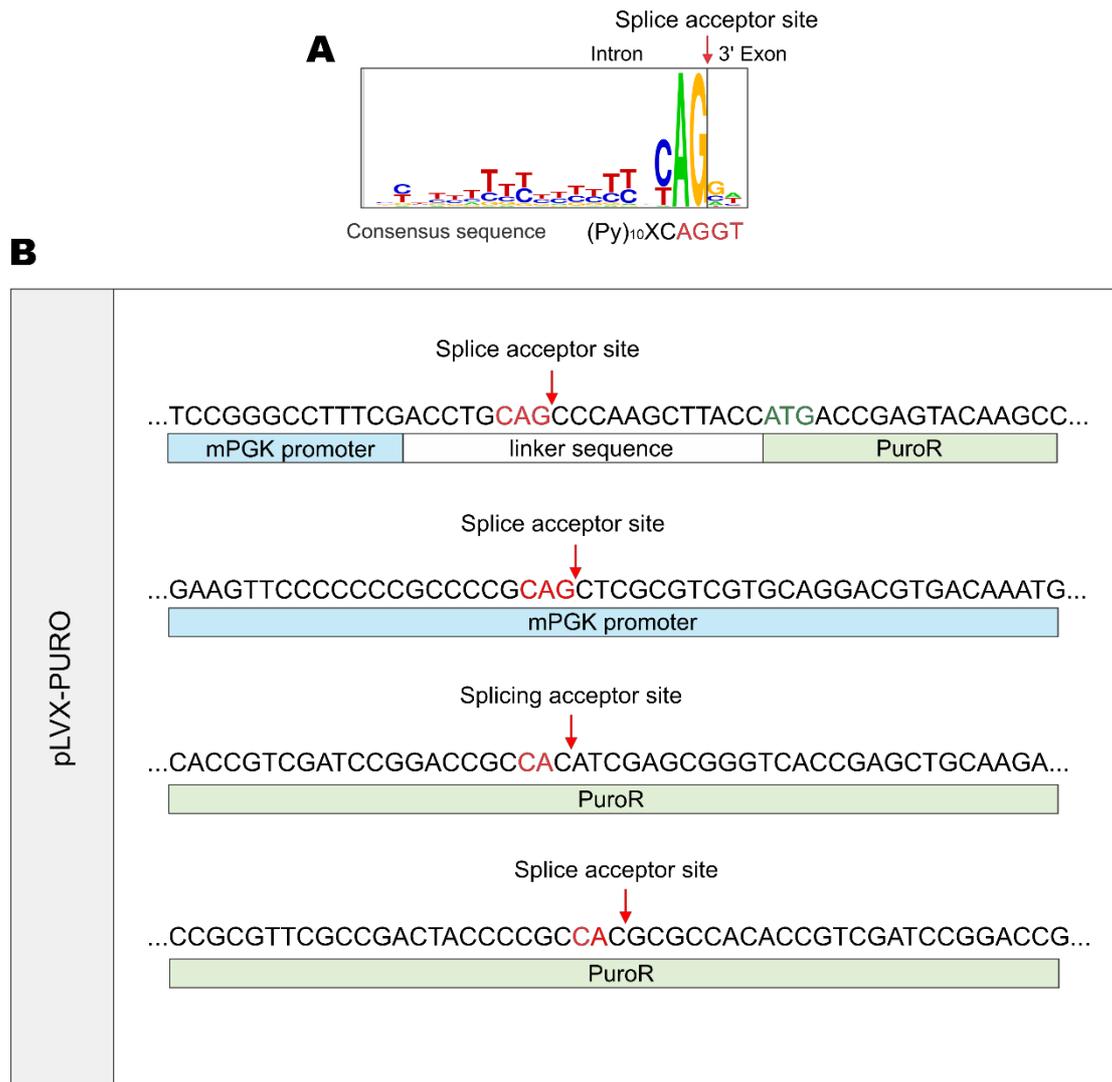
**Figure S1. Expression of large-sized hFACI proteins from different constructs. (A)** Immunoblotting. AML12 lines stably expressing V5-hFACI and mCherry were generated using pLVX-V5-hFACI and pLVX-mCherry-C1 (lanes 1-2). The AML12 stable cell line for inducible expression of V5-hFACI was generated based on pCW57-V5-hFACI (lanes 3-4). V5-hFACI expression in the three cell lines was detected by immunoblotting with anti-V5. Dox: doxycycline. **(B)** A list of human-mouse chimeric FACI constructs (left) and their protein expression (right). All chimeric constructs were cloned into the pLVX-Puro vectors with V5-tag. **(C)** A list of human-mouse chimeric FACI constructs were generated using pLVX-V5-mF1-4 (left). Protein expression from the chimeric constructs was detected by immunoblotting (right).



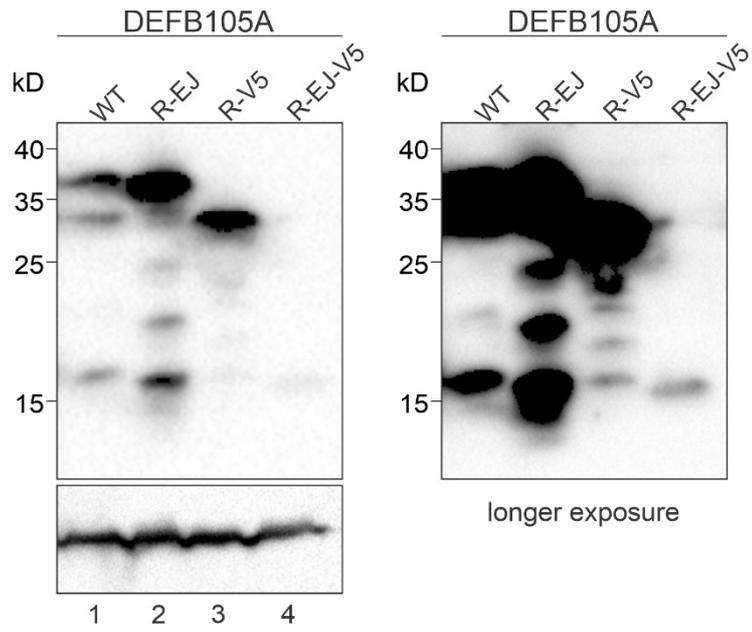
**Figure S2. Further analysis of aberrant splicing in the indicated GOIs.** Schematic diagrams and sequencing chromatograms of aberrantly spliced transcripts of 10 out of the 17 genes selected in Figure 3A. RS: the site of splice junction.



**Figure S3. Further analysis of aberrant splicing in the indicated GOIs.** Schematic diagrams and sequencing chromatograms of aberrantly spliced transcripts of the remaining 7 genes out of the 17 genes selected in Figure 3A. RS: the site of splice junction.



**Figure S4. Splice acceptor sites in pLVX-Puro.** (A) Consensus sequence of the splice acceptor site [4, 27]. (B) Four splice acceptor sites were identified on the pLVX-Puro vector by RT-PCR and sequencing. The splice acceptor site within the linker sequence between the mPGK promoter and PuroR mediates most aberrant splicing events. Rarely, two sequences in PuroR and a sequence in mPGK promoter could also function as splice acceptor sites.



**Figure S5. Western blot analysis of DEFB105A.** A longer exposure immunoblot image of DEFB105A in Figure 5D is presented to reveal faint bands.