medRxiv preprint doi: https://doi.org/10.1101/2021.03.10. 235; this version posted March 14, 2021. The copyright holder for this preprint (which was not certified by peer review) is the author/funder, who has granted medRxiv a license to display the preprint in perpetuity. It is made available under a CC-BY-NC-ND 4.0 International license .

Supplemental figures

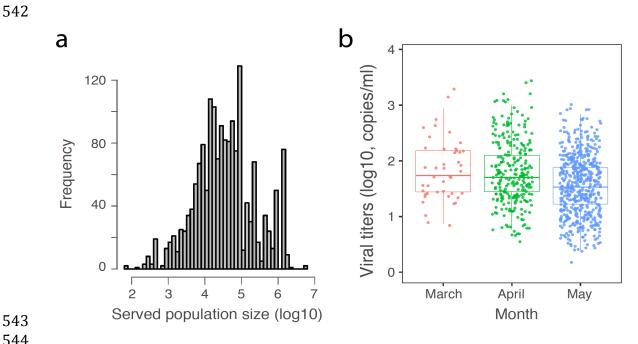


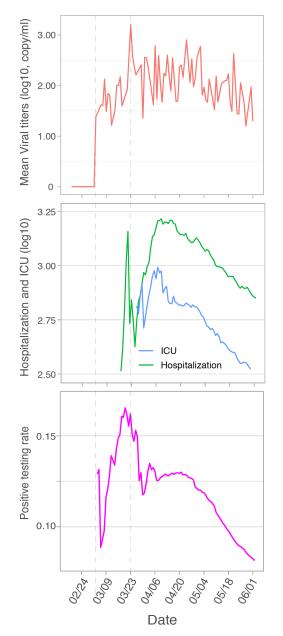
Fig. S1. Distribution of the wastewater treatment plant serving population size. and viral titers in positive wastewater samples by month. (a) Histogram of population sizes served by the sampled wastewater treatment plants. The median served population size is 31,745. (b) Viral titers in positive wastewater samples by month. The box represents the interguartile range of viral titers for each month, the horizontal line inside the box is the median. Significant difference was found between the mean viral titers in April and May (Welch's t-test, p-value = 0.025).

medRxiv preprint doi: https://doi.org/10.1101/2021.03.10.21253235; this version posted March 14, 2021. The copyright holder for this preprint (which was not certified by peer review) is the author/funder, who has granted medRxiv a license to display the preprint in perpetuity. It is made available under a CC-BY-NC-ND 4.0 International license.



Fig. S2. Temporal SARS-CoV-2 titers at the county level and catchment level. (a) Temporal viral titers in 12 representative counties. Each dot is a sample, and colored by the sampling catchments in the county. Red lines represent the daily new cases in the corresponding counties during the sampling period. (b) Temporal viral titers for samples collected in six different locations in the New Castle County of Delaware. Red lines represent the weighted daily new cases (daily new cases in the county multiplying the catchment served population size and divided by the county population size) during the sampling period.

medRxiv preprint doi: https://doi.org/10.1101/2021.03.10.21253235; this version posted March 14, 2021. The copyright holder for this preprint (which was not certified by peer review) is the author/funder, who has granted medRxiv a license to display the preprint in perpetuity. It is made available under a CC-BY-NC-ND 4.0 International license.



579

580 **Fig. S3.** Wastewater viral titers (top) and clinical COVID-19 surveillance

581 indicators including hospitalization and intensive care unit admissions (middle),

and the testing positive rates (bottom) from February to June. Positive

583 wastewater data from all the sampling locations were aggregated by date using

the mean function. Clinical data from the 40 sampled states were aggregated inthe same way.

586

587

588

medRxiv preprint doi: https://doi.org/10.1101/2021.03.10.21253235; this version posted March 14, 2021. The copyright holder for this preprint (which was not certified by peer review) is the author/funder, who has granted medRxiv a license to display the preprint in perpetuity. It is made available under a CC-BY-NC-ND 4.0 International license.

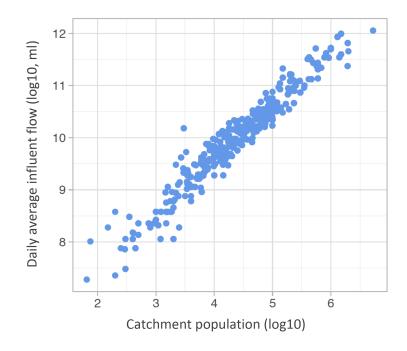
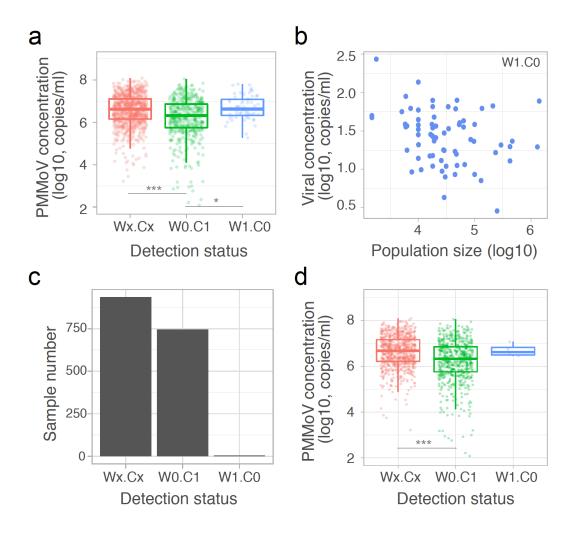


Fig. S4. Daily average influent flow at the wastewater treatment plant is

592 correlated with catchment population size.

medRxiv preprint doi: https://doi.org/10.1101/2021.03.10.21253235; this version posted March 14, 2021. The copyright holder for this preprint (which was not certified by peer review) is the author/funder, who has granted medRxiv a license to display the preprint in perpetuity. It is made available under a CC-BY-NC-ND 4.0 International license.



596 597

598 Fig. S5. PMMoV concentrations in wastewater samples, and detection status 599 with 7-day averages of new COVID-19 cases. (a) PMMoV concentrations in the 600 Wx.Cx (x = 0 or 1), W0.C1, and W1.C0 groups. W1.C1: SARS-CoV-2 detected in 601 Wastewater and new Clinical cases reported; W0.C0: no Wastewater detection 602 and no new Clinical cases reported; W0.C1: no Wastewater detection but new 603 Clinical cases reported: W1.C0: Wastewater detection but no new Clinical cases 604 reported. Significant differences of PMMoV concentrations were found between 605 W0.C1 and Wx.Cx or W1.C0 groups (Welch's t-test, and symbol *: p-value < 606 0.05; ***: p-value < 0.001). (b) Viral titers and the served population size for the 607 W1.C0 samples. Most of these samples were from catchments serving $10,000 \sim$ 608 100,000 population, with viral titers ranging from 10 to 100 copies per ml of 609 wastewater. (c-d) Detection status for wastewater data against 7-day averages of 610 new clinical cases (c), and PMMoV concentrations in the Wx.Cx, W0.C1, and 611 W1.C0 groups (d). Significant difference was found between Wx.Cx and W0.C1 612 groups (Welch's t-test, and symbol ***: p-value < 0.001).