

Supplementary Table 9. Reagents used in this study.

Antibodies

Antibody	Company	Catalogue number
α GM-CSF (PerCp/Cy5.5)	Biologend	502311
α IL-2 (APC)	Biologend	500310
α IL-17A (bv605)	Biologend	512325
α IL-22 (PE)	Biologend	366703
α IFN γ (APC/Cy7)	Biologend	502530
α IL-10 (bv421)	Biologend	501421
α FoxP3 (eFluor 660)	eBioscience	50-4777-42 (clone 236A/E7)
α TNF α (FITC)	Biologend	502906
α CD4 (PE-TR)	Invitrogen	MHCD0417
α CD25 (PC5)	Beckman Coulter	IM2646
α CD45RA (QD655)	Invitrogen	Q10069
α CCR7 (bv421)	Biologend	353208
α CD4 (FITC)	Beckman Coulter	A07750
α CD25 (PE)	Biologend	356103
α CD127 (PE-CF594)	BD Pharmingen	562397
α CCR6 (QD605)	Biologend	353420
α CXCR3 (APC)	BD Pharmingen	550967
α CCR4 (PE-Cy7)	BD Pharmingen	557864
α CXCR3 (AlexaFluor647)	Biologend	353712
α CCR6 (BV605)	Biologend	353420
α CD3 (TR66)	Recombinant antibody, provided by in-house facility	
α CD28	BD Pharmingen	555725
PE-Cy7 Mouse anti-NF- κ B p65 (pS529)	BD Biosciences	560335 (Clone K10-895.12.50 (RUO))
Anti-Human/Mouse phospho-I κ B α (S32/S36) eFluor® 660	eBioscience	50-9035-41
Western blot antibodies		
α DEC1 (S-8)	SantaCruz	sc-101023 (mouse)
α FLAG	Sigma-Aldrich	F1804
α GAPDH	Millipore	MAB374
α -rabbit-HRP (goat)	Sigma	A0545
α -mouse-HRP (goat)	Southern Biotech	1031-05
α NF- κ B p65 Antibody (C-20)	SantaCruz	sc-372
α NF- κ B p65 (phospho S536)	Abcam	ab76302

Primers for SYBR qPCR

Target gene	Primer fw	Primer rv
<i>CSF2</i>	5'-TGTGAATGCCATCCAGGAGG-3'	5'-GCTTGTACAGCTCCAGGCG-3'

<i>BHLHE40</i>	5'-ATTGCCCTGCAGAGTGGTTT-3'	5'-GATACTGAAGCACCTCCCGG-3'
<i>IL10</i>	5'-CCAAGACCCAGACATCAAGG-3'	5'-GGCCTTGCTCTTGTTCAC-3'
<i>IL22</i>	5'-TGCAAAAGCTGAAGGACACA-3'	5'-TTTGCTCTGGTCAAATGCAG-3'
<i>CTLA4</i>	5'-TTCATCCCTGTCTTCTGCAA-3'	5'-AGTGGCTTTGCCTGGAGAT-3'
<i>TIGIT</i>	5'-GTCATCTGCACAGCAGTCATC-3'	5'-TCCTGTCCAGCTGATTTTCTC-3'
<i>IL23R</i>	5'-TACTGGCAGCCTTGGAGTTC-3'	5'-TGCCCTGTAGAGATGGAAGC-3'
<i>IL2</i>	5'-CCCAGGGACTTAATCAGCAA-3'	5'-GGTTGCTGTCTCATCAGCAT-3'
<i>IFNG</i>	5'-CGAGATGACTTCGAAAAGCTG-3'	5'-CAGTTCAGCCATCACTTGA-3'
<i>IL17A</i>	5'-CCACCTCACCTTGGAAATCTC-3'	5'-TGGTAGTCCACGTTCCCATC-3'
<i>TNF</i>	5'-TCAGCCTCTTCTCCTTCCTG-3'	5'-GCCAGAGGGCTGATTAGAGA-3'
<i>ZC3H12D</i>	5'-TCCTGGCAGCATTGTCCTTA-3'	5'-TCTCCGGGTGGTAGAACTTG-3'
<i>ZC3H12A</i>	5'-CCACTCCAGAAGAGGAAAA-3'	5'-CGCTCCAGAAACCAGTTCAC-3'
Endogenous control		
<i>UBE2D2</i>	5'-GATCACAGTGGTCTCCAGCA-3'	5'-CGAGCAATCTCAGGCACTAA-3'

Taqman probes

Target genes	Probe ID	
<i>CSF2</i>	Hs00929873 m1	
<i>BHLHE40</i>	Hs01041212 m1	
<i>RORC</i>	Hs01076122 m1	
<i>AHR</i>	Hs00168233 m1	
<i>FOXP3</i>	Hs00203958 m1	
<i>TBX21</i>	Hs00894392 m1	
<i>IL23R</i>	Hs00332759 m1	
Endogenous control		
<i>TBP</i>	Hs99999910 m1	
Target miRNAs		
miR-146a	Hs 000468	
miR-181a	Hs 000480	
Endogenous controls		
RNU24	Hs 001001	
RNU48	Hs 001006	

Cloning primers

Name	sequence
BHLHE40 in Lenti-puro fw	5'-AATATTCTAGAATGGAGCGGATCCCCAGCGC-3'
BHLHE40 in Lenti-puro rv	5'-TTATAGAATTCGTCTTTGGTTTCTAAGTTTAAAGGGGGGATTGG-3'
BHLHE40 in Lenti-GFP fw	5'-AATATGAATTCATGGAGCGGATCCCCAGCGC-3'
BHLHE40 in Lenti-GFP rv	5'-TTATATCTAGATTAGTCTTTGGTTTCTAAGTTTAAAGGGGGGATTGG-3'

BHLHE40-1-297-stop fw	5'-GCAGCTTTCGGATGATtAAGGCCATTTtagAGCAGTGACCTGATC-3'
BHLHE40-1-297-stop rv	5'-GATCAGGTCACCTGCTctaGAAATGGCCTTaATCATCCGAAAGCTGC-3'
FlagHA into BHLHE40-GFP fw	5'-AATAAGAATTCGCCACCATGGACTACAAGGAC-3'
FlagHA into BHLHE40-GFP rv	5'-AATAAGAATTCTCCGGCGTAGTCGGGCAC-3'
miR146-fw-sponge PCR	5'-AACCCATGGACAGAGTTCTCACCGT-3'
miR146-rv-sponge PCR	5'-CTGTCCATGGGTACGGTGAGAACT-3'
BHLHE40_binding site	5'-GGTACCGAGCTCTTACGCGTGCTAGCAATCCTTGTGTCACGTGACA CAAGGATTGCTAGACGTCACGTGACACAAGGATTGCTAGCCCCGGGCTCGA GATCTGCGATCT-3'
ZC3H12D Promoter Nhe	5'-AATAAGCTAGCCTGATCATCTTCTCAGCTGGAGG-3'
ZC3H12D Promoter Xho	5'-TTATTCTCGAGCTCCTCCTGCCTCTGCCCG-3'
TNFa-3'UTR-pmirGLO NheI FW	5'-ATATGCTAGCGGAGGACGAACATCCAACCTTC-3'
TNFa-3'UTR-pmirGLO XbaI rv	5'-ATTCTAGATCTCGCCACTGAATAGTAGGGC-3'
IL22-3'UTR-pmirGLO EcoRI FW	5'-AAAGAATTCCAGAGCAAAGCTGAAAAATG-3'
IL22-3'UTR-pmirGLO XbaI rv	5'-ATCTCTAGATTAGGATATCCAAGTGTTTATTG-3'
ZC3H12D D95N mutagenesis FW	5'-GCGACCCATAGTGATTAATGGCAGCAACGTGGCGATG-3'
ZC3H12D D95N mutagenesis RV	5'-CATCGCCACGTTGCTGCCATTAATCACTATGGGTCGC-3'

miRNA mimics

name	Company	Sequence / catalogue number
siGLO green transfection indicator	Fisher Scientific	Catalogue number 11537220
miR146a mimic	Exiqon	Catalogue number 472124-001
miRNA mimic negative control	Exiqon	Catalogue number 479903-001

Plasmids

Plasmid name	Comment
empty LV	pScalps with puromycin cassette
empty GFP	pLVX-EF1 α -IRES-ZsGreen1 (Clontech 631982)

empty FlagHA-GFP	pLVX-EF1 α -FlagHA-IRES-ZsGreen1
BHLHE40-LV	pScalps- BHLHE40
BHLHE40-1-297-stop-LV	pScalps- BHLHE40-1-297
BHLHE40-GFP	pLVX-EF1 α - BHLHE40-IRES-ZsGreen1
FlagHA- BHLHE40-GFP	pLVX-EF1 α -FlagHA- BHLHE40-IRES-ZsGreen1
mCherry-miR146a/sponge-LV	pScalps-miR146a/sponge-mCherry
pLenti BHLHE40 plasmid	ABM LV089481, pLenti-GIII-EF1a
psPAX2	Addgene 12260
psMD2.G	Addgene 12259
pGL4.74[hRluc/TK] (TK Renilla)	Promega E6921
pGL3-promoter Luciferase	Promega E1761
pGL3-Luciferase-BHLHE40-4xbs	Containing 4 consensus binding sites for BHLHE40
pGL3-Basic Luciferase	Promega E1751
pGL3-Basic Luciferase-ZC3H12D-Promoter	Containing the <i>ZC3H12D</i> promoter
pGL3-Enhancer Luciferase	Promega E1771
pGL3-Enhancer Luciferase-ZC3H12D-Promoter	Containing the <i>ZC3H12D</i> promoter
pGL3-Basic Luciferase-ZC3H12D-Promoter MUT	Containing the <i>ZC3H12D</i> promoter mutated in two putative BHLHE40 binding sites
pmiRGLO Dual-Luciferase	Empty vector generated from Addgene 78131 plasmid
pmiRGLO Dual-Luciferase- <i>TNF</i> -3'UTR	Containing the 3'UTR of <i>TNF</i>
pmiRGLO Dual-Luciferase- <i>IL22</i> -3'UTR	Containing the 3'UTR of <i>IL22</i>
ZC3H12D-WT-eGFP	EF1 α -ZC3H12D-IRES-eGFP (Genecopoeia EX-Y4955-Lv165)
ZC3H12D-D95N-eGFP	Genecopoeia EX-Y4955-Lv165 containing ZC3H12D with D95N mutation

CRISPR-Cas9 deletion in primary T cells

Name	Guide RNA sequence	PAM
BHLHE40_Cas9_crRNA_gA	5'-CCCGTCTTGACTTGTACACT-3'	5'-TGG-3'
BHLHE40_Cas9_gRNA_gB	5'-CTGGAGCACGGAGACCTACC-3'	5'-AGG-3'
BHLHE40_Cas9_gRNA_gC	5'-GCCAAAGGTTTCGGAAGGTCC-3'	5'-TGG-3'
ZC3H12D_Cas9_crRNA_gA	5'-CCTGGTCAACGACGTGCTGC-3'	5'-AGG-3'
ZC3H12D_Cas9_crRNA_gB	5'-CACTATGGGTTCGAGAGAAC-3'	5'-TGG-3'

Primers for CRISPR-Cas9 knock-out screening

Name	Sequence
BHLHE40 Short-FW	5'-CTGGAGCACGGAGACCTACCAG-3'
BHLHE40 Short-Rev	5'-CCCGTCTTGACTTGTACACTTGG-3'
ZC3H12D Short-FW	5'-CTGGTCAACGACGTGCTGCAGGA-3'
ZC3H12D Short-Rev	5'-CGCAGAGAACTGGCCAGGGTT-3'
BHLHE40_T7EndoLong-FW	5'-AACTGAAGCTGCATCTCAAAGC-3'
BHLHE40_T7EndoLong-RV	5'-CAAAGGACTGGGTGGAAAGG-3'

ZC3H12D_T7EndoLong-FW	5'-AGTCTGAGAAACAAGAAACCTGTGT-3'
ZC3H12D_T7EndoLong-RV	5'-GTTAGGGACAGACTCCCAACAAG-3'

Primers for mutagenesis of two putative BHLHE40 sites in the *ZC3H12D* promoter

Name	Sequence
ZC3H12D site 1_FW	5'-CTCACGGAGGGTTTGTGTTTgATccGGTTGCTGAACTTTGATTTCG-3'
ZC3H12D site 1_RV	5'-CGAATCAAAGTTCAGCAACCggATcCAAAACAAACCTCCGTGAG-3'
ZC3H12D site 2_FW	5'-CCTCTGACTCCAGTGGgAtaTcGGGGCCTGGCCTCACT-3'
ZC3H12D site 2_RV	5'-AGTGAGGCCAGGCCCCgAtaTcCCACTGGAGTCAGAGG-3'