A

RPTEC/TERT1 tet on Cas9

FNIP1KO (+71,+71)

gFNIP1_2

WT

gFNIP2_3

FNIP2KO (-2,-7)

wt | FNIP1/2 KO clone

150 — FNIP1

37 — GAPDH

wt | FNIP1/2 KO clone

150 — FNIP2

37 — GAPDH

B

RPTEC/TERT1 tet on Cas9 FNIP1/2KO

TFE3 LAMP2

2% FCS | 0% FCS | 0% FCS minus AA | AA addback

2% FCS | 0% FCS | 0% FCS minus AA | AA addback

C

RPE/TERT1 tet on Cas9 TP53KO

wt | FLCN KO C1 | FLCN KO C2 | FLCN KO C3

100 — Vinculin

75 — FLCN

gFLCN_5

WT

FLCNKO C2 (+1)

D

RPTEC/TERT1 tet on Cas9 TP53KO

FLCNKO C3 | FLCNKO C3 + FLCN OE | FLCNKO C3 + FLCN iso 2 | FLCNKO C2 + FLCN OE | FLCNKO C2 + FLCN iso 2

100 — GPNMB

75 — FLCN

50 — Tubulin

E

RPTEC/TERT1 tet on Cas9 TP53KO FLCNKO C2 + FLCN OE

TFE3 LAMP2

2% FCS | 0% FCS | 0% FCS minus AA | AA addback

2% FCS | 0% FCS | 0% FCS minus AA | AA addback

F

RPTEC/TERT1 tet on Cas9 TP53KO

IFNγ CBA

IFNα ELISA

pg/ml

FLCNPOS | FLCNPOS + IFNγ | FLCNNEG C1 | FLCNNEG C2 | FLCNNEG C3

G

FLCNPOS

FLCNNEG

Normalized counts (RNA)

IFNAR1 | IFNAR2 | IFNGR1 | IFNGR2

H

RPTEC/TERT1 tet on Cas9 TP53KO

wt | FLCNKO C3 | FLCNKO C3 STAT1KO | wt | FLCNKO C3 | FLCNKO C3 STAT1KO

0 | 0 | 0 | 100 | 100 | 100 :IFNγ (IU/mL)

75 — pSTAT1_Y701

37 — GAPDH