

1 SUPPLEMENTARY FIGURE LEGENDS

2 **Fig. S1: Reproducibility between biological replicates of Spike in-SILAC**
3 **samples.** Scatter plot analysis of biological replicates at indicated time points p.i.
4 Displayed are $\log_2(L/H)$ values of Pan (upper panel) and Mal (lower panel) infected
5 cells, replicate 1 at y axis and replicate 2 at x axis. Numbers indicate number of
6 quantified protein groups (black) and the corresponding Pearson correlation (blue);
7 dots represent protein groups.

8 **Fig. S2: Biological processes executed by 16 differentially regulated proteins**
9 **(Pan vs. Mal infected cells).** Classification of 16 proteins with significant $\log_2(\text{fold}$
10 $\text{change}) >1$ or <-1 upon infection with Pan in comparison to Mal virus infected cells
11 at indicated time points according to their gene ontology terms “biological process”
12 (GOBP) using PANTHER (Protein analysis through evolutionary relationships)
13 Classification System Version 10.0.

14 **Fig. S3: siRNA transfection does not affect cell viability.** WST-1 assay was
15 performed 48 h post transfection by adding WST-1 reagent to the cells, followed by
16 incubation at 37 °C for 1.5 h. Absorbance was measured at 460 nm and at the
17 reference wavelength 590 nm. Non-targeting siRNA (AllStars) and siPLK1 were used
18 as positive and negative controls, respectively.

19 **Fig. S4: SAMHD1 expression levels are down-regulated in Pan infected cells.**
20 A549 cells were infected with Pan or Mal virus at an MOI of 0.5. At 4 h and 16 h p.i.
21 cells were fixed, permeabilized, stained with specific primary antibody against NP,
22 SAMHD1 and with DAPI. Images were representative for three independent
23 experiments. White arrows indicate NP positive cells with low SAMHD1 expression

24 levels, yellow arrows mark infected cells with high levels of SAMHD1. Scale bar: 10
25 μm . SAMHD1: green, NP: red, DAPI: blue.

26 **Supplementary Table 1:** All identified proteins summarized as protein groups.
27 Sequence coverage (%) and number of unique peptides were stated for all samples
28 (total) and separately for each individual sample. Displayed are also gene names as
29 well as L/H ratios, variability and count.

30 **Supplementary Table 2:** All peptides used for protein identification including their
31 sequence, charge, m/z, modifications, score, PEP, mass error (Da), retention time
32 and mass. Potential contaminants are also indicated as well as gene names,
33 proteins and protein group IDs.

34 **Supplementary Table 3:** Classification of peptide sequences as unique peptide for
35 a specific protein or protein group. Potential contaminants are also indicated as well
36 as gene names, proteins and protein group IDs.

37 **Supplementary Table 4:** Significantly regulated proteins in each individual sample
38 compared to 0 h time point. Listed are gene names, $-\text{Log}(\text{P-value})$, difference (of
39 two time points), Protein IDs, Protein names, Symbol color (see volcano plots – Fig.
40 4), GOBP name, GOMF name, GOCC name as well as $[\log_2(\text{L/H})]$ ratios of the
41 respective samples.

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