

A

fh

GCCCC **CGGTCCG**CATGTACCGCTCCGCTCGCTCCCTGCATCGCTTCAGCGCGAGTTTGTAGATCTGCGGGCCGCTCAGAGATCCATCAAAGCTCG

gsk3b

CCACAGTGGTGGCGACTCCTGGACA **GGACCTGACCGCCG**CAGGAGGTCAGCTACACTGACACCAAGGTCATCGG

apoea

GCTGCCAGGGGCGATTCTGTTTCA **GGATGAGCCAAGAAGCCG**CTGGGAAGAGGCCCGTGGATCAGTTCTGGAACCAT

B

Gene	Nuclease ^a	Somatic mutation rate ^b	# of founder identified / # of fish screened	Germline transmission rate
<i>fh</i>	TALENs	60.0%	3/3	33.3 - 41.7%
<i>fh</i>	RGN	52.7%	2/2	100%
<i>gsk3b</i>	TALENs	0.0%		
<i>gsk3b</i>	RGNs	27.1%	3/3	43.8 - 100%
<i>apoea</i>	TALENs	20.6%	2/2	8.3 - 58.3%
<i>apoea</i>	RGNs	24.1%	1/1	31.6%

^aTALENs were injected at 300 ng/ul; RGNs were injected at the dose combination of 12.5 ng/ul sgRNA:300 ng/ul of Cas9 mRNA.

^bFor TALENs, mutation rates were determined by colony sequencing as described in [1]. For RGNs, mutation rates were determined by T7EI assays [2].

Figure S3. TALENs and RGNs for the *fh*, *gsk3b* and *apoea* genes and their somatic and germline mutation efficiencies. (A) The genomic target sequences of TALENs and RGNs in *fh*, *gsk3b* and *apoea*. TALEN target sites are underlined, while the RGN target sites are highlighted in yellow or in green if the target site is on the reverse complement strand. (B) Comparison of the somatic and germline mutation rates of TALENs and RGNs at these three genes. Except for the founder screen data of the RGNs and the TALENs targeting *apoea*, the rest of the data have been reported previously [1,2].

References

1. Cade L, Reyon D, Hwang WY, Tsai SQ, Patel S, et al. (2012) Highly efficient generation of heritable zebrafish gene mutations using homo- and heterodimeric TALENs. *Nucleic Acids Res* 40: 8001-8010.
2. Hwang WY, Fu Y, Reyon D, Maeder ML, Tsai SQ, et al. (2013) Efficient genome editing in zebrafish using a CRISPR-Cas system. *Nat Biotechnol* 31: 227-229.