

SUPPLEMENTARY TABLES

Supplementary Table 1. PANoptosis-related genes.

PRGs	Source
<i>ADAR</i>	Karki, R., Sundaram, B., Sharma, B.R., et al. (2021). ADAR1 restricts ZBP1-mediated immune response and PANoptosis to promote tumorigenesis. <i>Cell Rep</i> 37(3), 109858. https://doi.org/10.1016/j.celrep.2021.109858 .
<i>AIM2</i>	Lee, S., Karki, R., Wang, Y., et al. (2021). AIM2 forms a complex with pyrin and ZBP1 to drive PANoptosis and host defence. <i>Nature</i> 597(7876), 415-419. https://doi.org/10.1038/s41586-021-03875-8 .
<i>CASP1</i>	Zheng, M., Karki, R., Vogel, P., et al. (2020). Caspase-6 Is a Key Regulator of Innate Immunity, Inflammasome Activation, and Host Defense. <i>Cell</i> 181(3), 674-687.e613. https://doi.org/10.1016/j.cell.2020.03.040 .
<i>CASP4</i>	Christgen, S., Zheng, M., Kesavardhana, S., et al. (2020). Identification of the PANoptosome: A Molecular Platform Triggering Pyroptosis, Apoptosis, and Necroptosis (PANoptosis). <i>Front Cell Infect Microbiol</i> 10, 237. https://doi.org/10.3389/fcimb.2020.00237 .
<i>CASP3</i>	Zheng, M., Karki, R., Vogel, P., et al. (2020). Caspase-6 Is a Key Regulator of Innate Immunity, Inflammasome Activation, and Host Defense. <i>Cell</i> 181(3), 674-687.e613. https://doi.org/10.1016/j.cell.2020.03.040 .
<i>CASP6</i>	Zheng, M., Karki, R., Vogel, P., et al. (2020). Caspase-6 Is a Key Regulator of Innate Immunity, Inflammasome Activation, and Host Defense. <i>Cell</i> 181(3), 674-687.e613. https://doi.org/10.1016/j.cell.2020.03.040 .
<i>CASP8</i>	Malireddi, R.K.S., Kesavardhana, S., and Kanneganti, T.D. (2019). ZBP1 and TAK1: Master Regulators of NLRP3 Inflammasome/Pyroptosis, Apoptosis, and Necroptosis (PAN-optosis). <i>Front Cell Infect Microbiol</i> 9, 406. https://doi.org/10.3389/fcimb.2019.00406 .
<i>FADD</i>	Malireddi, R.K.S., Kesavardhana, S., and Kanneganti, T.D. (2019). ZBP1 and TAK1: Master Regulators of NLRP3 Inflammasome/Pyroptosis, Apoptosis, and Necroptosis (PAN-optosis). <i>Front Cell Infect Microbiol</i> 9, 406. https://doi.org/10.3389/fcimb.2019.00406 .
<i>GSDMD</i>	Zheng, M., Karki, R., Vogel, P., et al. (2020). Caspase-6 Is a Key Regulator of Innate Immunity, Inflammasome Activation, and Host Defense. <i>Cell</i> 181(3), 674-687.e613. https://doi.org/10.1016/j.cell.2020.03.040 .
<i>GSDME</i>	Liu, J., Hong, M., Li, Y., et al. (2022). Programmed Cell Death Tunes Tumor Immunity. <i>Front Immunol</i> 13, 847345. https://doi.org/10.3389/fimmu.2022.847345 .
<i>IRF1</i>	Karki, R., Sharma, B.R., Lee, E., et al. (2020). Interferon regulatory factor 1 regulates PANoptosis to prevent colorectal cancer. <i>JCI Insight</i> 5(12). https://doi.org/10.1172/jci.insight.136720 .
<i>MAP3K7</i>	Samir, P., Malireddi, R.K.S., and Kanneganti, T.D. (2020). The PANoptosome: A Deadly Protein Complex Driving Pyroptosis, Apoptosis, and Necroptosis (PANoptosis). <i>Front Cell Infect Microbiol</i> 10, 238. https://doi.org/10.3389/fcimb.2020.00238 .
<i>MLKL</i>	Zheng, M., Karki, R., Vogel, P., et al. (2020). Caspase-6 Is a Key Regulator of Innate Immunity, Inflammasome Activation, and Host Defense. <i>Cell</i> 181(3), 674-687.e613. https://doi.org/10.1016/j.cell.2020.03.040 .
<i>NFS1</i>	Lin, J.F., Hu, P.S., Wang, Y.Y., et al. (2022). Phosphorylated NFS1 weakens oxaliplatin-based chemosensitivity of colorectal cancer by preventing PANoptosis. <i>Signal Transduct Target Ther</i> 7(1), 54. https://doi.org/10.1038/s41392-022-00889-0 .
<i>NLRC4</i>	Sundaram, B., and Kanneganti, T.D. (2021). Advances in Understanding Activation and Function of the NLRC4 Inflammasome. <i>Int J Mol Sci</i> 22(3). https://doi.org/10.3390/ijms22031048 .
<i>NLRP3</i>	Zheng, M., Karki, R., Vogel, P., et al. (2020). Caspase-6 Is a Key Regulator of Innate Immunity, Inflammasome Activation, and Host Defense. <i>Cell</i> 181(3), 674-687.e613. https://doi.org/10.1016/j.cell.2020.03.040 .
<i>PSTPIP2</i>	Samir, P., Malireddi, R.K.S., and Kanneganti, T.D. (2020). The PANoptosome: A Deadly Protein Complex Driving Pyroptosis, Apoptosis, and Necroptosis (PANoptosis). <i>Front Cell Infect Microbiol</i> 10, 238. https://doi.org/10.3389/fcimb.2020.00238 .

PYCARD	Zheng, M., Karki, R., Vogel, P., et al. (2020). Caspase-6 Is a Key Regulator of Innate Immunity, Inflammasome Activation, and Host Defense. <i>Cell</i> 181(3), 674-687.e613. https://doi.org/10.1016/j.cell.2020.03.040 .
RBCK1	Samir, P., Malireddi, R.K.S., and Kanneganti, T.D. (2020). The PANoptosome: A Deadly Protein Complex Driving Pyroptosis, Apoptosis, and Necroptosis (PANoptosis). <i>Front Cell Infect Microbiol</i> 10, 238. https://doi.org/10.3389/fcimb.2020.00238 .
RIPK1	Malireddi, R.K.S., Kesavardhana, S., and Kanneganti, T.D. (2019). ZBP1 and TAK1: Master Regulators of NLRP3 Inflammasome/Pyroptosis, Apoptosis, and Necroptosis (PAN-optosis). <i>Front Cell Infect Microbiol</i> 9, 406. https://doi.org/10.3389/fcimb.2019.00406 .
RIPK3	Malireddi, R.K.S., Kesavardhana, S., and Kanneganti, T.D. (2019). ZBP1 and TAK1: Master Regulators of NLRP3 Inflammasome/Pyroptosis, Apoptosis, and Necroptosis (PAN-optosis). <i>Front Cell Infect Microbiol</i> 9, 406. https://doi.org/10.3389/fcimb.2019.00406 .
RNF31	Samir, P., Malireddi, R.K.S., and Kanneganti, T.D. (2020). The PANoptosome: A Deadly Protein Complex Driving Pyroptosis, Apoptosis, and Necroptosis (PANoptosis). <i>Front Cell Infect Microbiol</i> 10, 238. https://doi.org/10.3389/fcimb.2020.00238 .
TNFAIP3	Samir, P., Malireddi, R.K.S., and Kanneganti, T.D. (2020). The PANoptosome: A Deadly Protein Complex Driving Pyroptosis, Apoptosis, and Necroptosis (PANoptosis). <i>Front Cell Infect Microbiol</i> 10, 238. https://doi.org/10.3389/fcimb.2020.00238 .
ZBP1	Malireddi, R.K.S., Kesavardhana, S., and Kanneganti, T.D. (2019). ZBP1 and TAK1: Master Regulators of NLRP3 Inflammasome/Pyroptosis, Apoptosis, and Necroptosis (PAN-optosis). <i>Front Cell Infect Microbiol</i> 9, 406. https://doi.org/10.3389/fcimb.2019.00406 .

Supplementary Table 2. Primers and siRNA target sequences.

Name	Sequences
Primers for real-time PCR	Primer: Forward primer CTCCTTCCCACAAGAGCCAC Reverse primer ACTCACTCCGACTGACCTGT
The target sites of siRNA	Si-1: target sequence: atggaactggccaacattaaaa guide (5'→3')UUAAUGUUGGCCAGUUUCCAU passenger (5'→3)GGAAACUGGCCAACAUUAAAA Si-2: target sequence: cagattaggaaggatgctaaca guide (5'→3')UUUAGCAUCCUCCUAAUCUG passenger (5'→')GAUUAGGAAGGAUGC UAAACA