## Supplementary materials:

Туре	Cell line	Description			
Breast cancer	MDA-MB-231	Her2 (-)			
	MDA-MB-453	Her2 (+)			
Cervical cancer	C-33A	Cervical adenocarcinoma (No infection)			
	Hala	Cervical adenocarcinoma (HPV-18			
	пета	infection)			
	0.11-	Cervical adenocarcinoma (HPV-16			
	ыпа	infection)			
Colorectal cancer	HCT-8	Ileocecal colorectal adenocarcinoma			
	SW480	Dukes' type B, colorectal adenocarcinoma			
Gastric cancer	AGS	Gastric carcinoma			
	MKN45	Gastric carcinoma, poorly differentiated			
Glioma	LN-18	Glioblastoma; glioma(IV)			
	SNB19	Astrocytoma cells			
	U118MG	Glioblastoma; astrocytoma (IV)			
	U251	Astrocytoma cells			
	U87MG	Glioblastoma; astrocytoma (IV)			
Hepatocellular	HepG2	Hepatocellular carcinoma (No infection)			
cancer	Hep3B	Hepatocellular carcinoma (HBV infection)			
Lung cancer	A549	Lung carcinoma			
	NCI-H460	Large cell lung cancer			
Ovarian cancer	Caov-3	adenocarcinoma (ovary)			
	SK-OV-3	adenocarcinoma (ovary: ascites)			
Pancreatic cancer	CFPAC-1	Ductal adenocarcinoma			
	PNAC-1	Pancreatic carcinoma			

Tahle	<b>S1</b>	Human	enithelial	cancer	cell	lines	used	in	this	study	J
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Table 52. Nucleic actus useu in tins study						
Name	Туре	Description				
5'ppp-dsRNA	dsRNA	a synthetic ligand for RIG-I [1].				
Poly(A:U)	A synthetic double	signals only through TLR3 [2].				
	stranded RNA					
Poly(I:C)	A synthetic double	recognized by TLR3 and cytosolic RNA helicases				
	stranded RNA	RIG-I and MDA-5 [3, 4].				
ISD Naked	Non-CpG oligomer from	mediated by the STING-TBK1-IRF3 signaling axis				
	the Listeria	[5, 6].				
	monocytogenes genome.					
HSV-60	A 60 bp oligonucleotide	induces IFN- $\beta$ in a TLR-independent, but STING-,				
Naked	containing viral DNA	TBK1- and IRF3-dependent manner. HSV-60 is				
	motifs	recognized by DDX41 and IFI16 [7, 8].				
VACV-70	A 70 bp oligonucleotides	induces IFN- $\beta$ in a TLR-independent, but STING-,				
Naked	containing viral DNA	TBK1- and IRF3-dependent manner. VACV-70 is				
	motifs	recognized by DDX41 and IFI16 [7, 8].				
Poly(dG:dC)	A synthetic analog of	recognized by cGAS, DAI, DDX41, IFI16,				
• • •	Z-DNA	LRRFIP1, and AIM2 [7-12].				
3'3'-cGAMP	A cyclic di-nucleotide	binds STING and subsequently induces				
		TBK1-IRF3-dependent production of IFN-β [13].				
Poly(dA:dT)	A synthetic analog of	recognized by several sensors, including cGAS,				
	B-DNA	DAI, DDX41, IFI16 and LRRFIP1, AIM2 and				
		RIG-I [7-12, 14-16].				

### Table S2. Nucleic acids used in this study

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#### Supplementary figure legends:

#### Figure S1. Nucleic acids transfected into human epithelial cancer cells

(A) MDA-MB-231, (B) MDA-MB-453, (C) C-33A, (D) HeLa, (E) SiHa, (F) HCT-8, (G) SW480, (H) AGS, (I) MKN45, (J) LN-18, (K) SNB19, (L) U87MG, (M) U118MG, (N) U251MG, (O) HepG2, (P) Hep3B, (Q) Caov-3, (R) SK-OV-3, (S) A549, (T) NCI-H460, (U) CFPAC-1, and (V) PANC-1 cells were transfected with 0.1  $\mu$ g/ml rhodamine-labeled poly(dA:dT) for 3 h, then imaged by fluorescence confocal microscope. (W) PANC-1 cells were cocultured with 0.1  $\mu$ g/ml rhodamine-labeled poly(dA:dT) for 3 h, then imaged by fluorescence as a negative control. Data are representative of at least three independent fields.

#### Figure S2. IFN-β is induced in dose-dependent manner

(A) 5'ppp-dsRNA, (B) poly(A:U), (C) poly(I:C)-HMW, (D) poly(I:C)-LMW, (E) ISD, (F) HSV60, (G) VACV-70, (H) poly(dG:dC), (I) 3'-3'cGAMP, and (J) poly(dA:dT) were transfected into PANC-1 cells at concentrations of 0.625, 1.25, 2.5, 5, 10, and 20  $\mu$ g/ml. Eighteen hours later, cell supernatants were harvested, and secretion of IFN- $\beta$  was detected by ELISA. Data are shown as mean  $\pm$  SD of at least three independent experiments (\* p < 0.05, \*\* p < 0.01).

# Figure S3. RNA polymerase III is not involved in the RIG-I signaling pathway in human epithelial cancer cells

293T (A), PANC-1 (B), and A549 (C) cells were treated with DMSO or ML-60218 for 12 h and then transfected with poly(dA:dT). After 18 h, total RNA was extracted, and the expression of IFN- $\beta$  determined.

#### Figure S4. Schema for stimuli of PRRs and their signaling pathways

The TLR3-TRIF, RIG-I-MAVS, and cGAS-STING pathways are activated by their ligands. Adaptors are then recruited, and TBK1 is phosphorylated. IRF3 is activated by phosphorylated TBK1. pIRF3 is translocated into the nucleus and binds to the IFN- $\beta$  promoter.