**eLife’s transparent reporting form**

We encourage authors to provide detailed information *within their submission* to facilitate the interpretation and replication of experiments. Authors can upload supporting documentation to indicate the use of appropriate reporting guidelines for health-related research (see [EQUATOR Network](https://www.equator-network.org)), life science research (see the [BioSharing Information Resource](http://www.biosharing.org)), or the [ARRIVE guidelines](https:// ARRIVEguidelines.org) for reporting work involving animal research. Where applicable, authors should refer to any relevant reporting standards documents in this form.

If you have any questions, please consult our Journal Policies and/or contact us: editorial@elifesciences.org.

**Sample-size estimation**

- You should state whether an appropriate sample size was computed when the study was being designed.
- You should state the statistical method of sample size computation and any required assumptions.
- If no explicit power analysis was used, you should describe how you decided what sample (replicate) size (number) to use.

Please outline where this information can be found within the submission (e.g., sections or figure legends), or explain why this information doesn’t apply to your submission:

For the phylogenetics, we used operational taxonomic units to cluster similar sequences (lines 404-416 in Methods). For live infections, 5 cells were imaged and analyzed per experiment with combined data from all three experiments presented (figure legend for figure 7).

**Replicates**

- You should report how often each experiment was performed.
- You should include a definition of biological versus technical replication.
- The data obtained should be provided and sufficient information should be provided to indicate the number of independent biological and/or technical replicates.
- If you encountered any outliers, you should describe how these were handled.
- Criteria for exclusion/inclusion of data should be clearly stated.
- High-throughput sequence data should be uploaded before submission, with a private link for reviewers provided (these are available from both GEO and ArrayExpress).

Please outline where this information can be found within the submission (e.g., sections or figure legends), or explain why this information doesn’t apply to your submission:

For the phylogenetics, replicates are described in lines 404-416 of the Methods. GenBank accession numbers for the strains analyzed can be found in Supplemental Tables 1-4. Replicates used in live infection are described in the figure legend of Figure 7 and in lines 441-442 and 469-470 of the Methods.
Statistical reporting

- Statistical analysis methods should be described and justified
- Raw data should be presented in figures whenever informative to do so (typically when N per group is less than 10)
- For each experiment, you should identify the statistical tests used, exact values of N, definitions of center, methods of multiple test correction, and dispersion and precision measures (e.g., mean, median, SD, SEM, confidence intervals; and, for the major substantive results, a measure of effect size (e.g., Pearson's r, Cohen's d)
- Report exact p-values wherever possible alongside the summary statistics and 95% confidence intervals. These should be reported for all key questions and not only when the p-value is less than 0.05.

Please outline where this information can be found within the submission (e.g., sections or figure legends), or explain why this information doesn’t apply to your submission:

Statistical analysis is detailed in the Methods section (lines 487 – 498) and the statistical tests used are indicated in each figure legend. Confidence intervals are indicated by dashed lines in Figures 3, 5, and 6. P values are reported in Figure 7.

(For large datasets, or papers with a very large number of statistical tests, you may upload a single table file with tests, Ns, etc., with reference to sections in the manuscript.)

Group allocation

- Indicate how samples were allocated into experimental groups (in the case of clinical studies, please specify allocation to treatment method); if randomization was used, please also state if restricted randomization was applied
- Indicate if masking was used during group allocation, data collection and/or data analysis

Please outline where this information can be found within the submission (e.g., sections or figure legends), or explain why this information doesn’t apply to your submission:

Masking was used during image analysis, as described in lines 471-486 of the Methods.

Additional data files (“source data”)

- We encourage you to upload relevant additional data files, such as numerical data that are represented as a graph in a figure, or as a summary table
- Where provided, these should be in the most useful format, and they can be uploaded as “Source data” files linked to a main figure or table
- Include model definition files including the full list of parameters used
- Include code used for data analysis (e.g., R, MatLab)
- Avoid stating that data files are “available upon request”

Please indicate the figures or tables for which source data files have been provided:

All code used for phylogenetics (Figures 2-6 and Supplemental Figures) have been provided (lines 499-501). Summary tables have been provided for Figures 2-7.