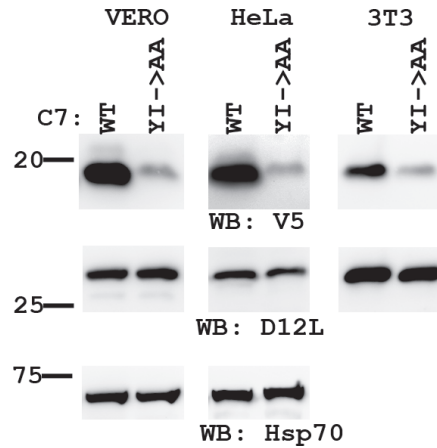
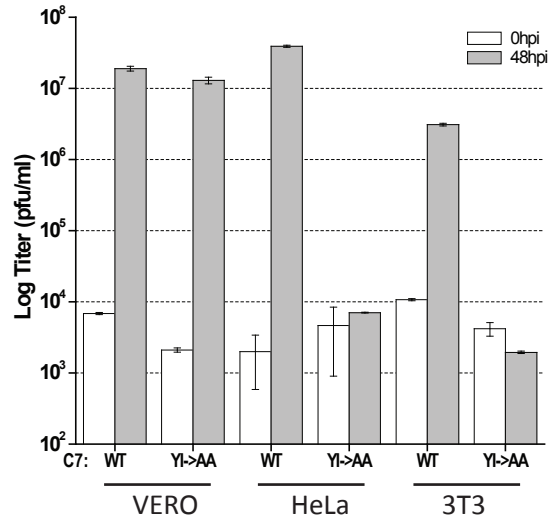


**A)**



**B)**



**Supplemental figure 1.** Tyr135 and Ile136 are critical for the function of VACV C7. **(A)**

A recombinant VACV was generated by introducing to vK1L-C7L a C7L with alanine substitutions at Tyr135 and Ile136. This virus (C7:YI->AA) or vVACV-C7L (C7:WT) were used to infect the indicated cell lines at a MOI of 5. At 8 hpi, Western blot were performed as described in legend to Fig. 2B. **(B)**. The cells were infected by the indicated viruses at a MOI of 0.5 PFU/cell. Virus titers at 0 and 48 hpi were determined by plaque assay on the permissive VERO cells.

**Supplementary Table 1.** Large scale ISG screens for VACV WT-GFP<sup>+</sup> and vK1L<sup>-</sup>C7L<sup>-</sup>/GFP<sup>+</sup>.

This table shows complete data sets from large scale ISG screens. For both viruses, replication levels, normalized to Fluc control, are shown with  $z$  scores. Data are ranked by differential  $z$  score, with genes preferentially inhibiting vK1L<sup>-</sup>C7L<sup>-</sup>/GFP<sup>+</sup> ranked first. The 100% normalization value was determined from the average of 3 independent preparations of Fluc.ires.TagRFP lentiviral pseudoparticles. Common ISG aliases: BST2 - tetherin, C19orf66 - FLJ11286, DDX58 - RIG-I, DDX60 - FLJ20035, DHX58 - LGP2, EIF2AK2 - PKR, IDO1 - INDO, IFI6 - G1P3 / 6-16, IFIH1 - MDA5, IFIT1 - ISG56, IFIT2 - ISG54, IFIT3 - ISG50, IFIT5 - ISG58, IFITM1 - 9-27, IFITM2 - 1-8D, IFITM3 - 1-8U, PSMB8 - LMP7, PSMB9 - LMP2, PSMB10 - MECL1, NAMPT - PBEF1, RSAD2 - viperin, ZBP1 - DAI, ZC3HAV1- ZAP.

Supplementary Table 1. Large scale ISG screen: VACV WT-GFP+ and vK1L-C7L-/GFP+ in Huh7 cells (Page 1 of 4)

Gene	VACV WT-GFP+		vK1L-C7L-/GFP+		Differential Z score	Gene	VACV WT-GFP+		vK1L-C7L-/GFP+		Differential Z score
	Percent Infected	Z score	Percent Infected	Z score			Percent Infected	Z score	Percent Infected	Z score	
TRIM25	140.9	3.13	73.7	-3.39	<b>-6.52</b>	KIAA1618	113.3	0.56	103.7	-0.37	<b>-0.93</b>
IRF1	120.4	1.22	87.7	-1.98	<b>-3.21</b>	CEBPD	107.3	0.00	98.1	-0.93	<b>-0.93</b>
IRF2	137.9	2.85	109.9	0.24	<b>-2.61</b>	DDIT4	100.7	-0.62	92.3	-1.52	<b>-0.90</b>
PRKD2	134.1	2.50	108.1	0.06	<b>-2.43</b>	DDX58	150.8	4.05	138.9	3.16	<b>-0.90</b>
THOC4	119.4	1.13	96.3	-1.11	<b>-2.24</b>	MT1G	98.4	-0.83	90.5	-1.70	<b>-0.87</b>
MT1F	118.6	1.06	96.1	-1.14	<b>-2.20</b>	CD69	118.4	1.03	109.1	0.17	<b>-0.87</b>
ERLIN1	117.6	0.96	97.1	-1.04	<b>-2.00</b>	Fluc	100.7	-0.62	92.8	-1.47	<b>-0.85</b>
STAT2	111.8	0.42	92.0	-1.55	<b>-1.97</b>	HERC6	111.0	0.35	102.5	-0.50	<b>-0.85</b>
FLJ39739	118.1	1.01	99.4	-0.81	<b>-1.82</b>	PUS1	111.0	0.35	102.7	-0.47	<b>-0.82</b>
CCNA1	124.5	1.60	105.5	-0.19	<b>-1.79</b>	PNPT1	104.7	-0.24	96.9	-1.06	<b>-0.82</b>
PPM1K	120.7	1.25	102.2	-0.52	<b>-1.77</b>	PI4K2B	102.4	-0.45	94.8	-1.27	<b>-0.81</b>
ADM	114.8	0.70	97.6	-0.99	<b>-1.69</b>	DHX58	116.1	0.82	107.6	0.01	<b>-0.81</b>
C1S	124.5	1.60	106.8	-0.06	<b>-1.66</b>	ABLIM3	113.1	0.54	104.8	-0.27	<b>-0.81</b>
RNF19B	112.8	0.51	96.3	-1.11	<b>-1.63</b>	MKX	117.6	0.96	109.1	0.17	<b>-0.80</b>
CCR1	122.7	1.43	106.0	-0.14	<b>-1.58</b>	OAS2	124.7	1.62	116.0	0.86	<b>-0.77</b>
UNC84B	113.1	0.54	97.4	-1.01	<b>-1.55</b>	NCF1	104.2	-0.29	97.1	-1.04	<b>-0.75</b>
IFITM3	115.6	0.77	99.9	-0.76	<b>-1.53</b>	BLVRA	102.4	-0.45	95.8	-1.16	<b>-0.71</b>
ZNF385B	117.9	0.99	102.5	-0.50	<b>-1.49</b>	IL15	100.7	-0.62	94.3	-1.32	<b>-0.70</b>
TAP1	125.2	1.67	109.6	0.22	<b>-1.45</b>	HEG1	120.4	1.22	112.7	0.52	<b>-0.70</b>
IL1RN	117.1	0.92	102.2	-0.52	<b>-1.44</b>	MCOLN2	107.5	0.02	100.7	-0.68	<b>-0.70</b>
C5orf39	120.2	1.20	105.3	-0.22	<b>-1.42</b>	HLA-C	98.7	-0.81	92.5	-1.50	<b>-0.69</b>
NOD2	111.6	0.40	97.4	-1.01	<b>-1.41</b>	CCND3	96.1	-1.04	90.2	-1.73	<b>-0.68</b>
MT1M	115.6	0.77	101.4	-0.60	<b>-1.38</b>	GMPR	108.5	0.11	102.0	-0.55	<b>-0.66</b>
ARHGEF3	116.1	0.82	102.0	-0.55	<b>-1.37</b>	SPSB1	113.8	0.61	107.1	-0.04	<b>-0.65</b>
HSH2D	122.9	1.46	108.6	0.11	<b>-1.34</b>	CYP1B1	111.8	0.42	105.3	-0.22	<b>-0.64</b>
TNFAIP3	114.8	0.70	101.2	-0.63	<b>-1.33</b>	IFI35	112.3	0.47	105.8	-0.17	<b>-0.63</b>
LIPA	109.0	0.16	96.1	-1.14	<b>-1.30</b>	FUT4	117.6	0.96	110.9	0.34	<b>-0.62</b>
TIMP1	122.2	1.39	108.3	0.09	<b>-1.30</b>	FNDC4	101.4	-0.55	95.8	-1.16	<b>-0.62</b>
SCARB2	115.9	0.80	102.5	-0.50	<b>-1.30</b>	FAM134B	106.2	-0.10	100.4	-0.70	<b>-0.60</b>
SAMD4A	114.3	0.66	101.2	-0.63	<b>-1.28</b>	DUSP5	107.0	-0.03	101.2	-0.63	<b>-0.60</b>
C5orf27	113.1	0.54	100.4	-0.70	<b>-1.24</b>	TBX3	130.5	2.17	123.1	1.57	<b>-0.59</b>
CRY1	96.1	-1.04	84.9	-2.26	<b>-1.22</b>	TNFAIP6	114.3	0.66	108.1	0.06	<b>-0.59</b>
NPAS2	116.6	0.87	104.0	-0.35	<b>-1.21</b>	SNN	121.2	1.29	114.4	0.70	<b>-0.59</b>
IRF7	113.6	0.59	101.4	-0.60	<b>-1.19</b>	NUP50	119.9	1.18	113.4	0.60	<b>-0.57</b>
UPP2	120.9	1.27	108.6	0.11	<b>-1.16</b>	STAP1	110.3	0.28	104.5	-0.29	<b>-0.57</b>
HLA-F	115.6	0.77	103.7	-0.37	<b>-1.15</b>	CRP	117.6	0.96	111.4	0.40	<b>-0.57</b>
C4orf32	119.4	1.13	107.3	-0.01	<b>-1.14</b>	LGALS9	111.8	0.42	106.0	-0.14	<b>-0.56</b>
G6PC	120.4	1.22	108.3	0.09	<b>-1.13</b>	GCH1	115.1	0.73	109.1	0.17	<b>-0.56</b>
RSAD2	107.3	0.00	96.3	-1.11	<b>-1.11</b>	P2RY6	116.4	0.84	110.4	0.29	<b>-0.55</b>
GBP4	108.3	0.09	97.4	-1.01	<b>-1.10</b>	ATF3	107.0	-0.03	101.7	-0.58	<b>-0.55</b>
IL6ST	124.7	1.62	112.7	0.52	<b>-1.10</b>	CD163	126.0	1.74	119.5	1.21	<b>-0.53</b>
PTMA	111.8	0.42	100.7	-0.68	<b>-1.10</b>	NAPA	100.2	-0.67	95.6	-1.19	<b>-0.52</b>
ULK4	118.6	1.06	107.3	-0.01	<b>-1.07</b>	CLEC2B	106.7	-0.05	101.7	-0.58	<b>-0.52</b>
MICB	103.5	-0.36	93.3	-1.42	<b>-1.06</b>	RGS1	103.7	-0.34	98.9	-0.86	<b>-0.52</b>
EHD4	101.9	-0.50	92.0	-1.55	<b>-1.05</b>	C10orf10	113.6	0.59	108.1	0.06	<b>-0.52</b>
NMI	100.7	-0.62	91.0	-1.65	<b>-1.03</b>	TAGAP	123.9	1.55	117.8	1.04	<b>-0.52</b>
PIM3	110.8	0.33	100.7	-0.68	<b>-1.00</b>	HK2	115.3	0.75	109.9	0.24	<b>-0.51</b>
KIAA0082	112.1	0.44	102.0	-0.55	<b>-0.99</b>	RIK2	119.6	1.15	113.9	0.65	<b>-0.50</b>
MTHFD2L	101.7	-0.52	92.5	-1.50	<b>-0.97</b>	APOL2	116.9	0.89	111.4	0.40	<b>-0.50</b>
CX3CL1	112.6	0.49	102.7	-0.47	<b>-0.96</b>	GLRX	120.2	1.20	114.4	0.70	<b>-0.50</b>

Supplementary Table 1. Large scale ISG screen: VACV WT-GFP+ and vK1L-C7L-/GFP+ in Huh7 cells (Page 2 of 4)

Gene	VACV WT-GFP+		vK1L-C7L-/GFP+		Differential Z score	Gene	VACV WT-GFP+		vK1L-C7L-/GFP+		Differential Z score
	Percent Infected	Z score	Percent Infected	Z score			Percent Infected	Z score	Percent Infected	Z score	
GBP5	105.0	-0.22	100.4	-0.70	-0.49	MT1H	91.3	-1.49	91.8	-1.57	-0.08
TMEM51	118.4	1.03	112.9	0.55	-0.48	SIRPA	101.4	-0.55	101.2	-0.63	-0.08
FAM125B	121.7	1.34	116.0	0.86	-0.48	CES1	108.5	0.11	108.1	0.06	-0.05
SLC16A1	108.8	0.14	104.0	-0.35	-0.48	FCGR1A	109.0	0.16	108.6	0.11	-0.05
ISG20	115.9	0.80	110.6	0.32	-0.48	UNC93B1	104.2	-0.29	104.2	-0.32	-0.03
GAK	104.5	-0.26	100.2	-0.73	-0.47	IL17RB	117.4	0.94	116.5	0.91	-0.03
LMO2	127.7	1.91	121.8	1.44	-0.46	HLA-E	101.2	-0.57	101.4	-0.60	-0.03
LRG1	89.5	-1.66	86.4	-2.11	-0.45	SERPINB9	91.6	-1.47	92.5	-1.50	-0.03
HES4	96.4	-1.02	93.0	-1.45	-0.43	COMMD3	108.3	0.09	108.1	0.06	-0.03
MAX	101.7	-0.52	98.1	-0.93	-0.41	TYMP	93.1	-1.33	94.1	-1.34	-0.02
NT5C3	99.9	-0.69	97.1	-1.04	-0.35	SLC25A28	100.2	-0.67	100.7	-0.68	-0.01
FAM46A	106.7	-0.05	103.5	-0.40	-0.35	ALDH1A1	109.0	0.16	109.1	0.17	0.01
SLC25A30	116.6	0.87	112.7	0.52	-0.34	Fluc	106.0	-0.12	106.3	-0.12	0.01
DNAPTP6	113.8	0.61	110.1	0.27	-0.34	TRIM14	97.6	-0.90	98.6	-0.88	0.02
IFITM3	112.1	0.44	108.6	0.11	-0.33	SERPINE1	99.9	-0.69	100.9	-0.65	0.04
STAT1	112.6	0.49	109.1	0.17	-0.33	HPSE	100.9	-0.59	102.0	-0.55	0.04
SLC15A3	99.7	-0.71	97.1	-1.04	-0.32	PDGFRL	105.2	-0.19	106.0	-0.14	0.05
CCL2	107.5	0.02	104.5	-0.29	-0.31	SECTM1	106.0	-0.12	106.8	-0.06	0.06
FBXO6	112.3	0.47	109.1	0.17	-0.30	WARS	96.4	-1.02	97.9	-0.96	0.06
HLA-G	103.5	-0.36	100.9	-0.65	-0.29	ATP10D	103.2	-0.38	104.2	-0.32	0.06
GBP2	107.3	0.00	104.5	-0.29	-0.29	LAP3	103.2	-0.38	104.2	-0.32	0.06
PRIC285	101.4	-0.55	99.2	-0.83	-0.28	AXUD1	110.3	0.28	110.9	0.34	0.07
CD38	111.6	0.40	108.6	0.11	-0.28	USP18	97.4	-0.93	98.9	-0.86	0.07
ADFP	101.9	-0.50	99.7	-0.78	-0.28	GBP3	107.3	0.00	108.1	0.06	0.07
JUNB	97.1	-0.95	95.3	-1.22	-0.27	GCA	107.3	0.00	108.1	0.06	0.07
PSCD1	97.4	-0.93	95.6	-1.19	-0.26	CFB	118.4	1.03	118.5	1.11	0.08
C4orf33	114.3	0.66	111.4	0.40	-0.26	IFNGR1	125.5	1.69	125.1	1.78	0.08
ZBP1	111.3	0.37	108.6	0.11	-0.26	PNRC1	97.4	-0.93	99.2	-0.83	0.09
MAFB	114.8	0.70	111.9	0.45	-0.26	IFIT1	99.7	-0.71	101.4	-0.60	0.11
C9orf91	103.2	-0.38	101.2	-0.63	-0.25	ANGPTL1	106.2	-0.10	107.6	0.01	0.11
RARRES3	106.0	-0.12	104.0	-0.35	-0.22	FAM46C	117.1	0.92	117.8	1.04	0.12
IRF9	109.8	0.23	107.6	0.01	-0.22	ISG15	94.6	-1.18	96.9	-1.06	0.12
C9orf19	136.3	2.71	132.3	2.49	-0.22	BST2	122.2	1.39	122.6	1.52	0.13
BTN3A3	112.1	0.44	109.9	0.24	-0.20	GJA4	109.8	0.23	111.1	0.37	0.14
CASP7	109.5	0.21	107.6	0.01	-0.20	AMPH	114.6	0.68	115.7	0.83	0.15
GTPBP2	107.0	-0.03	105.3	-0.22	-0.19	WHDC1	95.1	-1.14	97.6	-0.99	0.15
PSMB9	121.2	1.29	118.5	1.11	-0.18	TAP2	104.2	-0.29	106.3	-0.12	0.17
CSDA	99.2	-0.76	98.1	-0.93	-0.17	NRN1	105.7	-0.15	107.8	0.04	0.18
ARG2	105.7	-0.15	104.2	-0.32	-0.17	OPTN	97.1	-0.95	99.9	-0.76	0.19
B4GALT5	109.8	0.23	108.1	0.06	-0.17	VEGFC	94.1	-1.23	97.1	-1.04	0.20
RASSF4	107.3	0.00	105.8	-0.17	-0.16	BAG1	104.0	-0.31	106.3	-0.12	0.20
MARCKS	112.3	0.47	110.6	0.32	-0.15	ADAMDEC1	97.6	-0.90	100.4	-0.70	0.20
GK	107.0	-0.03	105.8	-0.17	-0.14	OASL	134.1	2.50	134.3	2.70	0.20
PCTK2	113.6	0.59	111.9	0.45	-0.14	LGALS3	99.2	-0.76	102.0	-0.55	0.21
SAMHD1	107.5	0.02	106.3	-0.12	-0.13	SSBP3	96.1	-1.04	99.2	-0.83	0.21
XAF1	115.3	0.75	113.7	0.63	-0.12	ETV7	92.3	-1.40	95.8	-1.16	0.23
STARD5	105.7	-0.15	104.8	-0.27	-0.12	MX1	113.3	0.56	115.5	0.81	0.24
AKT3	90.6	-1.56	91.0	-1.65	-0.09	PBEF1	108.3	0.09	111.1	0.37	0.28
ANKRD22	116.9	0.89	115.5	0.81	-0.09	IL28RA	99.4	-0.74	103.0	-0.45	0.29
RAB27A	107.5	0.02	106.8	-0.06	-0.08	JAK2	113.3	0.56	116.2	0.88	0.32

Supplementary Table 1. Large scale ISG screen: VACV WT-GFP+ and vK1L-C7L-/GFP+ in Huh7 cells (Page 3 of 4)

Gene	VACV WT-GFP+		vK1L-C7L-/GFP+		Differential Z score	Gene	VACV WT-GFP+		vK1L-C7L-/GFP+		Differential Z score
	Percent Infected	Z score	Percent Infected	Z score			Percent Infected	Z score	Percent Infected	Z score	
NFIL3	98.4	-0.83	102.5	-0.50	<b>0.33</b>	LAMP3	96.9	-0.97	104.2	-0.32	<b>0.65</b>
EIF3EIP	105.2	-0.19	108.8	0.14	<b>0.33</b>	FLJ23556	94.4	-1.21	102.0	-0.55	<b>0.66</b>
SLFN5	121.7	1.34	124.1	1.67	<b>0.33</b>	TXNIP	111.3	0.37	118.0	1.06	<b>0.69</b>
IMPA2	103.2	-0.38	107.1	-0.04	<b>0.34</b>	TREX1	96.1	-1.04	104.0	-0.35	<b>0.70</b>
IGFBP2	98.9	-0.78	103.2	-0.42	<b>0.36</b>	AGPAT9	90.3	-1.59	98.6	-0.88	<b>0.70</b>
SP110	104.0	-0.31	108.1	0.06	<b>0.37</b>	MAFF	117.6	0.96	124.1	1.67	<b>0.71</b>
STEAP4	97.6	-0.90	102.2	-0.52	<b>0.38</b>	MYD88	92.1	-1.42	100.4	-0.70	<b>0.72</b>
IFI44	100.9	-0.59	105.3	-0.22	<b>0.38</b>	ETV6	96.4	-1.02	104.5	-0.29	<b>0.72</b>
MAB21L2	110.8	0.33	114.4	0.70	<b>0.38</b>	EXT1	112.1	0.44	119.3	1.19	<b>0.74</b>
CCDC109B	104.7	-0.24	108.8	0.14	<b>0.38</b>	PSMB8	112.6	0.49	119.8	1.24	<b>0.75</b>
APOBEC3A	111.8	0.42	115.5	0.81	<b>0.38</b>	THBD	87.5	-1.85	96.6	-1.09	<b>0.76</b>
TNFSF10	122.4	1.41	125.4	1.80	<b>0.39</b>	SOCS1	97.4	-0.93	105.8	-0.17	<b>0.76</b>
CXCL9	107.8	0.04	111.9	0.45	<b>0.40</b>	PRAME	90.1	-1.61	99.2	-0.83	<b>0.78</b>
ADAR	111.3	0.37	115.2	0.78	<b>0.41</b>	SOCS2	106.5	-0.08	114.4	0.70	<b>0.78</b>
GALNT2	95.1	-1.14	100.2	-0.73	<b>0.41</b>	RBCK1	89.3	-1.68	98.6	-0.88	<b>0.80</b>
IFI30	95.9	-1.07	100.9	-0.65	<b>0.41</b>	PADI2	96.4	-1.02	105.3	-0.22	<b>0.80</b>
CD74	96.1	-1.04	101.2	-0.63	<b>0.42</b>	AIM2	97.4	-0.93	106.3	-0.12	<b>0.81</b>
CD80	116.1	0.82	119.8	1.24	<b>0.42</b>	ZNF313	128.5	1.98	135.3	2.80	<b>0.82</b>
CPT1A	99.2	-0.76	104.2	-0.32	<b>0.44</b>	BCL3	93.6	-1.28	103.2	-0.42	<b>0.86</b>
UBE2L6	101.4	-0.55	106.5	-0.09	<b>0.46</b>	CCL5	103.5	-0.36	112.4	0.50	<b>0.86</b>
PML	109.0	0.16	113.7	0.63	<b>0.47</b>	SPTLC2	103.5	-0.36	112.4	0.50	<b>0.86</b>
TMEM140	119.6	1.15	123.6	1.62	<b>0.47</b>	PMAIP1	97.6	-0.90	107.1	-0.04	<b>0.86</b>
C22orf28	100.2	-0.67	105.5	-0.19	<b>0.47</b>	S100A8	95.1	-1.14	104.8	-0.27	<b>0.87</b>
MSR1	114.8	0.70	119.3	1.19	<b>0.49</b>	ABTB2	118.6	1.06	126.7	1.93	<b>0.87</b>
TRIM5	92.1	-1.42	98.1	-0.93	<b>0.49</b>	EPAS1	119.9	1.18	128.0	2.06	<b>0.88</b>
TNFRSF10A	122.2	1.39	126.2	1.88	<b>0.49</b>	GBP1	101.4	-0.55	110.9	0.34	<b>0.89</b>
C6orf150	99.4	-0.74	105.0	-0.24	<b>0.49</b>	PFKFB3	99.4	-0.74	109.1	0.17	<b>0.90</b>
IFI27	106.5	-0.08	111.6	0.42	<b>0.50</b>	CXCL10	112.1	0.44	121.1	1.37	<b>0.92</b>
IFI44L	126.5	1.79	130.2	2.29	<b>0.50</b>	PXK	92.6	-1.37	103.0	-0.45	<b>0.93</b>
PPM1K	103.7	-0.34	109.1	0.17	<b>0.50</b>	PHF15	114.1	0.63	123.1	1.57	<b>0.94</b>
IFI6	107.5	0.02	112.9	0.55	<b>0.53</b>	TDRD7	111.8	0.42	121.1	1.37	<b>0.95</b>
IFIT5	95.1	-1.14	101.4	-0.60	<b>0.54</b>	DCP1A	118.9	1.08	127.7	2.03	<b>0.95</b>
TNSF13B	108.3	0.09	113.7	0.63	<b>0.54</b>	APOL1	97.4	-0.93	107.8	0.04	<b>0.96</b>
GTPBP1	99.7	-0.71	105.8	-0.17	<b>0.55</b>	SCO2	95.6	-1.09	106.3	-0.12	<b>0.97</b>
IL15RA	106.2	-0.10	111.9	0.45	<b>0.55</b>	LY6E	113.1	0.54	122.6	1.52	<b>0.98</b>
NDC80	112.8	0.51	118.0	1.06	<b>0.55</b>	ACSL1	101.4	-0.55	111.9	0.45	<b>0.99</b>
SLC1A1	99.9	-0.69	106.0	-0.14	<b>0.55</b>	MCL1	88.3	-1.77	100.2	-0.73	<b>1.04</b>
AQP9	91.6	-1.47	98.4	-0.91	<b>0.56</b>	KIAA0040	105.2	-0.19	116.0	0.86	<b>1.05</b>
TRIM21	95.1	-1.14	101.7	-0.58	<b>0.56</b>	B2M	97.6	-0.90	109.1	0.17	<b>1.07</b>
IFIT2	99.4	-0.74	105.8	-0.17	<b>0.57</b>	PLEKHA4	101.7	-0.52	113.2	0.57	<b>1.10</b>
CD274	87.3	-1.87	94.6	-1.29	<b>0.58</b>	MT1X	100.7	-0.62	112.4	0.50	<b>1.12</b>
SMAD3	98.4	-0.83	105.0	-0.24	<b>0.59</b>	IL1R	97.6	-0.90	109.6	0.22	<b>1.12</b>
VAMP5	103.0	-0.41	109.3	0.19	<b>0.60</b>	TRAFD1	92.8	-1.35	105.3	-0.22	<b>1.13</b>
LGMN	97.1	-0.95	104.0	-0.35	<b>0.60</b>	CCL8	96.6	-1.00	109.1	0.17	<b>1.16</b>
AHNAK2	97.9	-0.88	104.8	-0.27	<b>0.61</b>	OGFR	81.7	-2.39	95.3	-1.22	<b>1.17</b>
CLEC4E	83.2	-2.25	91.2	-1.62	<b>0.62</b>	CLEC4D	91.8	-1.44	104.8	-0.27	<b>1.17</b>
Fluc	93.3	-1.30	100.7	-0.68	<b>0.62</b>	GEM	96.4	-1.02	109.3	0.19	<b>1.21</b>
SAT3	108.8	0.14	115.2	0.78	<b>0.64</b>	ARNTL	75.9	-2.93	90.5	-1.70	<b>1.23</b>
NCOA3	121.9	1.36	127.4	2.01	<b>0.64</b>	CD9	108.8	0.14	121.1	1.37	<b>1.23</b>
GZMB	113.1	0.54	119.3	1.19	<b>0.65</b>	CCL19	107.3	0.00	119.8	1.24	<b>1.24</b>

Supplementary Table 1. Large scale ISG screen: VACV WT-GFP+ and vK1L-C7L-/GFP+ in Huh7 cells (Page 4 of 4)

Gene	VACV WT-GFP+		vK1L-C7L-/GFP+		Differential Z score	Gene	VACV WT-GFP+		vK1L-C7L-/GFP+		Differential Z score
	Percent Infected	Z score	Percent Infected	Z score			Percent Infected	Z score	Percent Infected	Z score	
GPX2	99.9	-0.69	113.2	0.57	<b>1.26</b>						
CTCF	99.2	-0.76	112.7	0.52	<b>1.28</b>						
CCDC75	113.8	0.61	126.4	1.90	<b>1.30</b>						
HESX1	107.8	0.04	120.8	1.34	<b>1.30</b>						
ENPP1	118.6	1.06	131.0	2.37	<b>1.31</b>						
TMEM49	94.1	-1.23	108.6	0.11	<b>1.35</b>						
TRIM34	92.6	-1.37	107.3	-0.01	<b>1.36</b>						
ANKFY1	132.0	2.31	144.0	3.67	<b>1.36</b>						
DEFB1	109.5	0.21	123.1	1.57	<b>1.36</b>						
PDK1	92.1	-1.42	107.1	-0.04	<b>1.38</b>						
CMAH	102.4	-0.45	116.7	0.93	<b>1.39</b>						
PMM2	97.1	-0.95	112.1	0.47	<b>1.42</b>						
SAA1	94.1	-1.23	109.3	0.19	<b>1.42</b>						
DYNLT1	101.9	-0.50	116.7	0.93	<b>1.43</b>						
LEPR	104.5	-0.26	119.5	1.21	<b>1.48</b>						
RPL22	103.5	-0.36	119.5	1.21	<b>1.57</b>						
CCL4	96.4	-1.02	113.2	0.57	<b>1.59</b>						
INDO	91.3	-1.49	108.8	0.14	<b>1.63</b>						
SERPING1	95.9	-1.07	113.9	0.65	<b>1.72</b>						
TRIM38	91.3	-1.49	110.1	0.27	<b>1.76</b>						
CDKN1A	112.3	0.47	129.7	2.24	<b>1.77</b>						
C15orf48	103.7	-0.34	122.1	1.47	<b>1.81</b>						
LINCR	90.6	-1.56	110.6	0.32	<b>1.88</b>						
IFITM2	89.8	-1.63	110.6	0.32	<b>1.95</b>						
CREB3L3	90.1	-1.61	111.6	0.42	<b>2.03</b>						
PLSCR1	108.3	0.09	128.7	2.14	<b>2.05</b>						
FKBP5	100.4	-0.64	122.6	1.52	<b>2.16</b>						
C2orf31	97.4	-0.93	122.3	1.50	<b>2.42</b>						
MAP3K5	109.8	0.23	137.6	3.03	<b>2.80</b>						
CHMP5	111.8	0.42	139.9	3.26	<b>2.84</b>						