

1 **Response to Mitchell et al – Relevant and Robust Data for Climate Change Risk**

2 **Assessment**

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16 Time-series studies, which are commonly used to analyse associations of temperature with
17 mortality, can inform short-term planning but have limited relevance to how changes in
18 temperature over longer-time periods influence human health. They also cannot account for
19 how populations adapt to their local climate, that for example people in Phoenix have adapted
20 to temperatures which Chicago would engender extreme situations and cause thousands of
21 deaths. Our study,¹ and another recent analysis,² used innovative designs and large quantities
22 of data to overcome these inherent limitations of time-series analysis in informing long-term
23 risk assessment. In particular, the anomaly metric used in our analysis is both intuitive and
24 consistent with how populations experience and respond to weather patterns in the long term.
25 Therefore, our study has made a major advance in making epidemiological analysis relevant

26 for long-term risk assessment. A second methodological innovation of our work is to leverage
27 variation over both space and time, versus each alone, to infer the size of associations. This
28 approach has been used in studies of air pollution^{3,4} and cardiovascular risk factors⁵ because
29 spatiotemporal inference is more efficient and robust than using variation in either dimension
30 alone.

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32 As incorrectly stated by Mitchell and colleagues, computational considerations had no
33 influence on the choice to analyse monthly mean temperature. Rather, in addition to anomaly
34 in mean temperature, we analysed and reported in the paper metrics of extreme temperature
35 including (1) the 90th percentile (°C) of daily mean temperatures; (2) number of days in a
36 month above the long-term 90th percentile of average temperature for each state and month;
37 and (3) number of episodes of 3+ day episodes above the long-term 90th percentile of average
38 temperature for each state and month. The estimated rate ratios of the temperature anomaly
39 based on daily means were robust to the addition of alternative measures of anomaly, whereas
40 the coefficients of the additional measures were generally not statistically significant and with
41 large credible intervals. This may be because empirically changes in heat waves are explained
42 by simple shifts in mean temperature.⁶

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44 We used a (log-)linear association, consistent with other analyses of injuries.^{2,7} Further, the
45 anomaly metric used in our paper has a much smaller size range (<8°C across all states and
46 months in our data) than the range of absolute temperatures (>40°C in our data). Finally, having
47 age- and month-specific coefficients gives our model the flexibility to infer different effect
48 sizes in cold and warm months and for different age groups, as seen in Figure 4 of our paper.
49 Our computer code is available on an open access basis (<http://globalenvhealth.org/code-data-download/>) and can be used to consider alternative functional forms.

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52 Our statistical model by design adjusts for environmental, social, and meteorological factors
53 that are specific to month, state and state-month through the use of corresponding random
54 intercepts. The potential unmeasured confounding factors would be those with anomalies
55 similar to those of the average monthly temperature in each state, such as air pollution, which
56 is not associated with injury mortality. As for other meteorological factors, anomalous
57 temperature in USA is not significantly correlated with anomalous cloud cover based on data
58 from ERA5, nor with precipitation.⁷ Further the association between temperature and transport
59 injury is robust to adjustment for precipitation.⁷ Therefore, the most plausible explanation for
60 our findings is a causal effect from anomalous temperature.

61
62 To illustrate the public health significance of the findings, we applied our estimated rate ratios
63 to a fixed temperature anomaly (1.5 and 2 degrees) and 2016 age- and sex-specific death rates,
64 carefully reported as an “anomalously warm year”. The choice of counterfactual exposure in
65 our illustrative example is similar to analyses for air pollution^{4,8} and for virtually every other
66 risk factor⁹ which use either a constant change in exposure or a constant level of exposure as
67 counterfactual exposure. Full climate change risk assessment, as done for other risk factors,¹⁰
68 requires three inputs: (i) spatially and temporally coherent projections of temperature; (ii)
69 spatially and temporally coherent projections of population; and (iii) spatially, temporally and
70 epidemiologically coherent projections of age-, sex- and cause-specific death rates,¹¹ because
71 death rates from each disease and injury change due to factors other than climate – as seen in
72 secular trends of deaths from various injuries in Figure 2 of our paper. The innovative scope
73 of our work, i.e. analysing the complete range of intentional and unintentional injuries by age
74 group and sex, and its methodological innovations provide a relevant and robust basis for such
75 analyses.

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